



US 20200255497A1

(19) **United States**

(12) **Patent Application Publication**
Ouzounov et al.

(10) **Pub. No.: US 2020/0255497 A1**

(43) **Pub. Date: Aug. 13, 2020**

(54) **RECOMBINANT COLLAGEN AND ELASTIN MOLECULES AND USES THEREOF**

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(21) Appl. No.: **16/839,044**

(22) Filed: **Apr. 2, 2020**

Related U.S. Application Data

(63) Continuation of application No. 16/144,914, filed on Sep. 27, 2018.

(60) Provisional application No. 62/657,591, filed on Apr. 13, 2018, provisional application No. 62/564,964, filed on Sep. 28, 2017.

Publication Classification

(51) **Int. Cl.**

C07K 14/78 (2006.01)

A61K 38/39 (2006.01)

A61K 8/65 (2006.01)

A61Q 17/04 (2006.01)

A61Q 19/08 (2006.01)

(52) **U.S. Cl.**

CPC *C07K 14/78* (2013.01); *A61K 38/39*

(2013.01); *A61Q 19/08* (2013.01); *A61Q*

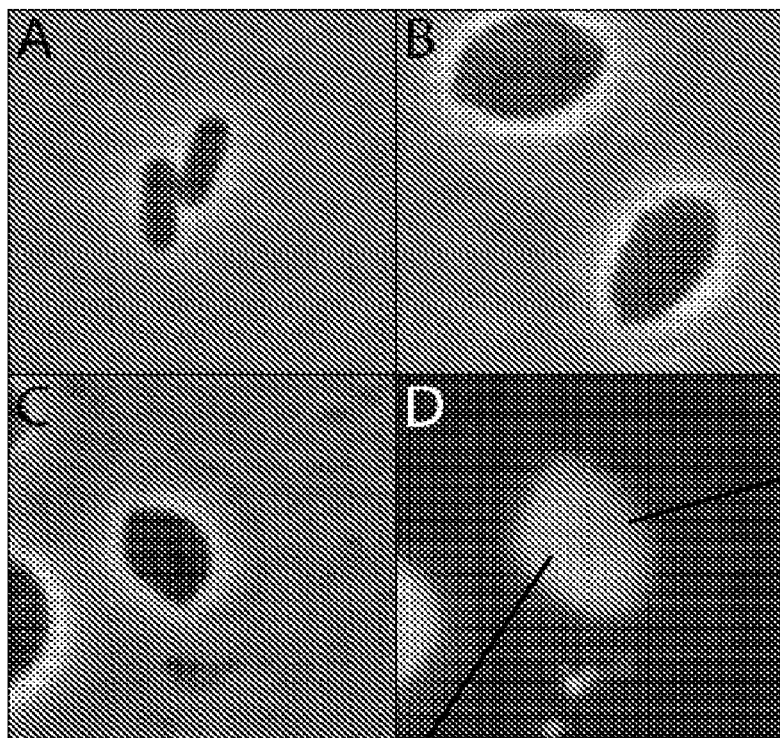
17/04 (2013.01); *A61K 8/65* (2013.01)

(57)

ABSTRACT

This disclosure provides non-naturally occurring collagen and elastin molecules. The non-naturally occurring collagens and elastins include truncated collagens, truncated elastins, as well as fusion proteins thereof. The non-naturally occurring collagen and elastin are useful in foods, cosmetics and many other products and uses.

Specification includes a Sequence Listing.



cytoplasmic RFP

periplasmic GFP

Fig. 1

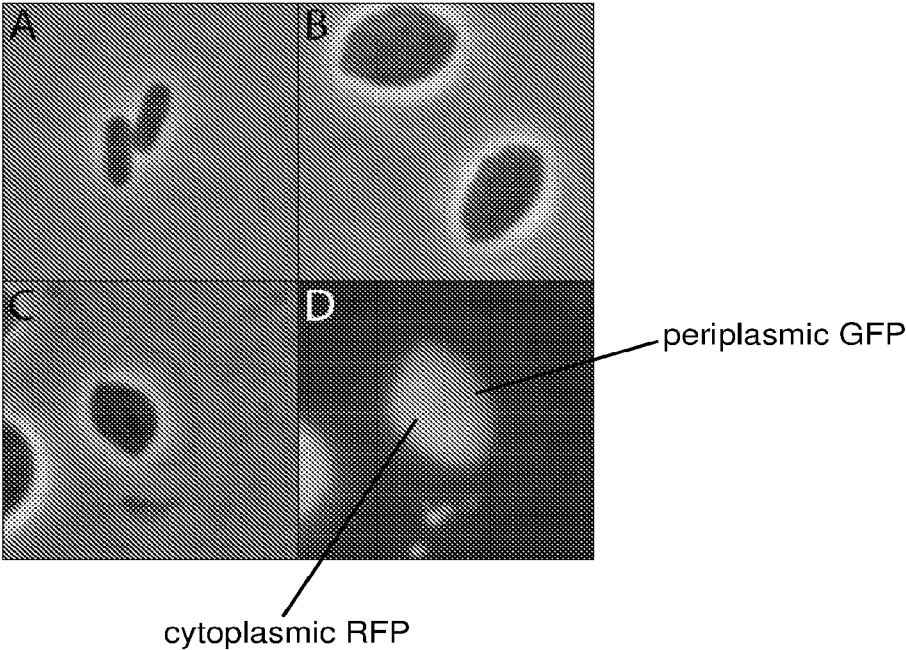


Fig. 2

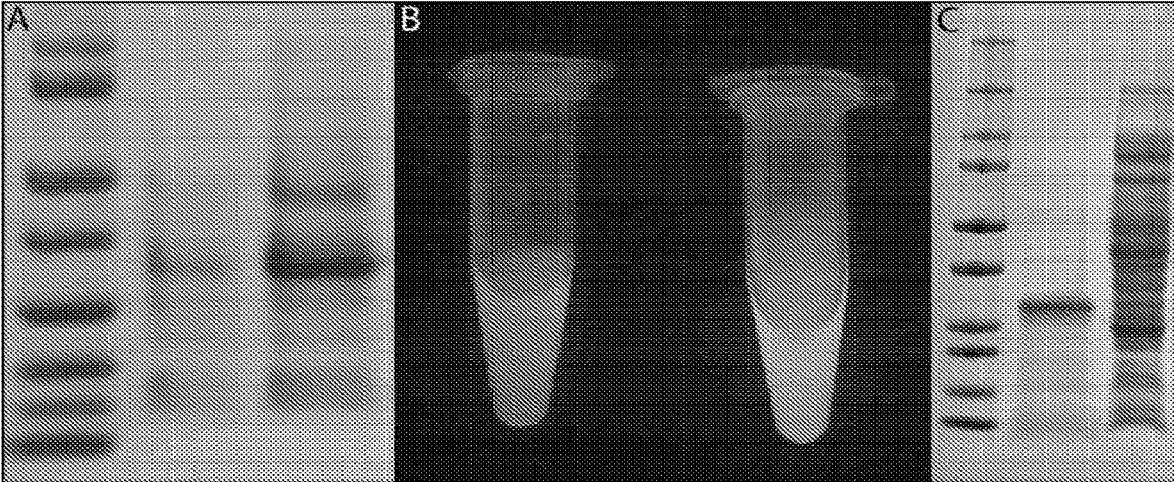


Fig. 3

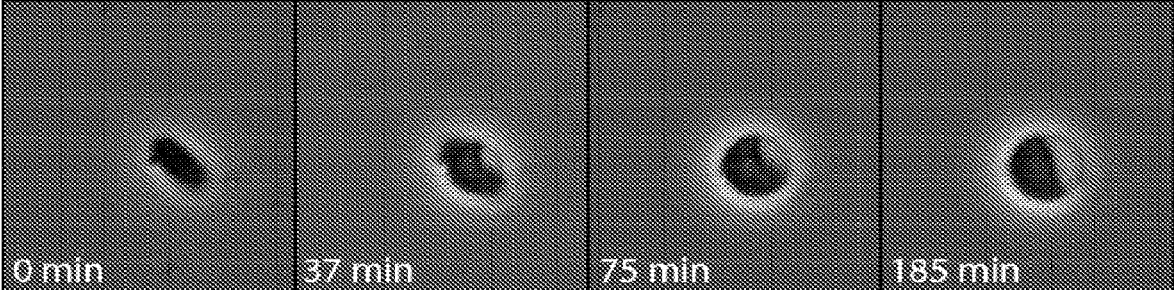
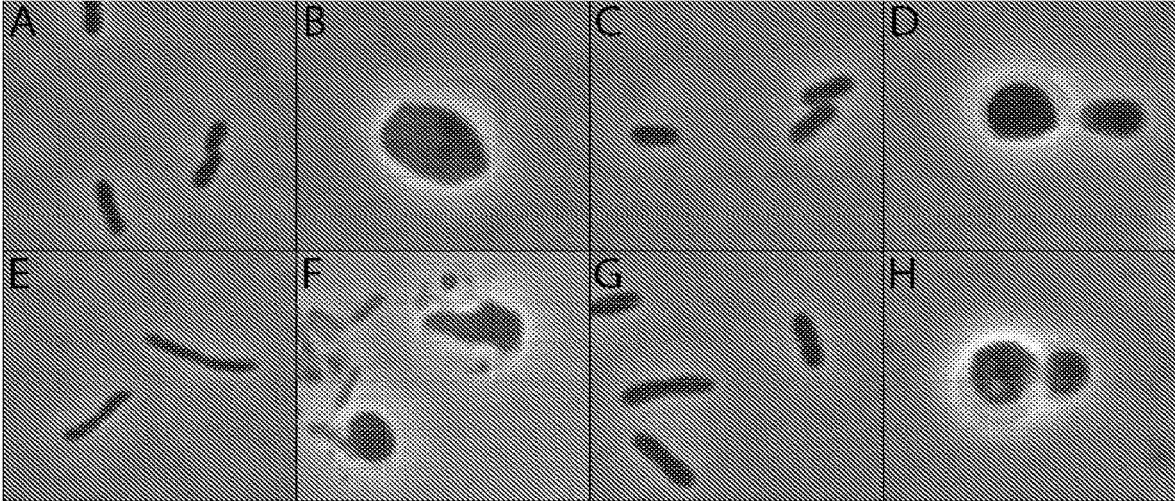


Fig. 4



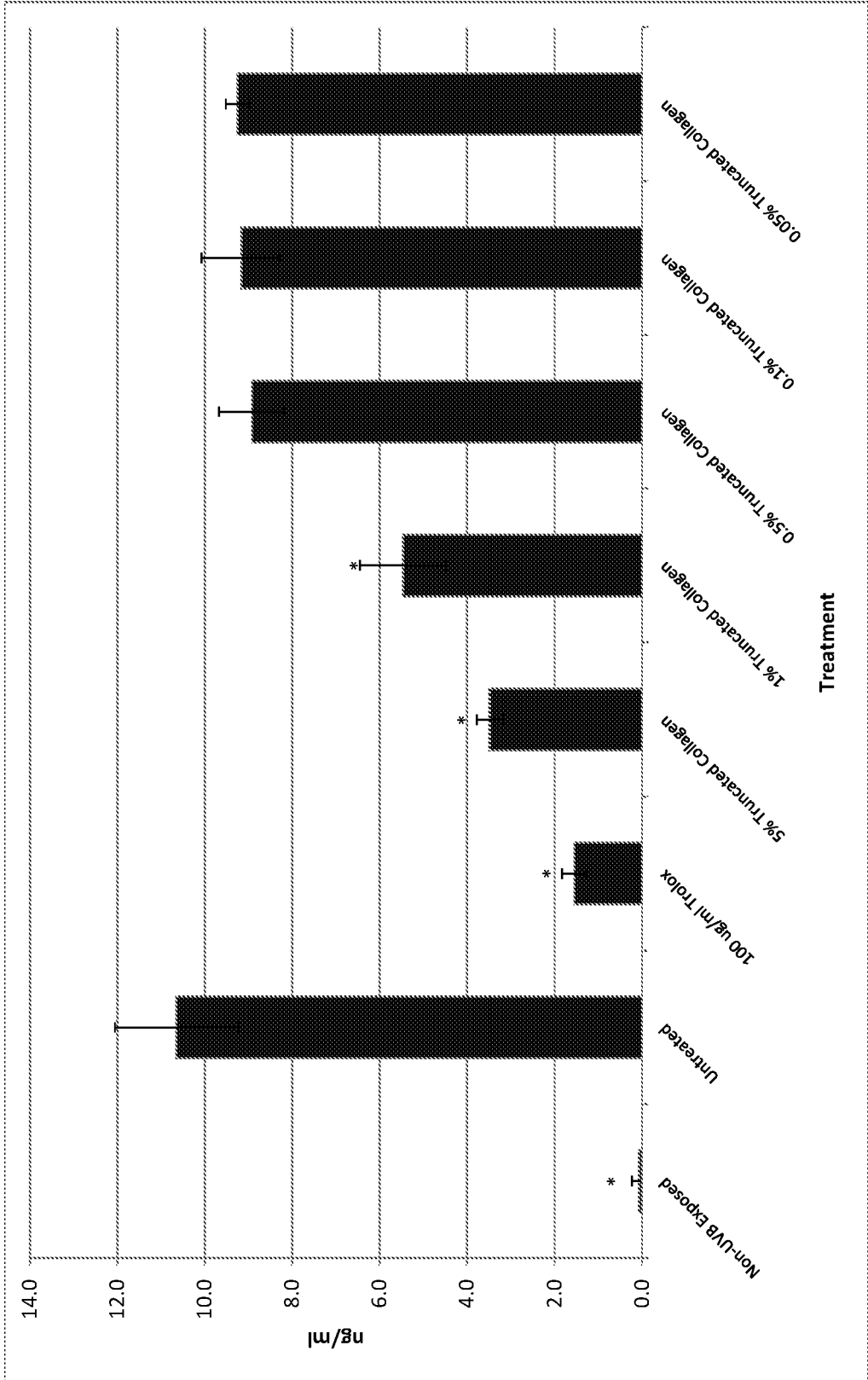


Fig. 5

RECOMBINANT COLLAGEN AND ELASTIN MOLECULES AND USES THEREOF

CROSS-REFERENCE

[0001] This application is a continuation application of U.S. patent application Ser. No. 16/144,914, filed Sep. 27, 2018, which application claims priority from U.S. Provisional Patent Application No. 62/564,964, filed Sep. 28, 2017, and U.S. Provisional Patent Application No. 62/657,591, filed Apr. 13, 2018, the disclosures of which are incorporated by reference herein in their entirety.

SEQUENCE LISTING

[0002] The instant application contains a Sequence Listing which has been submitted electronically in ASCII format and is hereby incorporated by reference in its entirety. Said ASCII copy, created on Nov. 21, 2018, is named 57607702SL.txt and is 320,570 bytes in size.

FIELD

[0003] The present disclosure relates to non-naturally occurring full-length and truncated collagen molecules and full-length and truncated elastin molecules and uses thereof.

BACKGROUND

[0004] Collagens and similar proteins are the most abundant proteins in the biosphere. Collagens and elastins are structural proteins found in the skin, connective tissue and bone of animals and other tissues. In humans, the amount of collagen present in the body is approximately one third of the total proteins and accounts for about three fourths of the dry weight of skin. Elastin is a highly elastic protein found in connective tissue and other types of tissue.

[0005] The structure of collagen is a triple helix in which three polypeptide strands together form a helical coil. The individual polypeptide strands are composed of repeating triplet amino acid sequences designated as GLY-X-Y. X and Y can be any amino acid and the third amino acid is glycine. The amino acids proline and hydroxyproline are found in high concentrations in collagen. The most common triplet is proline-hydroxyproline-glycine (Gly-Pro-Hyp) accounting for approximately 10.5% of the triplets in collagen.

[0006] Gelatin is a product obtained by partial hydrolysis of collagen. Typically, gelatin is produced by acid hydrolysis, alkaline hydrolysis, and enzymatic hydrolysis or by exposing collagen to heat in an aqueous solution (e.g., boiling the bones and skins of animal, boiling fish scales, etc.).

[0007] Gelatin is used in many products including cosmetics, foods, pharmaceuticals, medical devices, photographic films, adhesives, binders and many others. The physical and chemical properties of gelatin are tuned to the particular application. These physical/chemical properties include gel strength, melting point temperature, viscosity, color, turbidity, pH, isoelectric point and others.

[0008] Elastin is an elastic protein that is crucial for the proper functioning of arteries, lung, tendons, ligament, skin and other tissue. Elastin provides the tissues with the ability to stretch and return to its original shape. The protein tropoelastin is the building block of elastin. In contrast to collagen that include a family of genes, there is one tropoelastin gene in humans. When expressed, the single elas-

tin gene is spliced to produce different forms of the tropoelastin protein. Many tropoelastin molecules associate together to form elastin.

[0009] L-form bacteria, or L-forms, are bacterial strains derived from parent species (N-forms) that are able to grow as cell wall-deficient (spheroplast type) or as cell wall-less (protoplast type) cells. See, Madoff S (Ed): *The Bacterial L-Forms*. New York: Marcel Dekker Inc., 1986; Mattmann L H (Ed): *Cell Wall Deficient Forms*. Boca Raton: CRC Press; 1993; and Gumpert J, Taubeneck U: Characteristic properties and biological significance of stable protoplast type L-forms. In *Protoplasts, Lecture Proceedings of the 6th International Protoplast Symposium: Basel*. *Experientia* 1983, 46(suppl):227-241.

[0010] Protoplast type L-forms have been cultivated in the cell wall-less state and represent genetically stable mutants showing extreme pleiotropic changes, including the inability to form cell walls, capsules, flagella, pili, spores and mesosomes, altered colony and cell morphology, qualitative and quantitative changes in the lipid and protein components of the cytoplasmic membrane, the absence of extracellular proteolytic activities, resistance against bacteriophages and the incapability to propagate outside laboratory conditions. See, Gumpert and Taubeneck (supra); and Hoischen et al., Lipid and fatty acid composition of cytoplasmic membranes from *Streptomyces* hygroscopic and its stable protoplast type L-form. *J Bacteriol* 1997, 179:3430-3436.

SUMMARY

[0011] In one aspect, a non-naturally occurring collagen produced by a host cell is provided. The non-naturally occurring collagen is jellyfish (Hydrozoan) collagen, human collagen, *Chondrosia reniformis* (kidney sponge) collagen, or *Rhincodon typus* (whale shark) collagen. In an embodiment, the non-naturally occurring collagen is a full-length or a truncated collagen. In one embodiment, the collagen is truncated by an internal truncation of between 50 amino acids and 500 amino acids. In another embodiment, the truncation is at the C-terminal end or the N-terminal end of the collagen polypeptide. The non-naturally occurring collagens are SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 74, SEQ ID NO: 76, SEQ ID NO: 78, SEQ ID NO: 80, SEQ ID NO: 82, SEQ ID NO: 84, SEQ ID NO: 86, SEQ ID NO: 89, SEQ ID NO: 91, SEQ ID NO: 92, SEQ ID NO: 94, SEQ ID NO: 96, SEQ ID NO: 102, SEQ ID NO: 104, SEQ ID NO: 106, SEQ ID NO: 108, SEQ ID NO: 110, or SEQ ID NO: 112.

[0012] In another aspect, the non-naturally occurring collagen further comprises amino acid sequences including a secretion tag, a histidine tag, a green fluorescent protein tag, a protease cleavage site, a Beta-lactamase and/or GEK amino acid trimer repeats and/or GDK amino acid trimer repeats. When the non-naturally occurring collagen comprises one or more amino acid trimer repeats of the sequence glycine-glutamic acid-lysine (GEK) and/or glycine-aspartic acid-lysine (GDK), the number of GEK and/or GDK trimer repeats can range from 2 to 50 trimer repeats (SEQ ID NOS: 130-131, respectively). In one aspect, the secretion tag is DsbA, PelB, OmpA, TolB, MalE, lpp, TorA, or HylA, or a hybrid secretion tag that comprises a portion of one secretion tag fused to a portion of a second secretion tag. An exemplary secretion tag is DsbA.

[0013] In one aspect, provided are compositions that comprise between 0.005% and 30% w/w non-naturally occurring collagen. The compositions can further comprise at least one additional ingredient comprising a topical carrier or a preservative.

[0014] Compositions comprising non-naturally occurring collagen are in one aspect topical compositions for applying to skin. The topical compositions are used for decreasing skin damage or promoting the repair of damaged skin.

[0015] One aspect provides methods for decreasing skin damage or promoting the repair of damaged skin. The method comprises applying the composition comprising elastin to the skin of a subject. The method increases the viability of the fibroblast cells or keratinocytes of the skin of the subject. In another aspect the application of the composition increases the synthesis of procollagen by the fibroblast cells of the subject's skin. In another aspect the topical application of the composition protects skin or keratinocytes against UV damage. In yet another embodiment, thymine-thymine (TT) dimer formation is decreased by the collagens or elastins disclosed herein.

[0016] Another aspect provided herein are methods of increasing the viability of skin cells. The method comprises applying collagen or elastin molecules to the skin or skin cell. The collagen or elastin as provided increases the viability of keratinocytes and/or fibroblasts is increased upon exposure to UV radiation, urban dust or other damaging stimuli.

[0017] In another aspect provided herein are methods for decreasing the production of inflammatory cytokines in a skin cell. In one embodiment the skin cell is a keratinocyte. The method comprises applying a collagen or elastin molecule to a skin cell. The production of inflammatory cytokines including TNF α , IL-1 α , IL-1 β , IL-3, IL-6, IL-7, IL-8, IL-10, IL-18, and IL-1RA.

[0018] In another aspect, provided are methods of protecting skin cells against the effect of exposure to urban dust. The method comprises the step of applying the collagen or elastin disclosed herein to the skin cell. The exposure of skin cell to collagen or elastin increases the viability of the skin cell. In an embodiment, the skin cell is a keratinocyte or a fibroblast.

[0019] In one aspect, a non-naturally occurring elastin produced by a host cell is provided. The non-naturally occurring elastin is jellyfish elastin, human elastin, *Chondrosia reniformis* (kidney sponge) elastin, or *Rhincodon typus* elastin. In an embodiment, the non-naturally occurring elastin is a full-length or truncated elastin. In one embodiment, the elastin is truncated by an internal truncation of between 50 amino acids and 500 amino acids. In another embodiment, the truncation is at the C-terminal end or the N-terminal end of the elastin polypeptide. The non-naturally occurring elastins are SEQ ID NO: 31, SEQ ID NO: 33, SEQ ID NO: 35, SEQ ID NO: 37, SEQ ID NO: 39, SEQ ID NO: 41, SEQ ID NO: 43, SEQ ID NO: 45, SEQ ID NO: 47, SEQ ID NO: 49, SEQ ID NO: 51, SEQ ID NO: 53, SEQ ID NO: 55, SEQ ID NO: 57, SEQ ID NO: 59, SEQ ID NO: 61, SEQ ID NO: 63, SEQ ID NO: 65, SEQ ID NO: 67, SEQ ID NO: 69, SEQ ID NO: 71, SEQ ID NO: 98, or SEQ ID NO: 110.

[0020] In another aspect, the non-naturally occurring elastin further comprises amino acid sequences including a secretion tag, a histidine tag, a green fluorescent protein tag, a protease cleavage site, a Beta-lactamase and/or GEK amino acid trimer repeats and/or GDK amino acid trimer

repeats. When the non-naturally occurring collagen comprises one or more amino acid trimer repeats of the sequence glycine-glutamic acid-lysine (GEK) and/or glycine-aspartic acid-lysine (GDK), the number of GEK and/or GDK trimer repeats can range from 2 to 50 trimer repeats (SEQ ID NOS: 130-131, respectively). In one aspect, the secretion tag is DsbA, PelB, OmpA, TolB, MalE, Ipp, TorA, or HylA, or a hybrid secretion tag that comprises a portion of one secretion tag fused to a portion of a second secretion tag. An exemplary secretion tag is DsbA.

[0021] In another embodiment, compositions that comprise between 0.005% and 30% w/w non-naturally occurring elastin are provided. The compositions can further comprise at least one additional ingredient comprising a topical carrier or a preservative.

[0022] Compositions comprising non-naturally occurring elastin are in one aspect topical compositions for applying to skin. The topical compositions are used for decreasing skin damage or promoting the repair of damaged skin.

[0023] One embodiment provides methods for decreasing skin damage or promoting the repair of damaged skin. The method comprises applying the composition comprising elastin to the skin of a subject. The method increases the viability of the fibroblast cells of the skin of the subject. In another aspect the application of the composition increases the synthesis of procollagen by the fibroblast cells of the subject's skin. In another aspect the topical application of the compositions protects skin or keratinocytes against UV damage. In yet another embodiment, thymine-thymine (TT) dimer formation is decreased by the collagens or elastins disclosed herein.

[0024] Another embodiment provides polynucleotides that encode a non-naturally occurring collagen or a non-naturally occurring elastin. The polynucleotides encode collagen or elastin from jellyfish, human, *Chondrosia reniformis* (kidney sponge), or *Rhincodon typus*. The encoded collagen or elastin may be full length or truncated. In one embodiment, the collagen or elastin is truncated by an internal truncation of between 50 amino acids and 500 amino acids.

[0025] In one embodiment polynucleotides that encode fusion proteins comprising a secretion tag, a histidine tag, a green fluorescent protein tag, a protease cleavage site, a Beta-lactamase along and/or GEK amino acid trimer repeat and/or GDK amino acid trimer repeats together with collagen or elastin are provided. The non-naturally occurring collagen or elastin may comprise one or more amino acid trimer repeats of the sequence glycine-glutamic acid-lysine (GEK) and/or glycine-aspartic acid-lysine (GDK), the number of GEK and/or GDK trimer repeats can range from 2 to 50 trimer repeats (SEQ ID NOS: 130-131, respectively). In one aspect, the secretion tag is DsbA, PelB, OmpA, TolB, MalE, Ipp, TorA, or HylA, or a hybrid secretion tag that comprises a portion of one secretion tag fused to a portion of a second secretion tag. An exemplary embodiment secretion tag is DsbA.

[0026] The polynucleotides and vectors can be used to transform host cells and express the polynucleotides. Polynucleotides encoding a non-naturally occurring collagen, wherein the polynucleotide is SEQ ID NO: 9, SEQ ID NO: 11, SEQ ID NO: 13, SEQ ID NO: 15, SEQ ID NO: 17, SEQ ID NO: 19, SEQ ID NO: 73, SEQ ID NO: 75, SEQ ID NO: 77, SEQ ID NO: 79, SEQ ID NO: 81, SEQ ID NO: 83, SEQ ID NO: 85, SEQ ID NO: 90, SEQ ID NO: 93, SEQ ID NO:

95, SEQ ID NO: 97, SEQ ID NO: 103, SEQ ID NO: 105, SEQ ID NO: 107, SEQ ID NO: 109, SEQ ID NO: 111, SEQ ID NO: 113, or SEQ ID NO: 105 are provided. Polynucleotides encoding a non-naturally occurring elastin, wherein the polynucleotide is SEQ ID NO: 34, SEQ ID NO: 36, SEQ ID NO: 38, SEQ ID NO: 40, SEQ ID NO: 42, SEQ ID NO: 44, SEQ ID NO: 46, SEQ ID NO: 48, SEQ ID NO: 50, SEQ ID NO: 52, SEQ ID NO: 54, SEQ ID NO: 56, SEQ ID NO: 58, SEQ ID NO: 60, SEQ ID NO: 62, SEQ ID NO: 64, SEQ ID NO: 66, SEQ ID NO: 68, SEQ ID NO: 70, and SEQ ID NO: 72, SEQ ID NO: 99, or SEQ ID NO: 101 provided.

[0027] Host cells that express the polynucleotides of the invention are disclosed. Host cells can be any host cell including bacterial cells, yeast cells, fungal cells, insect cells, mammalian cells, plant cells and any other cells used to express exogenous polynucleotides.

[0028] Bacterial host cells in which the cells have been modified to inhibit cell division and the periplasmic space is increased are provided. An exemplary host cell is *E. coli*.

[0029] One embodiment provides a method of producing a non-naturally occurring collagen or a non-naturally occurring elastin. The method comprises the steps of inoculating a culture medium with a recombinant host cell comprising polynucleotides that encode the collagen or elastin, cultivating the host cell, and isolating the non-naturally occurring collagen or the non-naturally occurring elastin from the host cell.

BRIEF DESCRIPTION OF THE DRAWINGS

[0030] FIG. 1 depicts the physiological state difference between switched and unswitched cells. A) Unswitched *Escherichia coli* cells. B) Same *Escherichia coli* population as figure A but has undergone the physiological switch. C) Phase contrast of switched *Escherichia coli* cell containing cytoplasmic RFP and periplasmic GFP. D) Fluorescent imaging of cell in figure C illustrates targeted protein localization.

[0031] FIG. 2 depicts enhanced protein production in switched cells. A-B) Target protein for T7 inducible protein production is periplasmic expressed GFP, produced in *Escherichia coli* BL21. The same population of cells was used and induced at OD 1.1. A) Protein ladder (lane 1), IPTG induced protein production (lane 2), IPTG induced protein production with physiological switch (lane 3). B) Two vials of the cell GFP induced cultures with IPTG only on left and IPTG+Switch on right. C) Expression of a 22 kDa collagen using switched cells showing protein ladder (lane 1), supernatant after protein production (lane 2), cell pellet (lane 3).

[0032] FIG. 3 depicts a time-lapse of *Escherichia coli* cell switching over time.

[0033] FIG. 4 illustrates other organisms undergoing the physiological switch. A) *Agrobacterium tumefaciens* normal physiology. B) *Agrobacterium tumefaciens* switched physiology. C) *Pseudomonas aeruginosa* PAO1 normal physiology. D) *Pseudomonas aeruginosa* PAO1 switched physiology. E) *Brevundimonas diminuta* normal physiology. F) *Brevundimonas diminuta* switched physiology. G) *Agrobacterium tumefaciens* normal physiology. H) *Agrobacterium tumefaciens* switched physiology.

[0034] FIG. 4 illustrates other organisms undergoing the physiological switch. A) *Agrobacterium tumefaciens* normal physiology. B) *Agrobacterium tumefaciens* switched physiology. C) *Pseudomonas aeruginosa* PAO1 normal physiology. D) *Pseudomonas aeruginosa* PAO1 switched physi-

ogy. E) *Brevundimonas diminuta* normal physiology. F) *Brevundimonas diminuta* switched physiology. G) *Agrobacterium tumefaciens* normal physiology. H) *Agrobacterium tumefaciens* switched physiology.

[0035] FIG. 5 illustrates the reduction in TT dimer formation by treatment of human keratinocytes with truncated collagen.

DESCRIPTION

[0036] In the following description, certain specific details are set forth in order to provide a thorough understanding of various embodiments of the disclosure. However, one skilled in the art will understand that the disclosure may be practiced without these details.

[0037] As used herein the term “about” refers to +10%.

[0038] The term “consisting of” means “including and limited to”.

[0039] The term “consisting essentially of” means that the composition, method or structure may include additional ingredients, steps and/or parts, but only if the additional ingredients, steps and/or parts do not materially alter the basic and novel characteristics of the claimed composition, method or structure.

[0040] As used herein, the singular form “a”, “an” and “the” include plural references unless the context clearly dictates otherwise. For example, the term “a compound” or “at least one compound” may include a plurality of compounds, including mixtures thereof.

[0041] Throughout this application, various embodiments of this disclosure may be presented in a range format. It should be understood that the description in range format is merely for convenience and brevity and should not be construed as an inflexible limitation on the scope of the disclosure. Accordingly, the description of a range should be considered to have specifically disclosed all the possible subranges as well as individual numerical values within that range. For example, description of a range such as from 1 to 6 should be considered to have specifically disclosed sub-ranges such as from 1 to 3, from 1 to 4, from 1 to 5, from 2 to 4, from 2 to 6, from 3 to 6 etc., as well as individual numbers within that range, for example, 1, 2, 3, 4, 5, and 6. This applies regardless of the breadth of the range.

[0042] Whenever a numerical range is indicated herein, it is meant to include any cited numeral (fractional or integral) within the indicated range. The phrases “ranging/ranges between” a first indicate number and a second indicate number and “ranging/ranges from” a first indicate number “to” a second indicate number are used herein interchangeably and are meant to include the first and second indicated numbers and all the fractional and integral numerals there between.

[0043] As used herein the term “method” refers to manners, means, techniques and procedures for accomplishing a given task including, but not limited to, those manners, means, techniques and procedures either known to, or readily developed from known manners, means, techniques and procedures by practitioners of the chemical, pharmacological, biological, biochemical and medical arts.

[0044] The term “collagen” or “collagen-like” as used herein refers to a monomeric polypeptide that can associate with one or more collagen or collagen-like polypeptides to form a quaternary structure. Collagen can be treated with acid, base or heat to prepare gelatin. The quaternary structure of natural collagen is a triple helix typically composed

of three polypeptides. Of the three polypeptides that form natural collagen, two are usually identical and are designated as the alpha chain. The third polypeptide is designated as the beta chain. Thus a typical natural collagen can be designated as AAB, wherein the collagen is composed of two alpha (“A”) strands and one beta (“B”) strand. The term “procollagen” as used herein refers to polypeptides produced by cells that can be processed to naturally occurring collagen.

[0045] The terms “elastin” as used herein refers to a polypeptide that is elastic and functions to stretch or contract and return to its original shape. Elastin is found naturally in connective tissue.

[0046] The term “expression vector” or “vector” as used herein refers to a nucleic acid assembly which is capable of directing the expression of the exogenous gene. The expression vector may include a promoter which is operably linked to the exogenous gene, restriction endonuclease sites, nucleic acids that encode one or more selection markers, and other nucleic acids useful in the practice of recombinant technologies.

[0047] The term “fibroblast” as used herein refers to a cell that synthesizes procollagen and other structural proteins. Fibroblasts are widely distributed in the body and found in skin, connective tissue and other tissues.

[0048] The term “fluorescent protein” is a protein that is commonly used in genetic engineering technologies used as a reporter of expression of an exogenous polynucleotide. The protein when exposed to ultraviolet or blue light fluoresces and emits a bright visible light. Proteins that emit green light is green fluorescent protein (GFP) and proteins that emit red light is red fluorescent protein (RFP).

[0049] The term “gelatin” as used herein refers to collagen that has been further processed by exposure to acid, base or heat. While not wishing to be bound by theory or mechanism, treatment of collagen with acid, base or heat is thought to denature the collagen polypeptides. Aqueous denatured collagen solutions form reversible gels used in foods, cosmetics, pharmaceuticals, industrial products, medical products, laboratory culture growth media, and many other applications.

[0050] The term “gene” as used herein refers to a polynucleotide that encodes a specific protein, and which may refer to the coding region alone or may include regulatory sequences preceding (5' non-coding sequences) and following (3' non-coding sequences) the coding sequence.

[0051] The term “histidine tag” is a 2-30 contiguous series of histidine residues on a recombinant polypeptide.

[0052] The term “host cell” is a cell that is engineered to express an introduced exogenous polynucleotide.

[0053] The term “keratinocyte” is a cell that produces keratins found in the epidermal layer of the skin.

[0054] The term “lactamase” as used herein refer to enzymes that hydrolyze antibiotics that contain a lactam (cyclic amide) moiety. “Beta-lactamase” or “ β -lactamase” are enzymes that hydrolyze antibiotics that contain a β -lactam moiety.

[0055] The term “non-naturally occurring” as used herein refers to collagen or elastin that is not normally found in nature. The non-naturally occurring collagen or elastin are recombinantly prepared. The non-naturally occurring collagen or elastin is a recombinant collagen or recombinant elastin. The non-naturally occurring collagen is in one embodiment a truncated collagen. Other non-naturally occurring collagen polypeptides include chimeric collagens.

A chimeric collagen is a polypeptide wherein one portion of a collagen polypeptide is contiguous with a portion of a second collagen polypeptide. For example, a collagen molecule comprising a portion of a jellyfish collagen contiguous with a portion of a human collagen is a chimeric collagen. In another embodiment, the non-naturally occurring collagen comprises a fusion polypeptide that includes additional amino acids such as a secretion tag, histidine tag, green fluorescent protein, protease cleavage site, GEK repeats, GDK repeats, and/or beta-lactamase. The non-naturally occurring elastin in one embodiment a truncated elastin. Other non-naturally occurring elastin polypeptides include chimeric elastins. A chimeric elastin is a polypeptide wherein one portion of an elastin polypeptide is contiguous with a portion of a second elastin polypeptide. For example, a collagen molecule comprising a portion of a jellyfish elastin contiguous with a portion of a human elastin is a chimeric elastin. In another embodiment, the non-naturally occurring elastin comprises a fusion polypeptide that includes additional amino acids such as a secretion tag, histidine tag, green fluorescent protein, protease cleavage site and/or beta-lactamase. The chimeric gelatin or the chimeric elastin can comprise additional amino acids such as a secretion tag, histidine tag, green fluorescent protein, protease cleavage site, GEK repeats, GDK repeats, and/or beta-lactamase.

[0056] The term “protease cleavage site” is an amino acid sequence that is cleaved by a specific protease.

[0057] The term “secretion tag” or “signal peptide” refers to an amino acid sequence that recruits the host cell’s cellular machinery to transport an expressed protein to a particular location or cellular organelle of the host cell.

[0058] The term “truncated collagen” refers to a monomeric polypeptide that is smaller than a full-length collagen wherein one or more portions of the full-length collagen is not present. Collagen polypeptides are truncated at the C-terminal end, the N-terminal end, or truncated by removal of internal portion(s) of the full-length collagen polypeptide.

[0059] The term “truncated elastin” refers to a monomeric polypeptide that is smaller than a full-length elastin wherein one or more portions of the full-length elastin is not present. Elastin polypeptides are truncated at the C-terminal end, the N-terminal end, or truncated by removal of internal portion(s) of the full-length elastin polypeptide.

[0060] In co-owned application PCT/US17/24857, incorporated by reference, an expression system that uses modified bacterial cells (switched cells) in which cell division is inhibited and growth of the periplasmic space is greatly enhanced was disclosed. In this expression system, the expressed proteins are targeted to the periplasmic space. Recombinant protein production in these switched cells is dramatically increased compared with that in non-switched cells. Structurally, the cells comprise both inner and outer membranes but lack a functional peptidoglycan cell wall, while the cell shape is spherical and increases in volume over time. Notably, while the periplasmic space normally comprises only 10-20% of the total cell volume, the periplasmic compartment of the switched state described herein can comprise more than 20%, 30%, 40% or 50% and up to 60%, 70%, 80% or 90% of the total cell volume.

[0061] The modified bacterial cells of PCT/US17/24857 are derived from Gram-negative bacteria, e.g. selected from: gammaproteobacteria and alphaproteobacteria. In some embodiments, the bacterium is selected from: *Escherichia*

coli, *Vibrio natriegens*, *Pseudomonas fluorescens*, *Caulobacter crescentus*, *Agrobacterium tumefaciens*, and *Brevibacterium diminuta*. In specific embodiments, the bacterium is *Escherichia coli*, e.g. strain BL21(DE3).

[0062] In another aspect, the host bacterial cells have an enlarged periplasmic space in a culture medium comprising a magnesium salt, wherein the concentration of magnesium ions in the medium is at least about 3, 4, 5 or 6 mM. In further embodiments, the concentration of magnesium ions in the medium is at least about 7, 8, 9 or 10 mM. In some embodiments, the concentration of magnesium ions in the medium is between about 5 mM and 25 mM, between about 6 mM and/or about 20, 15 or 10 mM. In some embodiments, the magnesium salt is selected from: magnesium sulfate and magnesium chloride.

[0063] In other embodiments, the culture medium further comprises an osmotic stabilizer, including, e.g. sugars (e.g., arabinose, glucose, sucrose, glycerol, sorbitol, mannitol, fructose, galactose, saccharose, maltotriose, erythritol, ribitol, pentaerythritol, arabitol, galactitol, xylitol, iditol, maltotriose, and the like), betaines (e.g., trimethylglycine), proline, sodium chloride, wherein the concentration of the osmotic stabilizer in the medium is at least about 4%, 5%, 6%, or 7% (w/v). In further embodiments, the concentration of osmotic stabilizer is at least about 8%, 9%, or 10% (w/v). In some embodiments, the concentration of the osmotic stabilizer in the medium is between about 5% to about 20% (w/v).

[0064] In some embodiments, the cell culture may further comprise ammonium chloride, ammonium sulfate, calcium chloride, amino acids, iron(II) sulfate, magnesium sulfate, peptone, potassium phosphate, sodium chloride, sodium phosphate, and yeast extract.

[0065] The host bacterial cell may be cultured continuously or discontinuously; in a batch process, a fed-batch process or a repeated fed-batch process.

[0066] In some embodiments, the antibiotic is selected from: β -lactam antibiotics (e.g. penicillins, cephalosporins, carbapenems, and monobactams), phosphonic acid antibiotics, polypeptide antibiotics, and glycopeptide antibiotics. In particular embodiments, the antibiotic is selected from alafosfalin, amoxicillin, ampicillin, aztreonam, bacitracin, carbenicillin, cefamandole, cefotaxime, cefsulodin, cephalothin, fosmidomycin, methicillin, nafcillin, oxacillin, penicillin G, penicillin V, fosfomicin, primaxin, and vancomycin.

[0067] Without being bound by theory, the cell morphology that promotes recombinant protein production and inhibits cell division appears to be driven by the removal of the cell wall under the media conditions stated above. In some embodiments, the methods for removal/inhibition of cell wall synthesis can be through the use of antibiotics that inhibit peptidoglycan synthesis (such as ampicillin, carbenicillin, penicillins or fosfomicin), or other methods known in the art.

[0068] When having an appropriate periplasmic targeting signal sequence, recombinantly produced polypeptides can be secreted into the periplasmic space of bacterial cells. Joly, J. C. and Laird, M. W., in *The Periplasm* ed. Ehrmann, M., ASM Press, Washington D.C., (2007) 345-360. In the chemically oxidizing environment of the periplasm the formation of disulfide bonds and thereby the functionally correct folding of polypeptides is favored.

[0069] In general, the signal sequence may be a component of the expression vector, or it may be a part of the exogenous gene that is inserted into the vector. The signal

sequence selected should be one that is recognized and processed (i.e., cleaved by a signal peptidase) by the host cell. For bacterial host cells that do not recognize and process the native signal sequence of the exogenous gene, the signal sequence is substituted by any commonly known bacterial signal sequence. In some embodiments, recombinantly produced polypeptides can be targeted to the periplasmic space using the DsbA signal sequence. Dinh and Bernhardt, *J Bacteriol*, September 2011, 4984-4987.

[0070] In one aspect, a non-naturally occurring collagen or elastin is produced by a host cell is provided. The non-naturally occurring collagen or elastin is jellyfish collagen or elastin, human collagen or elastin, or *Chondrosia reniformis* (kidney sponge) collagen or elastin, or *Rhincodon typus* collagen or elastin. The non-naturally occurring collagen or elastin is a truncated collagen. The truncation is an internal truncation, a truncation at the N-terminal portion of the collagen or elastin, or a truncation at the C-terminal portion of the collagen or elastin. The collagen or elastin is truncated by a truncation of between 50 amino acids and 1000 amino acids, between 50 amino acids and 950 amino acids, between 50 amino acids and 900 amino acids, between 50 amino acids and 850 amino acids, between 50 amino acids and 800 amino acids, between 50 amino acids and 750 amino acids, between 50 amino acids and 700 amino acids, between 50 amino acids and 650 amino acids, between 50 amino acids and 600 amino acids, between 50 amino acids and 550 amino acids, between 50 amino acids and 500 amino acids, between 50 amino acids and 450 amino acids, between 50 amino acids and 400 amino acids, between 50 amino acids and 350 amino acids, between 50 amino acids and 300 amino acids, between 50 amino acids and 250 amino acids, between 50 amino acids and 200 amino acids, between 50 amino acids and 150 amino acids, or between 50 amino acids and 100 amino acids. In another embodiment, the collagen or elastin is truncated by 50, 60, 70, 80, 90, 100, 110, 120, 130, 140, 150, 160, 170, 180, 190, 200, 210, 220, 230, 240, 250, 260, 270, 280, 290, 300, 310, 320, 330, 340, 350, 360, 370, 380, 390, 400, 410, 420, 430, 440, 450, 460, 470, 480, 490, 500, 510, 520, 530, 540, 550, 560, 570, 580, 590, 600, 650, 700, 750, 800, 850, 900, 950, or 1000 amino acids. The non-naturally occurring collagen or elastin are encoded by a portion of a polynucleotide sequence or the entire polynucleotide sequence disclosed herein.

[0071] The non-naturally occurring collagen or elastin further comprises amino acid sequences including a secretion tag. The secretion tag directs the collagen or elastin to the periplasmic space of the host cell. In particular embodiments, the signal peptide is derived from DsbA, PelB, OmpA, TolB, MalE, Ipp, TorA, or HylA, or a hybrid secretion tag that comprises a portion of one secretion tag fused to a portion of a second secretion tag. In one aspect the secretion tag is attached to the non-naturally occurring collagen or elastin. In another aspect the secretion tag is cleaved from the non-naturally occurring collagen or elastin.

[0072] The non-naturally occurring collagen or the non-naturally occurring elastin or further comprises a histidine tag. The histidine tag or polyhistidine tag is a sequence of 2 to 20 histidine residues (SEQ ID NO: 117) that are attached to the collagen or elastin. The histidine tag comprises 2 to 20 histidine residues (SEQ ID NO: 117), 5 to 15 histidine residues (SEQ ID NO: 118), 5 to 18 histidine residues (SEQ

ID NO: 119), 5 to 16 histidine residues (SEQ ID NO: 120), 5 to 15 histidine residues (SEQ ID NO: 118), 5 to 14 histidine residues (SEQ ID NO: 121), 5 to 13 histidine residues (SEQ ID NO: 122), 5 to 12 histidine residues (SEQ ID NO: 123), 5 to 11 (SEQ ID NO: 124), 5 to 10 histidine residues (SEQ ID NO: 125), 6 to 12 histidine residues (SEQ ID NO: 126), 6 to 11 histidine residues (SEQ ID NO: 127), or 7 to 10 histidine residues (SEQ ID NO: 128). The histidine tags are useful in purification of proteins by chromatographic methods utilizing nickel based chromatographic media. Exemplary fluorescent proteins include green fluorescent protein (GFP) or red fluorescent protein (RFP). Fluorescent proteins are well known in the art. In one embodiment the non-naturally occurring collagen or the on-naturally occurring elastin comprises a GFP and/or RFP. In one embodiment a superfolder GFP is fused to the on-naturally occurring collagen or elastin. The superfolder GFP is a GFP that folds properly even when fused to a poorly folded polypeptide. In one aspect the histidine tag is attached to the non-naturally occurring collagen or elastin. In another aspect the histidine tag is cleaved from the non-naturally occurring collagen or elastin.

[0073] The non-naturally occurring collagen or non-naturally occurring elastin further comprises a protease cleavage site. The protease cleavage site is useful to cleave the recombinantly produced collagen or elastin to remove portions of the polypeptide. The portions of the polypeptide that may be removed include the secretion tag, the histidine tag, the fluorescent protein tag and/or the Beta-lactamase. The proteases comprise endoproteases, exoproteases, serine proteases, cysteine proteases, threonine proteases, aspartic proteases, glutamic proteases, and metalloproteases. Exemplary protease cleavage sites include amino acids that are cleaved by Thrombin, TEV protease, Factor Xa, Enteropeptidase, and Rhinovirus 3C Protease. In one aspect the cleavage tag is attached to the non-naturally occurring collagen or elastin. In another aspect the cleavage tag is removed by an appropriate protease from the non-naturally occurring collagen or elastin.

[0074] The non-naturally occurring collagen or non-naturally occurring elastin further comprises an enzyme that is a Beta-lactamase. The beta-lactamase is useful as a selection marker. In one aspect the beta-lactamase is attached to the non-naturally occurring collagen or elastin. In another aspect the beta-lactamase is cleaved from the non-naturally occurring collagen or elastin.

[0075] The non-naturally occurring collagen or non-naturally occurring elastin further comprises GEK amino acid trimer repeats and/or GDK amino acid trimer repeats. The GEK and the GDK trimer repeats facilitate the gelling of the collagen and/or the gelatin. In one embodiment, the non-naturally occurring collagen or the non-naturally occurring elastin comprises 2-50 GEK and/or 2-50 GDK trimer repeats (SEQ ID NOS: 130-131, respectively), 2-40 GEK and/or 2-40 GDK trimer repeats (SEQ ID NOS: 132-133, respectively), 2-30 GEK and/or 2-30 GDK trimer repeats (SEQ ID NOS: 134-135, respectively), 2-20 GEK and/or 2-20 GDK trimer repeats (SEQ ID NOS: 136-137, respectively), 2-15 GEK and/or 2-15 GDK trimer repeats (SEQ ID NOS: 138-139, respectively), 2-10 GEK and/or 2-10 GDK trimer repeats (SEQ ID NOS: 140-141, respectively), 2-9 GEK and/or 2-9 GDK trimer repeats (SEQ ID NOS: 142-143, respectively), 2-8 GEK and/or 2-8 GDK trimer repeats (SEQ ID NOS: 144-145, respectively), 2-7 GEK and/or 2-7 GDK

trimer repeats (SEQ ID NOS: 146-147, respectively), 2-6 GEK and/or 2-6 GDK trimer repeats (SEQ ID NOS: 148-149, respectively), 2-5 GEK and/or 2-5 GDK trimer repeats (SEQ ID NOS: 150-151, respectively), or 2-4 GEK and/or 2-4 GDK trimer repeats (SEQ ID NOS: 152-153, respectively). In one aspect the GEK trimer repeat or the GDK trimer repeat is attached to the non-naturally occurring collagen or elastin. In another aspect the GEK trimer repeat or the GDK trimer repeat is cleaved from the non-naturally occurring collagen or elastin.

[0076] Provided herein are compositions that comprise between 0.005% and 30% w/w non-naturally occurring collagen and/or non-naturally occurring elastin. The composition comprises between 0.005% and 20% w/w non-naturally occurring collagen and/or non-naturally occurring elastin, between 0.005% and 10% w/w non-naturally occurring collagen and/or non-naturally occurring elastin, between 0.005% and 5% w/w non-naturally occurring collagen and/or non-naturally occurring elastin, between 0.005% and 2% w/w non-naturally occurring collagen and/or non-naturally occurring elastin, between 0.005% and 1% w/w non-naturally occurring collagen and/or non-naturally occurring elastin, between 0.005% and 0.5% w/w non-naturally occurring collagen and/or non-naturally occurring elastin, and between 0.005% and 0.2% w/w non-naturally occurring collagen and/or non-naturally occurring elastin.

[0077] The compositions that comprise the between non-naturally occurring collagen and/or non-naturally occurring elastin are personal care products. In some embodiments the compositions are formulated for topical administration. The compositions can contain other cosmetic ingredients suitable for human use. The personal care products are useful for preventing or treating ultraviolet radiation damage to human skin or hair. The personal care products are applied to skin or hair. The compositions include, for example, masks, skin cleaners such as soap, cleansing creams, cleansing lotions, cleansing milks, cleansing pads, facial washes, hair shampoo, hair conditioner and body shampoos.

[0078] The compositions that comprise the non-naturally occurring collagen and/or non-naturally occurring elastin can further comprise at least one additional ingredient comprising a topical carrier or a preservative. The topical carrier comprises a topical carrier selected from the group consisting of liposome, biodegradable microcapsule, lotion, spray, aerosol, dusting powder, biodegradable polymer, mineral oil, triglyceride oil, silicone oil, glycerin, glycerin monostearate, alcohols, emulsifying agents, liquid petroleum, white petrolatum, propylene glycol, polyoxyethylene, polyoxypropylene, wax, sorbitan monostearate, polysorbate, cetyl ester wax, cetearyl alcohol, 2-octyldodecanol, benzyl alcohol, cyclomethicone, cyclopentasiloxane, and water. The preservative comprises a preservative selected from the group consisting of tocopherol, diiodomethyl-p-tolylsulfone, 2-Bromo-2-nitropropane-1,3-diol, cis isomer 1-(3-chloroallyl)-3,5,7-triaza-1-azoniaadamantane chloride, glutaraldehyde, 4,4-dimethyl oxazolidine, 7-Ethylbicyclooxazolidine, methyl paraben, sorbic acid, Germaben II, rosemary extract, and EDTA

[0079] Provided are methods of decreasing skin damage, promoting the repair of damaged skin, protecting skin against UV damage, protecting skin cells against the effects of exposure to urban dust. The method comprises the step of applying the composition comprising the non-naturally occurring collagen and/or non-naturally occurring elastin to

the skin of a subject. Without being bound to a particular theory or mechanism, the collagen and/or the elastin in the composition decrease skin damage by protecting against UV damage, and/or promotes the repair of damaged skin by increasing the viability of cells and/or increasing procollagen synthesis when applied to skin, and/or promotes the viability of skin cells. The collagens and elastins in one aspect decrease the formation of thymine-thymine (TT) dimer formation.

[0080] One aspect provides polynucleotides that encode a non-naturally occurring collagen or a non-naturally occurring elastin. The polynucleotides encode collagen or elastin from jellyfish, human, *Chondrosia reniformis* (kidney sponge), or *Rhincodon typus*. The polynucleotides encode for collagen or elastin that is full length or truncated.

[0081] Another aspect provides polynucleotides that encode collagen or elastin fusion proteins. The elastin or collagen fusion proteins comprise a secretion tag, a histidine tag, a fluorescent protein tag, a protease cleavage site, a Beta-lactamase along and/or GEK amino acid trimer repeats and/or GDK amino acid trimer repeats together with collagen or elastin.

[0082] The polynucleotides are in one aspect vectors used to transform host cells and express the polynucleotides. The polynucleotides further comprise nucleic acids that encode enzymes that permit the host organism to grow in the presence of a selection agent. The selection agents include certain sugars including galactose containing sugars or antibiotics including ampicillin, hygromycin, G418 and others. Enzymes that are used to confer resistance to the selection agent include β -galactosidase or a β -lactamase.

[0083] In one aspect host cells that express the polynucleotides of the invention are provided. Host cells can be any host cell including gram negative bacterial cells, gram positive bacterial cells, yeast cells, insect cells, mammalian cells, plant cells or any other cells used to express exogenous polynucleotides. An exemplary gram-negative host cell is *E. coli*.

[0084] Bacterial host cells in which the cells have been modified to inhibit cell division and the periplasmic space is increased are taught. As discussed herein and taught in example 1, Beta-lactam antibiotics are useful as a switch to convert wild-type bacterial cells to a modified bacterial cell in which cell replication is inhibited and the periplasmic space is increased. Exemplary Beta-lactam antibiotics including penicillins, cephalosporins, carbapenems, and monobactams.

[0085] The switched form of bacteria (L-form) are cultivated in culture media that include certain salts and other nutrients. Salts and media compositions that support the physiological switch physiology that have been tested are M63 salt media, M9 salt media, PYE media, and Luria-Bertani (LB) media. Any necessary supplements besides carbon, nitrogen, and inorganic phosphate sources may also be included at appropriate concentrations introduced alone or as a mixture with another supplement or medium such as a complex nitrogen source. In certain embodiments, the medium further comprises one or more ingredients selected from: ammonium chloride, ammonium sulfate, calcium chloride, casamino acids, iron(II) sulfate, magnesium sulfate, peptone, potassium phosphate, sodium chloride, sodium phosphate, and yeast extract.

[0086] Beta-lactamases are enzymes that confer resistance to lactam antibiotics in prokaryotic cells. Typically when

Beta-lactamases are expressed in bacterial host cells, the expressed Beta-lactamase protein also includes targeting sequences (secretion tag) that direct the Beta-lactamase protein to the periplasmic space. Beta-lactamases are not functional unless they are transported to the periplasmic space. Beta-lactamase targeted to the periplasmic without the use of an independent secretion tag that targets the enzyme to the periplasmic space are provided. By creating a fusion protein in which a periplasmic secretion tag added to the N-terminus of a protein such as GFP, collagen, or GFP/collagen chimeras, the functionality of the Beta-lactamase lacking a native secretion tag can be used to select for full translation and secretion of the N-terminal fusion proteins. Using this approach, we have used a DsbA-GFP-Collagen-Beta-lactamase fusion to select for truncation products in the target collagens that favor translation and secretion.

[0087] Another embodiment provides methods of producing a non-naturally occurring collagen or a non-naturally occurring elastin. The method comprises the steps of inoculating a culture medium with a recombinant host cell comprising polynucleotides that encode the collagen or elastin, cultivating the host cell, and isolating the non-naturally occurring collagen or the non-naturally occurring elastin from the host cell.

[0088] A process for fermentative preparation of a protein is provided. The process comprises the steps of:

[0089] a) culturing a recombinant Gram-negative bacterial cell in a medium comprising a magnesium salt, wherein the concentration of magnesium ions in the medium is at least about 6 mM, and wherein the bacterial cell comprises an exogenous gene encoding the protein;

[0090] b) adding an antibiotic to the medium, wherein the antibiotic inhibits peptidoglycan biogenesis in the bacterial cell; and

[0091] c) harvesting the protein from the medium.

[0092] The bacteria may be cultured continuously—as described, for example, in WO 05/021772—or discontinuously in a batch process (batch cultivation) or in a fed-batch or repeated fed-batch process for the purpose of producing the target protein. In some embodiments, protein production is conducted on a large-scale. Various large-scale fermentation procedures are available for production of recombinant proteins. Large-scale fermentations have at least 1,000 liters of capacity, preferably about 1,000 to 100,000 liters of capacity. These fermentors use agitator impellers to distribute oxygen and nutrients, especially glucose (the preferred carbon/energy source). Small-scale fermentation refers generally to fermentation in a fermentor that is no more than approximately 20 liters in volumetric capacity.

[0093] For accumulation of the target protein, the host cell is cultured under conditions sufficient for accumulation of the target protein. Such conditions include, e.g., temperature, nutrient, and cell-density conditions that permit protein expression and accumulation by the cell. Moreover, such conditions are those under which the cell can perform basic cellular functions of transcription, translation, and passage of proteins from one cellular compartment to another for the secreted proteins, as are known to those skilled in the art.

[0094] The bacterial cells are cultured at suitable temperatures. For *E. coli* growth, for example, the typical temperature ranges from about 20° C. to about 39° C. In one embodiment, the temperature is from about 20° C. to about

37° C. In another embodiment, the temperature is at about 30° C. In one embodiment, the host cells, in the non-switched state or switched state are cultivated at one temperature and switched to a different temperature to induce protein production. The host cells are cultivated first at one temperature to propagate the cells, then to induce protein production the cell are cultivated at a lower temperature. The first temperature is 23°, 24°, 25°, 26°, 27°, 28°, 29°, 30°, 31°, 32°, 33°, 34°, 35°, 36°, or 37° C. The second temperature is 20°, 21°, 22°, 23°, 24°, 25°, 26°, 27°, 28°, 29°, 30°, 31°, 32°, 33°, 34°, 35° or 36° C. The cultivation at the second temperature is conducted between 1 hour and 100 hours, between 5 hours and 90 hours, between 5 hours and 80 hours, between 5 hours and 70 hours, between 10 hours and 70 hours, between 15 hours and 70 hours, between 15 hours and 65 hours, between 15 hours and 60 hours, between 20 hours and 60 hours, between 20 hours and 55 hours, between 20 hours and 50 hours, between 24 hours and 50 hours, between 24 hours and 48 hours, between 30 hours and 50 hours, between 30 hours and 45 hours, or between 30 hours and 40 hours.

[0095] The pH of the culture medium may be any pH from about 5-9, depending mainly on the host organism. For *E. coli*, the pH is from about 6.8 to about 7.4, or about 7.0.

[0096] For induction of gene expression, typically the cells are cultured until a certain optical density is achieved, e.g., an OD600 of about 1.1, at which point induction is initiated (e.g., by addition of an inducer, by depletion of a repressor, suppressor, or medium component, etc.) to induce expression of the exogenous gene encoding the target protein. In some embodiments, expression of the exogenous gene is inducible by an inducer selected from, e.g. isopropyl- β -d-1-thiogalactopyranoside, lactose, arabinose, maltose, tetracycline, anhydrotetracycline, vavlycin, xylose, copper, zinc, and the like. The induction of gene expression can also be accomplished by decreasing the dissolved oxygen levels during fermentation. The dissolved oxygen levels of the fermentation during cell propagation is between 10% and 30%. To induce gene expression the dissolved oxygen level is reduced to below 10%, 9%, 8%, 7%, 6%, 5%, 4%, 3%, 2%, 1%, or 0%. In host cells, in either the physiological state or the switched state, protein production can be induced by lowering the temperature of the fermentation as disclosed herein.

[0097] After product accumulation, the cells are vortexed and centrifuged in order to induce lysis and release of recombinant proteins. The majority of the proteins is found in the supernatant but any remaining membrane bound proteins can be released using detergents (such as triton X-100).

[0098] In a subsequent step, the target protein, as a soluble or insoluble product released from the cellular matrix, is recovered in a manner that minimizes co-recovery of cellular debris with the product. The recovery may be done by any means, but in one embodiment, can comprise histidine tag purification through a nickel column. See, e.g., Purification of Proteins Using Polyhistidine Affinity Tags, *Meth-ods Enzymology*. 2000; 326: 245-254.

EXAMPLES

Example 1: Expression System

Materials and Methods:

Strains:

[0099] Tested Physiological Switch and Protein Production:

E. coli BL21(DE3)—From NEB, product # c2527

E. coli K12 NCM3722—From The *Coli* Genetic Stock Center, CGSC #12355

[0100] Tested Physiological Switch:

Gammaproteobacteria:

[0101] *Vibrio natriegens*—From ATCC, product #14048

Pseudomonas fluorescens—From ATCC, product #31948

Pseudomonas aeruginosa PAO1—From ATCC, product # BAA-47

Alphaproteobacteria:

Caulobacter crescentus—From ATCC, product #19089

Agrobacterium tumefaciens/Rhizobium radiobacter—From ATCC, product #33970

Brevundimonas diminuta—From ATCC, product #13184

Media Compositions:

[0104] 1 liter 5 \times m63 salts:

10 g (NH₄)₂SO₄—From P212121, product #7783-20-2

68 g KH₂PO₄—From P212121, product #7778-77-0

2.5 mg FeSO₄.7H₂O—From Sigma Aldrich, product # F7002

Bring volume up to 1 liter with milliQ water

Adjust to pH 7 with KOH (From P212121, product #1310-58-3)

Autoclave mixture

[0105] 1 liter of 1M MgSO₄:

246.5 g MgSO₄.7H₂O—From P212121, (Sigma Aldrich, product #10034-99-8) Bring volume up to 1 liter with milliQ water.

Autoclave mixture.

[0106] 1 Liter of Switch Media 1:

133.4 mL 5 \times m63 salts

10 mL 1M MgSO₄

[0107] 38.6 g Glucose—From P212121, product #50-99-7

66.6 g Sucrose—From P212121, product #57-50-1

8.33 g LB mix—From P212121, product # lb-miller

Bring volume up to 1 liter with milliQ water.

Filter sterilize mixture through a 0.22 μ m pore vacuum filter (Sigma Aldrich, product # CLS430517).

[0108] 1 Liter of Switch Media 2:

133.4 mL 5 \times m63 salts

10 mL 1M MgSO₄

[0109] 38.6 g Glucose—From P212121, product #50-99-7

66.6 g Sucrose—From P212121, product #57-50-1

10 g Yeast Extract—From FisherSci.com, product # J60287A1

Bring volume up to 1 liter with milliQ water.

Filter sterilize mixture through a 0.22 μ m pore vacuum filter (Sigma Aldrich, product # CLS430517).

[0110] For Bioreactor Growth:

5 liter of bioreactor media MGZ12:

1) Autoclave 1 L of Glucose at concentration of 500 g/L in DI water. (VWR, product #97061-170).

2) Autoclave 1 L of Sucrose at concentration of 500 g/L in DI water. (Geneseesci.com, product #62-112).

3) Autoclave in 3946 mL of DI water:

20 g (NH₄)₂HPO₄. (VWR, product #97061-932).

66.5 g KH₂PO₄. (VWR, product #97062-348).

22.5 g H₃C₆H₅O₇. (VWR, product # BDH9228-2.5KG).

2.95 g MgSO₄.7H₂O. (VWR, product #97062-134).

10 mL Trace Metals (Teknova), 1000x. (Teknova, product # T1001).

After autoclaving add 400 mL of (1) to (3), 65 mL of 10M NaOH (VWR, product #97064-480) to (3), and 666 mL of (2) to (3).

A feed of 500 g/L of glucose can be used during fermentation run as needed.

At induction add:

50 mL of 1M MgSO₄.7H₂O to a 5 L bioreactor

1 to 10 mM concentration of IPTG. (carbosynth.com, product # EI05931).

Add Fosfomycin (50 µg/mL or higher) and Carbenicillin (100 µg/mL or higher).

Physiological Switch:

[0111] The physiological switch is optimally flipped at an OD 600 of 1 to 1.1 for *E. coli* for growth in shake flasks at volumes up to 1 L. For the other species tested, cultures were grown in switch media and subcultured once cultures reached maximal OD 600. In all cases the physiological switch is flipped through the addition of 100-200 µg/mL Carbenicillin (From P212121, product #4800-94-6) and 50-100 µg/mL Fosfomycin (From P212121, product #26016-99-9). The majority of the population is in the switched state within a few hours. To confirm that cells underwent a physiological switch, cells were imaged on a Nikon Ti-E with perfect focus system, Nikon CFI60 Plan Apo 100xNA 1.45 objective, Prior automated filter wheels and stage, LED-CFP/YFP/mCherry and LED-DA/FI/TX filter sets (Semrock), a Lumencor Sola II SE LED illumination system, and a Hamamatsu Flash 4.0 V2 CMOS camera.

Image Analysis of Physiological Switch:

[0112] Images were analyzed using ImageJ to measure dimensions. In the switched state, the spherical outline of the outer membrane is treated as a sphere to calculate total volume ($V=(4/3)\pi r^3$). The cytoplasmic volume is calculated as an ellipsoid that exists within the sphere ($V=(4/3)\pi*(\text{longest radius})*(\text{short radius})^2$). To calculate the periplasmic volume, the cytoplasmic volume is subtracted from the total volume of the cell.

Protein Expression and Quantification:

[0113] *E. coli* BL21(DE3) (NEB product # c2527) containing pET28a (emd Millipore product #69864) and its derivatives carrying GFP or collagen derivatives were grown in a shaking incubator at 37° C. overnight in switch media containing 50 mg/mL kanamycin (p212121 product #2251180). Next day, subcultures are started with a 1:10 dilution of the overnight culture into fresh switch media containing 50 mg/mL kanamycin. The culture is then physi-

ologically switched and protein production is induced simultaneously at an OD 600 of 1 to 1.1 (Read on a Molecular Devices Spectramax M2 microplate reader). The physiologically switch and protein production are flipped through the addition of 100 µg/mL Carbenicillin, 50 µg/mL Fosfomycin, and 100 µg/mL IPTG (p212121 product #367-93-1). Protein expression is continued in the switched state from between 8 hours to overnight at room temperature (approximately 22° C.) on an orbital shaker. In order to quantify total protein levels, Quick Start™ Bradford Protein Assay was used on mixed portion of culture and standard curves are quantitated on a Molecular Devices Spectramax M2 microplate reader. In order to quantitate the relative intensity of target protein production relative to the rest of the protein population the mixed portion of the cultures were run on Mini-PROTEAN® TGX™ Gels and stained with Bio-Safe™ Coomassie Stain.

Induction of Protein Production:

[0114] Standard procedures have been followed to induce protein production in the physiological state. We have been using the strain BL21(DE3) containing the plasmid pET28a driving the IPTG/lactose inducible production of recombinant proteins and targeting them to the periplasmic space using the DsbA signal sequence. Using the GFP protein, targeted to the periplasmic space as described above, we have demonstrated the ability to gain and increase of 5-fold in protein production when compared to un-switched cell populations induced at the same optical density, for the same amount of time (figures). The induction was optimal at an OD600 of 1.1 and induction was continued for 10 hours at which point the protein produced was measured at about 200 mg/mL.

Example 2: Production of Full-Length Collagen

[0115] Full length jellyfish collagen was produced using the expression system discussed in Example 1 herein. The wild-type, full length amino acid sequence of *Podocoryna carnea* (jellyfish or Hydrozoan) collagen is provided in SEQ ID NO: 1.

(SEQ ID NO: 1)
 GPQG VV GADGKDGTPGKGEQGRGTGAAGKQSGPGADGARGPLGSGIQQGAR
 GEPGDPGSPGLRGDTGLAGVKGVAGPSGRPGQPANGPLGVNNGRGLRGKP
 GAKGIAGSDGEAGESGAPGQSGPTGPRQRGPGSGEDGNPGLQGLPGSDGEP
 GEEGQPGRSGQPQQGPRGSPGEVGRGSKGPGSGDRGDRGERGVPGQTGSA
 GNVGEDGEGGGKGVGDGASGSPGALGARGPPGSRGDTGAVGPPGPTGRSGLP
 GNAGQKGPSPGEPGSPGKAGSAGEQGPBGKDGSGNPEGSPGKEGERGLAGPP
 GPDGRRGETGSPGIAGALGKPGLEGPKGYPGLRGRDGTNGKRGEQGETGPD
 GVRGIPGNDGQSGKPGIDGIDGTNGQPGEAGYQGGRTGRQLGETGDVGVGN
 GDRGAPGPDGSKGSGRPGPLR
 //www.ncbi.nlm.nih.gov/protein/4379341?report=genbank&log\$=protalign&blast_rank=1&RID=T1N9ZEUW014

[0116] The non-codon optimized polynucleotide sequence encoding the full length jellyfish collagen is disclosed in SEQ ID NO: 2.

(SEQ ID NO: 2)

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ggaccacaaggtgttgtaggagctgatggcaaatggaacaccgggagag
aaaggtgagcaaggacgaaccggagctgcaggaaaaacaggaagccctgga
gcagatggagcaagaggccctcttggatcaatggacaacaaggtgctcgt
ggagaacctggatccaggatctcccggcttaagaggagatactggattg
gctggagtcaggagtagcaggaccatctggctgcacctggacaaccgggt
gcaaatggattacctgggtgtaatggcagagggcgtttgagaggcaaacct
ggtgctaaggaaatgctggcagtgatggagaagcgggagaatctggcgca
cctggacagtcaggacctaccggctccacgtggtaaccaggaccgaagtggt
gaggtggtaaatcctggattacagggtatgcctggctctgatggagagccc
ggagaggaaggacaacctggaagatctggtaaccaggacagcaaggacca
cgtggttccccggagaggtaggaccaagaggatctaagggtccatcagga
gatcgtggtgacaggggagagagaggtgtcctggacaacaggttcggct
ggaaatgtaggagaagatggagagcaaggaggcaaggtgctgatggagcg
agtggaccaagtgagctcttggctcgtggctccccaggaagttagaggt
gacaccggggcagtgaggacctcccggacctactgggctgatcgtgttacct
ggaaacgcaggacaacagggaccgaagtggtgaaccaggttagtcaggaaaa
gcaggatcagctggtgaacagggtcctcctggtaagacggatcaaatggt
gaacctggatctcctggcaagagggtgaacgtggtcttctgctggtccacca
ggtccagatggcagacgtggtgaaacgggatctccaggtatcgtggtgct
cttggtaaacagggtttggaaggacctaaagggttaccaggattaagagga
agagatggaaccaatggcaaacaggagagaacaaggagaaactggtcctgat
ggagtcagaggtatctcctggaaatgatggacaatctggcaaacagggtatt
gatggtatgacggaaacaaatggtcaaccagggtgaggctggataccaaggt
ggtagaggtacacgtggtcagttaggtgaaactggtgatgctggacagaat
ggagatcgaggagctcctggctcctgatggatctaagggttctgctggtaga
ccaggacttcgtgg
https://www.ncbi.nlm.nih.gov/nucleotide/
3355656?report=genbank&log$=nuclalign&
blast_rank=1&RID=TSYP7CMV014

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[0117] Two different codon optimized polynucleotide sequences encoding the wild-type, full-length jellyfish collagen were synthesized. The two polynucleotide sequences were slightly different due to slightly different codon optimization methods. In addition to the non-truncated, full-length jellyfish collagen, the polynucleotides also encoded a secretion tag, a 9 amino acid his tag (SEQ ID NO: 129), a short linker, and a thrombin cleavage site. The DsbA secretion tag is encoded by nucleotides 1-71. The histidine tag comprising 9 histidine residues (SEQ ID NO: 129) is encoded by nucleotides 73-99 and encodes amino acids 25-33. The linker is encoded by nucleotides 100-111. The thrombin cleavage tag is encoded by nucleotides 112-135 and encodes amino acids 38-45. The truncated collagen is encoded by nucleotides 136-1422. The two polynucleotides are disclosed below in SEQ ID NO: 3 and 4.

(SEQ ID NO: 3)

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ATGAAAAAGATTGGCTGGCGCTGGCTGGTTAGTTTTAGCGTTTAGCGCA
TCGGCGGCGCAGTATGAAGATCACCATCACCACCACCACCATCACCCTCT
GGCTCGAGCCTGGTCCC CGCGGCAGCCATATGGTCCCGCAGGGTGTGTGT
GGTGAGATGGTAAAGACGGTACCCCGGGTAAAAAGGAGAACAGGGACGT
ACAGGTGCAGCAGGTAAACAGGGCAGCCCGGGTGCCGATGGTGCCCGTGGC
CCGCTGGGTAGCATTGGTCAGCAGGGTGCAAGAGGCGAACCCGGCGATCCG
GGTAGTCCGGGCTGCGTGGTGTACGGGTCTGGCCGGTGTAAAGCGTGT
GCAGGTCTTCTGGTCGTCAGGTCAACCCGGTGCAAATGGTCTGCCGGGT
GTTAATGGTCGTGGCGGTCTGCGTGGCAAACCCGGGAGCAAAGGTATTGCA
GGTAGCGATGGAGAAGCCGGTAAAGCGGTGCCCGGGTCAGAGTGGTCCG
ACCGGTCCCGCGCGGTGAGCGTGGTCCGTCTGGTGAAGATGGCAATCCGGGT
CTGCAGGGTCTGCCCTGGTAGTGATGGCGAACCCAGGTGAAGAAGGTGAGCCG
GGTCTGTCAGGCCAGCCCGGCAGCAGGGCCCGCTGGTAGCCCGGGCGAA
GTTGGCCCGCGGGTAGTAAAGGTCTTAGTGGCGATCGCGGTGATCGTGGT
GAACCGGTGTCTCTGGTCAGACCGGTAGCGCAGGTAAATGTGGCGAAGAT
GGTGAACAGGGTGGCAAAGGTGTGATGGTGAAGCGGTCCGAGCGGTGCA
CTGGGTGCACGTGGTCTCCGGGCAGCCGTGGTGACACCGGTGCAGTTGGT
CCGCTGGCCCGACCCGGCTAGTGGCTTACCCGGTAAATGCAGGTGAGAAA
GGTCCGTGAGGTGAACCTGGCAGCCCTGGTAAAGCAGGTAGTCCGGTGAG
CAGGGTCCCGCGGGCAAAGATGGTAGTAATGGTGAAGCCGGGTAGCCCTGGC
AAAGAAGGTGAACCTGGTCTGGCAGGACCCCGGGTCTGATGGTCCCGCG
GGTGAACCGGGTTCACCCGGTATTGCGCGGTGCCCTGGGTAAACCAGGTCTG
GAAGGTCCGAAAGGTATCTCTGGTCTGCGCGGTGATGGTACCAATGGC
AAACGTGGCGAACAGGGCGAAACCCGGTCCAGATGGTGTTCGTGGTATTCCG
GGTAACGATGGTCAGAGCGGTAAACCCGGCATGATGGTATTGATGGCACCC
AATGGTCAGCTGGCGAAGCAGGTATCAGGGTGGTCCGGGTACCCGTGGT
CAGCTGGGTGAAACAGGTGATGTTGGTCAGAAATGGTGTGATCGCGGCGCACCC
GGTCCGGATGGTAGCAAAGGTAGCGCGGTGCTCCGGGTTTACGTaa

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(SEQ ID NO: 4)

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ATGAAAAAGATTGGCTGGCGCTGGCTGGTTAGTTTTAGCGTTTAGCGCA
TCGGCGGCGCAGTATGAAGATCACCATCACCACCACCACCATCACCCTCT
GGCTCGAGCCTGGTCCC CGCGGCAGCCATATGGTCCCGCAGGGTGTGTGT
GGTGAGATGGTAAAGACGGTACCCCGGGTAAAAAGGTGAACAGGGTGTG
ACCGGTGCAGCAGGTAAACAGGGCAGCCCGGGTGCCGATGGTGCCCGTGGC
CCGCTGGGTAGCATTGGTCAGCAGGGTGCAAGTGGCGAACCCGGCGATCCG
GGTAGCCCGGGCTGCGTGGTGTACGGGTCTGGCCGGTGTAAAGCGTGT
GCAGGTCTTCTGGTCGTCAGGTCAACCCGGTGCAAATGGTCTGCCGGGT
GTTAATGGTCGTGGCGGTCTGCGTGGCAAACCCGGTGCAAAGGTATTGCA
GGTAGCGATGGCGAAGCCGGTAAAGCGGTGCCCGGGTCAGAGCGGTCCG

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ACCGGTCCGCGCGGTACAGCGTGGTCCGTCTGGTGAAGATGGCAATCCGGGT
 CTGCAGGGTCTGCCTGGTAgcGATGGCGAACAGGTGAAGAAGTCCAGCCG
 GGTCTGTTCTGGCCAGCCGGCCAGCAGGGCCCGCGTGGTAGCCCGGGCGAA
 GTTGGCCCCGCGcGGTtcTAAAGGTCCTAGcGGCGATCGCGGTGATCGTGGT
 GAACCGCGTGTTCCTGGTcAGACCGGTAGCGCAGGTAATGTTGGCGAAGAT
 GGTGAACAGGGTGGCAAAGGTGTTGATGGTGAAGCGGTCCGAGCGGTGCA
 CTGGGTGCACGTGGTCTCCGGGCAGCCGTGGTGACACCGGTGCAGTTGGT
 CCGCTGGCCCCGACCGCCGTAGcGGCctgCCGGTAATGCAGGTcAGAAA
 GGTCCGTCTcGGTGAACCTGGCAGCCCTGGTAAAGCAGGTAGcGCCGGTGA
 CAGGGTCCCGCGGCAAAGATGGTAGcAATGGTGAAGCCGGTGGTCCCTGGC
 AAAGAAGGTGAACGTGGTCTGGCAGGTCCGCGGGTCTGTATGGTCCGCCG
 GGTGAACAGGGTTCCTCCGGTATTGCCGGTCCCTGGGTAACACAGGTCTG
 GAAGGTCCGAAAGGTTATCCTGGTCTGCGCGGTCTGTATGGTACCAATGGC
 AAACGTGGCGAACAGGGCGAAACCGGTCCAGATGGTGTTCGTGGTATTCCG
 GGTAAACGATGGTcAGAGCGGTAAACCGGCATTGATGGTATTGATGGCAC
 AATGGTcAGCCTGGCGAAGCAGGTTATCAGGGTGGTCCGCGGTACCCGTGGT
 CAGCTGGGTGAACcGGTATGTTGGTcAGAATGGTATCGCGCGCACCCG
 GGTCCGGATGGTAGCAAAGGTAGCGCCGGTCTCCGGTctgCGTtaa

[0118] The amino acid sequence encoded by the polynucleotides of SEQ ID NO: 3 and SEQ ID NO:4 is disclosed in SEQ ID NO:5 below. In SEQ ID NO: 5 the DsbA secretion tag is encoded by nucleotides 1-71 and encodes amino acids 1-24; the histidine tag comprising 9 histidine residues (SEQ ID NO: 129) is encoded by nucleotides 73-99 and encodes amino acids 25-33; the linker is encoded by nucleotides 100-111 and encodes amino acids 34-37; the thrombin cleavage tag is encoded by nucleotides 112-135 and encodes amino acids 38-45; the full-length collagen is encoded by nucleotides 136-1422 and encodes amino acids 46-474.

(SEQ ID NO: 5)

MKKIWLALAGLVLAFSASAAQYEDHHHHHHHSGSSLVPRGSHMGPGQGVV
 GADGKDGTPEKEQGRGTGAAGKQSGADGARGPLGSIQGGARGEPGDP
 GSPGLRGDTGLAGVKGVAGPSGRPGQPGANGLPGVNGRGLRGKPGAKGIA
 GSDGEAGESGAPGQSGPTGPRGQRPSPGEDGNPGLQGLPGSDGEPGEEGQP
 GRSGQPQQGPRGSPEVGPGRSGKPSGDRGDRGERGVPGQTGSAGNVGED
 GEQGGKGVGDGASGPSALGARGPPGSRGDTGAVGPPGPTGRSGLPLGNAGQK
 GPSGEPGSPGKAGSAGEQPPGKDGSNPEGSPGKEGERGLAGPPGPDGRR
 GETGSPGIAGALGKPGLEGPKGYPLGRDRDGTNGKRGEQGETGPDGVRGIP
 GNDGQSGKPGIDGIDGTNGQPEAGYQGGRTGRQLGETGDVGVQNGDRGAP
 GPDGSKGSAGRPLGR

[0119] The full length jellyfish collagen without the DsbA secretion tag, the histidine tag, linker and thrombin cleavage site is disclosed in SEQ ID NO: 89.

(SEQ ID NO: 89)

GPQGVVADGKDGTPGEKEQGRGTGAAGKQSGADGARGPLGSIQGGAR
 GEPGDPGSPGLRGDTGLAGVKGVAGPSGRPGQPGANGLPGVNGRGLRGK
 GAKGIAGSDGEAGESGAPGQSGPTGPRGQRPSPGEDGNPGLQGLPGSDGEP
 GEEGQPGRSGQPQQGPRGSPGEVGPGRSGKPSGDRGDRGERGVPGQTGSA
 GNVGEDGEQGGKGVGDGASGPSALGARGPPGSRGDTGAVGPPGPTGRSGLP
 GNAGQKPSGEPGSPGKAGSAGEQPPGKDGSNPEGSPGKEGERGLAGPP
 GPDGRRGETGSPGIAGALGKPGLEGPKGYPLGRDRDGTNGKRGEQGETGPD
 GVRGIPGNDGQSGKPGIDGIDGTNGQPEAGYQGGRTGRQLGETGDVGVQ
 GDRGAPGPDGSKGSAGRPLGR

[0120] The polynucleotides of SEQ ID NO: 3 and SEQ ID NO: 4 were synthesized by Gen9 DNA, now Ginkgo Bio-works internal synthesis. Overlaps between the pET28 vector and SEQ ID NO: 3 and SEQ ID NO: 4 were designed to be between 30 and 40 bp long and added using PCR with the enzyme PrimeStar GXL polymerase (http://www.clontech.com/US/Products/PCR/GC_Rich/PrimeSTAR_GXL_DNA_Polymerase?site=10020:22372:US). The opened pET28a vector and insert DNA (SEQ ID NO: 3 or SEQ ID NO: 4) were then assembled together into the final plasmid using SGI Gibson assembly (<https://us.vwr.com/store/product/17613857/gibson-assembly-hifi-1-step-kit-synthetic-genomics-inc>). Sequence of plasmid was then verified through sanger sequencing through Eurofins Genomics (www.eurofinsgenomics.com).

[0121] The transformed cells were cultivated in minimal media and frozen in 1.5 aliquots with glycerol at a ratio of 50:50 of cells to glycerol. One vial of this frozen culture was revived in 50 ml of minimal media overnight at 37° C., 200 rpm. Cells were transferred into 300 ml of minimal media and grown for 6-9 hours to reach an OD600 of 5-10.

[0122] Minimal media used in this example and throughout this application is prepared as follows. The minimal media (Table 1) was autoclaved in several separate fractions, Salts mix (Ammonium Phosphate dibasic, Potassium phosphate monobasic, Citric acid anhydrous, Magnesium sulfate heptahydrate), the Sucrose at 500 g/L, the Glucose at 55%, the Trace Metals TM5 (table 2), and Sodium Hydroxide 10M. The minimal media was then mixed together at the above concentrations post-autoclaving in the hood.

TABLE 1

Minimal media recipe for shake flask cultures			
chemical	Formula	MW	Conc (g/L)
Ammonium Phosphate dibasic	(NH ₄) ₂ HPO ₄	133	4
Potassium phosphate monobasic	KH ₂ PO ₄	137	13.3
Citric acid anhydrous	H ₃ C ₆ H ₅ O ₇	192.14	4.5
Magnesium sulfate heptahydrate	MgSO ₄ •7H ₂ O	246	0.59
Trace Metals TM5			2
Glucose	C ₆ H ₁₂ O ₆	500	40
Sodium Hydroxide 10M	NaOH	400	5.2
Sucrose 500 g/L	C ₁₂ H ₂₂ O ₁₁	500	66.6

TABLE 2

Trace Metals TMS composition			
chemical	Formula	MW	Conc (g/L)
Ferrous Sulfate Heptahydrate	FeSO ₄ •7H ₂ O	278.02	27.8
Calcium Chloride	CaCl ₂ •2H ₂ O	147	2.94
Manganese Chloride	MnCl ₂	125.84	1.26
Zinc Sulfate	ZnSO ₄ •H ₂ O	179.5	1.8
Nickel Chloride	NiCl ₂ •6H ₂ O	237.69	0.48
Sodium Molybdate	Na ₂ MoO ₄ •2H ₂ O	241.95	0.48
Sodium Selenite	Na ₂ SeO ₃	172.94	0.35
Boric Acid	H ₃ BO ₃	61.83	0.12

[0123] The harvested cells were disrupted in a homogenizer at 14,000 psi pressure in 2 passes. Resulting slurry contained the collagen protein along with other proteins.

[0124] The collagen was purified by acid treatment of homogenized cell broth. The pH of the homogenized slurry was decreased to 3 using 6M Hydrochloric acid. Acidified cell slurry was incubated overnight at 4° C. with mixing, followed by centrifugation. Supernatant of the acidified slurry was tested on a polyacrylamide gel and found to contain collagen in relatively high abundance compared to starting pellet. The collagen slurry thus obtained was high in salts. To obtain volume and salt reduction, concentration and diafiltration steps were performed using an EMD Millipore Tangential Flow Filtration system with ultrafiltration cassettes of 0.1 m² each. Total area of filtration was 0.2 m² using 2 cassettes in parallel. A volume reduction of 5x and a salt reduction of 19x was achieved in the TFF stage. Final collagen slurry was run on an SDS-PAGE gel to confirm presence of the collagen. This slurry was dried using a multi-tray lyophilizer over 3 days to obtain a white, fluffy collagen powder.

[0125] The purified collagen was analyzed on an SDS-PAGE gel and a thick and clear band was observed at the expected size of 42 kilodaltons. The purified collagen was also analyzed by mass spectrometry and it was confirmed that the 42 kilodalton protein was jellyfish collagen.

[0126] The fermentations were performed at various temperature ranging from 25° to 28° C. For some fermentations, the temperature of the fermentation was maintained at a constant temperature and immediately upon completion of fermentation (OD600 of 5-10) the collagen was purified. For other fermentations, the temperature of the fermentations was maintained for a desired period of time and when cell densities of OD600 of 5-10 were reached, the temperature was reduced to induce protein production. Typically, the temperature was reduced from 28° C. to 25° C. After the fermentation at 25° C. was continued for 40-60 hours, the collagen was isolated.

Additional Full Length Jellyfish Collagens

[0127] A full length jellyfish collagen without a His tag, linker, and thrombin cleavage site is disclosed below. Two codon-optimized nucleotide sequence encoding this collagen are provided in SEQ ID NO: 6 and SEQ ID NO: 7. The differences in the nucleotide sequences are due to different codon-optimization strategies but encode the same protein. The amino acid sequence is disclosed in SEQ ID NO: 8. The DsbA secretion tag is encoded by nucleotides 1-72 and encodes amino acids 1-24. The collagen sequence is encoded by nucleotides 73-1359 and encodes amino acids 25-453.

(SEQ ID NO: 6)
 ATGAAAAAGATTGGCTGGCGCTGGCTGGTTAGTTTTAGCGTTTAGCGCA
 TCGGCGGCGCAGTATGAAGATGGTCCGCAGGGTGTGTTGGTGCAGATGGT
 AAAGACGGTACCCCGGGTGAAAAAGGAGAACAGGGACGTACAGGTGCAGCA
 GGTAACAGGGCAGCCCGGGTGCCGATGGTCCCGTGGCCCGCTGGGTAGC
 ATTGGTCAGCAGGGTGCAAGAGGCGAACCGGGCGATCCGGGTAGTCCGGGC
 CTGCGTGGTGATACGGGTCTGGCCGGTGTAAAGGCGTTGCAGGTCCCTCA
 GGTCTGCCAGGTCAACCGGGTGCAAAATGGTCTGCCGGGTGTTAATGGTCTG
 GGCGGTCTGCGTGGCAAACCGGGAGCAAAAGGTATTGCAGGTAGCGATGGA
 GAAGCCGGTGAAAGCGGTGCCCGGGTCAGAGTGGTCCGACCGGTCCGCGC
 GGTACAGCGTGGTCCGTCTGGTGAAGATGGCAATCCGGGTCTGCAGGGTCTG
 CCTGGTAGTGATGGCGAACAGGTGAAGAAGGTGAGCCGGGTGTTTCAGGC
 CAGCCGGGCCAGCGGGCCCGGTGGTAGCCCGGGCGAAGTTGGCCCGCGG
 GGTAGTAAAGTCTAGTGGCGATCGCGGTGATCGTGGTGAACCGGTGTT
 CCTGGTCAGACCGGTAGCGCAGGTAATGTTGGCGAAGATGGTGAACAGGGT
 GGCAAAGGTGTTGATGGTGCAAGCGGTCCGAGCGGTGCACTGGGTGCACGT
 GGTCTCCGGGCAGCCGTGGTACACCGGTGCAGTTGGTCCGCCTGGCCCG
 ACCGGCCGTAGTGGCTTACCGGGTAATGCAGGTGAGAAAGGTCCGTGAGGT
 GAACCTGGCAGCCCTGGTAAAGCAGGTAGTCCCGGTGAGCAGGGTCCGCGG
 GGCAAAGATGGTAGTAAATGGTGAAGCGGGTAGCCCTGGCAAAGAAGGTGAA
 CGTGGTCTGGCAGGACCCCGGGTCTGATGGTCCCGCGGTGAAACGGGT
 TCACCGGGTATTGCCGGTGCCCTGGGTAAACAGGTCTGGAAGGTCCGAAA
 GGTATTCTGGTCTGCGCGGTGATGGTACCAATGGCAAACGTGGCGAA
 CAGGGCGAAACCGGTCCAGATGGTGTTCGTGGTATTCCGGGTAACGATGGT
 CAGAGCGGTAAACCGGGCATTGATGGTATTGATGGCAACCAATGGTCAAGCT
 GGCGAAGCAGGTATTAGGGTGGTCCCGGTACCGGTGGTCAAGTGGGTGAA
 ACAGGTGATGTTGGTGAATGGTATCGCGCGCACCGGGTCCGGATGGT
 AGCAAAGGTAGCGCCGGTCTGTCGGGTTTACGTTaa

(SEQ ID NO: 7)
 ATGAAAAAGATTGGCTGGCGCTGGCTGGTTAGTTTTAGCGTTTAGCGCA
 TCGGCGGCGCAGTATGAAGATGGTCCGCAGGGTGTGTTGGTGCAGATGGT
 AAAGACGGTACCCCGGGTGAAAAAGGtGAACAGGGtCGTACCGGTGCAGCA
 GGTAACAGGGCAGCCCGGGTGCCGATGGTCCCGTGGCCCGCTGGGTAGC
 ATTGGTCAGCAGGGTGCAcgtGGCGAACCGGGCGATCCGGGTAGcCGGGC
 CTGCGTGGTGATACGGGTCTGGCCGGTGTAAAGGCGTTGCAGGTCCCTCt
 GGTCTGCCAGGTCAACCGGGTGCAAAATGGTCTGCCGGGTGTTAATGGTCTG
 GGCGGTCTGCGTGGCAAACCGGGtGCAAAAGGTATTGCAGGTAGCGATGGc
 GAAGCCGGTGAAAGCGGTGCCCGGGTCAGAGcGGTCCGACCGGTCCGCGC
 GGTACAGCGTGGTCCGTCTGGTGAAGATGGCAATCCGGGTCTGCAGGGTCTG
 CCTGGTAgcGATGGCGAACCGGTGAAGAAGGTGAGCCGGGTGCTTcGGC

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CAGCCGGGCCAGCAGGGCCCGCGTGGTAGCCCGGGCGAAGTTGGCCCCGCGc
GGTtcTAAAGTCCTAGcGGCGATCGCGGTGATCGTGGTGAACCGGTGTT
CCTGGT CAGACCCGGTAGCGCAGGTAATGTTGGCGAAGATGGTGAACAGGGT
GGCAAAGGTGTTGATGGTGCAAGCGGTCCGAGCGGTGCACCTGGGTGCACGT
GGTCTCCGGGCAGCCGTGGTGACACCGGTGCAGTTGGTCCGCCCTGGCCCCG
ACCGGCCGTAGcGGCctgCCGGTAATGCAGGTCAGAAAGTCCCGTctGGT
GAACCTGGCAGCCCCTGGTAAAGCAGGTAGcGCCGGTGAGCAGGGTCCGCCG
GGCAAAGATGGTAGcAATGGTGAAGCGGTAGCCCTGGCAAAGAAGGTGAA
CGTGGTCTGGCAGGtCCGCCGGTCTGATGGTGCAGCGGTGAAACGGGT
TctCCGGGTATTGCCGGTGCCTGGGTAAACCAGGTCTGGAAGGTCCGAAA
GGTTATCTGGTCTCGCGGTCTGATGGTACCAATGGCAAACGTGGCGAA
CAGGGCGAAACCGGTCCAGATGGTGTTCGTGGTATTCCGGGTAACGATGGT
CAGAGCGGTAAACCGGGCATTGATGGTATTGATGGCACCAATGGTCAAGCT
GGCGAAGCAGGTTATCAGGGTGGTCCGGTACCCGTGGTCAAGTGGGTGAA
ACcGGTGATGTTGGTCAGAATGGTATCGCGCGCACCCGGTCCGGATGGT
AGCAAAGGTAGCCCGGTCTCGGGTctgCGTtaa

(SEQ ID NO: 8)

MKKIWLALAGLVLAFSASAAQYEDGPQGVVADGKDGTPGEKGEQGRGTGAA
GKQGSFGADGARGPLGSIQQQGARGEPDGPSPGLRGTGLAGVKGVAGPS
GRPGQPGANGLPGVNGRGLRGPAGKGIAGSDGEAGESGAPGQSGPTGPR
GQRGSPGEDGNPGLQGLPGSDGEPGEGQPGRSGQPQGGPRGSPGVEVGR
GSKGSPGDRDRGERGVPQGTSGAGNVGEGDGEQGGKGVGDGASGSPSALGAR
GPPGSRGDTGAVGPPGPTGRSLPGLNAGQKGPSEPGSPGKAGSAGEQPP
GKDGSNPEGSPGKEGERGLAGPPGPDGRRGETGSPGIAGALGKPGLEGPK
GYPGLRGRDGTNGKRGEQGETGPDVVRGIPGNDGQSGKPGIDGIDGTNGQP
GEAGYQGGRTGRLGETGQNGDRGAPGPDGSKGSAGRPLR

Example 3: Production of Truncated Collagen

[0128] A codon optimized DNA sequence, optimized for expression in E. coli, encoding a jellyfish collagen with a truncation of 240 internal amino acids was synthesized and expressed. The DNA sequence is shown below in SEQ ID NO: 9. In SEQ ID NO: 9, The DsbA secretion tag is encoded by nucleotides 1-72 and encodes amino acids 1-24 of SEQ ID NO: 10. The histidine tag comprising 9 histidine (SEQ ID NO: 129) residues is encoded by nucleotides 73-99 and encodes amino acids 25-33 of SEQ ID NO: 10. The linker is encoded by nucleotides 100-111 and encodes amino acids 34-37 of SEQ ID NO: 10. The thrombin cleavage site is encoded by nucleotides 112-135 and encodes amino acids 38-45 of SEQ ID NO: 10. The truncated collagen is encoded by nucleotides 136-822 and encodes amino acids 46-274 of SEQ ID NO: 10.

(SEQ ID NO: 9)

ATGAAAAAGATTGGCTGGCGCTGGCTGGTTAGTTTTCAGCTTTAGCGCA
TCGGCGCGCAGTATGAAGATCACCATCACCAACCACCATCACCACTCT

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GGCTCGAGCCTGGTGCCGCGCGCAGCCATATGGGTCCGCAGGGTGTGTT
GGTG CAGATGGTAAAGACCGGTACCCCGGTGAAAAAGGAGAACAGGGACGT
ACAGGTGCAGCAGGTAACAGGGCAGCCCGGTGCCGATGGTGCCCGTGGC
CCGCTGGGTAGCATTGGTCAGCAGGGTGAAGAGCGCAACCCGGCGATCCG
GGTAGTCCGGGCTCGTGGTGATACGGGTCTGGCCGGTGTAAAGGCGTT
GCAGGTCTTCAGGTCGTCCAGGTCAACCGGTGCAAATGGTCTGCCGGGT
GTTAATGGTCTGGCGGTCTGGAACGTGGTCTGGCAGGACCCCGGGTCTT
GATGGTCCGCGCGGTGAAACGGGTTCACCGGTATTGCCGGTGCCTGGGT
AAACAGGTCTGGAAGGTCCGAAAGGTTATCTGGTCTGCGCGGTCTGAT
GGTACCAATGGCAAACGTGGCGAACAGGGCGAAACCGGTCCAGATGGTGT
CGTGGTATTCCGGGTAACGATGGTCAAGCGGTAAACCGGGCATTGATGGT
ATTGATGGCACCAATGGTCAAGCTGGCGAAGCAGGTTATCAGGGTGGTCCG
GGTACCCTGGTCAAGTGGTGAACAGGTGATGTTGGTCAGAATGGTGTGAT
CGCGCGCACCCGGTCCGGATGGTAGCAAAGGTAGCGCCGGTCTCGGGT
TTACGTtaa

[0129] The truncated collagen is approximately 54% of the full length collagen and is disclosed below in SEQ ID NO: 10.

(SEQ ID NO: 10)

MKKIWLALAGLVLAFSASAAQYEDHHHHHHHHHSGSSLVPRGSHMGPQGVV
GADGKDGTPGEKGEQGRTGAAKQGSFGADGARGPLGSIQQQGARGEPDGP
GSPGLRGTGLAGVKGVAGPSGRPGQPGANGLPGVNGRGLRGLAGLPPGP
DGRRETGSPGIAGALGKPGLEGPKGYPGLRGRDGTNGKRGEQGETGPDGV
RGIPGNDGQSGKPGIDGIDGTNGQPGEAGYQGGRTGRLGETGQNGDRGAP
GPDGSKGSAGRPLR

[0130] The polynucleotide encoding the truncated jellyfish collagen without the DsbA secretion tag, the histidine tag, linker and thrombin cleavage site is disclosed in SEQ ID NO: 85

(SEQ ID NO: 85)

GTCCGCAGGGTGTGTTGGTGCAGATGGTAAAGACCGGTACCCCGGTGAAA
AAGGAGAACAGGGACGTACAGGTGCAGCAGGTAACAGGGCAGCCCGGGTG
CCGATGGTGCCCGTGGCCCGCTGGGTAGCATTGGTCAGCAGGGTGAAGAG
GCGAACCCGGCGATCCGGGTAGTCCGGGCTCGTGGTGATACGGGTCTGG
CCGGTGTAAAGGCGTTGCAGGTCTTCAGGTCGTCCAGGTCAACCGGGTG
CAAATGGTCTGCCGGTGTAAATGGTCTGGCGGTCTGGAACGTGGTCTGG
CAGGACCCCGGGTCTGATGGTCCGCGCGGTGAAACGGGTTCACCGGGTA
TTGCCGGTCCCTGGGTAAACAGGTCTGGAAGGTCCGAAAGGTTATCTGT
GTCTGCGCGGTCTGATGGTACCAATGGCAAACGTGGCGAACAGGGCGAAA

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CCGGTCCAGATGGTGTTCGTGGTATTCGGGTAACGATGGTCAGAGCGGTA
 AACCGGGCATTGATGGTATTGATGGCACCAATGGTCAGCCTGGCGGAAGCAG
 GTTATCAGGGTGGTCGCGGTACCCGTGGTCAGCTGGGTGAAACAGGTGATG
 TTGGTCAGAATGGTGTATCGCGGCGCACCGGGTCCGGATGGTAGCAAAGGTA
 GCGCCGGTCTCGCGGGTTACGTTaa

[0131] The truncated jellyfish collagen without the DsbA secretion tag, the histidine tag, linker and thrombin cleavage site is disclosed in SEQ ID NO: 86

(SEQ ID NO: 86)

GPQGVVADGKDGTPGEKGEQGRGTGAAGKQSGADGARGPLGSIQQGAR
 GEPGDPGSPGLRGDTGLAGVKVAGPSGRPGQPGANGLPGVNGRGLERGL
 AGPPGPDGRRGETGSPGIAGALGKPLEGPKGYGLRGRDGTNGKRGEQGE
 TGPDGVRRIPGNDGQSGKPGIDIGDGTNGQPEAGYQGGRRTRQGLGETGD
 VGQNGDRGAPGPDGSKGSAGRPLR

[0132] The polynucleotides of SEQ ID NO: 9 was codon optimized and synthesized by Gen9 DNA, now Ginkgo Bioworks internal synthesis. Overlaps between the pET28 vector and SEQ ID NO: 9 were designed to be between 30 and 40 bp long and added using PCR with the enzyme PrimeStar GXL polymerase (http://www.clontech.com/US/Products/PCR/GC_Rich/PrimeSTAR_GXL_DNA_Polymerase?site=10020:22372:US). The opened pET28a vector and insert DNA (SEQ ID NO: 9) was then assembled together into the final plasmid using SGI Gibson assembly (<https://us.vwr.com/store/product/i7613857/gibson-assembly-hifi-1-step-kit-synthetic-genomics-inc>). Sequence of plasmid was then verified through sanger sequencing through Eurofins Genomics (www.eurofinsgenomics.com).

[0133] The transformed cells were cultivated in minimal media and frozen in 1.5 aliquots with glycerol at a ratio of 50:50 of cells to glycerol. One vial of this frozen culture was revived in 50 ml of minimal media overnight at 37° C., 200 rpm. Cells were transferred into 300 ml of minimal media and grown for 6-9 hours to reach an OD600 of 5-10.

[0134] A bioreactor was prepared with 2.7 L of minimal media+glucose and 300 ml of OD600 of 5-10 culture was added to bring the starting volume to 3 L. Cells were grown at 28° C., pH7 with Dissolved Oxygen maintained at 20% saturation using a cascade containing agitation, air and oxygen. pH was controlled using 28% w/w Ammonium hydroxide solution. Fermentation was run in a fed-batch mode using a DO-stat based feeding algorithm once the initial bolus of 40 g/L was depleted around 13 hours. After 24-26 hours of initial growth, the OD600 reached above 100. At this point, 300 mL of 500 g/L sucrose was added and temperature was reduced to 25 C. High density culture was induced for protein production using 1 mM IPTG. Fermentation was continued for another 20-24 hours and cells were harvested using a bench top centrifuge at 9000 rcf, 15 C for 60 minutes. Cell pellet recovered from centrifugation was resuspended in a buffer containing 0.5M NaCl and 0.1M KH₂PO₄ at pH8 in a weight by weight ratio of 2x buffer to 1x cells.

[0135] The harvested cells were disrupted in a homogenizer at 14,000 psi pressure in 2 passes. Resulting slurry contained the collagen protein along with other proteins.

[0136] The fermentations were performed at various temperature ranging from 25° to 28° C. For some fermentations, the temperature of the fermentation was maintained at a constant temperature and immediately upon completion of fermentation (OD600 of 5-10) the collagen was purified. For other fermentations, the temperature of the fermentations was maintained for a desired period of time and when cell densities of OD600 of 5-10 were reached, the temperature was reduced to induce protein production. Typically, the temperature was reduced from 28° C. to 25° C. After the fermentation at 25° C. was continued for 40-60 hours, the collagen was isolated.

[0137] The collagen was purified by acid treatment of homogenized cell broth. Additionally, acid treatment was also performed on non-homogenized whole cells recovered from the bioreactor after centrifugation and resuspension in the buffer described above. The pH of either the homogenized slurry of the resuspended whole cells was decreased to 3 using 6M Hydrochloric acid. Acidified cell slurry was incubated overnight at 4° C. with mixing, followed by centrifugation. Supernatant of the acidified slurry was tested on a polyacrylamide gel and found to contain collagen in relatively high abundance compared to starting pellet. The collagen slurry thus obtained was high in salts. To obtain volume and salt reduction, concentration and diafiltration steps were performed using an EMD Millipore Tangential Flow Filtration system with ultrafiltration cassettes of 0.1 m² each. Total area of filtration was 0.2 m² using 2 cassettes in parallel. A volume reduction of 5x and a salt reduction of 19x was achieved in the TFF stage. Final collagen slurry was run on an SDS-PAGE gel to confirm presence of the collagen. This slurry was dried using a multi-tray lyophilizer over 3 days to obtain a white, fluffy collagen powder.

[0138] The purified truncated collagen obtained from homogenized cell broth or non-homogenized cells were analyzed on an SDS-PAGE gel and a thick and clear band was observed at the expected size of 27 kilodaltons. The purified collagen was also analyzed by mass spectrometry and it was confirmed that the 27 kilodalton protein was jellyfish collagen.

[0139] An alternative purification method of the full length and truncated collagens is provided below.

[0140] The fermentation broth was mixed with 0.3-0.5% w/v of Poly Ethyl Imine (PEI). After 15 minutes of incubation with PEI, the fermentation broth was centrifuged at 9000 rcf for 15 minutes to recover the supernatant, which contained the collagen protein. The pellet containing the cells was discarded and the PEI-treated collagen containing supernatant was mixed with Sodium Bentonite (0.2% final w/v) (Wyopure®, Wyoming Bentonite) and centrifuged. The bentonite containing pellet was discarded and the supernatant was recovered.

[0141] The Bentonite treated supernatant was concentrated between 3-6 fold on a tangential flow filtration system (TFF) (EMD Millipore) using a 5 kDa cassette. The collagen was retained with almost no losses in the permeate stream. To remove salts, the retentate from the concentration step was diafiltered using the same TFF set-up. Final conductivity of the protein solution was <10 milliSiemens. The typical conductivity was between 400 microsiemens and 1.5 millisiemens. Highly concentrated collagen solutions had higher conductivities approaching 4 milliSiemens. A skilled artisan will understand that conductivities higher than 10 milliSiemens may be observed depending on the concentra-

tion of the collagen. Next, the desalted and concentrated protein was subjected to treatment with activated carbon using the W-L 9000 10×40 granulated resin (Carbon Activated Corporation). 5% w/v of the carbon resin was mixed with collagen containing protein feed and mixed at 45-50° C. with mild agitation. The carbon-treated slurry was filtered using a Buchner funnel lined with an Ertel Filter Press Pad M-953 (Ertel Alsop) in presence or absence of a filtration aid such as Diatomaceous Earth (Sigma Aldrich). Post-filtration, the collagen solution was filtered through a 0.2 micron filter followed by one to several hours of treatment with Sodium Bentonite (0.2% w/v final) (Wyopure®, Wyoming Bentonite) and centrifuged at 9000 rcf, 15-30 minutes to obtain a highly pure, clear and particulate free collagen solution. When removal of endotoxin proteins was desired, the protein was passed through a chromatographic filter like Sartobind-Q (Sartorius-Stedim) to specifically remove endotoxin proteins.

[0142] The purified collagen was analyzed on an SDS-PAGE gel and a thick and clear band was observed at 30 kilodaltons. The upshift in size is due to the structure of the collagen molecule and the high glycine/proline amino acid content. The purified collagen was also analyzed by mass spectrometry and it was confirmed that the 30 kilodalton protein was the truncated collagen.

[0143] The truncated collagens were further analyzed by HPLC using an Agilent 1100 series HPLC. The column was the 50 mm Agilent PLRP-S reverse phase column with an inner diameter of 4.6 mm, M particle size and 1000 Angstrom pore size.

[0144] The sample was prepared by diluting 1:1 in a 0.04% sodium azide solution in HPLC-grade water. After dilution, the resulting mixture was filtered through a 0.45 um filter to remove any large particles that can clog the HPLC column. For analysis, the samples are diluted appropriately with a 20 mM ammonium acetate buffer in HPLC-grade water at a pH of about 4.5. After mixing the sample, it was transferred to a 300 µL microvial that was then placed in the autosampler. Using ChemStation, the software that operates the HPLC, the analysis parameters such as sample flowrate, column temperature, mobile phase flowrate, mobile phase composition, etc. can be altered. In one exemplary, but non-limiting analysis the parameters were: sample flow rate of 1 mL/min, column temperature of 80° C., column pressure of 60-70 bar, mobile phase composition of 97.9% water/1.9% acetonitrile with 0.2% trifluoroacetic acid; UV wavelength for analysis of 214.4 nm, injection volume of 10 µL, and sample run time of 10 minutes.

[0145] Under these conditions, the truncated jellyfish collagen of SEQ ID NO: 91 has an elution time of about 5.4 minutes. ChemStation quantifies the peak area of the elution peak and calculates the protein concentration using a calibration curve that directly relates peak area to protein concentration. The calibration curve is generated using a known collagen solution that is serially diluted to contain collagen concentration ranges of 0.06 mg/mL to 1.00 mg/mL.

Truncated Collagen without His Tag-Linker-Thrombin Cleavage Site

[0146] A truncated jellyfish collagen without a His tag, linker, and thrombin cleavage site is disclosed below. The codon-optimized nucleotide sequence encoding this collagen is provided in SEQ ID NO: 11. The amino acid sequence is disclosed in SEQ ID NO: 12. The DsbA secretion tag is

encoded by nucleotides 1-72 and encodes amino acids 1-24. The truncated collagen sequence is encoded by nucleotides 73-639 and encodes amino acids 25-213.

(SEQ ID NO: 11)
 ATGAAAAAGATTGGCTGGCGCTGGCTGGTTAGTTTTAGCGTTTAGCGCA
 TCGGCGGCGCAGTATGAAGATGGTCCGCAGGGTGTTGTTGGTGACAGATGGT
 AAAGACGGTACCCCGGGTAATGCAGGTGAGAAAGGTCCGTCAGGTGAACCT
 GGCAGCCCTGGTAAAGCAGGTAGTCCCGGTGAGCAGGGTCCGCCGGGCAA
 GATGGTAGTAATGGTGAGCCGGGTAGCCCTGGCAAAGAAGGTGAACGTGGT
 CTGGCAGGACCCTCGGGTCTGATGGTCCGCCGGTGAACCGGGTTCACCG
 GGTATTGCCGGTGCCCTGGGTAAACCAGGTCTGGAAGGTCCGAAAGGTTAT
 CCTGGTCTGCGCGGTCTGTGATGGTACCAATGGCAAACGTGGCGAACAGGGC
 GAAACCGGTCCAGATGGTTCGTGGTATTCCGGTAAACGATGGTCAGAGC
 GGTAAACCGGCATTGATGGTATTGATGGCACCAATGGTCAGCCTGGCGAA
 GCAGGTTATCAGGGTGGTCCGGTACCCGTGGTCTGGTGGTGAACAGGTG
 ATGTTGGTCAGAATGGTATCGCGCGCACCCGGTCCGGATGGTAGCAAAG
 GTAGCGCCGGTCTCGGGTTACGTTaa

(SEQ ID NO: 12)
 MKKIWLALAGLVLAFSASAAQYEDGPQGVVADGKDGTPGNAGQKGPSGEP
 GSPGKAGSAGEQPPGKDGSNGEPPGPKGEGERGLAGPPGDGRRRGETGSP
 GIAGALGKPGLEGPKGYPLGRDRDGTNGKRGEQGTGPDGVRGIPGNDGQS
 GKPGIDGIDGTNGQPGEAGYQGGRTGRQLGETGDVQNGDRGAPGPDGSK
 GSAGRPGLR

[0147] A polynucleotide encoding a truncated jellyfish collagen without a His tag, linker and thrombin cleavage site disclosed in SEQ ID NO: 90

(SEQ ID NO: 90)
 GGTCCGCAGGGTGTTGTTGGTGACAGATGGTAAAGACGGTACCCCGGGTAA
 GCAGGTGAGAAAGGTCCGTGAGGTGAACCTGGCAGCCCTGGTAAAGCAGGT
 AGTGCCGGTGAGCAGGGTCCGCCGGCAAAGATGGTAGTAATGGTGAGCCG
 GGTAGCCCTGGCAAAGAAGGTGAACGTGGTCTGGCAGGACCCTGGTCTCT
 GATGGTCCCGCGGTGAAACGGGTTACCCGGTATTGCCGGTCCCTGGGT
 AAACAGGTCTGGAAGGTCCGAAAGGTTATCTGCTGTCGCGCGTCTGTAT
 GGTACCAATGGCAAACGTGGCGAACAGGGCGAAACCGGTCCAGATGGTGT
 CGTGGTATTCCGGTAAACGATGGTTCAGAGCGGTAAACCGGCATTGATGGT
 ATTGATGGCACCAATGGTCAGCCTGGCGAAGCAGGTTATCAGGGTGGTCCG
 GGTACCCGTGGTCTGGTGAACAGGTGATGTTGGTCAGAATGGTGTAT
 CGCGCGCACCCGGTCCGGATGGTAGCAAAGGTAGCGCCGGTCTCGGGT
 TTACGTTaa

[0148] A truncated jellyfish collagen without a His tag, linker and thrombin cleavage site disclosed in SEQ ID NO: 91

(SEQ ID NO: 91)
 GPQGVVADGDKDTPGNAGQKGPSGEPSPGKAGSAGEQGPFGKDGNSNGEP
 GSPGKEGERGLAGPPGPDGRRGETGSPGIAGALGKPGLEGPKGYPLRGRD
 GTNGKRGEQGETGPDGVRGIPGNDGQSGKPGIDGIDGTNGQPGEAGYQGGR
 GTRGQLGETGDVGNQNGDRGAPGPDGSKGSAGRPLR

[0149] Truncated Collagen with GEK Repeats

[0150] A jellyfish collagen with GEK repeats is disclosed below. The codon-optimized nucleotide sequence encoding this collagen is provided in SEQ ID NO: 13. The amino acid sequence is disclosed in SEQ ID NO: 14. The DsbA secretion tag is encoded by nucleotides 1-72 and encodes amino acids 1-24. The GEK repeat is encoded by nucleotides 73-126 and encodes the GEK repeats of amino acids 25-42. The truncated collagen sequence is encoded by nucleotides 127-693 and encodes amino acids 43-231.

(SEQ ID NO: 13)
 ATGAAAAAGATTGGCTGGCGCTGGCTGGTTTAGTTTATAGCGTTTAGCGCA
 TCGGCGCGCAGTATGAAGATGGTAAAAAGGTGAaAAGGGCGAGAAAGGT
 GAGAAAGGCGAAAAGGGTGA AAAAGGTCCGCAGGGTGTGTGGTGCAGAT
 GGTAAGACGGTACCCCGGTAATGCAGGTGAGAAAGGTCCGTCAGGTGAA
 CCTGGCAGCCCTGGTAAAGCAGGTAGTCCCGGTGAGCAGGGTCCGCCGGGC
 AAAGATGGTAGTAATGGTGAAGCGGGTAGCCCTGGCAAGAAAGGTGAACGT
 GGTCTGGCAGACCCCGGTCCTGATGGTCCGCCGGTGA AACGGTTCA
 CCGGTATTGCGGGTCCCTGGGTA AACAGGTCTGGAAGGTCCGAAAGGT
 TATCTGGTCTGCGCGGTGATGGTACCAATGGCAACAGTGGCGAACAG
 GCGAAACCGGTCCAGATGGTGTCTGTGGTATCCCGGTAACGATGGTCAG
 AGCGGTAAACCGGCATTGATGGTATTGATGGCACC AATGGTCAGCCTGGC
 GAAGCAGGTTATCAGGGTGGTCCGCGGTACCCGTGGTCCAGTGGGTGAAACA
 GGTGATGTTGGT CAGAAATGGTATCGCGCGCACCCGGTCCGGATGGTAGC
 AAAGGTAGCGCCGGTCTCGGGTTTACGTtaa

(SEQ ID NO: 14)
 MKKIWLALAGLVLAFAASAAQYEDGKGEKGEKGEKGEKGEKGPQGVVAD
 GKDGTGPNAGQKGPSGEPSPGKAGSAGEQGPFGKDGNSNGEPSPGKEGER
 GLAGPPGPDGRRGETGSPGIAGALGKPGLEGPKGYPLRGRDGTNGKRGEQ
 GETGPDGVRGIPGNDGQSGKPGIDGIDGTNGQPGEAGYQGGRGRTRGQLGET
 GDVGNQNGDRGAPGPDGSKGSAGRPLR

[0151] The polynucleotides of SEQ ID NO: 13 was codon optimized and synthesized by Gen9 DNA, now Ginkgo Bioworks internal synthesis. Overlaps between the pET28 vector and SEQ ID NO: 13 was designed to be between 30 and 40 bp long and added using PCR with the enzyme PrimeStar GXL polymerase (http://www.clontech.com/US/Products/PCR/GC_Rich/PrimeSTAR_GXL_DNA_Polymerase?site=10020:22372:US). The opened pET28a vector and insert DNA (SEQ ID NO: 13) was then assembled together into the final plasmid using SGI Gibson assembly (<https://us.vwr.com/store/product/17613857/gibson-assembly-hifi-1-step-kit-synthetic-genomics-inc>). Sequence of

plasmid was then verified through Sanger sequencing through Eurofins Genomics (www.eurofinsgenomics.com).

[0152] The transformed cells were cultivated in minimal media and frozen in 1.5 aliquots with glycerol at a ratio of 50:50 of cells to glycerol. One vial of this frozen culture was revived in 50 ml of minimal media overnight at 37° C., 200 rpm. Cells were transferred into 300 ml of minimal media and grown for 6-9 hours to reach an OD600 of 5-10.

[0153] A bioreactor was prepared with 2.7 L of minimal media+glucose and 300 ml of OD600 of 5-10 culture was added to bring the starting volume to 3 L. Cells were grown at 28° C., pH7 with Dissolved Oxygen maintained at 20% saturation using a cascade containing agitation, air and oxygen. pH was controlled using 28% w/w Ammonium hydroxide solution. Fermentation was run in a fed-batch mode using a DO-stat based feeding algorithm once the initial bolus of 40 g/L was depleted around 13 hours. After 24-26 hours of initial growth, the OD600 reached above 100. At this point, 300 mL of 500 g/L sucrose was added and temperature was reduced to 25 C. High density culture was induced for protein production using 1 mM IPTG. Fermentation was continued for another 20-24 hours and cells were harvested using a bench top centrifuge at 9000 rcf, 15 C for 60 minutes. Cell pellet recovered from centrifugation was resuspended in a buffer containing 0.5M NaCl and 0.1M KH2PO4 at pH8 in a weight by weight ratio of 2× buffer to 1× cells.

[0154] The harvested cells were disrupted in a homogenizer at 14,000 psi pressure in 2 passes. Resulting slurry contained the collagen protein along with other proteins.

[0155] The collagen was purified by acid treatment whole cells recovered from bioreactor after centrifugation and resuspension in a buffer as described above. The pH of either the homogenized slurry or the resuspended suspension was decreased to 3 using 6M Hydrochloric acid. Acidified cell slurry was incubated overnight at 4° C. with mixing, followed by centrifugation. Supernatant of the acidified slurry was tested on a polyacrylamide gel and found to contain collagen in relatively high abundance compared to starting pellet. The collagen slurry thus obtained was high in salts. To obtain volume and salt reduction, concentration and diafiltration steps were performed using an EMD Millipore Tangential Flow Filtration system with ultrafiltration cassettes of 0.1 m² each. Total area of filtration was 0.2 m² using 2 cassettes in parallel. A volume reduction of 5× and a salt reduction of 19× was achieved in the TFF stage. Final collagen slurry was run on an SDS-PAGE gel to confirm presence of the collagen. This slurry was dried using a multi-tray lyophilizer over 3 days to obtain a white, fluffy collagen powder.

[0156] The purified collagen was analyzed on an SDS-PAGE gel and was observed to run at an apparent molecular weight of 35 kilodaltons. The 35 kilodalton band does not correspond to the expected size of 22 kilodaltons. The upshift between the expected size and the apparent size is thought to be due to the GEK repeats interacting with the gel matrix. The 35 kDa band was confirmed by mass spectrometry to be the correct collagen with the GEK repeats.

Truncated Collagen with GDK Repeats

[0157] A jellyfish collagen with GDK repeats is disclosed below. The codon-optimized nucleotide sequence encoding this collagen is provided in SEQ ID NO: 15. The amino acid sequence is disclosed in SEQ ID NO: 16. The DsbA secretion tag is encoded by nucleotides 1-72 and encodes

amino acids 1-24. The GDK repeat is encoded by nucleotides 73-126 and encodes the GDK repeats of amino acids 25-42. The truncated collagen sequence is encoded by nucleotides 127-693 and encodes amino acids 43-231.

(SEQ ID NO: 15)
 ATGAAAAAGATTGGCTGGCGCTGGCTGGTTAGTTTGTAGCGTTTAGCG
 CATCGCGCGCGCAGTATGAAGATGGTGATAAAGGTGATAAGGGCGACAA
 AGGTGACAAAGCGGATAAAGGTGATAAAGGTCCGAGGGTGTGTGGT
 GCAGATGGTAAAGACGGTACCCGGGTAATGCAGGTCAGAAAGGTCCGT
 CAGGTGAACCTGGCAGCCCTGGTAAAGCAGGTAGTGCCGGTGAGCAGGG
 TCCGCGCGGCAAGATGGTAGTAAATGGTGGAGCCGGTAGCCCTGGCAAA
 GAAGGTGAACGTGGTCTGGCAGGACCGCGGGTCTGATGGTCGCGCGG
 GTGAAACGGGTTACCCGGGTATTGCGGGTCCCTGGGTAACCCAGGTCT
 GGAAGGTCCGAAAGGTTATCTGGTCTGCGCGGTCTGATGGTACCAAT
 GGCAAAACGTGGCGAACAGGGCGAAACCGGTCCAGATGGTGTTCGTGGTA
 TTCCGGTAAACGATGGTCAAGCGGTAAACCGGGCATTGATGGTATTGA
 TGGCACCAATGGTCAGCCTGGCGAAGCAGGTTATCAGGGTGGTCGCGGT
 ACCCGTGGTCAGCTGGGTGAAACAGGTGATGTTGGTCAGAATGGTGATC
 GCGGCGCACCGGGTCCGGATGGTAGCAAAGGTAGCGCCGGTCTGTCGGG
 TTTACGTTaa

(SEQ ID NO: 16)
 MKKIWLALAGLVLAFSASAAQYEDGDKGDKGDKGDKGDKGPKQGVVG
 ADGKDGTPGNAGQKGPSPEPGSPGKAGSAGEQPPGKDGNSGEPGSPGK
 EGERLAGPPGPDGRRGETGSPGIAGALGKPLGEPKGYPLRGRDGTN
 GKRGEQGETGPDGVRGIPGNDGQSGKPGIDGIDGTNGQPEAGYQGGRR
 TRGQLGETGDVQNGDRGAPGPDGSKGSAGRPLR

[0158] The polynucleotides of SEQ ID NO: 15 was codon optimized and synthesized by Gen9 DNA, now Ginkgo Bioworks internal synthesis. Overlaps between the pET28 vector and SEQ ID NO: 15 was designed to be between 30 and 40 bp long and added using PCR with the enzyme PrimeStar GXL polymerase (http://www.clontech.com/US/Products/PCR/GC_Rich/PrimeSTAR_GXL_DNA_Polymerase?site=10020:22372:US). The opened pET28a vector and insert DNA (SEQ ID NO: 15) was then assembled together into the final plasmid using SGI Gibson assembly (<https://us.vwr.com/store/product/17613857/gibson-assembly-hifi-1-step-kit-synthetic-genomics-inc>). Sequence of plasmid was then verified through sanger sequencing through Eurofins Genomics (www.eurofinsgenomics.com).

[0159] The transformed cells were cultivated in minimal media and frozen in 1.5 aliquots with glycerol at a ratio of 50:50 of cells to glycerol. One vial of this frozen culture was revived in 50 ml of minimal media overnight at 37° C., 200 rpm. Cells were transferred into 300 ml of minimal media and grown for 6-9 hours to reach an OD600 of 5-10.

[0160] A bioreactor was prepared with 2.7 L of minimal media+glucose and 300 ml of OD600 of 5-10 culture was added to bring the starting volume to 3 L. Cells were grown at 28° C., pH7 with Dissolved Oxygen maintained at 20% saturation using a cascade containing agitation, air and

oxygen. pH was controlled using 28% w/w Ammonium hydroxide solution. Fermentation was run in a fed-batch mode using a DO-stat based feeding algorithm once the initial bolus of 40 g/L was depleted around 13 hours. After 24-26 hours of initial growth, the OD600 reached above 100. At this point, 300 mL of 500 g/L sucrose was added and temperature was reduced to 25 C. High density culture was induced for protein production using 1 mM IPTG. Fermentation was continued for another 20-24 hours and cells were harvested using a bench top centrifuge at 9000 rcf, 15 C for 60 minutes. Cell pellet recovered from centrifugation was resuspended in a buffer containing 0.5M NaCl and 0.1M KH₂PO₄ at pH8 in a weight by weight ratio of 2× buffer to 1× cells.

[0161] The harvested cells were disrupted in a homogenizer at 14,000 psi pressure in 2 passes. Resulting slurry contained the collagen protein along with other proteins.

[0162] The collagen was purified by acid treatment of whole cells recovered from bioreactor after centrifugation and resuspension in a buffer as described above. The pH of either the homogenized slurry was decreased to 3 using 6M Hydrochloric acid. Acidified cell slurry was incubated overnight at 4° C. with mixing, followed by centrifugation. Supernatant of the acidified slurry was tested on a polyacrylamide gel and found to contain collagen in relatively high abundance compared to starting pellet. The collagen slurry thus obtained was high in salts. To obtain volume and salt reduction, concentration and diafiltration steps were performed using an EMD Millipore Tangential Flow Filtration system with ultrafiltration cassettes of 0.1 m² each. Total area of filtration was 0.2 m² using 2 cassettes in parallel. A volume reduction of 5× and a salt reduction of 19× was achieved in the TFF stage. Final collagen slurry was run on an SDS-PAGE gel to confirm presence of the collagen. This slurry was dried using a multi-tray lyophilizer over 3 days to obtain a white, fluffy collagen powder.

[0163] The purified collagen was analyzed on an SDS-PAGE gel and was observed to run at an apparent molecular weight of 35 kilodaltons. The 35 kilodalton band does not correspond to the expected size of 22 kilodaltons. The upshift between the expected size and the apparent size is thought to be due to the GDK repeats interacting with the gel matrix. The 35 kDa band was confirmed by mass spectrometry to be the correct collagen with the GDK repeats.

Truncated Collagen with DsbA Secretion Tag-His Tag-Linker-Thrombin Cleavage Site and GFP Beta-Lactamase Fusion (Version 1):

[0164] A jellyfish collagen with DsbA secretion tag-His tag-Linker-Thrombin cleavage site and GFP Beta-lactamase fusion is disclosed below. The codon-optimized nucleotide sequence encoding this collagen is provided in SEQ ID NO: 17. The amino acid sequence is disclosed in SEQ ID NO: 18. The DsbA secretion tag is encoded by nucleotides 1-72 and encodes amino acids 1-24. The His tag is encoded by nucleotides 73-99 and encodes a 9 histidine tag (SEQ ID NO: 129) of amino acids 25-33. The linker is encoded by nucleotides 100-111 and encodes amino acids 34-37. The thrombin cleavage site is encoded by nucleotides 112-135 and encodes amino acids 38-45. The green fluorescent protein (GFP) with linker is encoded by nucleotides 136-873 and encodes amino acids 46-291. The truncated collagen sequence is encoded by nucleotides 874-1440 and encodes amino acids 292-480. The Beta-lactamase with linker is encoded by nucleotides 1441-2232 and encodes amino acids

481-744. The Beta-lactamase was properly targeted to the periplasmic space even though the polypeptide did not have an independent secretion tag. The DsbA secretion tag directed the entire transcript (Truncated Collagen with DsbA secretion tag-His tag-Linker-Thrombin cleavage site and GFP Beta-lactamase fusion protein) to the periplasmic space and the Beta-lactamase functioned properly.

(SEQ ID NO: 17)
 ATGAAAAAGATTGGCTGGCGCTGGCTGGTTAGTTTTAGCGTTTAGCG
 CATCGGCGGCGCAGTATGAAGATCACCATCACCACCACCACCATCACCA
 CTCTGGCTCGAGCCTGGTGGCCGCGCGGAGCCATATGTCTGGCTCGAGC
 AGTAAAGGTGAAGAAGTGTTCACCGGTGTTGTTCCGATCCTGGTTGAAC
 TGGATGGTGTGTTAACGGCCACAAATTCTCTGTTCTGGTGAAGGTGA
 AGGTGATGCAACCAACGGTAAACTGACCTGAAATTCATCTGCACATACC
 GGTAAACTGCCGGTTCATGGCCGACTCTGGTGACTACCTGACCTATG
 GTGTTTCAGTGTTCCTCGTTACCCGGATCACATGAAGCAGCATGATTT
 CTTCAAATCTGCAATGCCGGAAGGTTATGTACAGAGCGCACCATTTCT
 TTCAAAGACGATGGCACCTACAAAACCCGTCGACAGGTTAAATTTGAAG
 GTGATACTCTGGTGAACCGTATTGAACTGAAAGGCATTGATTTCAAAGA
 GGACGGCAACATCTCGGCCACAACTGGAATATAACTTCAACTCCCAT
 AACGTTTACATCACCGCAGACAAACAGAAAGCGGTATCAAAGCTAACT
 TCAAATTCGCCATAACGTTGAAGACGGTAGCGTACAGCTGGCGGACCA
 CTACCAGCAGAACACTCCGATCGGTGATGGTCCGGTCTGCTGCCGGAT
 AACCACTACCTGTCCACCCAGTCTaaCTGTCCAAAGACCCGAACGAAA
 AGCGCGACCACATGGTGTGCTGGAGTTTCGTTACTGCAGCAGGTATCAC
 GCACGGCATGGATGAACTCTACAAATCTGGCGCGCCGGCGGTCCGCAG
 GGTGTTGTTGGTGCAGATGGTAAAGACGGTACCCCGGTAATGCAGGTG
 AGAAAGGTCCGTCAGGTGAACCTGGCAGCCCTGGTAAAGCAGGTAGTGC
 CGGTGAGCAGGGTCCGCCGGGCAAAGATGGTAGTAATGGTGAAGCCGGT
 AGCCCTGGCAAAGAAGGTGAACGTGGTCTGGCAGGACCCCGGGTCTGTG
 ATGGTCCGCGGGTGAACGGGTTACCCGGGTTATGCGCGTCCCTGGG
 TAAACAGGTCTGGAAGGTCCGAAAGGTTATCCTGGTCTGCGCGGTCTG
 GATGTTACCAATGGCAAACGTGGCGAACAGGGCGAAACCGGTCCAGATG
 GTGTTCTGGTATTCCGGGTAACGATGGTGCAGAGCGGTAACCCGGCAT
 TGATGGTATTGATGGCACAATGGTTCAGCTGGCGAAGCAGGTATCAG
 GGTGGTCCGCGTACCCGTGGTTCAGCTGGGTGAAACAGGTGATGTTGGT
 AGAATGGTGTGATCGCGCGCACCGGTCCGGATGGTAGCAAAGGTAGCGC
 CGGTCTCCGGTTTACGTCacccagaaacgctggtgaaagttaaagat
 gctgaagatcagttgggtgcacagtggtttacatcgaaactggtatctca
 acagcggtaagatccttgagagttttcgccccgaagaacgttttccaat
 gatgagcacttttaaagttctgctatgtggcgcggtattatcccgtatt
 gacgccccggaagagcaactcggctcgccgcatacactattctcagaatg

-continued

acttggttgagtactcaccagtcacagaaaagcatcttacggatggcat
 gacagtaagagaattatgcagtgctgccataaacatgagtgataaacact
 gcgcccaacttacttctgacaacgatcggaggaccgaaggagctaacgg
 ctatagcacaacatggggatcatgtaactcgcccttgatcgttgggaac
 cggagctgaatgaagccataccaaacgacgagcgtgacaccacgatgcc
 tgtagcaatggcaacaacgctgcgcaactataactggcgaactactt
 actctagcttcccggcaacaattaatagactggatggaggcggataaag
 ttgcaggaccacttctgcgctcggccctcccggctggtggtttattgc
 tgataaatctggagccggtgagcgtgggtctcgcggtatcattgcagca
 ctggggccagatggttaagccctcccgtatcgtagttatctacacgacgg
 ggagtcaggcaactatggatgaacgaaatagacagatcgctgagatagg
 tgcctcactgattaagcattggtaa

(SEQ ID NO: 18)
 MKKIWLALAGLVLAFSASAAQYEDHHHHHHHHSSSLVPRGSHMSGSS
 SKGEELFTGVVPILEVELDGVNKHKFSVRGEGEGDATNGKLTCLKFICTT
 GKLPVWPVTLVTLTYGVQCFSRYPDHMKQHDFFKMSAMPEGYVQERTIS
 FKDDGTYKTRAEVKFEGDTLVNRIELKIDFKEDGNILGHKLEYNFNHSH
 NVYITADKQKNGIKANFKIRHNVEDGSVQLADHYQONTIGDGPVLLPD
 NHYLSLTSKLSKDPNEKRDHMLLEFVTAAGITHGMDELKYSKGGAPGGPQ
 GVVGADGKDGTPGNAGQKQKPSGEPGSGPKAGSAGEQGGPPGKDGNSGEPG
 SPGKEGERGLAGPPGPDGRRGETGSPGIAGALGKPGLEPKGYPGLRGR
 DGTNGKRGEQGETGPDGVRGIPGNDGQSGKPGIDGIDGTNGQPGGAGYQ
 GGRGTRGQLGETGVDVQNGDRGAPGPDGSKSAGRPLRHPETLVKVKID
 AEDQLGARVGYIELDLNSGKILESFRPEERPFMMSTFKVLLCGAVLSRI
 DAGQEQLGRRIHYSQNDLVEYSPVTEKHLTDGMTVRELCSAAITMSDNT
 AANLLLTIGGPKELTAFLNHMGDHRVRLDRWEPLENAIPNDRDRTTM
 PVAMATTLRKLTLGELLTLASRQQLIDWMEADKVGAPLLRSALPAGWFI
 ADKSGAGERGSRGIIAALGPDGKPSRIIVVIYTTGSQATMDER
 NRQIAEIGASLIKHW

[0165] The polynucleotides of SEQ ID NO: 17 was constructed by assembling several DNA fragments. The collagen containing sequence was codon optimized and synthesized by Gen9 DNA, now Ginkgo Bioworks internal synthesis. The GFP was also synthesized by Gen9. The Beta-lactamase was cloned out of the plasmid pKD46 (<http://cgsc2.biology.yale.edu/Strain.php?ID=68099>) using PCR with the enzyme PrimeStar GXL polymerase (http://www.clontech.com/US/Products/PCR/GC_Rich/PrimeSTAR_GXL_DNA_Polymerase?site=10020:22372:US). Overlaps between the pET28 vector, GFP, Collagen, and Beta-lactamase was designed to be between 30 and 40 bp long and added using PCR with the enzyme PrimeStar GXL polymerase. The opened pET28a vector and inserts were then assembled together into the final plasmid using SGI Gibson assembly (<https://us.vwr.com/store/product/17613857/gibson-assembly-hifi-1-step-kit-synthetic-ge>

nomics-inc). Sequence of plasmid was then verified through sanger sequencing through Eurofins Genomics (www.eurofinsgenomics.com).

[0166] The transformed cells were cultivated in minimal media and frozen in 1.5 aliquots with glycerol at a ratio of 50:50 of cells to glycerol. One vial of this frozen culture was revived in 50 ml of minimal media overnight at 37° C., 200 rpm. Cells were transferred into 300 ml of minimal media and grown for 6-9 hours to reach an OD600 of 5-10.

[0167] A bioreactor was prepared with 2.7 L of minimal media+glucose and 300 ml of OD600 of 5-10 culture was added to bring the starting volume to 3 L. Cells were grown at 28° C., pH7 with Dissolved Oxygen maintained at 20% saturation using a cascade containing agitation, air and oxygen. pH was controlled using 28% w/w Ammonium hydroxide solution. Fermentation was run in a fed-batch mode using a DO-stat based feeding algorithm once the initial bolus of 40 g/L was depleted around 13 hours. After 24-26 hours of initial growth, the OD600 reached above 100. At this point, 300 mL of 500 g/L sucrose was added and temperature was reduced to 25 C. High density culture was induced for protein production using 1 mM IPTG. Fermentation was continued for another 20-24 hours and cells were harvested using a bench top centrifuge at 9000 rcf, 15 C for 60 minutes. Cell pellet recovered from centrifugation was resuspended in a buffer containing 0.5M NaCl and 0.1M KH₂PO₄ at pH8 in a weight by weight ratio of 2× buffer to 1× cells.

[0168] The harvested cells were disrupted in a homogenizer at 14,000 psi pressure in 2 passes. Resulting slurry contained the collagen protein along with other proteins.

[0169] The collagen was purified by acid treatment of non-homogenized whole cells recovered from the bioreactor after centrifugation and resuspension in the buffer described above. The pH of the resuspended suspension was decreased to 3 using 6M Hydrochloric acid. Acidified cell slurry was incubated overnight at 4° C. with mixing, followed by centrifugation. The pH was then raised to 9 using 10N NaOH and the supernatant of the slurry was tested on a polyacrylamide gel and found to contain collagen in relatively high abundance compared to starting pellet. The collagen slurry thus obtained was high in salts. To obtain volume and salt reduction, concentration and diafiltration steps were performed using an EMD Millipore Tangential Flow Filtration system with ultrafiltration cassettes of 0.1 m² each. Total area of filtration was 0.2 m² using 2 cassettes in parallel. A volume reduction of 5× and a salt reduction of 19× was achieved in the TFF stage. Final collagen slurry was run on an SDS-PAGE gel to confirm presence of the collagen. This slurry was dried using a multi-tray lyophilizer over 3 days to obtain a white, fluffy collagen powder.

[0170] The purified collagen-GFP-Beta-lactamase fusion protein was analyzed on an SDS-PAGE gel and was observed to run at an apparent molecular weight of 90 kilodaltons. The expected size of the fusion protein is 85 kd. The 90 kDa band was confirmed by mass spectrometry to be the correct collagen fusion protein.

Truncated Collagen with DsbA Secretion Tag-His Tag-Linker-Thrombin Cleavage Site and GFP Beta-Lactamase Fusion (Version 2):

[0171] A jellyfish collagen with DsbA secretion tag-His tag-Linker-Thrombin cleavage site and GFP Beta-lactamase fusion is disclosed below. The codon-optimized nucleotide sequence encoding this collagen is provided in SEQ ID NO:

19. The amino acid sequence is disclosed in SEQ ID NO: 20. The DsbA secretion tag is encoded by nucleotides 1-72 and encodes amino acids 1-24. The His tag is encoded by nucleotides 73-99 and encodes a 9 histidine tag (SEQ ID NO: 129) of amino acids 25-33. The linker is encoded by nucleotides 100-111 and encodes amino acids 34-37. The thrombin cleavage site is encoded by nucleotides 112-135 and encodes amino acids 38-45. The green fluorescent protein (GFP) with linker is encoded by nucleotides 136-873 and encodes amino acids 46-291. The truncated collagen sequence is encoded by nucleotides 874-1440 and encodes amino acids 292-480. The Beta-lactamase with linker is encoded by nucleotides 1441-2232 and encodes amino acids 481-744.

(SEQ ID NO: 19)

```
ATGAAAAAGATTGGCTGGCGCTGGCTGGTTAGTTTACGCTTTAGCG
CATCGGCGGCGCAGTATGAAGATCACCATCACCACCACCACCATCACCA
CTCTGGCTCGAGCCTGGTGCCCGCGGAGCCATATGTCTGGCTCGAGC
AGTAAAGTGGAAGAACTGTTACCCGGTGTGTCCGATCCTGGTTGAAC
TGGATGGTGATGTTAACGGCCACAAATCTCTGTTCGTGGTGAAGGTGA
AGGTGATGCAACCACCGTAAACTGACCCCTGAAATTCATCTGCACTACC
GGTAAACTGCCGGTTCATGGCCGACTCTGGTGACTACCCCTGACCTATG
GTGTTTCAGTGTTTTTCTCGTTACCCGGATCACATGAAGCAGCATGATTT
CTTCAAATCTGCAATGCCGGAAGTTATGTACAGGAGCGCACCATTTCT
TTCAAAGACGATGGCACCTACAAAACCCGTGCAGAGGTTAAATTTGAAG
GTGATACTCTGGTGAAACCGTATTGAACTGAAAGGCATTGATTTCAAAGA
GGACGGCAACATCTCGGCCACAACTGGAATATAACTTCAACTCCCAT
AACGTTTACATCACCAGACAAACAGAAAGCGGTATCAAAGCTAACT
TCAAATTCGCCATAACGTGAAGACGGTAGCGTACAGCTGGCGGACCA
CTACAGCAGAACACTCCGATCGGTGATGGTCCGGTCTGTCTGCCGGAT
AACCCTACCTGTCCACCAGTCTaaCTGTCAAAGACCCGAAACGAA
AGCGCGACCACATGGTGTGTGGAGTTCGTACTGCAGCAGGTATCAC
GCACGGCATGGATGAACCTACAAATCTGGCGCGCCGGCGGTCCGCAG
GGTGTGTGTGGTGCAGATGGTAAAGACGGTACCCCGGTAATGCAGGTC
AGAAAGGTCGGTCAGGTGAACCTGGCAGCCCTGGTAAAGCAGGTAGTGC
CGGTGAGCAGGTCGCCCGGCAAGATGGTAGTAATGGTGGAGCCGGGT
AGCCCTGGCAAAGAGGTGAACGTGGTCTGGCAGGACCCCGGTCCTG
ATGGTCGCGCGGTGAAACGGTTTACCAGGTTATGCGCGTCCCTGGG
TAAACCGGTCGGAAGGTCGAAAGGTTATCCTGGTCTGCGCGGTCGT
GATGGTACCAATGGCAAACGTGGCGAACAGGGCGAAACCGGTCCAGATG
GTGTTCTGGTATTCCGGTAACGATGGTCAGAGCGGTAAACCGGGCAT
TGATGGTATTGATGGCACCATGGTCAGCCTGGCGAAGCAGGTTATCAG
GGTGGTCGCGGTACCCGTGGTGGTGGTAAACAGGTGATGTTGGTC
AGAATGGTGATCGCGCGCACCGGTCGGATGGTAGCAAAGGTAGCGC
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CGGTCGTCGGGTTTACGTcaccagaaacgctggtgaaagtaaaagat
 gctgaagatcagttgggtgcacgagtggttacatcgaactggtatctca
 acagcggtaagatccttgagagttttcgccccgaagaacgttttccaat
 gatgagcacttttaagttctgctatgtggcgcggtattatcccgtatt
 gacgcccggcaagagcaactcggtcgcccgcatacactattctcagaatg
 acttggttgagtactcaccagtcacagaaaagcatcttaccgatggcat
 gacagtaagagaattatgcagtgctgccataaccatgagtataaact
 gggccaacttacttctgacaacgatcggaggaccgaaggagctaaccg
 ctttttgcacaacatgggggatcatgtaactcgccttgatcgttggga
 accggagctgaatgaagccataccaaacgacgagcgtgacaccacgatg
 cctgtagcaatggcaacaacgttgcgcaaaactattaactggcgaactac
 ttactctagcttcccggcaacaattaatagactggatggaggcggataaa
 agttgcaggaccacttctgcgctcgcccttcggctggctggtttatt
 gctgataaatctggagccggtgagcgtgggtctcgcggtatcattgcag
 cactggggccagatggttaagccctcccgtatcgtagttatctacacgac
 ggggagtcaggcaactatggatgaacgaaatagacagatcgctgagata
 ggtgcctcactgattaagcattggttaa

(SEQ ID NO: 20)

MKKIWLALAGLVLAFSASAAQYEDHHHHHHHSSSLVPRGSHMSGSS
 SKGEEELFTGVVPIVELDGDVNGHKFSVRGEGEGDATNGKLTLFICTT
 GKLVPWPPTLVTTLYGVQCFSRYPDHMKQHDFFKSAMPEGYVQERTIS
 FKDDGTYKTRAEVKFEGDTLVNRIELKIDFKEDGNILGHKLEYNFNSH
 NVYITADKQKNGIKANFKIRHNVEDGSVQLADHYQQNTPIGDGPVLLPD
 NHYLSTQS KLSKDPNEKRDMVLEFVTAAGITHGMDELYKSGAPGGP
 QGVVGADKGDGTPGNAGQKGPSGEPGSPGKAGSAGEQGPPKDGSDNGEP
 GSPGKEGERGLAGPPGPDGRRGETGSPGIAGALGKPGLEGPKGYPLGRG
 RDGTNGKRGEQGETGPDGVRGIPGNDGQSGKPGIDGIDGTNGQPGEAGY
 QGGRGTRGQLGETGDVGVQNGDRGAPGPDGSKGSAGRPLRHPETLVKVK
 DAEDQLGARVGYIELDLNSGKILESFRPEERFPMSTFKVLLCGAVLSR
 IDAQEQELGRRIHYSQNDLVEYSPVTEKHLTDGMTVRELCSAAITMSDN
 TAANLLLTITIGPKELTAFLHNMGDHVTRLDRWEPELNEAIPNDERDTT
 MPVAMATTLRLLTGELLTLASRQQLIDWMEADKVGAPLLRSALPAGWF
 IADKSGAGERGSRGIIAALGPDGKPSRIVVIYTTGSQATMDERNRQIAE
 IGASLIKHW

Example 4: Production of Full-Length Elastin

[0172] Full length human elastin were expressed as described below. The wild-type, full length amino acid sequence of human elastin is provided below.

(SEQ ID NO: 21)

MAGLTAAAPRPGVLLLLLSILHPSRPGGVPGAIPGGVPGGVFYPGAGLG
 ALGGGALGPGGKPLKVPVPGLAGAGLGAGLGAFFAVTFPGALVPGGVAD
 AAAAYKAAKAGAGLGGVPGVGGGLGVSAGAVVQPAGVAGKPKVPGVGLP
 GVYPGGVLPGARFPVGVLPVPTGAGVKPKAPGVGGAFAGIPGVGPF
 GPQPGVPLGYPIKAPKLPGGYGLPYTTGKLPYGYGPGGVAGAAGKAGYP
 TGTGVGPQAAAAAAKAAAKFGAGAAGVLPVGGAGVPGVPGAIPIGIGG
 IAGVGTAAAAAAKAAKYGAAAGLVPGGPGFPGVGVGAGVPG
 VGVPGAGIPVVPAGIPGAAVPGVVSPEAAKAAKAAKYGARPGVGVG
 GIPTYGVGAGGFPFGVGGIPGVAGVPGVGGVPGVGGVPGVISP
 QAAAAKAAKYGAAGAGVLGGLVPGPQAAVPGVPTGGVPGVGTAAAA
 AKAAKAAQFGLVPGVGVAPGVGVPAGVGVAPGVLAPGVGVPAGVVA
 PGVGVAPGIPGGVAAAASAAKVAQAQLRAAAGLGAGIPGLGVGVG
 PGLGVGAGVPLGVGAGVPGFAGADEGVRRLSPELREGDPSSSQHLP
 STPSSPRVPGALAAKAAKYGAAVPGVLGGLGALGGVGIPIGGVVGAGPA
 AAAAAKAAKAAQFGLVGAAGLGGLVGGVPGVGGVGGVGGVGGVGGV
 AAKYGAAGLGGVGGAGQFPLGGVAARPGFGLSPIFPGGACLKKAACGRK
 RK
<http://www.uniprot.org/uniprot/P15502>

[0173] The non-codon optimized polynucleotide sequence encoding the full length elastin is disclosed below. In SEQ ID NO: 22, nucleotides 1-78 encode the DsbA secretion tag and nucleotides 79-2358 encode the full length human elastin.

(SEQ ID NO: 22)

ATGGCGGCTCTGACGGCGCGCCCGCCGCGCCGAGTCCCTCTGCTCC
 TGCTGTCCATCCTCCACCCCTCTCGGCCCTGGAGGGGTCCCTGGGCCAT
 TCCTGGTGGAGTTCTGGAGGAGTCTTTTATCCAGGGGCTGGTCTCGGA
 GCCCTTGAGGAGGAGGAGCGCTGGGGCTGGAGGCAACCTCTTAAGCCAG
 TTCCCGGAGGGCTTGGGGTCTGGCCCTTGGGGCAGGGCTCGGCCCTT
 CCCCAGTTACCTTTCCGGGGCTCTGGTGCCTGGTGGAGTGGCTGAC
 GCTGTGTCAGCCTATAAAGCTGCTAAGGCTGGCGTGGGCTTGGTGGTG
 TCCAGGAGTTGGTGGCTTAGGAGTGTCTGCAGGTGCGGTGGTTCCTCA
 GCCTGGAGCCGAGTGAAGCCTGGGAAAGTCCCGGTGTGGGGCTGCCA
 GGTGTATAACCCAGGTGGCGTGTCTCCAGGAGCTCGGTTCCCGGTGTGG
 GGTGCTCCCTGGAGTCCCACTGGAGCAGGAGTTAAGCCCAAGGCTCC
 AGGTGTAGGTGGAGCTTTTGTGGAAATCCAGGAGTTGGACCTTTGGG
 GGACCGCAACCTGGAGTCCCACTGGGGTATCCCATCAAGGCCCCCAAGC
 TGCCCTGGTGGCTATGGACTGCCCTACACCACAGGAAACTGCCCTATGG
 CTATGGGCCCGGAGGAGTGGTGGTGCAGCGGGCAAGGCTGGTTACCCA
 ACAGGGACAGGGGTTGGCCCCAGGCAGCAGCAGCAGCGGCAGCTAAAG
 CAGCAGCAAAGTTCCGGTGTGGAGCAGCCGAGTCCCTCCCTGGTGTGG

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AGGGGCTGGTGTTCCTGGCGTGCCTGGGGCAATTCTGGAATTGGAGGC
 ATCGCAGGCGTTGGGACTCCAGCTGCAGCTGCAGCTGCAGCAGCAGCCG
 CTAAGGCAGCCAAAGTATGGAGCTGCTGCAGGCTTAGTGCCTGGTGGGCC
 AGGCTTTGGCCCGGAGTAGTTGGTGTCCCAGGAGCTGGCGTTCCAGGT
 GTTGGTGTCCCAGGAGCTGGGATTCAGTTGTCCAGGTGCTGGGATCC
 CAGGTGCTGCGGTTCCAGGGTTGTGTCAACAGAAGCAGCTGCTAAGGC
 AGCTGCAAAGGCAGCCAAATACGGGGCCAGGCCCGGAGTCGGAGTTGGA
 GGCATTCTACTTACGGGTTGGAGCTGGGGCTTCCCGGCTTTGGTG
 TCGGAGTCGGAGGTATCCCTGGAGTCGAGGTGTCCCTGGTGTGGAGG
 TGTTCCCGGAGTCGGAGGTGTCCCGGAGTTGGCATTTCCTCCGAAGCT
 CAGGCAGCAGCTGCCGCCAAGGCTGCCAAGTACGGTGTGCAGGAGCAG
 GAGTGTGGGTGGGCTAGTGCAGGTCCCCAGGCGGCAGTCCCAGGTGT
 GCCGGGCACGGGAGGAGTGCCAGGAGTGGGGACCCAGCAGCTGCAGCT
 GCTAAAGCAGCCGCCAAAGCCGCCAGTTTGGGTTAGTTCCTGGTGTGCG
 GCGTGGCTCCTGGAGTTGGCGTGGCTCCTGGTGTGGTGTGGCTCCTGG
 AGTTGGCTTGCTCCTGGAGTTGGCGTGGCTCCTGGAGTTGGTGTGGCT
 CCTGGCGTTGGCGTGGCTCCCGGCATTGGCCCTGGTGGAGTTGCAGCTG
 CAGCAAAATCCGCTGCCAAGGTGGCTGCCAAAGCCAGCTCCGAGCTGC
 AGCTGGGCTTGGTGTGGCATCCCTGGACTTGGAGTTGGTGTGGCGTGC
 CCTGGACTTGGAGTTGGTGTGGTGTCTCTGGACTTGGAGTTGGTGTGCTG
 GTGTTCTGGCTTCGGGGCAGGTGCAGATGAGGGAGTTAGCGGAGCCT
 GTCCCTGAGCTCAGGGAAGGAGATCCCTCCTCCTCAGCACCTCCCC
 AGCACCCCTCATCACCAGGTTACTGGAGCCCTGGTGCCTGCTAAAG
 CAGCCAAATATGGAGCAGCAGTGCCTGGGGTCTTGGAGGGCTCGGGGC
 TCTCGGTGGAGTAGGCATCCAGGCGGTGGTGGGAGCCGGACCCGCC
 GCCGCGCTGCCGACGCAAGCTGCTGCCAAGCCGCCAGTTTGGCC
 TAGTGGGAGCCGCTGGGCTCGGAGGACTCGGAGTCGGAGGGCTTGGAGT
 TCCAGGTGTTGGGGCCTTGGAGGTATACCTCAGCTGCAGCCGCTAAA
 GCAGCTAAATACGGTGTGCTGGCCCTTGGAGGTGTCTAGGGGTGCGG
 GGCAGTCCCCTTGGAGGAGTGGCAGCAAGACCTGGCTTCGGATTGTC
 TCCCATTTCCAGGTGGGGCTGCCTGGGAAAGCTTGTGGCCGGAAG
 AGAAAATGA

Codon Optimized Elastin with DsbA Secretion Tag-His Tag-Linker-Thrombin Cleavage Site

[0174] The codon optimized polynucleotide sequence encoding the full length human elastin with DsbA secretion tag-His tag-Linker-Thrombin cleavage site is disclosed below. In SEQ ID NO: 23: nucleotides 1-72 encode the DsbA secretion tag encoding amino acids 1-24 of SEQ ID NO: 24; nucleotides 73-99 encode the 9 His tag (SEQ ID NO: 129) encoding amino acids 25-33 of SEQ ID NO: 24; nucleotides 100-111 encode the linker encoding amino acids 34-37 of SEQ ID NO: 24; nucleotides 112-135 encode the thrombin cleavage tag encoding amino acids 38-45 of SEQ

ID NO: 24; nucleotides 136-2415 encode the amino acids 46-805 of the full length human elastin of SEQ ID NO: 24.

(SEQ ID NO: 23)

ATGAAAAAGATTGGCTGGCGCTGGCTGGTTAGTTTTAGCGTTTAGCG
 CATCGCGGGCGCAGTATGAAGATCACCATCACCACCACCACCATCACCA
 CTCTGGCTCGAGCCTGGTGC CGCGCGCAGCCATATGGGTGGCGTACCA
 GGCGCAATTCTGGGGGTGTCCCAGGCGGTTTTTTTATCCGGGCGCCG
 GTCTGGCGCACTGGGTGGCGGTGCATGGGGCCGGGCGCAAAACCGT
 GAAACCGGTACCAGGTGGTTAGCAGGCGCGGCTTAGGCGCAGGTCTG
 GGAGCATTTCGGCAGTTACCTTCCAGGGGCACGTGGTTCTGGAGGTG
 TGGCCGATGCAGCCCGGCATATAAAGCCGCTAAAGCCGGTGGGGTTT
 AGGAGGCGTCCCAGGTGTGGTGGCTGGGTGTAGCGCCGGTGCAGTT
 GTTCCGACGCGGGAGCAGGGGTAAACCTGGTAAAGTCCCGGGAGTAG
 GTCTGCCAGGCGTTTATCCTGGTGGTGTTTTGGCGGGTCCCGTTTCC
 GGGCGTTGGTGTCTTCCAGGCGTGCAGCCGGAGCCGGTGTAAACCG
 AAAGCCCCCGGTGTTGGAGGTGCATTGCAGGCATCCCGGGAGTTGGCC
 CGTTTGGTGGTCCGCAACCTGGGGTCCGTTAGGTTATCCGATTAAGC
 ACCGAAACTGCCCGCGGTTATGGTCTGCCGTACACAACCGGTAACCTG
 CCGTATGGTTATGGCCCGGGTGGAGTTGGCGGTGCAGCAGGTAAGCGG
 GTTATCCTACCGGAACCGGTAGGTCCGAGGCCGCTGTGCCCGCCG
 CGCAAAAGCAGCGGCTAAATTTGGCGCCGGAGCAGCGGGTGTCTGCCT
 GGAGTTGGTGGTGGCGGCGTCCAGGGGTACCTGTTGCAATTCCCGGGTA
 TTGGTGGTATTGCCGGTGTGGCACCCCGCCGCGGCAGCTGCCGCGC
 GCGCGCTGCCAAAGCTGCTAAATACGGTGC CGCGCGCGGCTGTTGCCA
 GGAGTCCGGGTTTTGGTCCGGAGTGGTGGCGTGCCTGGCGCAGGGC
 TTCTGGTGTGGGCGTCCAGGTGCAGGATTCTGTTGTGCTGGTGC
 CGGTATTCGGCGCGGCGTTCGGGGGTGGTTAGCCCGGAAGCCGCA
 GCGAAGGCTGCGGCAAAAGGCAGCAAGATGGCGCACGCCAGGAGTCG
 GCGTGGGTGGTATCCCGACTATGGGGTGGGCGCAGGGGGTTTTCTGG
 TTTCCGGCTAGGTGTAGGAGGTATACCGGGCTGGCCGGTGTACCAGGG
 GTTGGTGGCGTCCCTGGTGTGGCGGTGTGCCAGGTGTTGGTATTTTAC
 CGGAAGCACAGGCGCAGCCGAGCTAAGGCAGCAAAATATGGTGGCCG
 CGGCGCAGGAGTTTTAGTGGGCTGGTTCCGGGCCCGCAGGCGAGTGTG
 CCGGGGGTCCAGGCACCGGTGGTGTCCCTGGAGTCGGTACGCCGGCTG
 CAGCGGCAGCCAAAGCGGTGCGAAGCAGCACAGTTTGGCTTAGTACC
 GGGTGTGGGAGTTGCCCGCGGTTGGCGTGTCCAGGGGTGGGTGTT
 GCTCCTGGCGTCGGTCTGGCTCCTGGAGTGGGCGTAGCACCCGGTGTGG
 GGGTGGCCCGGGTGTGGGGTGTGCCCGGGTATCGTTCGGGCGGGTGT
 CGCAGCAGCAGCTAAAGCCGGCGAAAGTTGGCGCAAGCCCAACTG
 CGCGCCGCGCGGGCTCGGTGCAGGTATCCGGGGCTGGGTGTGGAG

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TTGGAGTCCCAGGTTTGGGCGTGGGCGCGGGAGTTCGGGACTGGGAGT
GGGTGCCGGAGTTCCTGGCTTTGGTGCAGGCGCAGATGAAGTGTTCGT
CGTAGCCTGAGTCCGGAAGTGCCTGAAGGTGATCCGAGTAGCAGCCAGC
ATCTGCCGAGCACCCGAGCAGCCCGCTGTTCGGGTGCATTAGCTGC
AGCAAAGCCGCCAAGTATGGTGCAGCCGTGCCGGGCGTCTTAGGTGGT
CTGGGCGCCCTGGGTGGTGTAGGCATTCGGGAGGTGTGTGGGTGCAG
GACCGCCCGCCGAGCTGCGGCGCCAAAGCAGCTGCAAAGCCGGCCCA
GTTTGGTTTAGTGGGCGCCGAGGTTTAGGCGGTTTAGGTGTGGGTGGA
CTGGTGTACCTGGCGTAGGCGGTCTGGGTGGAATTCCGCCCGCAGCGG
CCGCGAAAGCGGCAAAATATGGCGCGCAGGCTGGGCGGCGTGTGGG
TGGGGCAGGTGAGTTCCGCTGGGCGGGTTGCCGCACGTCCGGATTT
GGTCTGAGCCGATTTCCCTGGCGGCGCATGTCTGGGTAAGCATGTG
GTCGTAAACGTAATaa

(SEQ ID NO: 24)

MKKIWLALAGLVLAFSASAAQYEDHHHHHHHHSGLVPRGSHMGGVP
GAIPIGGVPGVFPYAGLGLGGGALPGGKPLKVPVGGLAGLAGL
GAPPAVTFPGALVPGVADAAAAYKAAKAGAGLGGVPGVGLVGSAGAV
VPQPGAGVKPKVPVGLPGVYPGVLPGARFPVGVLPVPTGAGVKP
KAPVGGAFAGIPGVFPGGPQPGVPLGYPIKAPKLPGGYGLPYTTGKL
PYGYPGGVVAGAGKAGYPTGTGVPQAAAAAAYKAAKFGAGAAGVLP
GVGGAGVPGVPAIPIGGIAGVGTAAAAAAYKAAKYGAAAGLVP
GGPFGPGVGVPGAGVPGVPGAGIPVVPAGIPGAAPGVVSPVPEAA
AKAAAKAAKYGARPGVGVGGIPTYGVGAGGFPFGVGVGGIPGVAGVPG
VGGVPGVGGVPGVISPAAQAAAAAAYKAAKYGAAAGVGLGLVPGQAAV
PGVPTGGVPGVGTAAAAAAYKAAKAAQFGLVPGVGVAPGVVAPGVV
APGVGLAPGVGAPGVVAPGVVAPGIPGGVAAAAYKAAKAAKAAQL
RAAAGLGAAGIPGLGVGVVPLGVGAGVPLGVGAGVPGFAGADEGVR
RSLSPVPELREGDPSSQHLPTSPSSPRVPGALAAAAYKAAKYGAAVPGVGG
LGALGGVGIPIGGVAGVAAAAAAYKAAKAAQFGLVGAAGLGLGVGG
LGVPVGGVGGIPAAAAAAYKAAKYGAAAGLGGVPLGGVAAAPGPF
GLSPIFPGGACLGKACGRKRK

[0175] The polynucleotide encoding the full length human elastin without the native sequence tag is disclosed in SEQ ID NO: 87.

(SEQ ID NO: 87)

GGTGGCTACCAGGCGCAATTCCTGGGGTGTCCAGGCGGTGTTTTTT
ATCCGGGCGCCGCTTGGCGCACTGGGTGGCGGTGCACTGGGCCCGGG
CGGCAACCGCTGAAACCGGTACCAGGTGGTTTAGCAGGCGCCGGCTTA
GGCGCAGGTCTGGGAGCATTTCCGGCAGTTACCTTTCCAGGGCACTGG
TTCTTGAGGTGTGGCCGATGCAGCCGCGCATATAAAGCCGCTAAAGC

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CGGTGCCGGTTTAGGAGGCGTCCCAGGTGTCGGTGGCCTGGGTGTTAGC
GCCGGTGCAGTTGTTCCGCAGCCGGGAGCAGGGTTAAACCTGGTAAAG
TGCCGGGAGTAGGCTGCCAGGCGTTTATCTGTTGGTGTGTTTTCGGG
TGCCCGTTTTCCGGGCGTGGTGTCTTCCAGGCGTGGCAGCCGGAGCC
GGTGTAAACCGAAAGCCCGGTGTTGGAGGTGCATTTGCAGGCATCC
CGGGAGTTGGCCGTTTGGTGGTCCGCAACCTGGGTTCCGTTAGGTTA
TCCGATTAAGCACCAGAACTGCCCGGCGTTATGGTGTGCCGTACACA
ACCGGTAAACTGCCGTATGGTTATGGCCCGGGTGGAGTTGCCGGTGCAG
CAGGTAAAGCGGTTATCCTACCGAAACCGTGTAGGTCCGAGGCCGC
TGCTGCCCGCCCGCAAAGCAGCGGCTAAATTTGGCGCGGAGCAGCG
GGTGTCTGCCCTGGAGTTGGTGGTCCGGGCGTGCAGGGGTACCTGGTG
CAATTCGGGTATTGGTGGTATTGCCGTTGTCGACCCCGCCCGCGG
AGCTGCCGCGAGCGGCTGCCAAAGCTGCTAAATACGGTCCCGCGGCG
GGTCTGGTGCAGGAGTCCGGTTTTGGTCCGGGAGTGGTGGCGTGC
CTGGCGCAGGCGTCTCTGGTGTGGGCGTCCAGGTGCAGGATTCCTGT
TGTGCTGTTGCCGTTATCCCGCGCGGCGTCCGGGGTGGTTAGC
CCGGAAGCCGAGCGAAGGCTGCCGCAAGGCAGCAAAGTATGGCGCAC
GCCAGGAGTCCGGTGGTGGTATCCCGACCTATGGGTGGGCGCAGG
GGGTTTTCTGGTTCCGCGTAGGTGTAGGAGGTATACCGGCGCTGGCC
GGGTACCAGGGTGGTGGCGTCCCTGGTGTGGCGGTGTGCCAGGTG
TTGGTATTTACCCGGAAGCACAGGCGAGCCGCAAGTAAAGCAGCGAA
ATATGGTCCCGCGCGCAGGAGTTTTAGGTGGGCTGGTCCCGGCCCG
CAGGCAGCTGTCCGGGGTCCAGGCACCGGTGGTGTCCCTGGAGTCCG
GTACGCCGCTGCAGCGGCGCAAGCGGCTGCCAAAGCAGCACAGTT
TGGCTTAGTACCGGTTGGGAGTTGCCCGGCGTGGCGTTGCTCCA
GGGTGGGTGTTGCTCTGGCGTCCGCTGGCTCTCGAGTGGGCGTAG
CACCAGGTGTGGGGTGGCCCGGTTGGGTTGCACCGGTATCCGG
TCCGGGCGGTGTCCGAGCAGCAGTAAAGCGCGCGCAAAGTTGCCGGC
AAAGCCCACTGCCGCGCCCGCGGGCTCCGTGCAGGTATTCGGGGC
TGGGTGTCGAGTTGGAGTCCCGGTTTTGGGCGTGGGCGCGGAGTTCC
GGGACTGGGAGTGGTGGCGGAGTTCTGGCTTTGGTGCAGCGCGAGAT
GAAGGTGTTCTGTCGTAGCTGAGTCCGGAAGTGCATGAGGTGATCCGA
GTAGCAGCCAGCATCTGCCGAGCACCAGCAGCCCGGTGTTCCGGG
TGCATTAGCTGCAGCAAAGCCGCAAGTATGGTGCAGCCGTGCCGGG
GTCTTAGGTGGTCTGGGCGCCCTGGGTGGTGTAGGCATTCGGGAGGTG
TTGTGGGTGCAGGACCGGCGCCGCGAGTGCAGGCGCAAAGCAGCTGC
AAAAGCGGCCAGTTTGGTTTAGTGGGCGCCGAGGTTTAGCGGTTTA
GGTGTGGGTGGACTGGGTGTACCTGGCGTAGGCGGTCTGGGTGGAATTC
CGCCCGCAGCGGCGCAAAGCGGCAAATATGGCGCGGCGAGGCTGGG

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CGGCGTGTGGGTGGGGCAGGTCAGTTTCCGCTGGGCGGGTTGCCGCA
CGTCCGGGATTTGGTCTGAGCCGATTTTCCCTGGCGGCATGTCTGG
GTAAAGCATGTGGTGTAAACGTAAAt aa

[0176] The full length human elastin sequence without the native sequence tag is disclosed in SEQ ID NO: 88.

(SEQ ID NO: 88)

GGVPGAIPGGVPGVFYFPGAGLGGALGPGGKPLKPVPGGLAGAGL
GAGLGAFPAVTFPGALVPGGVADAAAAYKAAKAGAGLGGVPGVGLGVS
AGAVVPQPGAGVKPKGVPVGLPGVYPPGGVLPGARFPVGVLPVPTGA
GVKPKAPGVGGAFAGIPGVGPFPGPQPGVPLGYPIKAPKLPGGYGLPYT
TGKLPYGYPGGVVAGAAGKAGYPTGTGVPQAAAAAAKAAKAFGAGAA
GVLPGVGGAGVPGVPAIPGI GGIAGVGTAAAAAATAAAYKAAKYGAAA
GLVPGGPGFPGVGVVPGVAGVPGVPGAGIPVVPAGVPGVAVPGVVS
PEAAAKAAKAAKYGARPGVGVGGIPTYGVGAGFPFPGVGVGGIPGVA
GVPVGGVPGVGGVPGVGI SPEAQAAAAKAAKYGAAGAGVLGGLVPGP
QAAVPGVPGTGGVPGVGTAAAAKAAKAAQFGLVPGVGVAPGVVAP
GVGVAPVGLAPGVVAPGVVAPGVVAPGIPGGVAAAASAAKVAA
KAQLRAAAGL GAGI PGLGVGVVPLGLVGVAGVPLGLVAGVPGFAGAD
EGVRRSLSPPELREGDPSSSHLPSTPSSPRVPGALAAKAAKYGAAVPG
VLGGLGALGGVGI PGGVVGAGPAAAAAATAAAYKAAKAAQFGLVGAAGLGLL
GVGGLGVPGVGGVGGIPPAKAAKAAKYGAAGLGGVGLGAGQFPLGGVAA
RPGFGLSPIFPGGACLKACGRKRK

Codon Optimized Elastin with DsbA Secretion Tag

[0177] The codon optimized polynucleotide sequence encoding the full length human elastin with a DsbA secretion tag is disclosed in SEQ ID NO: 25. In SEQ ID NO: 25: nucleotides 1-72 encode the DsbA secretion tag encoding amino acids 1-24 of SEQ ID NO: 26; nucleotides 73-2355 encode the amino acids 25-785 of the full length human elastin of SEQ ID NO: 26.

(SEQ ID NO: 25)

ATGAAAAAGATTTGGCTGGCGTGGCTGGTGTAGTTTAGCTTTAGC
GCATCGGCGGCGCAGTATGAAGATATGGGTGGCGTACCAGGCGCAATT
CCTGGGGTGTCCAGGCGGTGTTTTTATCCGGGCGCGGTCTTTGGC
GCACTGGGTGGCGGTGCACTGGGCCCGGGCGCAAAACCGCTGAAACCG
GTACCAGGTGGTTTTAGCAGGCGCCGGCTTAGGCGCAGGTCTGGGAGCA
TTTTCCGGCAGTTACCTTTCCAGGGGCACTGGTCTCGGAGGTGTGGCC
GATGCAGCCCGGCATATAAAGCGCTAAAGCCGGTGCGGTTTTAGGA
GGCGTCCAGGTGTCCGTGGCTGGGTGTTAGCGCCGGTGCAGTTGTT
CCGCAGCCGGGAGCAGGGTTAAACCTGGTAAAGTGCCGGGAGTAGGT
CTGCCAGGCGTTTATCCTGGTGGTGTGTTGCCGGGTGCCGTTTTCCG

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GGCGTTGGTGTCTTCCAGGCGTGCCGACCGGAGCCGGTGTAAACCG
AAAGCCCCCGGTGTTGGAGGTGCATTTGCAGGCATCCCGGGAGTTGGC
CCGTTTGGTGGTCCGCAACCTGGGTTCCGTTAGGTATCCGATTAAA
GCACCGAAACTGCCCGGCGTTATGGTCTGCCGTACACAACCGGTAAA
CTGCCGTATGGTTATGGCCCGGGTGGAGTTGCCGGTGCAGCAGGTAAA
GCGGGTTATCCTACCGGAACCGGTGAGGTCCGCAGGCCGCTGCTGCC
GCCCGCGAAAAGCAGCGGCTAAATTTGGCGCCGAGCAGCGGGTGT
CTGCCCTGGAGTTGGTGGTGGGGCGTCCAGGGGTACCTGGTGCAT
CCGGGTATTGGTGGTATTGCCCGGTGTCGCACCCCGGCGCGCAGCT
GCGGCAGCGCGGCTGCCAAAGCTGCTAAATACGGTCCCGCGCGGGT
CTGGTGCCAGGAGTCCGGTTTTGGTCCGGGAGTGGTGGCGTCCCT
GGCGCAGGCGTTCCTGGTGGGGCGTCCAGGTGCAGGATTCCTGTT
GTGCTGGTGGCGTATTCGCCGCGCGGCGTTCGGGGTGGTTAGC
CCGAAGCCGCAGCAAGGCTGCCGCAAGGCAGCAAGTATGGCGCA
CGCCAGGAGTCGGCGTGGTGGTATCCCGACCTATGGGGTGGCGCA
GGGGTTTTCTGGTTTTCGGCGTAGGTGATAGGAGTATACCGGGCGTG
GCCGTTGTAACAGGGTGGTGGCGTCCCTGGTGTGGCGGTGTC
GGTGTGGTATTTACCGGAAGCACAGGCAGCAGCCGAGCTAAGGCA
GCGAAATATGGTCCCGCGCGCAGGAGTTTTAGTGGGCTGGTCCG
GGCCCGCAGGCGTGTGCCGGGGTTCAGGCACCGTGGTGTCCCT
GGAGTCGGTACGCCGGCTGCAGCGGCAGCCAAAGCGGTGCAGAAAGCA
GCACAGTTTGGCTTAGTACCGGGTGGGAGTTGCCCGCGGTGGC
GTTGCTCCAGGGTGGGTGTTGCTCCTGGCGTCCGCTGGCTCCTGGA
GTGGCGTAGCACCCGGTGGGGTGGCCCCGGTGTGGGGTTGCA
CCGGTATCGGTCCGGCGGTGTCGCAGCAGCAGCTAAAAGCGCGCG
AAAGTTGCGCCAAAGCCAACTGCGCGCCGCGGGGCTCGGTGCA
GGTATTCGGGGTGGGTGTCGGAGTTGGAGTCCCGGTTTTGGCGTG
GGCGGGGAGTTCGGGACTGGGAGTGGTCCCGGAGTTCCTGGCTTT
GGTGCAGGCGCAGATGAAGGTGTTGTCGTAGCCTAGTCCGGAAGT
CGTGAAGGTGATCCGAGTAGCAGCCAGCATCTGCCGAGCACCCGAGC
AGCCCGCGTGTTCGGGTGCATTAGTGCAGCAAAAGCCGCAAGTAT
GGTGCAGCCGTGCCGGCGTCTTAGTGGTCTGGCGCCCTGGGTGGT
GTAGGCATTCGGGAGGTGTTGTGGGTGCAGGACCGGCCGCGCAGCT
GCGGCCGCAAAAGCAGCTGCAAAAGCGGCCAGTTTGGTTTTAGTGGC
GCCGAGGTTTTAGCGGTTTTAGGTGTTGGTGGACTGGGTGTACCTGGC
GTAGCGGTCTGGGTGGAATTCGCCCGCAGCGCGCGCAAGCGGCA
AAATATGGCGCGCAGGCTGGGGCGGCTGCTGGTGGGGCAGGTGAG
TTTCCGCTGGGCGGGTGGCGCAGTCCGGATTTGGTCTGAGCCCC

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ATTTTCCTGGCGGCGCATGTCTGGGTAAGCATGTGGTCGTAAACGT
 AAAt aa
 (SEQ ID NO: 26)
 MKKIWLALAGLVLAFSASAAQYEDMGVPGAI PGGVPGGVFYPGAGLGA
 LGGGALGPGGKPLKVPVGLAGAGLGAGLGAFFAVTFPGALVPGGVADA
 AAAYKAAKAGAGLGGVPGVGLGVSAVAVPQPGAGVKPGKVPVGLPG
 VYPGGVLPGARFPVGVLPVPTGAGVKPKAPGVGGAFAGIPGVGPFGG
 PQGVPLGYPIKAPKPLGGYGLPYTTGKLPYGYGPGGVAGAAGKAGYPT
 GTGVGPQAAAAA KAAKFGAGAAGVLPVGGAGVPGVPAIPIGIGGI
 AGVGT PAAAAA KAAKYGAAAGLVPGGPGFPGVGVPGAGVPGV
 GVPGAGIPVVPAGAGIPGAAVPGVVSPEAAKAAKAAKYGARPGVGVGG
 IPTYGVGAGGPFPGVGVGGIPGVAGVPGVGGVPGVGGVPGVGSPEAQ
 AAAAAKAAKYGAAGAGVLGGLVPGQAAVPGVPTGGVPGVGTAAAAA
 KAAKAAQFGLVPGVGVAPGVGVAPGVGVAPGVGLAPGVGVAPGVGVAP
 GVGVPAGIPGGVAAAAKSAKVAAKAQLRAAAGLGAAGIPGLGVGVGP
 GLGVGAGVPLGVGAGVPGFAGADEGVRRLSPELREGDPSSSQHLPS
 TPSSPRVPGALAAKAAKYGAAVPGVGLGGLGALGGVGI PGVVGAGPAA
 AAAAAKAAKAAQFGLVGAAGLGLGVGGLVPGVGGVGGIPPPAAAAKA
 AKYGAAGLGGVGGAGQFPLGGVAARPFGLSPI FPGGACLKACGRKR
 K

[0178] The polynucleotides of SEQ ID NO: 22 was codon optimized and synthesized by Gen9 DNA, now Ginkgo Bioworks internal synthesis. Overlaps between the pET28 vector and SEQ ID NO: 22 was designed to be between 30 and 40 bp long and added using PCR with the enzyme PrimeStar GXL polymerase (http://www.clontech.com/US/Products/PCR/GC_Rich/PrimeSTAR_GXL_DNA_Polymerase?site=10020:22372:US). The opened pET28a vector and insert DNA (SEQ ID NO: 22) was then assembled together into the final plasmid using SGI Gibson assembly (<https://us.vwr.com/store/product/17613857/gibson-assembly-hifi-1-step-kit-synthetic-genomics-inc>). Sequence of plasmid was then verified through sanger sequencing through Eurofins Genomics (www.eurofinsgenomics.com).

[0179] The transformed cells were cultivated in minimal media and frozen in 1.5 aliquots with glycerol at a ratio of 50:50 of cells to glycerol. One vial of this frozen culture was revived in 50 ml of minimal media overnight at 37° C., 200 rpm. Cells were transferred into 300 ml of minimal media and grown for 6-9 hours to reach an OD600 of 5-10.

[0180] A bioreactor was prepared with 2.7 L of minimal media+glucose and 300 ml of OD600 of 5-10 culture was added to bring the starting volume to 3 L. Cells were grown at 28° C., pH7 with Dissolved Oxygen maintained at 20% saturation using a cascade containing agitation, air and oxygen. pH was controlled using 28% w/w Ammonium hydroxide solution. Fermentation was run in a fed-batch mode using a DO-stat based feeding algorithm once the initial bolus of 40 g/L was depleted around 13 hours. After 24-26 hours of initial growth, the OD600 reached above 100. At this point, 300 mL of 500 g/L sucrose was added and

temperature was reduced to 25 C. High density culture was induced for protein production using 1 mM IPTG. Fermentation was continued for another 20-24 hours and cells were harvested using a bench top centrifuge at 9000 rcf, 15 C for 60 minutes. Cell pellet recovered from centrifugation was resuspended in a buffer containing 0.5M NaCl and 0.1M KH2PO4 at pH8 in a weight by weight ratio of 2x buffer to 1x cells.

[0181] The fermentations were performed at various temperature ranging from 25° to 28° C. For some fermentations, the temperature of the fermentation was maintained at a constant temperature and immediately upon completion of fermentation (OD600 of 5-10) the elastin was purified. For other fermentations, the temperature of the fermentations is maintained for a desired period of time and when cell densities of OD600 of 5-10 are reached, the temperature is reduced to induce protein production. Typically, the temperature is reduced from 28° C. to 25° C. After the fermentation at 25° C. is continued for 40-60 hours, the elastin is isolated.

[0182] The harvested cells were disrupted in a homogenizer at 14,000 psi pressure in 2 passes. Resulting slurry contained the collagen protein along with other proteins.

[0183] The supernatant from the homogenized cells was analyzed on an SDS-PAGE gel and a clear band was observed at around 70 kilodaltons corresponding to the expected size of 68 kilodaltons. The purified elastin is analyzed by mass spectrometry.

Full Length Elastin with DsbA Secretion Tag-His Tag-Linker-Thrombin Cleavage Site and GFP Beta-Lactamase Fusion

[0184] A human elastin with DsbA secretion tag-His tag-Linker-Thrombin cleavage site and GFP Beta-lactamase fusion is disclosed below. The codon-optimized nucleotide sequence encoding this elastin is provided in SEQ ID NO: 27. The amino acid sequence is disclosed in SEQ ID NO: 28. The DsbA secretion tag is encoded by nucleotides 1-72 and encodes amino acids 1-24. The His tag is encoded by nucleotides 73-99 and encodes a 9 histidine tag (SEQ ID NO: 129) of amino acids 25-33. The linker is encoded by nucleotides 100-111 and encodes amino acids 34-37. The thrombin cleavage site is encoded by nucleotides 112-135 and encodes amino acids 38-45. The green fluorescent protein (GFP) with linker is encoded by nucleotides 136-873 and encodes amino acids 46-291. The full-length elastin sequence is encoded by nucleotides 874-3153 and encodes amino acids 292-1051. The Beta-lactamase with linker is encoded by nucleotides 3154-3945 and encodes amino acids 1052-1315.

(SEQ ID NO: 27)
 ATGAAAAAGATTGGCTGGCGCTGGCTGGTTAGTTTTAGCGTTTAGCG
 CATCGGCGGCGCAGTATGAAGATCACCATCACCACCACCACCATCACCA
 CTCTGGCTCGAGCCTGGTCCCGCGCGCAGCCATATGTCTGGCTCGAGC
 AGTAAAGGTGAAGAACTGTTCCACGGTGTGTTCGGATCCTGGTTGAAC
 TGGATGGTGATGTTAACGGCCACAAATCTCTGTTCTGGTGAAGGTGA
 AGGTGATGCAACCAACGGTAAACTGACCCGTAAATTCATCTGCACTACC
 GGTAACCTGCCGGTCCATGGCCGACTCTGGTGACTACCCTGACCTATG

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GTGTTTCAGTGTTTTTCTCGTTACCCGGATCACATGAAGCAGCATGATTT
CTTCAAATCTGCAATGCCGGAAGTTATGTACAGGAGCGCACCATTCT
TTCAAAGACGATGGCACCTACAAAACCCGTGCAGAGTTAAATTTGAAG
GTGATACTCTGGTGAACCGTATTGAACTGAAAGGCATTGATTTCAAAGA
GGACGGCAACATCTGGGCCACAACTGGAATATAACTTCAACTCCCAT
AACGTTTACATCACCGCAGACAACAGAAAGCGGTATCAAAGCTAACT
TCAAAAATTCGCATAACGTTGAAGACGGTAGCGTACAGCTGGCGGACCA
CTACCAGCAGAACTCCGATCGGTGATGGTCCGGTCTGTCTGCCGGAT
AACCCTACCTGTCCACCCAGTCTTaaCTGTCCAAAGACCCGAACGAAA
AGCGCGACACATGGTGTCTGTGGAGTTCGTTACTGCAGCAGGTATCAC
GCACGGCATGGATGAACCTTACAACTTGGCGCGCCGGCGGTGGCGTA
CCAGGCGCAATTCCTGGGGGTGCCAGGCGGTGTTTTTATCCGGGCG
CCGGTCTTGGCGCACTGGGTGGCGGTGCACTGGGCCCGGGCGGCAACC
GCTGAAACCGGTACAGGTGGTTTAGCAGGCGCCGGCTTAGGCGCAGGT
CTGGGAGCATTTCCGGCAGTTACCTTTCCAGGGGCACTGGTTCCTGGAG
GTGTGGCCGATGCAGCCCGGCATATAAAGCCGCTAAAGCCGGTGGCGG
TTTAGGAGCGCTCCAGGTGTCTGGTGGCTGGGTGTAGCAGCGGTGCA
GTTGTTCCGCAGCCGGGAGCAGGGTTAAACCTGGTAAAGTCCGGGAG
TAGGCTGCCAGGCGTTTATCTGGTGGTGTTTTCCGGGTGCCCGTTT
TCCGGCGGTTGGTGTCTTCCAGGCGTCCGACCCGGAGCCGGTGTAAA
CCGAAAGCCCCGGTGTGGAGGTGCATTTGCAGGCATCCCGGAGTTG
GCCCCGTTGGTGGTCCGCAACTGGGGTCCGTTAGGTTATCCGATTAA
AGCACCGAACTGCCCGCGGTTATGGTCTGCCGTACACAACCGGTA
CTGCCGTATGGTTATGGCCCGGTGGAGTTGCCGGTGCAGCAGGTAAAG
CGGGTTATCTACCGGAACCGGTGTAGGTCGCGAGCCGCTGTCTCCGC
CGCCGCAAAAGCAGCGGTAATTTGGCGCCGAGCAGCGGGTGTCTTG
CCTGGAGTTGGTGGTGGCGGTGCCAGGGGTACCTGGTGAATTCGGG
GTATTTGGTGGTATTGCCGGTGTCCGACCCCGCCGCGGAGCTGCCGC
AGCGGCGGCTGCCAAAGCTGTAAATACCGTCCCGCGGGGTCTGGTG
CCAGGAGTCCGGGTTTTGGTCCGGGAGTGGTTGGCGTCCCTGGCGCAG
GCGTTCCTGGTGTGGGCGTCCAGGTGCAGGGATTCCTGTTGTGCCTGG
TGCCGGTATTCGGCGCGCGCCGTTCCGGGGTGGTTAGCCCGGAAGCC
GCAGCGAAGGCTGCCGCAAGGCGAGCAAGTATGGCGCACGCCAGGAG
TCGGCGTGGGTGGTATCCCGACCTATGGGGTGGGCGCAGGGGTTTTCC
TGGTTTCGGCGTAGGTGTAGGAGGTATACCGGGCGTGGCCGGTGTACCA
GGGGTTGGTGGCGTCCCTGGTGTGGCGGTGTGCCAGGTGTGGTATTT
CACCGGAAGCACAGGCAGCAGCCGAGCTAAGGCAGCGAAATATGGTGC
CGCCGGCGCAGGAGTTTTAGTGGGCTGGTTCGGGCCCGCAGGCAGCT
GTGCCGGGGTTCCAGGCACCGGTGGTCCCTGGAGTCCGTACGCCGG

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CTGCAGCGGCAGCCAAAGCGGCTGCGAAAGCAGCACAGTTTGGCTTAGT
ACCGGGTGTGGGAGTTGCCCGCGGTTGGCGTTGCTCCAGGGGTGGGT
GTTGCTCCTGGCGTCCGTCTGGCTCCTGGAGTGGGCGTAGCACCCGGT
TGGGGTGGCCCCGGGTGTGGGGTGCACCCGGTATCCGTCCGGGCGG
TGTCCGAGCAGCAGCTAAAGCGCGGCAAGTTGCGGCCAAAGCCCAA
CTGCGCGCCCGCGGGCTCCGTGCAGGTATTCGGGGCTGGGTGTGCG
GAGTTGGAGTCCCGGTTTGGGCGTGGGCGCGGGAGTTCCGGGACTGGG
AGTGGGTGCCGAGTCTCTGGCTTTGGTGCAGCGCAGATGAAGGTGTT
CGTCTGAGCCTGAGTCCGGAAGTGCCTGAAGGTGATCCGAGTAGCAGCC
AGCATCTGCCGAGCACCCGAGCAGCCCGCGTGTTCGGGTGCATTAGC
TGCAGCAAAGCCGCAAGTATGGTGCAGCCGTGCCGGGCGTCTTAGGT
GGTCTGGGCGCCCTGGGTGGTGTAGGCATTCCGGGAGGTGTGTGGGTG
CAGGACCGGCCCGCAGCTGCCGCCGCAAGCAGCTGCAAAAGCGGC
CCAGTTTGGTTAGTGGCGCCGCGAGTTTAGCGGTTTAGGTGTGGGT
GGACTGGGTGTACTGGCGTAGGCGGTCTGGGTGGAATTCGCCCGCAG
CGGCCGCAAAAGCGGCAAAATATGGCGCGGAGGCTGGGCGCGGTGCT
GGGTGGGCGAGTCACTTCCGCTGGGCGGGTTGCCGACGTCGGGGA
TTTGGTCTGAGCCGATTTCCCTGGCGCGCATGCTGGGTAAAGCAT
GTGGTCTGTAACGTAAAcaccagaaacgctggtgaaagtaaaagatgc
tgaagatcagttgggtgcaagagtggttacatcgaactggtatctcaac
agcggtaaagatccttgagagtttccgcccgaagaacgctttccaatga
tgagcacttttaagttctgctatgtggcgggatattatcccgattga
cgccgggcaagagcaactcggtcgcccatacactattctcagaatgac
ttggttgagtagtaccagtcacagaaagcatcttacggatggcatga
cagtaagagaattatgagtgctgccataaccatgagtgataaactgc
ggccaacttactctgacaacgatcggaggaccgaaggagtaaccgct
tttttgcaacatgggggatcatgtaactcgccttgatcgttgggaac
cggagctgaatgaagccatacacaacgacgagcgtgacaccacgatgcc
tgtagcaatggcaacaacgcttgccgcaactataactggcgaactactt
actctagcttcccggcaacaataatagactggatggaggcggataaag
ttgcaaggaccacttctcgctcggccctccggctgggtggtttattgc
tgataaatctggagccggtgagcgtgggtctcggcgatcattgcagca
ctggggccagatggtgagccctccgctatcgtagttatctacacgagcg
ggagtcaggcaactatggatgaacgaaatagacagatcgctgagatagg
tgcctcactgattaagcattggtaa

(SEQ ID NO: 28)

MKKIWLALAGLVLAFSASAAQYEDHHHHHHHHSGLVPRGSHMSGSS
SKGELFTGVVPILEVELDGVNKHKFSVRGEGEGDATNGKLTCLKFICTT
GKLPVWPPLVTTLYGVQCFSRYPDHMKQHDFFKSAMPEGYVQERTIS
FKDDGTYKTRAEVKFEGDVLVNRIELKIDFKEDGNILGHKLEYNFNH

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NVYITADKQKNGIKANFKIRHNVEDGSVQLADHYQONTPIGDGPVLLPD
 NHYLSTQS KLSKDPNEKRDMVLEFVTAAGITHGMDELYKSGAPGGGV
 PGAI PGGVPGGVFYPGAGL GALGGGALPGGKPLKPVPGGLAGAGLGAG
 LGAFP AVTFPGALVPGGVADAAAAAYKAAKAGAGLGGVPGVGLGVSAGA
 VVPQPGAGVKPGKVPVGLPGVYPGGVLPGARFPVGVLPVPGVPTGAGVK
 PKAPGVGGAFFAGIPGVGPFPGPQPGVPLGYPIKAPKLPGGYGLPYTTGK
 LPYGYPGGVVAGAAKAGYPTGTGVGPQAAAAAAKAAAKFAGAGAGLV
 PGVGGAGVPGVPGAIPGIGGIAGVGTAAAAAAKAAKAYGAAAGLV
 PGGPFGPGVGVPGAGVPGVPGAGIPVVPAGIPGA AVPGVVSPEA
 AAKAAKAAKAYGARPGVGVGGIPTYGVGAGGFPFGVGVGGIPGVAGVP
 GVGVPVGGVPGVGISPEAQAAAAKAAKYGAAGAGVLGGLVPGQAA
 VPGVPGTGGVPGVGTAAAAKAAKAAQFGLVPGVGVAPGVGVPAGVGV
 VAPGVGLAPGVGAPGVGAPGVGAPGIGPGGVAAAAKSAKVAAKAQ
 LRAAAGL GAGIPGLGVGVPGLVGAGVPLGVGAGVPGFAGAGDEGV
 RRSLSPELREGD PSSSQHLPS TPSSPRVPGALAAKAAKYGA AVPGVLG
 GLGALGGVGI PGGVVAGPAAAAAAKAAKAAQFGLVGAAGLGLGVLG
 GLGVPGVGLGGIPAAAAKAAKYGAAGLGGVLLGGAGQFPLGGVAARPG
 FGLSPIFPGGACL GKACGRKRKHPE TLVKVDAEDQLGARVYIELDLN
 SGKILESFRPEERFPM MSTFKVLLCGAVLSRIDAGQEQLGRRRIHYSQND
 LVEYSPVTEKHLTDGMTVRELCSAAITMSDNTAANLLTTIGGPKELTA
 FLHMGD HVTRLRDRWEP ELNEAI PNDERDTMPVAMATLRKLLTGELL
 TLASRQQLIDWMEADK VAGPLLR SALPAGWFIADKSGAGERGSRGIIAA
 LGPDGKPSRIVVIYTTGSQATMDERNRQIAEIGASLIKHW

Example 5: Production of Truncated Elastin

[0185] Truncated human elastin is produced using the expression system as described in Example 4. The full length amino acid sequence lacking the native secretion tag is disclosed in SEQ ID NO: 29.

(SEQ ID NO: 29)

GGVPGAIPGGVPGGVFYPGAGL GALGGGALPGGKPLKPVPGGLAGAGL
 GAGLGAFFAVTFPGALVPGGVADAAAAAYKAAKAGAGLGGVPGVGLGVS
 AGAVVPQPGAGVKPGKVPVGLPGVYPGGVLPGARFPVGVLPVPGVPTGA
 GVKPKAPGVGGAFFAGIPGVGPFPGPQPGVPLGYPIKAPKLPGGYGLPYT
 TGKLPYGYPGGVVAGAAKAGYPTGTGVGPQAAAAAAKAAAKFAGAGAA
 GVLPGVGGAGVPGVPGAIPGIGGIAGVGTAAAAAAKAAKAYGAAA
 GLVPGGPGFPGVGVPGAGVPGVPGAGIPVVPAGIPGA AVPGVVS
 PEAAKAAKAAKAYGARPGVGVGGIPTYGVGAGGFPFGVGVGGIPGVA
 GVPVGGVPGVGGVPGVGISPEAQAAAAKAAKYGAAGAGVLGGLVPGP
 QAAVPGVPGTGGVPGVGTAAAAKAAKAAQFGLVPGVGVAPGVGVP

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GVGVPAGVGLAPGVGAPGVGVPAGVPGVAPGIGPGGVAAAAKSAKVA
 KAQLRAAAGL GAGIPGLGVGVPGLVGAGVPGLVGAGVPGFAGAGAD
 EGVRRSLSPELREGD PSSSQHLPS TPSSPRVPGALAAKAAKYGA AVPG
 VLGGLGALGGVGI PGGVVAGPAAAAAAKAAKAAQFGLVGAAGLGLG
 GVGGLGVPGVGLGGIPAAAAKAAKYGAAGLGGVLLGGAGQFPLGGVAA
 RPFGLSPIFPGGACL GKACGRKRK

[0186] The codon optimized polynucleotide sequence encoding the full length human elastin lacking the native secretion tag is disclosed in SEQ ID NO: 30.

(SEQ ID NO: 30)

GGTGGCGTACCAGGCGCAATTCCTGGGGTGTCCAGGCGGTGTTTTTT
 ATCCGGGCGCCGGTCTTGGCGCACTGGGTGGCGGTGCACTGGGCCCGG
 CGGCAAAACCGCTGAAACCGGTACCAGGTGGTTTAGCAGGCGCCGGCTTA
 GGCGCAGGTCTGGGAGCATTTCCGGCAGTTACCTTTCCAGGGCCTGCG
 TTCTGGAGGTGGCCGATGCAGCCGGCATATAAAGCCGCTAAAGC
 CGGTGCGGGTTTAGGAGGCGTCCCAGGTGTCGGTGGCCTGGGTGTTAGC
 GCCGGTGCAGTTGTTCCGCAGCCGGGAGCAGGGTTAAACCTGGTAAAG
 TGCCGGGAGTAGGTCTGCCAGGCGTTTATCTGGTGGTGTGTTTTCGGG
 TGCCCGTTTTCCGGGCGTGGTGTCTTCCAGGCGTGCCGACCCGGAGCC
 GGTGTTAAACCGAAAGCCCCGGTGTGGAGGTGCATTTGCAGGCATCC
 CGGGAGTTGGCCCGTTTGGTGGTCCGCAACCTGGGGTTCCGTTAGGTTA
 TCCGATTAAGCACCAGAACTGCCCGGCGTTATGGTCTGCCGTACACA
 ACCGGTAAACTGCCGTATGGTTATGGCCCGGTGGAGTTCGGGTGCAG
 CAGGTAAAGCGGTTATCTACCGGAACCGGTGTAGGTCCGCAGGCCGC
 TGCTGCCCGCCGCAAAAGCAGCGGTAAATTTGGCCCGGAGCAGCG
 GGTGTTCTGCCTGGAGTTGGTGGTGGGGCGTCCAGGGTACCTGGT
 CAATTCGGGTATTGGTGGTATTGCCGTGTCCGCACCCCGCCCGCGC
 AGCTGCGCAGCGCGGCTGCCAAAGCTGCTAAATACGGTCCCGCGGCG
 GGTCTGGTCCAGGAGTCCGGTTTTGGTCCGGGAGTGGTGGCGTGC
 CTGGCGCAGGCTTCTGGTGTGGCGTTCAGGTGCAGGATTCTCTGT
 TGTGCTGGTCCCGTATCCCGCGCGGCGTTCCGGGGGTGGTTAGC
 CCGGAAGCCGCAGCGAAGCTGCGCAAGGCAGCAAAGTATGGCGCAC
 GCCCAGGAGTCCGGTGGTGGTATCCCGACCTATGGGTGGGCGCAGG
 GGGTTTTCTGGTTTTCCGGCTAGGTGTAGGAGGTATACCGGGCGTGGCC
 GGTGTACCAGGGTTGGTGGCGTCCCTGGTGTGGCGGTGTGCCAGGTG
 TTGGTATTTACCGGAAGCACAGGCAGCAGCCGACTAAGGCAGCGAA
 ATATGGTCCCGCGCGCAGGAGTTTTAGGTGGGCTGGTTCGGGCCCG
 CAGGCAGCTGTGCCGGGGTCCAGGCACCGGTGGTGTCCCTGGAGTCG
 GTACGCGGCTGCAGCGCAGCCAAAGCGGCTGCCAAAGCAGCACAGTT

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TGGCTTAGTACCGGGTGTGGGAGTTGCCCCCGCGTGGCGTTGCTCCA
GGGGTGGGTGTGCTCCTGGCGTCGGTCTGGCTCCTGGAGTGGGCGTAG
CACCCGGTGTGGGGTGGCCCCGGGTGTGGGGTTGCACCCGGTATCGG
TCCGGCGGTGTGCGCAGCAGCAGCTAAAAGCGCGCGAAAGTTGCGGCC
AAAGCCCAACTGCGCGCCCGCGGGCCTCGGTGCAGGTATTCCGGGGC
TGGGTGTCGGAGTTGGAGTCCCGGTTTGGGCGTGGGCGCGGAGTTCC
GGGACTGGGAGTGGGTGCCGGAGTTCTGGCTTTGGTGCAGCGCAGAT
GAAGGTGTCGTCGTAGCTGAGTCCGGAAGTGCCTGAAGTGTATCCGA
GTAGCAGCCAGCATCTGCCGAGCACCCGAGCAGCCCGTGTTCGGG
TGCATTAGCTGCAGCAAAAGCCCAAGTATGGTGCAGCCGTGCCGGC
GTCTTAGGTGGTCTGGGCGCCCTGGGTGGTGTAGGCATTCCGGGAGGTG
TTGTGGGTGCAGGACCGCGCCCGCAGCTGCGGCCGCCAAAGCAGCTGC
AAAAGCGGCCAGTTTGGTTTAGTGGGCGCGCAGGTTTAGCGGTTTA
GGTGTGGGTGGACTGGGTGTACCTGGCGTAGGCGGTCTGGGTGGAATTC
CGCCt aa

[0187] The amino acid sequence of a 60.7 kDa human elastin truncated at the C-terminal is disclosed in SEQ ID NO: 31. The 60.7 kDa truncated elastin has amino acids 706-761 deleted from the full length elastin.

(SEQ ID NO: 31)

GGVPGAIPGGVPGVFYFPGAGLGGALGPGGKPLKVPVPGLAGAGL
GAGLGAFFAVTFPGALVPGGVADAAAAYKAAKAGAGLGGVPGVGLGVS
AGAVVPQPGAGVKPKGVPGVLPVPGVLPGARFPVGVLPVPTGA
GVKPKAPGVGGAFAGIPGVPGFPGVQPGVPLGYPIKAPKLPGGYGLPYT
TGKLPYGYPGGVVAGAAGKAGYPTGTGVPQAAAAAANKAAKFGAGAA
GVLPGVGGAGVPGVPGAIPGI GGIAGVGTAAAAAANKAAKYGAAA
GLVPGGPGFPGVGVPGAGVPGVPGAGIPVVPAGIPGAAVPGVVS
PEAAAKAAAKAAYGARPGVGVGGIPTYGVGAGFPFPGVGVGGIPGVA
GVPVGGVPGVGGVPGVGI SPEAQAAAAAANKAAYGAAGVGLGLVPGP
QAAVPGVPGTGGVPGVGTAAAAAANKAAKAAQFGLVPGVGVAPGVVAP
GVGVAPGVGLAPGVVAPGVVAPGVVAPGIPGGVAAAASAAKVAA
KAQLRAAAGL GAGI PGLGVGVPGVPLGVGAGVPLGVGAGVPGFAGAD
EGVRRSLSPELREGDPSSSHLPSTPSSPRVPGALAAAKAAYGAAVPG
VLGGLGALGGVGI PGGVGVGAGPAAAAAANKAAKAAQFLVGAAGLGLL
GVGGLGVPGVGLGGIPP

[0188] The codon optimized polynucleotide sequence encoding the truncated 60.7 kDa human elastin is disclosed in SEQ ID NO: 32.

(SEQ ID NO: 32)

GGTGGCGTACCAGGCGCAATTCTGGGGGTGTCCAGGCGGTGTTTTT
ATCCGGGCGCCGGTCTTGGCGCACTGGGTGGCGGTGCACTGGGCCCCGG
CGGCAACCGCTGAAACCGGTACCAGGTGGTTTAGCAGGCGCCGGCTTA
GGCGCAGGTCTGGGAGCATTTCCGGCAGTTACCTTCCAGGGGCACTGG
TTCTGGAGGTGTGGCCGATGCAGCCCGGCATATAAAGCCGCTAAAGC
CGGTGCGGGTTTAGGAGCGTCCCAGGTGTGGTGGCTGGGTGTAGC
GCCGGTGCAGTTGTTCCGCAGCCGGGAGCAGGGTTAAACCTGGTAAAG
TGCCGGGAGTAGGTCTGCCAGGCGTTTATCCTGGTGGTGTGTTGCCGGG
TGCCCGTTTCCGGGCGTGGTGTCTTCCAGCGCTGCCGACCGGAGCC
GGTGTAAACCGAAAGCCCCGGTGTGGAGGTGCATTGCAGGCATCC
CGGGAGTTGGCCCGTTGGTGGTCCGCAACCTGGGTTCCGTTAGGTTA
TCCGATTAAAGCACCGAAACTGCCCGCGGTTATGGTGTGCCGTACACA
ACCGGTAAACTGCCGTATGGTTATGGCCCGGTGGAGTTGCCGGTGCAG
CAGGTAAAGCGGGTTATCCTACCGAACCGGTGTAGGTCCGCAGGCCGC
TGCTGCCGCCCGCGCAAAGCAGCGGCTAAATTTGGCGCCGGAGCAGCG
GGTGTCTGCCTGGAGTTGGTGGTGCGGCGTGCAGGGGTACCTGGTG
CAATTCGGGTATTGGTGGTATTGCCGGTGTCCGACCCCGGCCCGGC
AGCTGCCGAGCGGCGGCTGCCAAAGCTGCTAATAACGGTCCCGCGGC
GGTCTGGTGCCAGGAGGTCGGGTTTGGTCCGGGAGTGGTGGCGTGC
CTGGCGCAGGCGTCTGGTGTGGGCGTCCAGGTGCAGGGATTCTGT
TGTGCTGTGCGGTTATCCCGCGCGCCGTTCCGGGGTGGTTAGC
CCGGAAGCCGAGCGAAGGCTCGCGCAAAGCAGCAAAGTATGGCGCAC
GCCAGGAGTCCGGTGGGTGGTATCCCGACCTATGGGGTGGGCGCAGG
GGGTTTTCTGTTTTCGGCGTAGGTGTAGGAGGTATACCGGCGTGGCC
GGTGTACCAGGGGTTGGTGGCGTCCCTGGTGTGGCGGTGCCAGGTG
TTGGTATTTACCGGAAGCACAGGCAGCAGCCGACGCTAAGGCAGCGAA
ATATGGTCCCGCCGCGCAGGAGTTTTAGTGGGCTGGTTCGGGCCCG
CAGGCAGCTGTCCGGGGTTCCAGGCACCGGTGGTGTCCCTGGAGTCG
GTACGCCGGCTGCAGCGCAGCCAAAGCGGCTGCGAAAGCAGCACAGTT
TGGCTTAGTACCGGGTGTGGGAGTTGCCCCCGCGTGGCGTTGCTCCA
GGGGTGGGTGTGCTCCTGGCGTCCGTCTGGAGTGGGCGTAG
CACCCGGTGTGGGGTGGCCCCGGGTGTGGGGTTGCACCGGTATCCGG
TCCGGGCGGTGTCGCAGCAGCAGCTAAAAGCGCGCGCAAAGTTGCCGGC
AAAGCCCAACTGCGCGCCCGCGGGCCTCGGTGCAGGTATTCCGGGGC
TGGGTGTCGGAGTTGGAGTCCCGGTTTGGGCGTGGGCGGGGAGTTCC
GGGACTGGGAGTGGGTGCCGGAGTTCTGGCTTTGGTGCAGCGCAGAT
GAAGGTGTCGTCGTAGCTGAGTCCGGAAGTGCCTGAAGTGTATCCGA
GTAGCAGCCAGCATCTGCCGAGCACCCGAGCAGCCCGGTGTTCCGGG
TGCATTAGCTGCAGCAAAAGCCCAAGTATGGTGCAGCCGTGCCGGC

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GTCTTAGGTGGTCTGGGCGCCCTGGGTGGTGTAGGCATTCGGGGAGGTG
TTGTGGGTGCAGGACCGGCGCCCGCAGCTGCGGCCGCAAGCAGCTGC
AAAAGCGGCCAGTTTGGTTTGTAGTGGGCGCCGAGGTTTAGCGGGTTTA
GGTGTGGGTGGACTGGGTGTACCTGGCGTAGGCGGTCTGGGTGGAATTC
CGCCt aa

[0189] The amino acid sequence of a 58.8 kDa human
elastin truncated at the N-terminal is disclosed in SEQ ID
NO: 33. The 58.8 kDa truncated elastin has amino acids 2-85
deleted from the full length elastin.

(SEQ ID NO: 33)

GLGGVPGVGLVSVAGAVVPPGAGVKPKGKVPVGLPGVYPGGVLPGARF
PGVGLVPGVPTGAGVKPKAPGVGGAFAGIPGVGPFPGPQPGVPLGYPIKA
PKLPGGYLPHYTTGKLPYGYGPGVAGAAAGKAGYPTGTGVPQAAAAA
KAAAKFGAAGAVLPGVGGAGVPGVPAIPGIIGIAGVGTAAAAA
AKAAKYGAAAGLVPGGPGFPGVGVPGAGVPGVPGAGIPVVPAGIP
GAAVPGVVSPAAAAKAAKAAKYGARPGVGVGGIPTYGVGAGGFPFGVG
VGGIPGVAGVPGVGGVPGVGGVPGVGI SPEAQAAAAKAAKYGAAGAVL
GGLVPGPQAAVPGVPGTGGVPGVGTAAAAA KAAKAAQFGLVPGVGVAP
GVGVAPGVVAPVGLAPGVVAPGVVAPGVVAPGI GPGVAAAAKSA
AKVAAKAQLRAAAGL GAGI PGLGVGVPGVGLGVGAGVPGVGVG
AGADEGVRRLSPELREGDPS SSQHLPSTPSSPRVPGALAAKAAKYGAA
VPGVGLGALGGVGI PGGVVAGPAAAAA KAAKAAQFGLVGAAGLG
GLGVGGLVPGVGGVGGIPAAAAA KAAKYGAAGLGGVGLGAGQPPLGGVA
ARPGFGLSPIFPGGACLKACGRKRK

[0190] The codon optimized polynucleotide sequence
encoding the 58.8 kDa truncated human elastin is disclosed
in SEQ ID NO: 34.

(SEQ ID NO: 34)

GGTTTAGGAGGCGTCCAGGTGTCCGGTGGCCTGGGTGTAGCCCGGT
GCAGTTGTTCCGCAGCCGGGAGCAGGGTTAAACTGGTAAAGTCCCG
GGAGTAGGTCTGCCAGGCGTTTATCCTGGTGGTGTTCGCCGGTGCC
CGTTTTCCGGGCGTTGGTGTCTTCCAGGCGTGCCGACCGGAGCCGGT
GTTAAACCGAAAGCCCCGGTGTGGAGGTGCATTTGCAGGCATCCCG
GGAGTTGGCCCGTTTGGTGGTCCGCAACTGGGGTTCGGTTAGGTTAT
CCGATTAAGCACCAGAACTGCCCGCGGTTATGGTCTGCCGTACACA
ACCGGTAAACTGCCGTATGGTTATGGCCCGGGTGGAGTTGCGGGTGCA
GCAGGTAAAGCGGGTTATCCTACCGAAACCGGTGTAGTCCGAGGCC
GCTGTGCCCGCCGCAAAAGCAGCGGCTAAATTTGGCGCCGGAGCA
GCGGGTGTTCGCCGTGGAGTTGGTGGTGGGGCGTGCCAGGGTACCT
GGTGCAATCCGGGTATTGGTGGTATTGCCGGTGTCCGACCCCGCC

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GCGGCAGCTGCGGCAGCGCGGCTGCCAAAGCTGCTAAATACGGTGCC
GCGGCGGGTCTGGTGCCAGGAGGTCGGGTTTGGTCCGGGAGTGGTT
GGCGTGCCGTGGCGCAGGCGTTCCTGGTGTGGCGGTTCCAGGTGCAGGG
ATTCTGTGTGCTGGTGCCGGTATTCGCCGCGCGGCGTTCGGGG
GTGGTTAGCCCGAAGCCGCGAGCGAAGGCTGCGGCAAAGGCAGCAAAG
TATGGCGCACGCCAGGAGTCGGCGTGGTGGTATCCCGACCTATGGG
GTGGGCGCAGGGGGTTTTCTGGTTTCGGCGTAGGTGTAGGAGGTATA
CCGGGCGTGGCCGGTGTACCAGGGGTTGGTGGCGTTCCTGGTGTGGC
GGTGTGCCAGGTGTTGGTATTTACCGGAAGCACAGGCGAGCCGCA
GCTAAGGCAGCGAAATATGGTGCCCGCGCAGGAGTTTTAGGTGGG
CTGGTTCCGGGCCCCGAGGCAGCTGTGCCGGGGTTCCAGGCACCGGT
GGTGTCCCTGGAGTCGGTACGCCGGTGCAGCGGCGCAAAAGCGGCT
GCGAAAGCAGCACAGTTTGGCTTAGTACCGGGTGTGGGAGTTGCCCC
GGCGTTGGCGTTGCTCCAGGGGTTGGTGTGCTCCTGGCGTCCGTTCTG
GCTCCTGGAGTGGGCGTAGCACCCGGTGTGGGGTGGCCCCGGGTGT
GGGGTTGCACCGGGTATCGGTCCGGGCGGTGTGCGCAGCAGCAGTAAA
AGCGCGGCGAAAGTTGCGGCCAAAGCCCAACTGCGCGCCGCGCGGGC
CTCGGTGCAGGTATTCGGGGCTGGGTGTGCGAGTGGAGTCCCGGGT
TTGGGCGTGGGCGCGGGAGTTCCGGGACTGGGAGTGGGTGCCGGAGTT
CCTGGCTTTGGTGCAGGCGCAGATGAAGGTGTTCTGTCGTAGCCTGAGT
CCGGAAGTGCAGTGAAGGTGATCCGAGTAGCAGCCAGCATCTGCCGAGC
ACCCCGAGCAGCCCGGTGTTCCGGGTGCATTAGCTGCAGCAAAGCC
GCCAAGTATGGTGCAGCCGTGCCGGGCGTCTTAGGTGGTCTGGGCGCC
CTGGGTGGTGTAGGCATTCGGGAGGTGTTGTGGGTGCAGGACCGGCC
GCCGAGCTGCGGCCGCAAGCAGCTGCAAAAGCGGCCAGTTTGGT
TTAGTGGGCGCCGAGGTTTAGGCGGTTTAGGTGTGGGTGGACTGGGT
GTACCTGGCGTAGGCGGTCTGGGTGGAATTCGCCCGCAGCGCCCGC
AAAGCGCAAATATGGCGCGCAGGCTGGGCGCGGTGCTGGGTGGG
GCAGGTCAGTTTCCGCTGGGCGGGTTGCCGACGTCCGGGATTTGGT
CTGAGCCGATTTTCCCTGGCGGCGCATGTCTGGGTAAAGCATGTGGT
CGTAAACGTAAAt aa

[0191] The amino acid sequence of a 57 kDa human
elastin truncated at the C-terminal is disclosed in SEQ ID
NO: 35. The 57 kDa truncated elastin has amino acids
661-761 deleted from the full length elastin.

(SEQ ID NO: 35)

GGVPGAIPGGVPGVFPYAGLGLGGGALGPGGKPLKVPVGGLAGG
LGAGLGAFFPAVTFPGALVPGGVADAAAAYKAAKAGAGLGGVPGVGLG
VSAGAVVPPGAGVKPKGKVPVGLPGVYPGGVLPGARFPVGVVLPVGP

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TGAGVKPKAPGVGGAFAGIPGVGPFGGPQGVPLGYPIKAPKLPGGYG
 LPYTTGKLPYGYGPGGVAGAAGKAGYPTGTGVGPQAAAAAAKAAKF
 GAGAAGVLPVGGAGVPGVPGAIPIGGIAGVGTAAAAAAKAAKAA
 KYGAAAGLVPGGPGFPGVVGVPAGVPGVPGAGIPVVPAGIPGA
 AVPGVVSPEAAKAAKAAKYGARPGVVGVIPTYGVGAGGFPFGVGV
 VGGIPGVAGVPGVGGVPGVGGVPGVVISPEAQAAAAKAAKYGAAGAG
 VLGGLVPGPQAAVPGVPGTGGVPGVGTAAAAKAAKAAQFGLVPGV
 GVAPGVGAPGVGAPVGLAPGVGAPGVGAPGVGAPGIPGGVA
 AAKSAAKVAKAQLRAAAGLGAGIPGLGVGVPGVGLVGVGAPGLV
 GAGVPGFAGADEGVRRSLSPELREGDPSSSQHLPTSPSPRVPGALA
 AAKAAKYAAVPGVGLGALGGVPIPGGVGAGP

[0192] The codon optimized polynucleotide sequence encoding the 57 kDa truncated human elastin is disclosed in SEQ ID NO: 36

(SEQ ID NO: 36)

GGTGGCGTACCAGGCGCAATCCTGGGGTGTCCAGGCGGTGTTTTT
 TATCCGGCGCCGGTCTTGGCGCACTGGGTGGCGGTGCACTGGGCCG
 GCGGCAAAACCGCTGAAACCGTACCAGGTGTTTAGCAGGCGCCGGC
 TTAGCGCAGGCTCTGGGAGCATTCCGGCAGTTACCTTCCAGGGCA
 CTGGTTCCTGGAGGTGTGGCCGATGCAGCCCGGCATATAAAGCCGCT
 AAAGCCGGTGCAGGGTTTAGGAGGCGTCCAGGTGTCCGGTGGCCTGGGT
 GTTAGCGCCGGTGCAGTGTTCGCGAGCCGGAGCAGGGTTAAACCT
 GGTAAGTGCAGGGAGTAGGTCTGCCAGGCGTTTATCCTGGTGGTGT
 TTGCCGGGTGCCCGTTTCCGGGCGTGGTGTCTTCCAGGCGTGCCG
 ACCGGAGCCGGTGTAAACCGAAAGCCCGGTTGGAGGTGCATTT
 GCAGGCATCCCGGAGTTGGCCGTTTGGTGGTCCGCAACCTGGGGTT
 CCGTTAGGTTATCCGATTAAGCACCGAAACTGCCCGGCGTTATGGT
 CTGCCGTACACACCGTAACCTGCCGTATGGTTATGGCCCGGTGGA
 GTTGCAGGTGCAGCAGGTAAGCGGGTTATCCTACCGGAACCGGTGTA
 GGTCGCGAGGCGCTGTGTCGCCCGCCGCAAAAGCAGCGGCTAAATTT
 GGCGCCGAGCAGCGGGTGTCTGCTGGAGTTGGTGGTGCAGGGCGTG
 CCAGGGGTACCTGGTGCAATCCGGGTATTGGTGGTATTGCCGGTGTCT
 GGCACCCCGGCCCGGCAGCTGCCGCGAGCGCGGCTGCCAAAGCTGCT
 AAATACGGTGCAGCGCGGGTCTGGTGCAGGAGGTCCGGTTTTGGT
 CCGGAGTGGTGGCGTGCCTGGCGAGGCGTTCTGGTGGTGGCGGTT
 CCAGGTGCAGGATTCCTGTTGTGCTGGTGCAGGATTCCTCCGGCGCG
 GCCGTTCCGGGGTGGTTAGCCCGGAAGCCGAGCGAAGGCTGCCGCA
 AAGGAGCAAAAGTATGGCGCACCCAGGAGTCCGGTGGTGGTATC
 CCGACCTATGGGGTGGCGCAGGGGTTTTCTGGTTTTCCGGCGTAGGT
 GTAGGAGTATACCGGCGTGGCGGTTACCAGGGTGGTGGCGCT

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CCTGGTGTGGCGGTGTGCCAGGTGTGGTATTTACCAGGAAAGCACAG
 GCAGCAGCCGCGAGCTAAGGCAGCGAAATATGGTCCCGCCGCGCAGGA
 GTTTTAGGTGGGCTGGTTCGGGGCCCGCAGGCAGCTGTGCCGGGGTT
 CCAGGCACCGGTGGTGTCCCTGGAGTCGGTACGCCGGGTGCAGCGCA
 GCCAAAGCGGCTGCCAAAGCAGCACAGTTTGGCTTAGTACCGGGTGTG
 GGAGTTGCCCCCGGCTGGCGTTGCTCCAGGGTGGTGTGCTCCT
 GCGCTCGGTCTGGCTCCTGGAGTGGGCGTAGCACCCGGTGTGGGGTG
 GCCCCGGTGTGGGGTGCACCGGATTCGGTCCGGGCGGTGTGCGCA
 GCAGCAGCTAAAAGCGCGCGAAAGTTCCGGCCAAAGCCCAACTGCGC
 GCCCGCGGGCCTCGGTGCAGGTATTCGGGGCTGGGTGTGGAGTT
 GGAGTCCCGGGTTTGGGCGTGGGCGGGAGTTCCGGGACTGGGAGTG
 GGTCCCGAGTTCCTGGCTTGGTGCAGGCGCAGATGAAGGTGTTCTGT
 CGTAGCCTGAGTCCGGAAGTGCAGGTGATCCGAGTAGCAGCCAG
 CATCTGCCGAGCACCAGCAGCAGCCCGGTGTTCGGGTGCATTAGCT
 GCAGCAAAAGCCGCAAGTATGGTGCAGCCGTGCCGGGCGTCTTAGGT
 GGTCTGGGCGCCCTGGGTGGTGTAGGCATTCGGGAGGTGTGTGGGT
 GCAGGACCGTaa

[0193] The amino acid sequence of a 53.9 kDa human elastin truncated at the C-terminal is disclosed in SEQ ID NO: 37. The 53.9 kDa truncated elastin has amino acids 624-761 deleted from the full length elastin.

(SEQ ID NO: 37)

GGVPGAIIPGVPGVFPYFPGAGLGLGALGPGGKPLKPVPGGLAGAG
 LGAGLGAFFPAVTFPGALVPGGVADAAAAYKAAKAGAGLGGVPGVGLG
 VSAGAVVPQPGAGVKPGKVPVGLPGVYPGVLPGARFPVPGVLPVGP
 TGAGVKPKAPGVGGAFAGIPGVGPFGGPQGVPLGYPIKAPKLPGGYG
 LPYTTGKLPYGYGPGGVAGAAGKAGYPTGTGVGPQAAAAAAKAAKF
 GAGAAGVLPVGGAGVPGVPGAIPIGGIAGVGTAAAAAAKAAKAA
 KYGAAAGLVPGGPGFPGVVGVPAGVPGVPGAGIPVVPAGIPGA
 AVPGVVSPEAAKAAKAAKYGARPGVVGVIPTYGVGAGGFPFGVGV
 VGGIPGVAGVPGVGGVPGVGGVPGVVISPEAQAAAAKAAKYGAAGAG
 VLGGLVPGPQAAVPGVPGTGGVPGVGTAAAAKAAKAAQFGLVPGV
 GVAPGVGAPGVGAPVGLAPGVGAPGVGAPGVGAPGIPGGVA
 AAKSAAKVAKAQLRAAAGLGAGIPGLGVGVPGVGLVGVGAGVPGLV
 GAGVPGFAGADEGVRRSLSPELREGDPSSSQHLPTSPSPRVPGA

[0194] The codon optimized polynucleotide sequence encoding the 53.9 kDa truncated human elastin is disclosed in SEQ ID NO: 38

(SEQ ID NO: 38)

GGTGGCGTACCAGGCGCAATTCTGGGGGTGTCCAGGCGGTGTTTTT
TATCCGGGCGCCGGTCTTGGCGCACTGGGTGGCGGTGCACTGGGCCCG
GGCGGCAAACCGCTGAAACCGGTACCAGGTGGTTTAGCAGGCGCCGGC
TTAGCGCAGGTCTGGGAGCATTTCGCGCAGTTACCTTCCAGGGGCA
CTGGTTCCTGGAGGTGTGGCCGATGCAGCCGCGGCATATAAAGCCGCT
AAAGCCGGTGGCGGTTTAGGAGCGCTCCAGGTGTCTGGTGGCCTGGGT
GTTAGCGCCGGTGCAGTTGTTCCGCGAGCCGGGAGCAGGGTTAAACCT
GGTAAAGTGCCGGGAGTAGGTCTGCCAGGCGTTTATCCTGGTGGTGT
TTGCCGGGTGCCCGTTTTCCGGCGCTTGGTGTCTTCCAGGCGTGCCG
ACCGGAGCCGGTGTAAACCGAAAGCCCGGCTGTGGAGGTGCATTT
GCAGGCATCCCGGAGTTGGCCCGTTTGGTGGTCCGCAACCTGGGGTT
CCGTTAGGTTATCCGATTAAGCACCGAACTGCCCGCGGTTATGGT
CTGCCGTACACAACCGGTAACCTGCCGTATGGTTATGGCCCGGTTGGA
GTTGCGGGTGCAGCAGGTAAAGCGGTTATCCTACCGGAACCGGTGTA
GGTCCGACGGCCGCTGCTGCCGCCCGCAAAGCAGCGGCTAAATTT
GGCGCCGAGCAGCGGGTGTCTGCCCTGGAGTTGGTGGTGGCGCGT
CCAGGGGTACCTGGTGCAATTCCGGGTATTGGTGGTATTGCCCGGTGTC
GGCACCCCGGCGCGGCGAGCTGCCGCGCGGCTGCCAAAGCTGCT
AAATACGGTGCAGCGCGGGTCTGGTGCAGGAGTCCGGGTTTTGGT
CCGGGAGTGGTTGGCGTGCTGGCGCAGGCGTTCCTGGTGTGGCGTT
CCAGGTGCAGGATTCCTGTTGTGCTGGTGCCTGATTCGCCGCGCG
GCCGTTCCGGGGTGGTTAGCCGGAAGCCGCGAGCAAGGCTGCAGCA
AAGGCAGCAAAGTATGGCGCACGCCAGGAGTCCGCGTGGGTGGTATC
CCGACCTATGGGGTGGCGCAGGGGTTTTCTGGTTTTCGCGTAGGT
GTAGGAGGTATACCGGCGTGGCCGGTGTACCAGGGTTGGTGGCGT
CCTGGTGTGGCGGTGTCAGGTTGGTATTTACCGGAAGCACAG
GCAGCAGCCGAGCTAAGCAGCGAAATATGGTGGCCGCGCGCAGGA
GTTTTAGGTGGGCTGGTTCCGGGCCCGCAGGCGAGTGTGCCGGGGTT
CCAGGCACCGGTGGTGTCCCTGGAGTCGGTACGCCGGCTGCAGCGGCA
GCCAAAGCGGCTGCAGAAAGCAGCACAGTTTGGCTTAGTACCGGGTGTG
GGAGTTGCCCGCGGTTGGCGTTGCTCCAGGGTGGGTGTGCTCCT
GGCGTCCGCTGGCTCCTGGAGTGGCGGTAGCACCCGGTGTGGGGGTG
GCCCCGGGTGTGGGGTGCACCGGATATCGGTCCGGCGGTGTGCGA
GCAGCAGCTAAAGCGCGGCGAAAGTTGCGGCCAAAGCCAACTGCGC
GCCGCCCGGGCTCGGTGCAGGATTCGGGGCTGGGTGTCCGAGTT
GGAGTCCCGGTTTTGGCGTGGCGCGGGAGTTCGGGACTGGGAGTG
GGTCCCGAGTTCCTGGCTTGGTGCAGGCGCAGATGAAGGTGTTTCGT
CGTAGCCTGAGTCCGGAACGCGTGAAGGTGATCCGAGTAGCAGCCAG
CATCTGCCGAGACCCCGAGCAGCCCGGCTTCCGGGTGCAtaa

[0195] The amino acid sequence of a 45.3 kDa human elastin truncated at the C-terminal is disclosed in SEQ ID NO: 39. The 45.3 kDa truncated elastin has amino acids 529-761 deleted from the full length elastin.

(SEQ ID NO: 39)

GGVPGAIPGGVPGGVFYPGAGLGLGGGALPGGKPLKVPVGGLAGAG
LGAGLGAFPAVTFPGALVPGGVADAAAAYKAAKAGAGLGGVPGVGGLG
VSAGAVVPQPGAGVKPGKVPVGLPGVYPGGVLPGARFPVGVLPVGP
TGAGVKPKAPGVGGAFAGIPGVGPFGGPQPGVPLGYPIKAPKLPGGYG
LPYTTGKLPYGYGPGGVAGAAGKAGYPTGTGVPQAAAAAATAKAAAF
GAGAAGVLPVGGAGVPGVPGAIPGIGGIAGVGTAAAAAATAKAA
KYGAAAGLVPGGPFGPGVGVPGAGVPGVPGAGIPVVPVPGAGIPGA
AVPGVVSPEAAAKAAAKAAKYGARPGVGVGPIPTYGVGAGGFPFGV
VGGIPGVAGVPGVGGVPGVGGVPGVVISPEAQAAAAAATAKAAAGAG
VLGGLVPGPQAAPVPGVGTGGVPGVGTAAAAAATAKAAQFGLVPGV
GVAPGVGAPVGVGAPVGLAPGVGAPGVGAPVGVGAPVGGV

[0196] The codon optimized polynucleotide sequence encoding the 45.3 kDa truncated human elastin is disclosed in SEQ ID NO: 40

(SEQ ID NO: 40)

GGTGGCGTACCAGGCGCAATTCTGGGGGTGTCCAGGCGGTGTTTTT
TATCCGGGCGCCGGTCTTGGCGCACTGGGTGGCGGTGCACTGGGCCCG
GGCGGCAAACCGCTGAAACCGGTACCAGGTGGTTTAGCAGGCGCCGGC
TTAGCGCAGGTCTGGGAGCATTTCGCGCAGTTACCTTCCAGGGGCA
CTGGTTCCTGGAGGTGTGGCCGATGCAGCCGCGGCATATAAAGCCGCT
AAAGCCGGTGCGGGTTTTAGGAGGCGTCCAGGTGTCGGTGGCCTGGGT
GTTAGCGCCGGTGCAGTTGTTCCGCGAGCCGGGAGCAGGGGTTAAACCT
GGTAAAGTGCCGGGAGTAGGTCTGCCAGGCGTTTATCCTGGTGGTGT
TTGCCGGGTGCCCGTTTTCCGGCGCTGGTGTCTTCCAGGCGTGCCG
ACCGGAGCCGGTGTAAACCGAAAGCCCGGTTGGAGGTGCATTT
GCAGGCATCCCGGAGTTGGCCCGTTTTGGTGGTCCGCAACCTGGGGTT
CCGTTAGGTTATCCGATTAAGCACCGAAACTGCCCGGCGGTTATGGT
CTGCCGTACACAACCGTAAACTGCCGTATGGTTATGGCCCGGTTGGA
GTTGCGGGTGCAGCAGGTAAAGCGGGTTATCCTACCGGAACCGGTGTA
GGTCCGAGGCGCTGCTGCCCGCGCAAAGCAGCGGCTAAATTT
GGCGCCGAGCAGCGGGTGTCTGCCCTGGAGTTGGTGGTGGCGGGCTG
CCAGGGGTACCTGGTGCATTCGGGTATTGGTGGTATTGCCGGTGTG
GGCACCCCGGCGCGGCGAGCTGCCGCGCGGCGCAAAGCAGCGGCTGCT
AAATACGGTGCAGCGCGGGTCTGGTGCAGGAGTCCGGGTTTTGGT
CCGGGAGTGGTTGGCGTGCTGGCGCAGGCGTTCCTGGTGTGGCGTT
CCAGGTGCAGGATTCCTGTTGTGCTGGTGGTGGTATTCCCGGCGC

[0200] The codon optimized polynucleotide sequence encoding the 40.4 kDa truncated human elastin is disclosed in SEQ ID NO: 44

(SEQ ID NO: 44)

GGTCTGGTCCAGGAGGTCGGGTTTGGTCCGGAGTGGTTGGCGTG
 CCTGGCGCAGGCGTTCCCTGGTGTGGGCGTTCCAGGTGCAGGGAATTCCT
 GTTGTGCCTGGTGCCTGATTCCCGCGCGCGCCTCCGGGGTGGTT
 AGCCCGGAAGCCCGCAGCAAGGCTCGGCAAGGCAGCAAGTATGGC
 GCACGCCAGGAGTCGGCGTGGTGGTATCCCGACCTATGGGGTGGG
 GCAGGGGGTTTCCCTGGTTCCGGCTAGGTGTAGGAGGTATACCGGG
 GTGGCCGGTGTACCAGGGTGGTGGCGTCCCTGGTGTGGCGGTGTG
 CCAGGTGTGGTATTTCCCGGAAGCACAGGCAGCAGCCGACGTAAG
 GCAGCGAAATATGGTGCCTGGCGCGCAGGAGTTTGGTGGGCTGGTT
 CCGGGCCCGCAGGCAGCTGTGCCGGGGTCCAGGCACCGTGGTGTG
 CCTGGAGTCGTTACGCCGGTGCAGCGGCAGCAAGCGGCTGCGAAA
 GCAGCACAGTTTGGCTTAGTACCGGGTGTGGAGTTGCCCGCGCGTT
 GGCGTTGCTCCAGGGTGGTGTGCTCCTGGCGTGGTCTGGCTCCT
 GGAGTGGGCGTAGCACCGGTGTGGGGTGGCCCGGGTGTGGGGTT
 GCACCGGGTATCGTCCGGGCGGTGCGCAGCAGCAGTAAAGCGCG
 GCGAAAGTTGCGGCCAAAGCCCACTGCGCGCCCGCGGGCTCGGT
 GCAGGTATCCGGGCTGGGTGTCGGAGTTGGAGTCCCGGGTTTGGG
 GTGGCGCGGGAGTTCCGGGACTGGGAGTGGGTGCGGAGTTCTGGC
 TTTGGTGCAGGCGCAGATGAAGGTTCGTGCTAGCCTGAGTCCGGAA
 CTGCGTGAAGGTATCCAGTAGCAGCCAGCATCTGCCGAGCACCCCG
 AGCAGCCCGCGTGTCCGGGTGCATTAGCTGCAGCAAAAGCCGCAAG
 TATGGTGCAGCCGTGCCGGGCTTTAGGTGGTCTGGGCGCCCTGGGT
 GGTGTAGGCATCCCGGAGGTGTGTGGGTGCAGGACCGGCGCCGCA
 GCTGCGGCCGCAAAAGCAGCTGCAAAAGCGGCCAGTTGGTTTAGTG
 GGCGCCGAGGTTTAGCGGTTTAGGTGTGGGTGGACTGGGTGTACCT
 GGCGTAGGCGGTCTGGGTGGAATCCGCCCCGAGCGGCGCGAAAGCG
 GCAAAATATGGCGCGCAGGCTGGGCGGCGTGTGGTGGGGCAGGT
 CAGTTTCCGCTGGGCGGGTTGCCGACGTCCGGATTGGTCTGAGC
 CCGATTTCCCTGGCGGCGCATGTCTGGGTAAAGCATGTGGTCTGATAA
 CGTAAAtaa

[0201] The amino acid sequence of a 39.8 kDa human elastin truncated at the C-terminal is disclosed in SEQ ID NO: 45. The 39.8 kDa truncated elastin has amino acids 462-761 deleted from the full length elastin.

(SEQ ID NO: 45)

GGVPGAIPGGVPGVVFYFAGLGLGGLGPGKPLKVPVGGLAGAG
 LGAGLGAFPAVTFPGALVPGGVADAAAAYKAAKAGAGLGGVPGVGGGLG

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VSAGAVVPQPGAGVKPGKVPVGLPGVYPGGVLPGARFPVGVLPVGV
 TGAGVKPKAPGVGGAFAGIPGVPPFGPQPGVPLGYPIKAPKLPGGYG
 LPYTTGKLPYGYPGGVAGAAGKAGYPTGTGVGPQAAAAAATAAAAF
 GAGAAGVLPVGGAGVPGVPGAIPGIIGIAGVGTAAAAAATAAAAF
 KYGAAAGLVPGGPGFPGVGVVPGAGVPGVPGAGIPVVPAGIPGA
 AVPGVVSPEAAAKAAKAAKYGARPGVGVGGIPTYGVGAGGFPFGVGV
 VGGIPGVAGVPGVGGVPGVGGVPGVGISPEAQAAAAAATAAAKAAAGAG
 VLGGLVPGPQAAPVPGVPGTGGVPGVGTG

[0202] The codon optimized polynucleotide sequence encoding the 39.8 kDa truncated human elastin is disclosed in SEQ ID NO: 46

(SEQ ID NO: 46)

GGTGGCGTACCAGGCGCAATTCCTGGGGTGTCCAGGCGGTGTTTTT
 TATCCGGGCGCGGTCTTGGCGCACTGGTGGCGGTGCACTGGGCCCC
 GGGCGCAAACCGTGAACCGGTACCAGTGGTGTAGCAGGCGCCCGC
 TTAGCGCAGGTCTGGGAGCATTTCCGCGAGTTACCTTTCCAGGGCA
 CTGGTTCCTGGAGGTGTGGCCGATGCAGCCCGGCATATAAAGCCGT
 AAAGCCGTTGCGGGTTTAGGAGCGTCCAGGTGTGGTGGCTGGGT
 GTTAGCGCCGGTGCAGTTGTTCCGCGAGCCGGAGCAGGGTTAAACCT
 GGTAAAGTGCCTGGGAGTAGGTCTGCCAGGCGTTTATCCTGGTGGTGT
 TTGCGGGTGCCTGTTTTCCGGGCGTGGTGTCTTCCAGGCGTGGCG
 ACCGGAGCCGGTGTAAACCGAAAGCCCGGTGTGGAGGTGCATTT
 GCAGGCATCCCGGGAGTTGGCCCGTTTGGTGGTCCGCAACCTGGGGT
 CCGTTAGGTTATCCGATTAAGCACCGAAACTGCCCGGCGTTATGGT
 CTGCCGTACACAACCGGTAAACTGCCGTATGGTTATGGCCCGGTGGA
 GTTGGGGTGCAGCAGGTAAAGCGGTTATCCTACCGGAACCGGTGTA
 GGTCCGCGAGCCGCTGCTGCCCGCCGCAAAAGCAGCGGCTAAATTT
 GGCGCCGAGCAGCGGGTCTGCTGCTGGAGTTGGTGGTGCAGGCGGT
 CCAGGGGTACTGGTGAATTCGGGTATTGGTGGTATTGCCGGTGTG
 GGCACCCCGGCGCGGCGAGCTGCCGCGAGCGGCGGTGCCAAAGCTGCT
 AAATACCGTGCCTGGCGGGGTCTGGTGCAGGAGTCCGGGTTTGGT
 CCGGGAGTGGTGGCGTGCCTGGCGCAGGCGTTCCTGGTGTGGCGGTT
 CCAGGTGCAGGATTCTGTTGTGCTGGTGGCGGTTATCCCGGCGCG
 GCCGTTCCGGGGTGGTTAGCCCGGAAGCCGAGCGAAGGCTGCCGCA
 AAGGCGCAAAAGTATGGCGCACGCCAGGAGTCCGGTGGGTGGTATC
 CCGACCTATGGGGTGGCGCAGGGGTTTTCTGGTTTTCCGGCTAGGT
 GTAGGAGGTATACCGGGCGTGGCCGGTGTACCAGGGTGGTGGCGT
 CCTGGTGTGGCGGTGTGCCAGGTGTGGTATTTCCCGGAAGCACAG
 GCAGCAGCCGCGAGTAAAGCAGCGAAATATGGTGCCTGGCGCGCAGGA

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GTTTTAGGTGGGCTGGTTCCGGGCCCGCAGGCAGCTGTGCCGGGGTT
CCAGGCACCGGTGGTGTCCCTGGAGTCGGTACGCCGtaa

[0203] The amino acid sequence of a 36.1 kDa human elastin truncated at the C-terminal is disclosed in SEQ ID NO: 47. The 36.1 kDa truncated elastin has amino acids 418-761 deleted from the full length elastin.

(SEQ ID NO: 47)

GGVPGAIPGGVPGGVFYPGAGLGLGGGALPGGKPLKVPVPGGLAGAG
LGAGLGAFFAVTFPGALVPGGVADAAAAAYKAAKAGAGLGGVPGVGGGLG
VSAGAVVPQPGAGVKPGKVPVGLPGVYPGGVLPGARFPVGVLPVGP
TGAGVKPKAPGVGGAFAGIPGVGPFPGPQGVPLGYPIKAPKLPGGYG
LPYTTGKLPYGYGPGGVAGAAGKAGYPTGTGVGPQAAAAAAKAAAFK
GAGAAGVLPVGGAGVPGVPGAIPGIGGIAGVGTAAAAAAKAAKAA
KYGAAAGLVPGGPGFPGVGVPGAGVPGVPGAGIPVVPAGIPGA
AVPGVVSPEAAKAAKAAKYGARPGVGVGGIPTYGVGAGGFPFGVGV
VGGIPGVAGVPGVGVPGVGGVPGVGVISPEAQ

[0204] The codon optimized polynucleotide sequence encoding the 36.1 kDa truncated human elastin is disclosed in SEQ ID NO: 48

(SEQ ID NO: 48)

GGTGGCGTACCAGGCGCAATTCCTGGGGGTGTCCAGGCGGTGTTTTT
TATCCGGGCGCCGGTCTTGGCGCACTGGGTGGCGGTGCACTGGGCCCC
GGCGGCAAACCGCTGAAACCGGTACAGGTGGTTTAGCAGGCGCCGGC
TTAGGCGCAGGTCTGGGAGCATTCCCGGCAGTTACCTTCCAGGGGCA
CTGGTTCCTGGAGGTGTGGCCGATGCAGCCGCGCATATAAAGCCGCT
AAAGCCGGTGCGGGTTTAGGAGGCGTCCAGGTGTCGGTGGCCTGGGT
GTTAGCGCCGGTGCAGTTGTTCCGAGCCGGGAGCAGGGGTTAAACCT
GGTAAAGTGCCGGGAGTAGGTCTGCCAGGCGTTTATCCTGTGTGTGT
TTGCCGGGTGCCCGTTTTCCGGCGTGTGGTCTTCCAGGCGTGCCG
ACCGGAGCCGGTGTAAACCGAAAGCCCCGGTGTGGAGGTGCATTT
GCAGGCATCCCGGAGTTGGCCCGTTTGGTGGTCCGCAACCTGGGGTT
CCGTTAGGTTATCCGATTAAGCACCGAAACTGCCCGCGGTTATGGT
CTGCCGTACACAACCGGTAAACTGCCGTATGGTTATGGCCCGGGTGG
GTTGCGGGTGCAGCAGGTAAGCGGGTTATCCTACCGGAACCGGTGTA
GGTCCGCAAGCCGCTGCTGCCCGCCGCAAAAGCAGCGGCTAAATTT
GGCGCCGGAGCAGCGGGTGTCTGCCCTGGAGTTGGTGGTGGCGGGCGTG
CCAGGGTACCTGGTGCAATCCGGGTATTGGTGGTATTGCCGGTGTG
GGCACCCCGGCGCGGAGCTGCCGAGCGGCGGCTGCCAAAGCTGCT
AAATACGGTGCCCGCGGGTCTGGTGCAGGAGTCCGGGTTTTGGT
CCGGGAGTGGTTGGCGTCTGCCGAGCGGCTTCTGTGTGGCGGT

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CCAGGTGCAGGGATTCTGTTGTGCCTGGTGCCGGTATCCCGGCGCG
GCCGTTCCGGGGGTGGTTAGCCCGGAAGCCGAGCGAAGGCTGCGGCA
AAGGCAGCAAGTATGGCGCACGCCAGGAGTCGGCGTGGGTGGTATC
CCGACCTATGGGGTGGGCGCAGGGGGTTTTCTGGTTTCGCGTAGGT
GTAGGAGGTATACCGGGCGTGGCCGGTGTACCAGGGGTTGGTGGCGTC
CCTGGTGTGGCGGTGTGCCAGGTGTTGGTATTTACCAGGAAGCACAG
taa

[0205] The amino acid sequence of a 34.9 kDa human elastin truncated at the N-terminal is disclosed in SEQ ID NO: 49. The 34.9 kDa truncated elastin has amino acids 2-360 deleted from the full length elastin.

(SEQ ID NO: 49)

RPVGVGGIPTYGVGAGFPFGVGVGGIPGVAGVPGVGGVPGVGGVPGV
GISPEAQAAAAKAAKYGAAGAGVLGGLVPGVQAAPVPGTGGVPGVGT
PAAAAKAAKAAQFGLVPGVGVAPGVGVPVPGVGLAPGVGVAPG
VGVAPGVGVPVGIIPGGVAAAASAAKVAQAQLRAAAGLGAIPGLGVG
VGVPLGVGAGVPLGVGAGVPGFAGADEGVRRLSPELREGDPSSSQH
LPSTPSSPRVPGALAAKAAKYGAAPVGLGGLGAGVPIPGVGGVAGP
AAAAAAKAAKAAQFGLVGAAGLGLGVGGLVPGVGGVGGIPAAAAAK
AAKYGAAGLGGVGGAGQFPLGGVAARPGFGLSPIFPGGACLGKACGRKR
K

[0206] The codon optimized polynucleotide sequence encoding the 34.9 kDa truncated human elastin is disclosed in SEQ ID NO: 50

(SEQ ID NO: 50)

CGCCAGGAGTCGGCGTGGGTGGTATCCCGACCTATGGGGTGGGCGCAGG
GGGTTTTCTGGTTTCGGCGTAGGTGTAGGAGGTATACCGGGCTGGCCG
GTGTACCAGGGTTGGTGGCGTCCCTGGTGTGGCGGTGTGCCAGGTGTT
GGTATTTACCAGGAGCACAGGCAGCAGCCGAGCTAAGGCAGCGAAATA
TGGTGCCCGCGCAGGAGTTTTAGGTGGGCTGGTCCCGGGCCCGCAGG
CAGCTGTGCCGGGGTTCAGGCACCGGTGGTGTCCCTGGAGTCGGTACG
CCGGCTGCAGCGGCAGCCAAAGCGGCTGCGAAAGCAGCACAGTTGGCTT
AGTACCGGGTGTGGAGTTGCCCGCGGCTGGCGTTGCTCCAGGGTGG
GTGTGCTCCTGGCGTCCGCTGGCTCCTGGAGTGGGCGTAGCACCCGGT
GTGGGGTGGCCCCGGGTGGGGTTGCGCCGGTATCGGTCCGGGCGG
TGTCGCAGCAGCAGCTAAAGCGCGGCAAGTTGCGGCCAAAGCCCAAC
TGCGCGCCCGCGGGCTCGGTGCAGGTATTCGGGGCTGGGTGTCCGGA
GTTGGAGTCCCGGGTTGGCGTGGGCGCGGAGTTCCGGGACTGGGAGT
GGGTGCCGGAGTCTTGGCTTTGGTGCAGGCGCAGATGAAGGTGTTCTGTC
GTAGCCTGAGTCCGGAAGTGCAGGTGATCCGAGTAGCAGCCAGCAT

-continued

CTGCCGAGCACCCCGAGCAGCCCGCGTGTCCGGGTGCATTAGCTGCAGC
AAAAGCCGCCAAGTATGGTGCAGCCGTGCCGGCGTCTTAGGTGGTCTGG
GCCCCCTGGGTGGTGTAGGCATTCCGGGAGGTGTTGTGGGTGCAGGACCG
GCCGCCGAGCTGCCGCCGCCAAAGCAGCTGCAAAGCGGCCAGTTTGG
TTTAGTGGGCGCCGAGGTTTAGGCGGTTTAGGTGTGGGTGGACTGGGTG
TACCTGGCGTAGGCGGTCTGGGTGGAATTCGCCCGCAGCGGCCGCGAAA
CCGGCAAATATGGCGCGCAGGCCCTGGCGCGCTGCTGGGTGGGCGAGG
TCAGTTTCCGCTGGGCGGGGTTGCCGCACGTCGGGATTTGGTCTGAGCC
CGATTTTCCCTGGCGCGCATGTCTGGGTAAGCATGTGGTCTGTAACGT
AAAt aa

[0207] The amino acid sequence of a 32 kDa human elastin truncated at the C-terminal is disclosed in SEQ ID

NO: 51. The 32 kDa truncated elastin has amino acids 373-761 deleted from the full length elastin.

(SEQ ID NO: 51)
GGVPGAIPGGVPGGVFYPGAGLGLGGGALPGGKPLKPVPGGLAGAGLGA
GLGAFPAVTFPGALVPGGVADAAAAYKAAKAGAGLGGVPGVGGGLGVSAGAV
VPQPAGVKGKVPVGLPGVYPGGVLPGARFPGVGLPGVPTGAGVKPKA
PGVGGAFAGIPGVGPFGGPQPGVPLGYPIKAPKLPGGYGLPYTTGKLPYGY
GPGGVAGAAGKAGYPTGTGVPQAAAAAAKAAAKFGAGAAGVLPVGGGAG
VPGVPGAIPGIGGIAGVGTAAAAAAKAAKAAKYGAAAGLVPGGPGFPGG
VVGVPGAGVPGVPGAGIPVVPGAGIPGAAPGVVSPAAAAKAAKAAKY
GARPGVGVGGIPTY

[0208] The codon optimized polynucleotide sequence encoding the 32 kDa truncated human elastin is disclosed in SEQ ID NO: 52

(SEQ ID NO: 52)
GGTGGCGTACCAGGCGCAATTCCTGGGGGTGTCCAGGCGGTGTTTTTATCCGG
GCCCGGTCTTGGCGCACTGGGTGGCGGTGCACTGGGCCGGGCGCAAACCGC
TGAAACCGGTACCAGGTGGTTTAGCAGGCGCCGGCTTAGCGCAGGTCTGGGAG
CATTTCGGCAGTTACCTTCCAGGGGCACTGGTTCCTGGAGGTGTGGCCGATGC
AGCCCGGCATATAAAGCCGCTAAAGCCGGTGCGGGTTTAGGAGGCGTCCAGG
TGTCCGTGGCCTGGGTGTTAGCGCCGGTGCAGTTGTTCCGCAGCCGGGAGCAGGG
GTTAAACCTGGTAAAGTGCCGGGAGTAGGTCTGCCAGGCGTTTATCTGGTGGTG
TTTTTCCGGGTGCCCGTTTTCCGGCGTTGGTGTCTTCCAGGCGTCCCGACCGGA
GCCGGTGTAAACCGAAAGCCCCGGTGTGGAGGTGCATTTGCAGGCATCCCGG
GAGTTGGCCCCGTTTGGTGGTCCGCAACCTGGGGTTCGGTTAGGTTATCCGATTAA
AGCACCGAAACTGCCCGCGGTTATGGTCTGCCGTACACAACCGGTAAACTGCCG
TATGGTTATGGCCCGGTGGAGTTGCGGGTGCAGCAGGTAAGCGGGTTATCCTA
CCGGAACCGGTGTAGTCCGCAGGCCGCTGCTGCCCGCCGCAAAGCAGCCGG
CTAAATTTGGCGCGGAGCAGCGGTGTTCTGCCTGGAGTTGGTGGTGCGGGCGT
GCCAGGGTACCTGGTGAATTCGGGTATTGGTGGTATTGCCGGTGTCCGCACC
CCGGCCCGCAGCTGCCGCAGCGCGGCTGCCAAAGCTGCTAAATACGGTGC
CGGGCGGTCTGGTGCAGGAGTCCGGGTTTGGTCCGGGAGTGGTTGGCGTG
CTGGCGAGGCGTTCCTGGTGTGGGCGTTCAGGTGCAGGATTCCTGTTGTGCC
TGGTGCCTGATTCCCGCGCGCGGTTCCGGGGTGGTTAGCCGGAAGCCGCA
GCCAAGGCTGCCGCAAAGGCAGCAAAGTATGGCGCACGCCAGGAGTCCGGCGTG
GGTGGTATCCCGACCTATt aa

[0209] The amino acid sequence of a 29.9 kDa human elastin truncated at the C-terminal is disclosed in SEQ ID NO: 53. The 60.7 kDa truncated elastin has amino acids 347-761 deleted from the full length elastin.

(SEQ ID NO: 53)

GGVPGAIPGGVPGGVFYPGAGLGGALGPGGKPLKVPVGGLAGAGLG
 AGLGAFPAVTFPGALVPGGVADAAAAYKAAKAGAGLGGVPGVGGVGSAG
 AVVPQPGAGVKPGKVPVGLPGVYVPGVLPGARFPVGVLPVPTGAGVK
 PKAPGVGGAFAGIPGVGPFGGPQVGVPLGYPIKAPKLPGGYGLPYTTGKL
 PYGYPGGVGAGAAGKAGYPTGTGVGPQAAAAAAKAAKFGAGAAGVLPG
 VGGAGVPGVPGAIPGIGGIAGVTPAAAAAAKAAKYGAAAGLVPGG
 PGFPGVVGVPVAGVPGVPGAGIPVVPVAGIPGAAVPGVVSPE

[0210] The codon optimized polynucleotide sequence encoding the 29.9 kDa truncated human elastin is disclosed in SEQ ID NO: 54

(SEQ ID NO: 54)

GGTGGCGTACCAGGCGCAATTCCTGGGGTGTCCAGGCGGTGTTTTTATCCGG
 GCGCCGGTCTTGGCGCACTGGTGGCGGTGCACTGGGCCGGCGCAAACCGC
 TGAAACCGGTACCAGGTGGTTTAGCAGGCGCCGGCTTAGGCGCAGGTCTGGGAG
 CATTTCGGCAGTTACCTTTCAGGGGCACTGGTTCCTGGAGGTGTGGCCGATGC
 AGCCCGGCATATAAAGCCGCTAAAGCCGGTGCAGGTTTAGGAGCGTCCCAGG
 TGTCCGGTGGCCTGGGTGTAGCAGCGGTGCAAGTTGTTCCGACCCGGAGCAGGG
 GTTAAACCTGGTAAAGTGCAGGAGTAGGTCTGCCAGGCGTTTATCCTGGTGGTG
 TTTTCCGGGTGCCCGTTTCCGGGCGTTGGTGTCTTCCAGGCGTGCCGACCGGA
 GCGGTGTGTTAAACGAAAGCCCGGTGTGGAGGTGCATTTGCAGGCATCCCGG
 GAGTTGGCCCGTTTGGTGGTCCGCAACCTGGGGTTCCTTAGGTTATCCGATTAA
 AGCACCGAAACTGCCCGCGGTTATGGTCTGCCGTACACAACCGGTAAACTGCCG
 TATGGTTATGGCCCGGTGGAGTTGCGGGTGCAGCAGGTAAAGCGGTTATCCTA
 CCGGAACCGGTGTAGGTCCGACAGCCGCTGCTGCCGCCCGCAAAGCAGCGG
 CTAATTTGGCCCGGAGCAGCGGTGTTCTGCCGAGTTGGTGGTGGCGGCGT
 GCCAGGGGTACCTGGTGAATTCGGGTATTGGTGGTATTGCCGGTGTCCGCACC
 CCGGCCCGGCAGCTGCCGAGCGGCGGCTGCCAAAGCTGCTAAATACGGTGCC
 GCGGCGGGTCTGGTGCCAGGAGGTCCGGGTTTGGTCCGGGAGTGGTTGGCGTGC
 CTGGCGCAGGCGTTCTGGTGTGGGCGTTCCAGGTGCAGGATTCTGTGTGCC
 TGGTCCCGGTATTCGGCGCGCCGTTCCGGGGTGGTTAGCCCGAAtaa

[0211] The amino acid sequence of a 29.4 kDa human elastin truncated at the N-terminal is disclosed in SEQ ID NO: 55. The 29.4 kDa truncated elastin has amino acids 2-425 deleted from the full length elastin.

(SEQ ID NO: 55)

KYGAAGAGVLGGLVPGPQAAVPGVPGTGGVPGVGTAAAAAKAAKAAQF
 GLVPGVGVAPGVGVAPGVGLAPGVGVAPGVGVAPGVGVAPGIGP
 GGVAATAAKSAKVAQAQLRAAAGLGGAGIPGLGVGVGVPGLGVGAGVPLG
 VVGAGVPGFAGADEGVRRLSPELREGDPSSQHLPTSPSPRVPGALA
 AAKAAKYGAAVPGVGLGGLGALGGVGI PGGVVGAGPAAAAAAKAAKAAQ
 FGLVGAAGLGGVGGVPGVGGVGGI PPAATAAKKYGAAGLGGVGG
 AGQFPLGGVAARPGFGLSP IFFGGACLKACGRKRK

[0212] The codon optimized polynucleotide sequence encoding the 29.4 kDa truncated human elastin is disclosed in SEQ ID NO: 56

(SEQ ID NO: 56)

AAATATGGTGCCCGCGCAGGAGTTTTAGGTGGGCTGGTTCCGGGCCCGCAG
 GCAGCTGTGCCGGGGTTCCAGGCACCGGTGGTGTCCCTGGAGTCGGTACGCCGG
 CTGCAGCGGCAGCCAAAGCGGCTGCGAAAGCAGCACAGTTTGGCTTAGTACCGG
 GTGTGGGAGTTGCCCGCGGTTGGCGTTGCTCCAGGGTGGGTGTTGCTCCTGG
 CGTCGGTCTGGCTCCTGGAGTGGGCGTAGCACCCGGTGTGGGGTGGCCCCGGT
 GTTGGGGTGCACCGGTATCGGTCCGGCGGTGTGCGAGCAGCAGCTAAAAGC
 GCGCGAAAAGTTGCGGCCAAAGCCAACTGCGCGCCGCGGGGCTCGGTGCA
 GGTATCCCGGGCTGGGTGTCGGAGTTGGAGTCCCGGGTTTGGCGTGGCGCGG
 GAGTTCGGGACTGGGAGTGGGTGCCGGAGTTCCTGGCTTTGGTGCAGGCGCAG
 ATGAAGGTGTTCTCGTAGCCTGAGTCCGGAAGTGCAGTGAAGGTGATCCGAGTAG
 CAGCCAGCATCTGCCGAGCACCCGAGCAGCCCGCGTGTCCGGGTGCATTAGCT
 GCAGCAAAGCCGCAAGTATGGTGCAGCCGTGCCGGCGTCTTAGGTGGTCTG
 GCGGCCCTGGGTGGTGTAGGCATTCCGGGAGGTGTTGTGGGTGCAGGACCGGCC
 GCCGCGAGTGCAGCCGCAAGCAGCTGCAAAGCGGCCAGTTTGGTTTAGTG
 GCGCGCCGAGTTTAGGCGTTTAGGTGTGGGTGGACTGGGTGTACCTGGCGTAG
 GCGGTCTGGGTGGAATCCGCCCGCAGCGCCGCAAAGCGGCAAAAATAGCGG
 CGGCAGGCCTGGGCGCGTGTGGGTGGGGCAGGTCAGTTCCCGTGGGCGGGG
 TTGCCGACGTCGGGATTTGGTCTGAGCCGATTTCCCTGGCGCGCATGTCTG
 GGTAAAGCATGTGGTCGTAAACGTAAAtaa

[0213] The amino acid sequence of a 25.3 kDa human elastin truncated at the N-terminal is disclosed in SEQ ID NO: 57. The 25.3 kDa truncated elastin has amino acids 2-473 deleted from the full length elastin.

(SEQ ID NO: 57)

QFGLVPGVGVAPGVVAPGVVAPGVVAPGVVAPGVVAPGVVAPGVVAPGVVAPGI
 GPGGVAAAASAAKVAQAQLRAAAGLGGIPGLGVGVVPLGLGVAGVVP

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GLGVGAGVPGFAGADEGVRRSLSPELREGDPSSSQHLPSTPSSPRVPGA
 LAAAKAAKYGAAVPGVLGGLGALGGVGI PGGVVGAGPAAAAAKAAAKA
 AQPFLVGAAGLGGLVGGLVPGVGGGLGIPAAAAAKAAKYGAAGLGGVL
 GGAGQFPLGGVAARPGFGLSPIFPGGACLKAKCRKRK

[0214] The codon optimized polynucleotide sequence encoding the 25.3 kDa truncated human elastin is disclosed in SEQ ID NO: 58

(SEQ ID NO: 58)

CAGTTTGGCTTAGTACCGGGTGTGGGAGTTGCCCGCGGCTTGGCGTTGCTCCAG
 GGGTGGGTGTGCTCCTGGCGTCGGTCTGGCTCCTGGAGTGGGCGTAGCACCCGG
 TGTGGGGTGGCCCCGGTGTGGGGTGCACCGGTATCGGTCCGGCGGTGTG
 GCAGCAGCAGCTAAAAGCGCGCGAAAAGTTGCGGCCAAAGCCAACTGCGCGCC
 GCCGCGGGCCTCGGTGCAGGTATTCCGGGGCTGGGTGTCGGAGTTGGAGTCCCGG
 GTTTGGGCGTGGGCGCGGAGTTCCGGGACTGGGAGTGGGTGCCGGAGTTCCTG
 GCTTGGTGCAGGCGCAGATGAAGGTGTTCTCGTAGCCTGAGTCCGGAAGTCCG
 TGAAGGTGATCCGAGTAGCAGCCAGCATCTGCCGAGCACCCGAGCAGCCCGG
 GTTCCGGGTGCATTAGCTGCAGCAAAGCCGCAAGTATGGTGCAGCCGTGCCG
 GCGCTTAGGTGGTCTGGGCGCCCTGGGTGGTGTAGGCATTCCGGGAGGTGTTG
 TGGTGCAGGACCGCCGCGCAGCTGCGGCCGCAAAGCAGCTGCAAAGCGG

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CCCAGTTTGGTTTAGTGGGCGCCGAGGTTTAGGCGGTTAGGTGTGGGTGGACT
 GGGTGTACTCTGGCGTAGGCGGTCTGGGTGGAATCCGCCCGCAGCGGCCGCGAA
 AGCGGCAAAATATGGCGCGGCAGGCCTGGGCGCGTGTGGTGGGGCAGGTCA
 GTTTCCGCTGGGCGGGTTGCCGCACGTCCGGATTGGTCTGAGCCCGATTTTC
 CCTGGCGGCGCATGTCTGGGTAAAGCATGTGGTCTGTAACGTAATaa

[0215] The amino acid sequence of a 24.1 kDa human elastin truncated at the C-terminal is disclosed in SEQ ID NO: 59. The 24.1 kDa truncated elastin has amino acids 277-761 deleted from the full length elastin.

(SEQ ID NO: 59)

GGVPGAIPGGVPGGVFYPGAGLGGALGPGGKPLKPVPGGLAGAGLG
 AGLGAFPAVTFPGALVPGGVADAAAAYKAAKAGAGLGGVPGVGGGLGVSAG
 AVVPQPGAGVKPGKVPVGLPGVYVPGVLPGARFPVGVLPVPTGAGVK
 PKAPGVGAFAGIPGVGPFGGPQPGVPLGYPIKAPKLPGGYGLPYTTGKL
 PYGYPGGVVAGAAGKAGYPTGTGVPQAAAAAAKAAKFGAGAAGVLPG
 VGGAGVPGVPGAIPGIGGIAGVGTG

[0216] The codon optimized polynucleotide sequence encoding the 24.1 kDa truncated human elastin is disclosed in SEQ ID NO: 60

(SEQ ID NO: 60)

GGTGGCGTACCAGGCGCAATCCTGGGGTGTCCAGGCGGTGTTTTTATCCGG
 GCGCCGCTCTGGCGCACTGGGTGGCGGTGCACTGGGCCCGGGCGCAACCGC
 TGAACCGGTACCAGGTGGTTTAGCAGGCGCCGGCTTAGGCGCAGGTCTGGGAG
 CATTTCGGCAGTTACCTTCCAGGGGCACTGGTTCCTGGAGGTGTGGCCGATGC
 AGCCCGGCATATAAAGCCGCTAAAGCCGTTGCGGGTTTAGGAGCGTCCCAGG
 TGTCCGGTGGCTGGGTGTAGCGCCGGTGCAGTTGTTCCGCAGCCGGGAGCAGGG
 GTTAAACCTGGTAAAGTGCCGGGAGTAGGTCTGCCAGGCGTTTATCCTGGTGGT
 TTTTCCGGGTGCCCGTTTTCCGGCGTTGGTGTCTTCCAGGCGTCCGACCGGA
 GCGGTGTAAACCGAAAGCCCGGTGTGGAGGTGCATTTGCAGGCATCCCGG
 GAGTTGGCCCGTTTTGGTGGTCCGCAACCTGGGGTTCGGTTAGGTTATCCGATTA
 AGCACCGAAACTGCCCGCGGTTATGGTCTGCCGTACACAACCGGTAAACTGCCG
 TATGGTTATGGCCCGGTGGAGTTGCGGGTGCAGCAGGTAAGCGGGTTATCCTA
 CCGGAACCGGTGTAGGTCCGCGAGCCGCTGCTGCCCGCCCGCAAAAGCAGCGG
 CTAAATTTGGCGCCGAGCAGCGGTGTTCTGCCTGGAGTTGGTGGTCCGGCGT
 GCCAGGGGTACTGGTGAATTCGGGTATTGGTGGTATTGCCGGTGTCCGCACC
 CCGTaa

[0217] The amino acid sequence of a 20.3 kDa human elastin truncated at the C-terminal is disclosed in SEQ ID NO: 61. The 20.3 kDa truncated elastin has amino acids 229-761 deleted from the full length elastin.

(SEQ ID NO: 61)

GGVPGAIPGGVPGGVFYPGAGLGGALGPGGKPLKPVPGGLAGAGLGA
 GLGAFPAVTFPGALVPGGVADAAAAYKAAKAGAGLGGVPGVGGGLGVSAGAV
 VPQPGAGVKPGKVPVGLPGVYVPGVLPGARFPVGVLPVPTGAGVKPKA
 PGVGGAFAGIPGVGPFGGPQPGVPLGYPIKAPKLPGGYGLPYTTGKLPYGY
 GPGGVAGAAGKAGYPTGTGVPQ

[0218] The codon optimized polynucleotide sequence encoding the 20.3 kDa truncated human elastin is disclosed in SEQ ID NO: 62

(SEQ ID NO: 62)

GGTGGCGTACCAGGCGCAATTCCTGGGGGTGCCAGGCGGTGTTTTTATCCGG
GCGCCGGTCTTGGCGCACTGGGTGGCGGTGCACTGGGCCCGGGCGGCAAACCGC
TGAAACCGGTACCAGGTGGTTTAGCAGGCGCCGGCTTAGGCGCAGGTCTGGGAG
CATTTCCGGCAGTTACCTTCCAGGGGCACTGGTTCCTGGAGGTGTGGCCGATGC
AGCCCGGCATATAAAGCCGCTAAAGCCGGTGCAGGGTTTAGGAGGCGTCCAGG
TGTGGTGGCCCTGGGTGTAGCGCCGGTGCAGTTGTTCGCGAGCCGGGAGCAGGG
GTAAACCTGGTAAAGTGCAGGGAGTAGGTCTGCCAGGCGTTTATCCTGGTGGTG
TTTTGCCGGGTGCCCGTTTCCGGGCGTTGGTGTCTTCCAGGCGTCCGACCCGGA
GCCGTGTAAACCGAAAGCCCCGGTGTGGAGGTGCATTTGCAGGCATCCCGG
GAGTTGGCCCGTTTGGTGGTCCGCAACCTGGGGTTCGGTTAGGTTATCCGATTAA
AGCACCGAAACTGCCCGGCGGTTATGGTCTGCCGTACACAACCGTAAACTGCCG
TATGGTTATGGCCCGGTGGAGTTGCGGGTGCAGCAGGTAAAGCGGGTTATCCTA
CCGGAACCGGTGAGGTCCGCAGt aa

[0219] The amino acid sequence of a 19.6 kDa human elastin truncated at the N-terminal is disclosed in SEQ ID NO: 63. The 19.6 kDa truncated elastin has amino acids 2-542 deleted from the full length elastin.

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CCGCTGGGCGGGTTGCCGCACGTCCGGGATTGGTCTGAGCCCATTTC
CCTGGCGCGCATGTCTGGGTAAAGCATGTGGTCGTAACGTAAAt aa

(SEQ ID NO: 63)

QLRAAAGLGAGIPLGLGVGVPLGLVGVAGVPLGLVGVAGVPGFAGADEGV
RRSLSPELREGDPSSSQHLPS TPSSPRVPGALAAKAAKYGA AVPGVLGG
LGALGGVGI PGGVVGAGPAAAAA KAAKAAQFGLVGAAGLGLGVGGL
GVPVGGGLGGIP PAAAAA KAAKYGAAGLGGV LGGAGQFPLGGVAARPGFGL
SPIFPGGA CLGKACGRKRK

[0220] The codon optimized polynucleotide sequence encoding the 19.6 kDa truncated human elastin is disclosed in SEQ ID NO: 64

(SEQ ID NO: 64)

CAACTGCGCGCCGCCCGGGCTCGGTGCAGTATCCGGGGCTGGGTGTC
GGAGTTGGAGTCCCGGTTTGGCGTGGCGCGGGAGTCCGGGACTGGGA
GTGGGTGCCGAGTTCCTGGCTTGGTGCAGCGCAGATGAAGGTGTTTCGT
CGTAGCCTGAGTCCGGAAGTGCAGGTGATCCGAGTAGCAGCCAGCAT
CTGCCAGCACCCGAGCAGCCCGGTGTTCCGGGTGCATTAGTGCAGCA
AAAGCCGCAAGTATGGTGCAGCCGTGCCGGCGTCTTAGTGGTCTGGGC
GCCCTGGGTGGTGTAGGCATCCGGGAGGTGTGTGGGTGCAGGACCGGCC
GCCGAGCTGCGGCCGCAAGCAGCTGCAAAAGCGGCCAGTTTGGTTTA
GTGGGCGCCGAGGTTTAGGCGGTTTAGGTGTGGGTGGACTGGGTGTACCT
GGCGTAGGCGGTCTGGGTGGAATCCGCCCGCAGCGGCCGCAAGCGGCA
AAATATGGCGCGCAGGCTGGGCGCGTGTGGTGGGCGAGTTCAGTTT

[0221] The amino acid sequence of a 11 kDa human elastin truncated at the N-terminal is disclosed in SEQ ID NO: 65. The 11 kDa truncated elastin has amino acids 2-635 deleted from the full length elastin.

(SEQ ID NO: 65)

VPGVLGGLGALGGVGI PGGVVGAGPAAAAA KAAKAAQFGLVGAAGLGG
LVGVLGVPVGGGLGGIP PAAAAA KAAKYGAAGLGGV LGGAGQFPLGGVAAR
PGFGLSPIFPGGA CLGKACGRKRK

[0222] The codon optimized polynucleotide sequence encoding the 11 kDa truncated human elastin is disclosed in SEQ ID NO: 66

(SEQ ID NO: 66)

GTGCCGGCGTCTTAGTGGTCTGGGCGCCCTGGTGGTGTAGGCATTCG
GGAGGTGTGTGGGTGCAGGACCGGCCCGCCGAGCTGCGGCCGCCAAAGCA
GCTGC AAAAGCGGCCAGTTTGGTTTAGTGGGCGCCGAGGTTTAGGCGGT
TTAGTGTGGGTGGACTGGGTGTACCTGGCGTAGGCGGTCTGGGTGGAAT
CCGCCCGCAGCGGCCGCAAGCGGCAAAATATGGCGCGGAGGCTGGGC
GGCGTGTGGGTGGGCGAGTTCAGTTCCCGTGGCGGGGTGCGCCAGCT
CCGGATTGGTCTGAGCCGATTTCCCTGGCGCGCATGTCTGGGTAA
GCATGTGGTCGTAACGTAAAt aa

[0223] The amino acid sequence of a 7.9 kDa human elastin truncated at the N-terminal is disclosed in SEQ ID NO: 67. The 7.9 kDa truncated elastin has amino acids 2-674 deleted from the full length elastin.

(SEQ ID NO: 67)
 QFGLVGAAGLGGGLGVGGLGVPGVGGGLGGIPPPAAAAKAAKYGAAGLGGVLGG
 AGQFPLGGVAARPGFGLSPIFPGGACLGKACGRKRK

[0224] The codon optimized polynucleotide sequence encoding the 7.9 kDa truncated human elastin is disclosed in SEQ ID NO: 68

(SEQ ID NO: 68)
 CAGTTTGGTTTAGTGGGCGCCGAGGTTAGGCGGTTAGGTGTGGTGGA
 CTGGTGTACCTGGCGTAGGCGGTCTGGGTGGAATCCGCCCGCAGCGGCC
 GCGAAAGCGGCAAAATATGGCGCGGCGAGGCTGGGCGGCGTCTGGGTGGG
 GCAGGTCAGTTTCCGCTGGGCGGGGTTGCCGCACGTCGGGATTTGGTCTG
 AGCCCGATTTTCCCTGGCGGCGCATGTCTGGGTAAAGCATGTGGTCGTAA
 CGTAAAtaa

[0225] The amino acid sequence of a 6.3 kDa human elastin truncated at the C-terminal is disclosed in SEQ ID NO: 69. The 6.3 kDa truncated elastin has amino acids 74-761 deleted from the full length elastin.

(SEQ ID NO: 69)
 GGVPGAIPGGVPGGVFYPGAGLGGALGPGGKLPVPGGLAGAGLGA
 GLGAFPAVTFPGALVPGGVAD

[0226] The codon optimized polynucleotide sequence encoding the 6.3 kDa truncated human elastin is disclosed in SEQ ID NO: 70:

(SEQ ID NO: 70)
 GGTGGCGTACCAGGCGCAATTCCTGGGGTGTCCAGGCGGTGTTTTTAT
 CCGGGCGCCGGTCTTGGCGCACTGGGTGGCGGTGCACTGGCCCGGGCGGC
 AAACCGCTGAAACCGGTACCAGGTGTTTAGCAGGCGCGGCTTAGGCGCA
 GGTCTGGGAGCATTTCGGCAGTTACCTTCCAGGGGCACTGTTCTGGA
 GGTGTGGCCGATaa

[0227] The amino acid sequence of a 4.3 kDa human elastin truncated at the N-terminal is disclosed in SEQ ID NO: 71. The 4.3 kDa truncated elastin has amino acids 2-717 deleted from the full length elastin.

(SEQ ID NO: 71)
 LGGVGLGGAGQFPLGGVAARPGFGLSPIFPGGACLGKACGRKRK

[0228] The codon optimized polynucleotide sequence encoding the 4.3 kDa truncated human elastin is disclosed in SEQ ID NO: 72

(SEQ ID NO: 72)
 GGCCCTGGGCGGCGTCTGGGTGGGCGAGTCAGTTTCCGCTGGGCGGGTT
 GCCGCACGTCGGGATTTGGTCTGAGCCGATTTTCCCTGGCGGCGCATGT
 CTGGGTAAAGCATGTGGTCGTAAACGTAAAtaa

Truncated Human Elastin 1 with DsbA Secretion and FLAG Tag

[0229] The amino acid sequence of truncated human elastin 1 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 98. The DsbA secretion tag is encoded by nucleotides 1-57 of SEQ ID NO: 99 and the amino acid sequences are amino acids 1-19 of SEQ ID NO: 98. The elastin nucleotide sequences are nucleotides 58-657 of SEQ ID NO: 99 and the amino acid sequences are amino acids 20-219 of SEQ ID NO: 98. The FLAG nucleotide sequences are nucleotides 658-684 of SEQ ID NO: 99 and the amino acid sequences are amino acids 220-228 of SEQ ID NO: 98.

(SEQ ID NO: 98)
 MKKIWLALAGLVLAFSASAGGVPGAIPGGVPGGVFYPGAGLGGALGGP
 GGKLPKVPGGLAGAGLGAFAVTFPGALVPGGVADAAAAYKAAKAG
 AGLGGVPGVGLGVSAGAVVPQPGAGVKPKVPGVGLPGVYPGGVLPGARF
 PGVGVLPGVPTGAGVKPKAPGVGGAFAGIPGVGPPGGPQPGVPLGYPKAP
 KLPGGYGLPYTTGKLDYKDDDDK

[0230] The nucleic acid sequence of truncated human elastin 1 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 99.

(SEQ ID NO: 99)
 ATGAAAAAGATTTGGCTGGCGCTGGCTGGTTAGTTTTAGCGTTTAGCGCA
 TCGGCGGGTGGCGTACCAGGCGCAATTCCTGGGGTGTCCAGGCGGTGTT
 TTTTATCCGGGCGCCGGTCTTGGCGCACTGGGTGGCGGTGCACTGGCCCG
 GGCGGCAAAACCGCTGAAACCGGTACCAGGTGTTTAGCAGGCGCGGCTTA
 GGCGCAGGTCTGGGAGCATTTCGGCAGTTACCTTCCAGGGGCACTGGTT
 CCTGGAGGTGTGGCCGATGCAGCCGCGGCATATAAAGCCGCTAAAGCCGGT
 GCGGGTTTAGGAGGCGTCCAGGTGTCCGTGGCCGTTGGTGTAGCGCGGT
 GCAGTTGTCCCGAGCCGGGAGCAGGGGTTAAACCTGGTAAAGTCCGGGA
 GTAGGTCTGCCAGGCGTTTATCCTGGTGGTGTGTTTCCGGGTGCCGTTTT
 CCGGGCGTGGTGTCTTCCAGGCGTGCAGCCGAGCGGCGGTGTTAAACCG
 AAAGCCCCCGGTGTTGGAGGTGCATTTGCAGGCATCCCGGGAGTTGGCCCG
 TTTGGTGGTCCGCAACCTGGGGTCCGTTAGGTATCCGATTAAGCACC
 AAACCTGCCCGGCGTTATGGTCTGCCGTACACAACCGGTAAACTGGGTGAC
 TACAAAGACGACGACGACAAAtaa

[0231] The polynucleotide of SEQ ID NO: 99 was subcloned into vector pET28a, expressed host *E. coli* cells and the truncated elastin was purified as described herein. The purified elastin produced a clear band on SDS-PAGE and an anti-FLAG western was observed at around 25 kilodaltons. There were no existing bands that appear at that location on the gel in the absence of expression of this protein.

Truncated Human Elastin 2 with DsbA Secretion and FLAG Tag

[0232] The amino acid sequence of truncated human elastin type 2 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 100. The DsbA secretion tag is encoded by nucleotides 1-57 of SEQ ID NO: 101 and the amino acid sequences are amino acids 1-19 of SEQ ID NO: 100. The

elastin nucleotide sequences are nucleotides 58-657 of SEQ ID NO: 101 and the amino acid sequences are amino acids 20-219 of SEQ ID NO: 100. The FLAG nucleotide sequences are nucleotides 658-684 of SEQ ID NO: 101 and the amino acid sequences are amino acids 220-228 of SEQ ID NO: 100.

(SEQ ID NO: 100)

MKKIWLALAGLVLAFSASAPYGYGPGGVAGAAAGKAGYPTGTGVGPQAAAAA
 AAKAAAKFGAGAAGVLPVGGAGVPGVPGAIPGIGGIAGVGTAAAAA
 AAKAAKYGAAAGLVPGGPGFPGVGVPGAGVPGVGVGAGIPVVPAGAGIP
 GAAVPGVVSPEAAAKAAAKAAYGARPGVGVGGIPTYGVGAGGFPFGVGV
 GGIPGVAGVPGVGGVGDYKDDDDK

[0233] The nucleic acid sequence of truncated human elastin 2 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 101.

(SEQ ID NO: 101)

ATGAAAAAGATTGGCTGGCGCTGGCTGGTTAGTTTTCAGCTTTAGCGCA
 TCGGCGCCGTATGGTTATGGCCCGGGTGGAGTTGCGGGTGCAGCAGGTAA
 GCGGGTTATCTACCGGAACCGGTGAGGTCCGCAGGCCGCTGCTGCCGCG
 GCCGCAAAGCAGCGGCTAAATTTGGCGCCGAGCAGCGGGTGTCTGCCT
 GGAGTTGGTGGTGCAGGGTGCAGGGGTACCTGGTGAATTCGGGTATT
 GGTGTATTGCCGGTGTCCGCCACCCCGGCCGCGCAGCTGCAGCAGCGGCG
 GCTGCCAAAGCTGCTAAATACGGTCCCGCGCGGGTCTGGTCCAGGAGGT
 CCGGGTTTTGGTCCGGGAGTGGTTGGCGTGCTGGCGCAGCGTTCCTGGT
 GTGGGCGTTCAGGTGCAGGATTCCTGTTGTGCTGGTCCGGTATTCCCG
 GCGCGCGCGTTCGGGGTGGTTAGCCCGGAAGCCGCAGCGAAGGCTGCG
 GCAAAGGCAGCAAAGTATGGCGCACGCCAGGAGTCGGCGTGGTGGTATC
 CCGACCTATGGGGTGGCGCAGGGGTTTTCTGGTTTTCGGCGTAGGTGTA
 GGAGGTATACCGGGCGTGCCGGTGTACCGGGGTTGGTGGCGTGGTGC
 TACAAAGACGACGACGACAAAaa

[0234] The polynucleotide of SEQ ID NO: 101 was subcloned into vector pET28a, expressed host *E. coli* cells and the truncated elastin was purified as described herein. The purified elastin produced a clear band on SDS-PAGE and an anti-FLAG western was observed at around 25 kilodaltons. There were no existing bands that appear at that location on the gel in the absence of expression of this protein.

Example 6: Effect of Truncated Collagen on Fibroblast Cell Viability, Procollagen Synthesis, and Elastin Synthesis

[0235] A human fibroblast cell culture was used to assess the ability of the truncated jellyfish collagen molecule of Example 2 to determine its effect on procollagen, and elastin synthesis. The human fibroblast cell culture was also used to determine the increased viability of the human fibroblast cells after exposure to the truncated jellyfish collagen.

[0236] A stock solution of 2% w/w truncated collagen was prepared from the histidine tagged truncated collagen of

example 3. Aliquots from the 2% stock truncated collagen solution were then used in the experiments described below.

Preparation of Fibroblasts

[0237] Fibroblasts were seeded into the individual wells of a 24-well plate in 0.5 ml of Fibroblast Growth Media (FGM) and incubated overnight at $37\pm 2^\circ\text{C}$. and $5\pm 1\%$ CO_2 . On the following day the media was removed via aspiration to eliminate any non-adherent cells and replaced with 0.5 ml of fresh FGM. The cells were grown until confluent, with a media change every 48 to 72 hours. Upon reaching confluency the cells were treated for 24 hours with DMEM supplemented with 1.5% FBS to wash out any effects from the growth factors included in the normal culture media. After the 24-hour wash out period the cells were treated with the truncated jellyfish collagen at specified concentrations dissolved in FGM with 1.5% FBS. Transforming Growth Factor Beta ($\text{TGF-}\beta$) (20 ng/ml) was used as a positive control for collagen and elastin synthesis. Untreated cells (negative controls) just received DMEM with 1.5% FBS. The cells were incubated for 48 hours and at the end of the incubation period cell culture medium was collected and either stored frozen (-75°C .) or assayed immediately. Materials were tested in triplicate.

MTT Assay

[0238] The MTT (3-(4,5-Dimethylthiazol-2-yl)-2,5-diphenyltetrazolium bromide, a tetrazole) assay is a colorimetric assay used to determine the metabolic activity of cells. Changes in cell number were assessed via an MTT assay. When cells are exposed to MTT, reduction of MTT by mitochondria in viable cells results in the formation of insoluble purple formazin crystals that are extracted from the cells with isopropanol and quantified spectrophotometrically. Non-living cells cannot reduce MTT and therefore cannot produce the purple formazin crystals. The intensity of the purple color is directly proportional to the number of living cells (metabolically active cells). The intensity of the purple color is directly proportional to the metabolic activity of the cells and is inversely proportional to the toxicity of the test material.

[0239] After the 2-day incubation discussed above, the cell culture medium was removed (see above) and the fibroblasts were washed twice with PBS to remove any remaining jellyfish glycogen molecules. After the final wash, 500 μl of DMEM supplemented with 0.5 mg/ml MTT was added to each well and the cells were incubated for 1 hour at $37\pm 2^\circ\text{C}$. and $5\pm 1\%$ CO_2 . After the incubation, the DMEM/MTT solution was removed and the cells were washed again once with PBS and then 0.5 ml of isopropyl alcohol was added to the well to extract the purple formazin crystals. Two hundred microliters of the isopropyl extracts was transferred to a 96-well plate and the plate was read at 540 nm using isopropyl alcohol as a blank.

[0240] The mean MTT absorbance value for the negative control cells was calculated and used to represent 100% cell viability. The individual MTT absorbance values from the cells undergoing the various treatments were then divided by the mean value for the negative control cells and expressed as a percent to determine the change in cell viability caused by each treatment.

[0241] In Tables 1, 2 and 3 of this example, the experiments were performed by using the designated aliquots of

the 2% stock truncated collagen solution in the assays. For example, in the samples that tested the “10% Collagen Solution,” an aliquot of the 2% truncated collagens in an amount sufficient to provide 10% of the assay volume was used. For a total assay volume of 1.0 ml, 100 μ l of the 2% stock truncated collagen solution was used. In Tables 1, 2 and 3, “10% Collagen Solution” is 0.2% collagen, “5% Collagen Solution” is 0.1% collagen, “1% Collagen Solution” is 0.02% collagen, “0.5% Collagen Solution” is 0.01% collagen, “0.1% Collagen Solution” is 0.002% collagen, “0.05% Collagen Solution” is 0.001% collagen, “0.01% Collagen Solution” is 0.0002% collagen, “0.005% Collagen Solution” is 0.0001% collagen, f

[0242] The results for the MTT assay are presented in Table 3. The values are presented as the mean percent viability \pm the deviation from the mean.

TABLE 3

MTT Assay	
Untreated	100 \pm 6.1
20 ng/ml TGF-B	110 \pm 2.9
10% Collagen Solution	131 \pm 7.8*
5% Collagen Solution	140 \pm 8.3*
1% Collagen Solution	116 \pm 0.9
0.5% Collagen Solution	105 \pm 4.8
0.1% Collagen Solution	102 \pm 1.1
0.05% Collagen Solution	106 \pm 3.4
0.01% Collagen Solution	112 \pm 1.9
0.005% Collagen Solution	103 \pm 3.9

*Denotes values that are significantly different from the Untreated group ($p < 0.05$).

[0243] The histidine tagged truncated jellyfish collagen showed protective effect by increasing the cell viability of human fibroblast cells. As can be seen in Table 3, the highest value in the MTT assay was observed when the fibroblast cells were exposed to 0.02% to 0.2% truncated jellyfish collagen.

Procollagen Synthesis

[0244] Fibroblasts are the main source of the extracellular matrix peptides, including the structural proteins collagen and elastin. Procollagen is a large peptide synthesized by fibroblasts in the dermal layer of the skin and is the precursor for collagen. As the peptide is processed to form a mature collagen protein, the propeptide portion is cleaved off (type I C-peptide). Both the mature collagen protein and the type I C-peptide fragment are then released into the extracellular environment. As collagen is synthesized the type I C-peptide fragment accumulates into the tissue culture medium. Since there is a 1:1 stoichiometric ratio between the two parts of the procollagen peptide, assaying for type I C-peptide will reflect the amount of collagen synthesized. Type I C-peptide can be assayed via an ELISA based method.

[0245] A series of type I C-peptide standards was prepared ranging from 0 ng/ml to 640 ng/ml. Next, an ELISA microplate was prepared by removing any unneeded strips from the plate frame followed by the addition of 100 μ l of peroxidase-labeled anti procollagen type I-C peptide antibody to each well used in the assay. Twenty (20) μ l of either sample (collected tissue culture media) or standard was then added to appropriate wells and the microplate was covered and allowed to incubate for 3 \pm 0.25 hours at 37° C. After the incubation the wells were aspirated and washed three times with 400 μ l of wash buffer. After the last wash was removed

100 μ l of peroxidase substrate solution (hydrogen peroxide+ tetramethylbenzidine as a chromagen) was added to each well and the plate was incubated for 15 \pm 5 minutes at room temperature. After the incubation 100 μ l of stop solution (1 N sulfuric acid) was added to each well and the plate was read using a microplate reader at 450 nm.

[0246] To quantify the amount of each substance present, a standard curve was generated using known concentrations of each substance. A regression analysis was performed to establish the line that best fits these data points. Absorbance values for the test materials and untreated samples were used to estimate the amount of each substance present in each sample.

[0247] The results for the ELISA assays are presented in Table 4.

TABLE 4

Type I Collagen Assay. The values presented are mean concentration (ng/ml) \pm the deviation from the mean.	
Treatment	Type I C-Peptide (ng/ml)
Untreated	1718 \pm 94
20 ng/ml TGF-B	3028 \pm 332*
10% Collagen Solution	1940 \pm 100
5% Collagen Solution	2394 \pm 125*
1% Collagen Solution	1773 \pm 183
0.5% Collagen Solution	1127 \pm 19*
0.1% Collagen Solution	1158 \pm 10*
0.05% Collagen Solution	1416 \pm 64
0.01% Collagen Solution	1835 \pm 404
0.005% Collagen Solution	1551 \pm 149

*Denotes values that are significantly different from the Untreated group ($p < 0.05$).

[0248] The truncated histidine tagged jellyfish collagen was observed to have a biphasic effect on collagen synthesis. At the 1%, 5% and the 10% levels, collagen synthesis increased. At the 5% concentration truncated jellyfish collagen significantly increased collagen synthesis with a p-value of less than 0.05.

Elastin Synthesis

[0249] Elastin is the main component of a network of elastic fibers that give tissues their ability to recoil after a transient stretch. This protein is released by fibroblasts (soluble elastin) into the extracellular space where it is then cross-linked to other elastin proteins to form an extensive network of fibers and sheets (insoluble elastin). Soluble elastin can be readily measured from cell culture medium via an ELISA based method.

[0250] Soluble α -elastin was dissolved in 0.1 M sodium carbonate (pH 9.0) at a concentration of 1.25 μ g/ml. 150 μ l of this solution was then applied to the wells of a 96-well maxisorp Nunc plate and the plate was incubated overnight at 4° C. On the following day the wells were saturated with PBS containing 0.25% BSA and 0.05% Tween 20. The plate was then incubated with this blocking solution for 1 hour at 37° C. and then washed two times with PBS containing 0.05% Tween 20.

[0251] A set of α -elastin standards was generated ranging from 0 to 100 ng/ml. 180 μ l of either standard or truncated jellyfish collagen was then transferred to a 650 μ l microcentrifuge tube. An anti-elastin antibody solution was prepared (the antibody was diluted 1:100 in PBS containing 0.25% BSA and 0.05% Tween 20) and 20 μ l of the solution

was added to the tube. The tubes were then incubated overnight at $4\pm 2^\circ\text{C}$. On the following day, 150 μl was transferred from each tube to the 96-well elastin ELISA plate, and the plate was incubated for 1 hour at room temperature. The plate was then washed 3 times with PBS containing 0.05% Tween 20. After washing, 200 μl of a solution containing a peroxidase linked secondary antibody diluted in PBS containing 0.25% BSA and 0.05% Tween 20 was added, and the plate was incubated for 1 hour at room temperature. After washing the plate three times, 200 μl of a substrate solution was added and the plate was incubated for 10 to 30 minutes in the dark at room temperature. After this final incubation the plate was read at 460 nm using a plate reader.

TABLE 5

The values are also presented as mean concentration (ng/ml) \pm deviation from the mean.	
Treatment	Elastin (ng/ml)
Untreated	79 \pm 19
20 ng/ml TGFB1	243 \pm 35*
10% Collagen Solution	68 \pm 18
5% Collagen Solution	99 \pm 13
1% Collagen Solution	126 \pm 21
0.5% Collagen Solution	145 \pm 21*
0.1% Collagen Solution	76 \pm 14
0.05% Collagen Solution	58 \pm 6
0.01% Collagen Solution	53 \pm 5
0.005% Collagen Solution	56 \pm 24

*Denotes values that are significantly different from the Untreated group ($p < 0.05$).

[0252] Truncated his-tagged jellyfish collagen significantly increased elastin production when it was used at the 0.5% concentration as shown in Table 5.

Example 7: Effect of Truncated Collagen on Keratinocyte Proliferation and UVB Protection

[0253] A human keratinocyte cell culture model was used to assess the ability of the test materials to exert an effect on cell proliferation. In addition, the impact of the test materials on the cell viability after an exposure to UVB was also assessed.

[0254] A stock solution of 2% w/w truncated collagen was prepared from the truncated collagen of example 1. Aliquots from the 2% stock truncated collagen solution was then used in the experiments described below.

[0255] This study was conducted in two parts. In the first part cultured keratinocytes were incubated with the test materials for 48 hours, after which changes in the number of viable cells were assessed using an MTT assay. In the second part of the study cultured keratinocytes were irradiated with UVB and then treated with the test materials for 48 hours. At the end of the 48 hour period the number of viable cells was again assessed via an MTT assay.

[0256] Changes in cell number of viable cells can be determined using an MTT (3-(4,5-Dimethylthiazol-2-yl)-2,5-diphenyltetrazolium bromide, a tetrazole) assay. The MTT assay is a colorimetric analysis of the metabolic activity of the cell, which is a reflection of the number of viable cells. Reduction of MTT by mitochondria in viable cells results in the formation of insoluble purple formazin crystals that are extracted from the cells with isopropanol and quantified

spectrophotometrically. The intensity of the purple color is directly proportional to the number of metabolically active cells.

Proliferation Assay

[0257] For the proliferation assay the keratinocytes were seeded into 96-well plates using without growth factors and incubated for 24 hours at $37\pm 2^\circ\text{C}$. and $5\pm 1\%$ CO_2 . After this initial incubation the media was replaced with media supplemented with the test materials. Normal media (with growth factors) was used as a positive control. After the addition of the test materials the cells were cultured for 48 hours as described above. At the end of the incubation period changes in the number of viable cells was determined using an MTT assay.

UVB Protection Assay

[0258] For the UVB protection assay the keratinocytes were seeded into 96-well plates using normal media and incubated for 24 hours at $37\pm 2^\circ\text{C}$. and $5\pm 1\%$ CO_2 . After this initial incubation the media was replaced with 100 μl of phosphate buffered saline (PBS) and the cells were exposed to UVB (40 mJ/cm²). After the UVB exposure, the PBS was replaced with fresh media supplemented with the test materials (ascorbic acid at 100 $\mu\text{g/ml}$ served as the positive control) and the cells were cultured for 48 hours at $37\pm 2^\circ\text{C}$. and $5\pm 1\%$ CO_2 . At the end of the 48 hour incubation cell viability was determined using an MTT assay.

MTT Assay

[0259] After the 48-hour incubation the cell culture medium was removed and replaced with 200 μl of culture media supplemented with 0.5 mg/ml MTT. The well plates were incubated for 1 hour at $37\pm 2^\circ\text{C}$. and $5\pm 1\%$ CO_2 . After the incubation, the MTT solution was removed and the cells were washed once with phosphate buffered saline and then 200 μl of isopropyl alcohol was added to the well to extract the purple formazin crystals. The 96-well plate was read at 540 nm using isopropyl alcohol as a blank.

[0260] The mean absorbance value for the cells not treated with test materials (proliferation assay: Untreated Group) or not exposed to UVB (UVB protection assay: Non-UVB Exposed Group) was calculated and used to represent 100% cell viability. The individual absorbance values from the cells undergoing the various treatments were then divided by the mean absorbance value representing 100% cell viability and expressed as a percent to determine the change in cell viability caused by each treatment.

[0261] The results for the Proliferation assay using the his-tagged truncated jellyfish collagen are presented in Table 6. The results for the UVB Protection assay using the his-tagged truncated jellyfish collagen are presented in Table 7. The values for both assays are presented as mean viability \pm standard deviation.

TABLE 6

Proliferation Assay	
Untreated	100 \pm 3.4
Positive Control (Growth Factors)	139 \pm 3.8*
10% Collagen Solution	103 \pm 9.4
5% Collagen Solution	97 \pm 7.3
1% Collagen Solution	94 \pm 5.0

TABLE 6-continued

Proliferation Assay	
0.5% Collagen Solution	93 ± 7.0
0.1% Collagen Solution	95 ± 2.5
0.05% Collagen Solution	99 ± 6.0
0.01% Collagen Solution	96 ± 6.4
0.005% Collagen Solution	96 ± 2.8

*Denotes values which are significantly different from the Untreated Group (p < 0.05)

TABLE 7

UVB Protection Assay	
Non-UVB Exposed	100 ± 1.7*
Untreated	77 ± 1.8
100 ug/ml Trolox	92 ± 2.0*
10% Collagen Solution	76 ± 8.6
5% Collagen Solution	92 ± 3.9*
1% Collagen Solution	91 ± 2.9*
0.5% Collagen Solution	100 ± 4.5*
0.1% Collagen Solution	86 ± 4.8
0.05% Collagen Solution	91 ± 1.6*
0.01% Collagen Solution	83 ± 7.5
0.005% Collagen Solution	82 ± 4.7

*Denotes values which are significantly different from the Untreated Group (p < 0.05)

[0262] For the proliferation assay the Untreated group was used to represent 100% cell viability. Values above 100% reflect an increase in the number of viable cells and hence are indicative of cell proliferation. In this study, the test material was not observed to promote cell proliferation.

[0263] In addition, the keratinocyte proliferation assay was performed using the truncated collagen of SEQ ID NO: 91. A 1% and 0.5% collagen solution of a 5% stock solution was prepared according to Example 8 and tested. The truncated collagen of SEQ ID NO: 91 had keratinocyte cell viability assay values of 102±2.9 and 102±2.0, respectively. The observed values were statistically significant (p<0.05).

[0264] In addition to its effects on cell proliferation, the test material was also screened to determine if it had an impact on cell recovery after UVB exposure. In this study, exposure to UVB was observed to result in a significant reduction in the number of viable cells 48 hours post exposure. However, treatment with the test material prevented this decrease in cell viability. The effect was evident within a concentration range between 0.05% and 5% of the test material, with an optimal effect at a concentration of 0.05%. Within this range of concentrations cell viability was significantly greater than the untreated group (with the lone exception of the 0.01% concentration), demonstrating that the material has UVB protective effect. Since this material was added after the UVB exposure it could be acting to reduce the damaging effects of the UVB irradiation, or it could be helping the damaged cells to recover at a faster rate. With respect to the latter, then truncated collagen is beneficial when applied topically to the skin and has a regenerative effect on skin cells damaged by UVB.

[0265] In addition, the UVB protection assay was performed using the truncated collagen of SEQ ID NO: 91. A 1% and 0.5% collagen solution of SEQ ID NO: 91 had keratinocyte cell viability assay values of 80±4.6 and 78±2.5, respectively. The observed values were statistically significant (p<0.05).

Example 8: Effect of Truncated Collagen on Thymine Dimer Formation

[0266] Upon exposure to ultraviolet radiation the thymine dimer (TT dimer) content in DNA present in cells increases. Increases in TT dimer formation are correlated with skin damage and certain types of cell proliferative diseases including skin cancer.

[0267] The polynucleotide of SEQ ID NO: 11 was expressed in the expression system of Example 1 and purified as described in this example. The encoded polypeptide includes the DsbA secretion tag. As the polypeptide is processed through the secretion pathway, the DsbA tag, amino acids 1-24 of SEQ ID NO: 12 is cleaved by the host cell. The truncated collagen without the DsbA secretion tag is provided in SEQ ID NO: 91.

[0268] The truncated collagen of SEQ ID NO: 91 was tested to determine if it could reduce TT dimer formation in human epidermal keratinocytes. For this study, the cells were exposed to UVB (25 mJ/cm²). Following the exposures cells were treated with the test materials or Trolox (100 ug/ml) and incubated overnight. On the following day cellular DNA was extracted and assayed for thymine dimer content using an ELISA based method.

[0269] Human keratinocytes were seeded into 12-well plates using normal media and incubated for 24 hours at 37±2° C. and 5±1% CO₂. After this initial incubation the media was replaced with 100 µl of phosphate buffered saline (PBS) and the cells were exposed to UVB (25 mJ/cm²). After the UVB exposure, the PBS was replaced with fresh media supplemented with the test materials or Trolox (100 µg/ml, this served as the positive control) and the cells were cultured overnight at 37±2° C. and 5±1% CO₂. At the end of the incubation cellular DNA was extracted.

[0270] After the overnight incubation the cell culture media was removed from the wells and replaced with 200 µl of PBS and 20 µl of Proteinase K. After swirling the plate to mix the PBS and Proteinase K, 200 µl of buffer AL was added to each well. After again swirling the plate to mix the reagents, the plates were incubated for 10 minutes at 55±2° C. After cooling the plate to room temperature, the DNA was precipitated by the addition of 200 µl of 100% ethanol. The precipitated DNA mixtures were then transferred to DNEasy Spin Columns in 2 ml collection tubes and centrifuged at 8,000 RPM for 1 minute. The flow through and collection tubes were discarded, and 500 µl of Wash Buffer One was added to the spin column and the column was placed into a new collection tube and centrifuged at 8,000 RPM for 1 minute. The flow through and collection tube were again discarded, and 500 µl of Wash Buffer Two was added to the spin column and the column was placed into a new collection tube and centrifuged at 14,000 RPM for 3 minutes. The spin column was then placed into a new 1.5 ml centrifuge tube and 110 µl of ultrapure water was added to the column. The column was incubated for 1 minute at room temperature and then centrifuged at 8,000 RPM for 1 minute.

[0271] Extracted DNA was quantified via a fluorometric assay. A 2 µl aliquot of the DNA sample was mixed with 100 µl TE buffer in a 96-well plate. A series of DNA standards was also transferred to wells in a 96-well plate (in duplicate). Finally, 100 µl of dilute Cyquant Green dye was added to each well and the fluorescence intensity of each well was determined using an excitation wavelength of 480 nm and an emission wavelength of 520 nm.

[0272] Thymine Dimer Detection was determined using an OxiSelect™ UV-Induced DNA Damage ELISA Kit)

[0273] Aliquots of genomic DNA samples or standards were converted to single stranded DNA by incubating the samples at 95° C. for 10 minutes and then chilled on ice. 100 µl or each sample or standard was transferred to a DNA binding ELISA plate and incubated overnight at 4° C. On the following day the wells were rinsed once with 100 µl of PBS and then blocked with 150 µl of Assay Diluent for one hour at room temperature. After removing the Assay Diluent, 100 µl of anti-CPD antibody was added to each well and the plate was incubated for one hour at room temperature. After this incubation, the plate was washed three times with 250 µl of wash buffer per well, and then 150 µl of Blocking Reagent was added to the plate. The plate will be blocked again for one hour at room temperature, and then washed three times as described before. 100 µl of Secondary Antibody was then added to each well and the plate was incubated for 1 hour at room temperature. After washing the plate again, 100 µl of substrate was added to each well and the plate was incubated for 5-20 minutes to allow for color generation in the plate. The color generation reaction was stopped by the addition of 100 µl of stop solution and the plate was read at 460 nm using a plate reader.

[0274] To quantify the amount of DNA present, a standard curve was generated using known concentrations of DNA and their respective fluorescence intensity (measured in RFUs or relative fluorescence units). A regression analysis was performed to establish the line that best fits these data points. The Relative Fluorescence Units (RFU) for each unknown sample was then used to estimate the amount of DNA.

[0275] A series of DNA standards with known amounts of thymine dimer content were used to generate a standard curve. This standard curve was used to determine the amount of DNA damage in the sample DNA. Means for each treatment group were calculated and compared using an ANOVA. In table 8 and FIG. 5, a standard 5% collagen solution (5 g truncated collagen in 95 ml deionized water was further diluted with phosphate buffered saline (PBS) to the indicated percent solution.

TABLE 8

Thymine Dimer Assay	
Treatment	Thymine Dimer, ng/ml
Non-UVB Exposed	0.1 ± 0.2*
Untreated	10.7 ± 1.4*
100 µg/ml Trolox	1.6 ± 0.3*
5% Truncated Collagen	3.5 ± 0.3*
1% Truncated Collagen	5.5 ± 1.0*
0.5% Truncated Collagen	8.9 ± 0.8
0.1% Truncated Collagen	9.2 ± 0.9
0.05% Truncated Collagen	9.2 ± 0.3

*Denotes values which are statistically significantly different from untreated with P < 0.05

[0276] Table 8 shows that the 5% and the 1% truncated collagen solution reduced TT dimer formation with statistical significance (p<0.05). The data is presented graphically in FIG. 5.

[0277] The experiment was repeated with a different lot of truncated collagen (SEQ ID NO: 91). The amount of TT dimer in ng/ml in non-UVB exposed cells was 1.3±1.2, untreated cells was 18.1±0.4, 100 µg/ml Trolox treated cells

was 7.9±0.3, 5% collagen was 13.1±0.2, and 1% collagen was 17.4±0.7. The reduction in TT dimer formation for non-UVB exposed cells, Trolox treated cells, 5% collagen treated cells and 1% collagen treated cells was statistically significant compared to untreated cells (p<0.05).

Example 9: Human Collagens

Truncated Human Collagen Type 21 Alpha 1

[0278] A truncated human collagen type 21 alpha 1 without a His tag, linker, and thrombin cleavage site is disclosed below. The codon-optimized nucleotide sequence encoding this collagen and the amino acid sequence are disclosed below. In SEQ ID NOs: 73 and 74, the DsbA secretion tag is encoded by nucleotides 1-72 and encodes amino acids 1-24. In SEQ ID NOs: 73 and 74, the truncated collagen sequence is encoded by nucleotides 73-633 and encodes amino acids 25-211.

[0279] The codon-optimized nucleotide sequence encoding this collagen is provided in SEQ ID NO: 73.

(SEQ ID NO: 73)
 ATGAAAAAGATTGGCTGGCGCTGGCTGGTTAGTTTTAGCGTTTAGCGCA
 TCGGCGGCGCAGTATGAAGATGCAGGTTTTCCGGGTCTGCCTGGTCCGGCA
 GGCGAACCCGGTTCGTATGGTAAAGATGGTCTGATGGGTAGTCCGGGTTTT
 AAAGGTGAAGCAGGTTCCACCGGTGCACCTGGTCAGGATGGCACCCGTGGT
 GAACCGGTATTCCGGGATTTCCGGGTAATCGTGGCCTGATGGGTGAGAAA
 GGTGAAATTGGTCCGCCTGGT CAGCAGGGTAAAAAAGGCGCACCCGGGTATG
 CCAGGACTGATGGGTTCAAATGGCAGTCCGGGT CAGCCAGGCACACCCGGT
 TCAAAAGGTAGCAAAGGCGAACCTGGTATT CAGGGTATGCCTGGTGCAAGC
 GGCTGAAAGGCGAGCCAGGTGCCACCGGTTCTCCGGGTGAACCAGGTTAT
 ATGGTCTGCCAGGTATCCAAGGCAAAAAGGTGATAAAGGTAATCAGGGC
 GAAAAAGGCATTCAGGGCCAGAAAGGCGAAAATGGCCGT CAGGGTATTCCA
 GGCCAGCAGGCATCCAGGGT CATCATGGTCAAAGGTGAACGTGGTGA
 AAGGGCGAACCCAGGTGTTTCGTtaa

[0280] The amino acid sequence is disclosed in SEQ ID NO: 74.

(SEQ ID NO: 74)
 MKKIWLALAGLVLAFSASAAQYEDAGFPPLPGPAGEPRRHGKDLGMLGSPGF
 KGEAGSPGAPGQDGRGEPGIPGFPNRLMLGQKGEIGPPGQQKKGAPGM
 PGLMGSNGSPGPGTPGSKGSKGEPGIQGMPPASGLKGEPEGATGSPGEPGY
 MGLPGIQGKKGDKGNQGEKGIQGGKGENGRQGIPIGQQGIQHHGAKGERGE
 KGEPGVR

[0281] The codon-optimized nucleotide sequence encoding the truncated human collagen type 21 alpha 1 without the DsbA secretion tag collagen is provided in SEQ ID NO: 75.

(SEQ ID NO: 75)
 TGCAGGTTTTCCGGGTCTGCCTGGTCCGGCAGGCGAACCGGTCTGCATG
 GTAAAGATGGTCTGATGGGTAGTCCGGGTTTTAAAGGTGAAGCAGGTTCA
 CCGGGTGCACCTGGTCAGGATGGCACCCGGTGGTGAACCGGTATTCCGGG
 ATTTCCGGGTAATCGTGGCCTGATGGGTGAGAAAGGTGAAATTTGGTCCGC
 CTGGTCAGCAGGGTAAAAAAGGCGCACCCGGTATGCCAGGACTGATGGG
 TTCAATGGCAGTCCGGGTGAGCCAGGCACCCGGTTCAAAGGTAGCA
 AAGGCGAACCTGGTATTGAGGTATGCCTGGTGAAGCGGTCTGAAAGGC
 GAGCCAGGTGCCACCGGTTCTCCGGGTGAACAGGTTATATGGGTCTGCC
 AGGTATCCAAGGCAAAAAGGTGATAAAGGTAATCAGGGCGAAAAGGC
 ATTCAGGCGCCGAAAAGGCGAAAATGGCCGTGAGGTATTCCAGGCGCAGC
 AGGGCATCCAGGTCATCATGGTGCAAAGGTGAACGTGGTGAAGGG
 CGAACAGGTGTTCTGTTaa

[0282] The amino acid sequence of truncated human collagen type 21 alpha 1 without the DsbA secretion tag is disclosed in SEQ ID NO: 76.

(SEQ ID NO: 76)
 AGFPLPGPAGPEPRHGKDGMLGSPGFKGEAGSPGAPGQDTRGEPGIPG
 FPGNRGLMQKGEIGPPGQQGKKGAPGMPGLMGSNGSPGQPGTPGSKGSK
 GEPGIQGMPPGASGLKGEPEGATGSPGEPGYMGLPGIQGKKGDKGNQGEKGI
 QGQKGENGRQGIPIGQQGIQHHGAKGERGEKGEPEGVR

[0283] A truncated human collagen type 1 alpha 2 without a His tag, linker, and thrombin cleavage site is disclosed below. The codon-optimized nucleotide sequence and the amino acid sequences are disclosed below. In SEQ ID NOs: 78 and 79, The DsbA secretion tag is encoded by nucleotides 1-72 and encodes amino acids 1-24. The truncated collagen sequence is encoded by nucleotides 73-636 and encodes amino acids 25-212.

[0284] The codon-optimized nucleotide sequence encoding this collagen is provided in SEQ ID NO: 77.

(SEQ ID NO: 77)
 ATGAAAAAGATTGGCTGGCGCTGGCTGGTTTGTATTTAGCGTTTAGCGC
 ATCGCGCGCAGTATGAAGATATGGGTCGCGCTGGTAGCCGTGGTGCAA
 GTGGTCCGGCAGGCGTTTCGTGGTCCGAATGGTATGACAGGTCGTCCGGGT
 GAACCGGTCCTGATGGTCTCTGCTGGTCTGCTGGTTCACCGGTAATAT
 TGGTCTCAGGTAAGAAGGTCGCGTTGGTCTGCCAGGTATTGATGGCC
 GTCCGGTCCGATTGGTCCAGCCGGTGCACGTGGTGAACCTGGCAATATT
 GGTTTTCCGGTCTTAAAGGTCGACCGGTGATCCGGTAAAATGGTGA
 TAAAGGTCATGCAAGTCTGGCAGGCGCACCGGTGCACCTGGTCCGGATG
 GTAATAATGGTGCACAGGTCACCCGGTCCGACGGGTGTTCAAGGTGGT
 AAAGGCGAACAGGTCCTCCGGTCTCCGGTTTTTCCAGGACTGCTGG
 TCCGAGCGGTCTGCGGGTGAAGTTGGTAAACCTGGTGAACCGGTCTGC

-continued

ATGGTGAATTTGGCCTGCCTGGGCTGCAGGTCGCGGTGGCGAACGTGGT
 CCGCCAGGTGAAAGCGGTGCAGCAGGTCGACAGGTTaa

[0285] The amino acid sequence is disclosed in SEQ ID NO: 78.

(SEQ ID NO: 78)
 MKKIWLALAGLVAFASAAQYEDMGPPGSRGASGPAGVRGPNGDAGRPG
 EPGLMGPRPLPGSPGNI GPAGKEGVPVGLPGIDGRPGPIGPAGARGEPGNI
 GFPGPKGPTGDPGKNGDKGHAGLAGARGAPGPDGNNGAQGGPPGQVQGG
 KGEQGPAGPPGFQGLPGPSGPAGEVKGKPERGLHGEFGLPGPAGPRGERG
 PPGESGAAGPTG

[0286] The nucleic acid sequence of truncated human collagen type 1 alpha 2(1) without the DsbA secretion tag is disclosed in SEQ ID NO: 79.

(SEQ ID NO: 79)
 ATGGTCCGCTGGTAGCCGTTGCAAGTGGTCCGGCAGGCGTTCTGTTG
 TCCGAATGGTATGACAGGTCGTCCGGGTGAACCGGTCCTGATGGGTCTCT
 GTGGTCTGCCTGGTTCACCGGTAATATTGGTCTGCAGGTAAGAAGGT
 CCGGTTGGTCTGCCAGGTATTGATGGCCGTCGCGGTCCGATTGGTCCAGC
 CGGTGCACGTGGTGAACCTGGCAATATTGGTTTTTCCGGTCTTAAAGGTC
 CGACCGGTGATCCGGTAAAATGGTGAAGAAGTATGACAGGTCCTGGCA
 GGCGCACGCGGTGCACCTGGTCCGGATGGTAATAATGGTGCACAGGGTCC
 ACCGGTCCGACGGGTGTTCAAGGTGTAAGGCGAACAGGTCCTGCCG
 GTCTCCGGTTTTTCCAGGACTGCCTGGTCCGAGCGGTCCTGCGGGTGA
 GTTGGTAAACCTGGTGAACCGGTCTGCATGGTGAATTTGGCTGCCTGG
 GCCTGCAGGTCGCGTGGCGAACGTGGTCCGCGAGGTGAAAGCGGTGCAG
 CAGGTCCGACAGGTTaa

[0287] The amino acid sequence of truncated human collagen type 1 alpha 2(1) without the DsbA secretion tag is disclosed in SEQ ID NO: 80.

(SEQ ID NO: 80)
 MGPPGSRGASGPAGVRGPNGDAGRPEGLMGPRPLPGSPGNI GPAGKEG
 PVGLPGIDGRPGPIGPAGARGEPGNI GFPGPKGPTGDPGKNGDKGHAGLA
 GARGAPGPDGNNGAQGGPPGQVQGGKGEQGPAGPPGFQGLPGPSGPAGE
 VGKPERGLHGEFGLPGPAGPRGERGPPGESGAAGPTG

Truncated Human Collagen Type 1 Alpha 2 (2)

[0288] A truncated human collagen type 1 alpha 2 without a His tag, linker, and thrombin cleavage site is disclosed below. The codon-optimized nucleotide sequence and the amino acid sequences are disclosed below. In SEQ ID NOs: 82 and 83, the DsbA secretion tag is encoded by nucleotides 1-72 and encodes amino acids 1-24. The truncated collagen sequence is encoded by nucleotides 73-609 and encodes amino acids 25-203.

[0289] The codon-optimized nucleotide sequence encoding this collagen is provided in SEQ ID NO: 81.

(SEQ ID NO: 81)
ATGAAAAAGATTGGCTGGCGCTGGCTGGTTTAGTTTTAGCGTTTAGCGC
ATCGGCGGCGCAGTATGAAGATGGTTTTTCAGGGTCTGCCGGTGAACCGG
GTGAACCTGGTCAGACAGGTCCGGCAGGCGCACGTGGTCTGCAGGTCTT
CCTGGTAAAGCCGGTGAAGATGGTCATCCGGTAAACCGGGTCTGCTCTGG
TGAACGTGGTGTGTGGTCCGAGGGTCCCGTGGTTTTCCGGTACTC
CGGGTCTGCCAGGTTTTAAAGGTATTCGTGGTCATAATGGTCTGGATGGT
CTGAAAGGTACGCTGGTGCACCGGGTGTAAAGGTGAACAGGTGCTCC
GGGTGAAAATGGCACACCGGGTCCAGACCGGTGCGCGTGGTCTGCCTGGCG
AACCGGGTCTGTTGGTGCACCTGGTCCAGCCGGTGCACCGGTAGTGAT
GGTAGCGTTGGTCCGGTGGTCCAGCGGGTCCGATTGGTAGCGCAGGTCC
ACCGGGTTTTCCAGCGGCACCGGGTCCGAAAGGTGAAATGGTGCAGTTG
GTAATGCAGGCCCTGCCGGTCCAGCAGGACCGGTGGTGAAGTTGGCCTG
CCTGGTCTGt aa

[0290] The amino acid sequence is disclosed in SEQ ID NO: 82.

(SEQ ID NO: 82)
MKKIWLALAGLVLAFSASAAQYEDGFQGPAGEPGEQGTGPAGARGPAGP
PGKAGEDGHPGKPRPGERGVVGPQARGFPPTPLPGFKGIRGHNLG
LKGQPGAPGVKGEPEGAPGENGTPGQTGARGLPGERGRVGPAGARGSD
GSVGPVGPAGPIGSAGPPFPAGPKGEIGAVGNAGPAGPAGPRGEVGL
PGL

[0291] The nucleic acid sequence of truncated human collagen type 1 alpha 2(2) without the DsbA secretion tag is disclosed in SEQ ID NO: 83.

(SEQ ID NO: 83)
GGTTTTCAGGGTCTGCCGGTGAACCGGGTGAACCTGGTCAGACAGGTCC
GGCAGGCGCACGTGGTCTGCAGGTCTCTCGTTAAAGCCGGTGAAGATG
GTCATCCGGTAAACCGGGTCTCTGGTGAACGTGGTGTGTTGGTCCG
CAGGGTGCCTGGTTTTCCGGTACTCCGGTCTGCCAGGTTTTAAAGG
TATTCGTGGTCATAATGGTCTGGATGGTCTGAAAGGTGAGCTGGTGCAC
CGGGTGTAAAGGTGAACAGGTGCTCCGGGTGAAAATGGCACACCGGGT
CAGACCGGTGCGCGTGGTCTGCCTGGCGAACCGGTCTGTTGGTGCACC
TGGTCCAGCCGGTGCACCGGTAGTGATGGTAGCGTTGGTCCGGTGGTCC
CAGCGGGTCCGATTGGTAGCGCAGGTCCACCGGGTTTTCCAGGCGCACCG
GGTCCGAAAGGTGAAATGGTGCAGTTGGTAAATGCAGGCCCTGCCGGTCC
AGCAGGACCGGTGGTGAAGTTGGCTGCCTGGTCTGt aa

[0292] The amino acid sequence of truncated human collagen type 1 alpha 2(2) without the DsbA secretion tag is disclosed in SEQ ID NO: 84.

(SEQ ID NO: 84)
GFQGPAGEPGEQGTGPAGARGPAGPPGKAGEDGHPGKPRPGERGVVGP
QGARGFPPTPLPGFKGIRGHNLGDLKQPGAPGVKGEPEGAPGENGTPG
QTGARGLPGERGRVGPAGARGSDGSVGPVGPAGPIGSAGPPFPAGP
GPKGEIGAVGNAGPAGPAGPRGEVGLPGL

[0293] The polynucleotides of SEQ ID NO: 73, 77 or 81 were subcloned in vector pET28a as described herein to prepare a transformation vector. Host cells were transformed with the vector the polynucleotides were expressed as described in Example 2.

[0294] After the fermentation was completed, the truncated human collagen was purified from the fermentation broth using the procedures disclosed in Example 3. The purified truncated human collagens were analyzed using SDS-PAGE and HPLC as disclosed in Example 3.

[0295] All three truncated human collagens ran at the expected molecular weights in the SDS-PAGE analysis. In analyzing the truncated human collagens using HPLC, a standard curve using the jellyfish collagen of Example 3 was utilized. The retention times of the human collagens were slightly different than the jellyfish collagen. The retention time of SEQ ID NO: 76 was 5.645 minutes, the retention time of SEQ ID NO: 80 was 5.631 minutes, and SEQ ID NO: 84 ran at two peaks and the retention times were 5.531 and 5.7 minutes.

Truncated Human Collagen Type 1 Alpha 2 Truncation 5 with DsbA Secretion and FLAG Tag

[0296] The amino acid sequence of truncated human collagen type 1 alpha 2 truncation 5 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 92. The DsbA secretion tag is encoded by nucleotides 1-57 of SEQ ID NO: 93 and the amino acid sequences are amino acids 1-19 of SEQ ID NO: 92. The collagen nucleotide sequences are nucleotides 58-657 of SEQ ID NO: 93 and the amino acid sequences are amino acids 20-219 of SEQ ID NO: 92. The FLAG nucleotide sequences are nucleotides 658-684 of SEQ ID NO: 93 and the amino acid sequences are amino acids 220-228.

(SEQ ID NO: 92)
MKKIWLALAGLVLAFSASAGDQGPVGRTEVAVGPPGFAGEKGPSGEAG
TAGPPGTPGPQGLLGPGLGAPGILGLPGRGERGLPGVAVAGVGPGLGIAGPP
GARGPPGAVGSPGVNAPGEAGRDGNPNDGPPGRDQPGHKGERGYPGN
IGPVGAAGAPGPHGVPVGPAGKHGNERGETGPSGPVGPAGAVGPRGSPGQG
IRGDKGEPGEKPRGLPLGLDYKDDDDK

[0297] The nucleic acid sequence of truncated human collagen type 1 alpha 2 truncation 5 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 93.

(SEQ ID NO: 93)
ATGAAAAAGATTGGCTGGCGCTGGCTGGTTAGTTTTAGCGTTTAGCGC
ATCGGCGGGTGATCAGGGTCCGGTGGTCTGACCGGTGAAGTTGGTGCAG
TCGGGCCCGGGTTTTGCGGGTGA AAAAGGCCCGTCAAGTGAAGCAGGC
ACCGCTGGCCCTCTGGCACGCTGGCCACAGGGTTTTACTGGGCGCAC

-continued

TGGAATTCGGACTGCCGGGCAGCCGTGGAGAACCGGTTTACCAGGTG
 TTGCCGGTGCCGTTGGTGAACCTGGTCCACTGGGCATTGCAGGGCCGCT
 GGCGCACGGGGACCCTGGTGTCTGGTAGTCCGGTGTGAATGGTGC
 TCCGGTGAAGCCGTCGTGACGGTAATCCGGGAATGACGGCCCGCCAG
 GCCGCGATGGTCAGCCGGGTATAAAGGTGAGCGTGGTTACCAGGTAAT
 ATTGGTCCAGTCGGTCCGCCGGTCCGCCGGTCTCATGGCCCTGTCCG
 TCCAGCCGGTAAACATGGTAATCGCGGTGAGACAGGTCCGTGAGACCAG
 TGGGCCCTGTGGCGCAGTCGGTCCCGCGGGCCGAGTGGCCCTCAGGGT
 ATTCGTGGCGATAAAGGGGAACCGGGCGAAAAGGGCCCGGGGTCTGCC
 AGGCCTGGGTGACTACAAAGACGACGACGACAAAtaa

[0298] The polynucleotide of SEQ ID NO: 93 was subcloned into vector pET28a, expressed host *E. coli* cells and the truncated collagen was purified as described herein. The purified collagen produced a clear band on SDS-PAGE and an anti-FLAG western was observed at around 100 kilodaltons. There were no existing bands that appear at that location on the gel in the absence of expression of this protein.

Truncated Human Collagen Type 1 Alpha 2 Truncation 6 with DsbA Secretion and FLAG Tag

[0299] The amino acid sequence of truncated human collagen type 1 alpha 2 truncation 6 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 94. The DsbA secretion tag is encoded by nucleotides 1-57 of SEQ ID NO: 95 and the amino acid sequences are amino acids 1-19 of SEQ ID NO: 94. The collagen nucleotide sequences are nucleotides 58-657 of SEQ ID NO: 95 and the amino acid sequences are amino acids 20-219 of SEQ ID NO: 94. The FLAG nucleotide sequences are nucleotides 658-684 of SEQ ID NO: 95 and the amino acid sequences are amino acids 220-228 of SEQ ID NO: 94.

(SEQ ID NO: 94)

MKKIWLALAGLVLAFSASAKGHNLQGLPGIAGHHGDQAGPSVGPAGPR
 GPAGPSGPAGKDRGTGHPGTVGPAGIRGPQGHQGPAGPPGPPGPPGV
 SGGGYDFGYDGFYRADQPRSAAPSLRPKDYEV DATLKSLNNQIETLLTPE
 GSRKNPARTCDLRLLSHPPEWSSGYWIDPNQGTMDAIKVYCDFSTGETC
 IRAQPENIPAKNWRSSKDGDKDDDDK

[0300] The nucleic acid sequence of truncated human collagen type 1 alpha 2 truncation 6 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 95.

(SEQ ID NO: 95)

ATGAAAAAGATTTGGCTGGCGCTGGCTGGTTTGTATTTAGCGTTTAGCCG
 ATCGGCAAAAGTCAACAATGGACTGCAAGGCCTGCCAGGTATTGCAGTTC
 ATCATGGTGATCAAGGTGCCCGGGAAGCGTTGGTCCGGCGGGCCGAGA
 GGCCCTGCGGGACCTTCAGGTCCGGCAGGCAAGATGGTCCGACAGGCCA
 TCCGGCACCCGTTGGCCCTGCAGGAATTCGTGGACCGCAGGTCATCAGG

-continued

GACCTGCTGGTCCGCCAGGTCCCCGGGCCCTCCGGGACCACCGGGTGT
 AGTGGTGGTGGTTATGATTTTGGCTATGATGGTGTATTTTATCGTGCAGA
 TCAGCCCGTAGCGCACCGAGCCTGCGTCTTAAAGATTATGAAGTTGATG
 CAACCTGAAAAGCCTGAATAATCAGATTGAAACACTGCTGACACCGGAA
 GGTAGCCGTAAAAATCCGGCCGTACCTGCTGCTGATCTGCGTCTGAGCCA
 CCGGAATGGAGCAGCGGTATTATTGGATTGATCCGAATCAAGGTTGTA
 CCATGGATGCAATTAAAGTTTATTGTGATTTTAGCACAGGTGAAACATGT
 ATCCGTGCACAGCCGAAAATATTCGGCCAAAATTGGTATCGTAGTAGT
 CAAAGATGGTACTACAAAGACGACGACGACAAAtaa

[0301] The polynucleotide of SEQ ID NO: 94 was subcloned into vector pET28a, expressed host *E. coli* cells and the truncated collagen was purified as described herein. The purified collagen produced a clear band on SDS-PAGE and an anti-FLAG western was observed at around 25 kilodaltons. There were no existing bands that appear at that location on the gel in the absence of expression of this protein.

Truncated Human Collagen Type 1 Alpha 2 Truncation 7 with DsbA Secretion and FLAG Tag

[0302] The amino acid sequence of truncated human collagen type 1 alpha 2 truncation 7 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 96. The DsbA secretion tag is encoded by nucleotides 1-57 of SEQ ID NO: 97 and the amino acid sequences are amino acids 1-19 of SEQ ID NO: 96. The collagen nucleotide sequences are nucleotides 58-759 of SEQ ID NO: 96 and the amino acid sequences are amino acids 20-253 of SEQ ID NO: 96. The FLAG nucleotide sequences are nucleotides 760-786 of SEQ ID NO: 97 and the amino acid sequences are amino acids 254-262 of SEQ ID NO: 96.

(SEQ ID NO: 96)

MKKIWLALAGLVLAFSASAYEVDATLKSLNNQIETLLTPEGSRKNPARTC
 RDLRLSHPPEWSSGYWIDPNQGTMDAIKVYCDFSTGETCIRAQPENIPA
 KNWYRSSKDKKHVWLGETINAGSQPEYINVEGVTSKEMATQLAFMRLLANY
 ASQNIITYHCKNSIAYMDEETGNLKKAVILQGSNDVELVAEGNSRFTYTVL
 VDGCSKKTNEWGKTIIEYKTNKPSRLPFLDIAPLDIGGADQEFFVDIGPV
 CFKGDYKDDDDK

[0303] The nucleic acid sequence of truncated human collagen type 1 alpha 2 truncation 7 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 97.

(SEQ ID NO: 97)

TGAAAAAGATTTGGCTGGCGCTGGCTGGTTTGTATTTAGCGTTTAGC
 GCATCGCGTATGAAGTTGATGCAACCTGAAAAGCCTGAATAATCA
 GATTGAAACACTGCTGACACCGGAAGGTAGCCGTAATAATCCGGCC
 GTACCTGTGCTGATCTGCGTCTGAGCCACCCGGAATGGAGCAGCGGT
 TATTATTGGATTGATCCGAATCAAGGTTGTACCATGGATGCAATTAA
 AGTTTATTGTGATTTTAGCACAGGTGAAACATGTATCCGTGCACAGC

- continued

CGGAAAATATTCCGGCCAAAATTTGGTATCGTAGTAGCAAAGATAAA
 AAACATGTGTGGCTGGGTGAAACCATTAATGCAGGTAGCCAGTTTGA
 ATACAATGTTGAAGGTGTTACCAGCAAAGAAATGGCAACACACAGCTGG
 CATTATGCGTCTGCTGGCAAATTATGCAAGCCAGAATATTACATAT
 CATTGTAAAAATAGCATTGCATATATGGATGAAGAAACCGGTAATCT
 GAAAAAGCAGTTATTCTGCAGGGTAGCAATGATGTTGAACCTGGTTG
 CGGAAGGTAATAGCCGTTTTACATATACCGTTCTGGTTGATGGTTGT
 AGCAAAAAACCAATGAATGGGGTAAAACCATCATTGAATATAAAAC
 CAACAACCGAGCCGCTCTCCGCTTTCTGGATATCGCTCCGCTGGATA
 TTGGTGGTGCCGATCAGGAATTTTTTGTGATATCGGTCTGTGTGT
 TTTAAAGGTGACTACAAAGACGACGACGACAAAat aa

[0304] The polynucleotide of SEQ ID NO: 97 was subcloned into vector pET28a, expressed host *E. coli* cells and the truncated collagen was purified as described herein. The purified collagen produced a clear band on SDS-PAGE and an anti-FLAG western was observed at around 30 kilodaltons. There were no existing bands that appear at that location on the gel in the absence of expression of this protein.

Example 10: Protective Effect of Truncated Human Collagen on Fibroblasts

[0305] The effect of truncated human collagen on fibroblast cell viability, procollagen synthesis, and elastin Synthesis is determined according to the methods of Example 6.

[0306] The effect of truncated human collagen on Keratinocyte proliferation and UVB protection is determined according to the methods of Example 7.

[0307] The effect of truncated collagen on thymine dimer formation after exposure to UV radiation is determined according to the methods of Example 8.

[0308] It is understood that the examples and embodiments described herein are for illustrative purposes only and that various modifications or changes in light thereof will be suggested to persons skilled in the art and are to be included within the spirit and purview of this application and scope of the appended claims. All publications, patents, and patent applications cited herein are hereby incorporated by reference in their entirety for all purposes.

Example 11: Effect of Truncated Collagen on Inflammatory Cytokines

[0309] Keratinocytes and dermal fibroblasts play an important role in the immune response of the skin. In response to irritating chemicals or UV radiation (pro-inflammatory/pro-irritation stimuli), keratinocytes can release a vast array of cytokines. These cytokines are thought to help engage immune cells to the site of inflammation. Cytokines released by the keratinocytes include TNF α , IL-1 α , IL-1 β , IL-3, IL-6, IL-7, IL-8, IL-10, IL-18, and IL-1RA.

[0310] The testing model used for this study was the MatTek EpiDerm. This skin model consists of normal human-derived epidermal keratinocytes that have been cultured to form a multilayered, highly differentiated model of the human epidermis. Ultrastructural analysis has revealed

the presence of keratohyalin granules, tonofilament bundles, desmosomes, and a multi-layered stratum corneum containing intercellular lamellar lipid layers arranged in patterns characteristic of in vivo epidermis. Markers of mature epidermis specific differentiation such as pro-filaggrin, the K1/K10 cytokeratin pair, involucrin, and type I epidermal transglutaminase have been localized in this model. The MatTek EpiDerm is also mitotically and metabolically active.

[0311] The MatTek EpiDerm tissues were used to assess the ability of various test materials to inhibit the release the inflammatory mediator IL-1 α . Test materials were compared to an over the counter topical hydrocortisone preparation (positive control) as well as to untreated tissues (negative control 1) and untreated, non-inflamed tissues (negative control 2). This test was also used to assess the viability of the tissues after exposure to the test materials.

[0312] IL-1 α , IL-6 and IL-8 are synthesized and stored in keratinocytes and have been identified as a mediators of skin irritation and inflammation. Release of these cytokines can be directly measured in tissue culture media via a colorimetric based enzyme linked immunosorbent assay (ELISA). Briefly, antibodies covalently linked to a solid support will bind IL-1 α , IL-6 or IL-8 present in spent culture media samples. A second antibody that is covalently attached to an acetylcholinesterase enzyme will in turn detect the specific bound cytokines. Upon addition of an appropriate color substrate the acetylcholinesterase enzyme will generate a colored end product that can be measured spectrophotometrically.

[0313] MatTek EpiDerm Tissues were purchased from MatTek corporation and were stored at 4° C. until used. Prior to use, the tissues to be used were removed from the agarose-shipping tray and placed into a 6-well plate containing 0.9 ml of hydrocortisone free assay medium (37 \pm 2° C.). The tissues were allowed to incubate overnight at 37 \pm 2° C. and 5 \pm 1% CO₂. After this initial incubation, the assay medium was replaced with 0.9 ml of fresh hydrocortisone free medium (37 \pm 2° C.). Three tissues were prepared for each test material.

[0314] An inflammatory response in the tissues was initiated via UV irradiation (UVB). A UV lamp was used to give a 300 mJ/cm² dose of UVB radiation to the tissues. Immediately after the application of the inflammatory stimuli 50 μ l or mg of test material was applied directly onto the surface of the tissue. An over the counter hydrocortisone cream was used as a positive control. For a negative control tissues were exposed to the inflammatory stimuli but were not treated with any type of anti-inflammatory material. One additional set of tissues was left without exposure to the inflammatory stimuli to provide a baseline measurement for the cytokines. The tissues were incubated at 37 \pm 2° C. and 5 \pm 1% CO₂ for 24 hours after exposure to the inflammatory stimuli. After the 24-hour incubation the cell culture medium was collect and stored at -75° C. until analyzed for cytokines.

[0315] The ELISA plates were prepared by diluting the appropriate capture antibody in PBS. Next, 100 μ l of the diluted capture antibody was added to the wells of a 96-well ELISA plate and the plate was incubated overnight at room temperature. On the following day the plate was washed three times with 300 μ l wash buffer (0.05% Tween 20 in PBS) and then blocked by adding 300 μ l of blocking buffer (1% BSA in PBS) to each well. The plate was incubated with

the blocking buffer for at least one hour. After the incubation the blocking buffer was removed and the plate was washed three times as described above.

[0316] A series of standards was prepared and 100 µl of each of these standards was dispensed into two wells (duplicates) in the appropriate 96-well plate. Subsequently, 100 µl of each sample was added to additional wells and the plate was incubated for two hours at room temperature. After the incubation the plate was washed three times as described above. Once the last wash was removed, 100 µl of a biotin conjugated detection antibody was added. After incubating the plate for two hours at room temperature the plate was washed again as described above. 100 µl of HRP-streptavidin was then added to each well and the plate was incubated for 20 minutes at room temperature. Once the last wash was removed, 100 µl of substrate solution (hydrogen peroxide+tetramethylbenzidine as a chromagen) was added to each well. Once a sufficient level of color development had occurred, 50 µl of stop solution (2N sulfuric acid) was added to each well and the plate was read at 460 nm.

[0317] After the 24 hour incubation, the tissues were rinsed twice with at least 100 µl of phosphate buffered saline to remove the test material and then transferred to a 6-well plate containing 1.0 ml of assay medium supplemented with MTT (1 mg/ml) and allowed to incubate for 3±0.25 hours at 37±2° C. and 5±1% CO₂. After the incubation, the tissues were rinsed at least twice with 100 µl of phosphate buffered saline, blotted dry, and then placed into a 24-well plate containing 2 ml of isopropanol per well. The 24-well plate was covered and allowed to incubate at room temperature for at least 2 hours on a rocking platform to extract the reduced MTT from the tissues. After the extraction, a 200 µl sample of the isopropanol/MTT mixture was transferred to a 96-well plate and the absorbance of the sample was read at 540 nm with a plate reader using 200 µl of isopropanol as the blank. The MTT assay is described in Example 6 herein. The cell viability results of the MTT assay were similar to the results obtained in Example 6

[0318] The results of the IL-1a assay are shown in Table 9 below. A 2% stock solution of the jellyfish collagen of SEQ ID NO: 91 is Sample 4 and Sample 3 is a 2% stock solution of the truncated jellyfish collagen of SEQ ID NO: 10. In Table 9 below, the indicated percentage is the percent dilution of the stock solution used for the test. For example, the 1% Sample 4 treatment is a 1% solution of the 2% stock truncated collagen solution. The untreated cells produced 18.2 pg/ml of IL-1a. Upon treatment with truncated collagen, all samples showed decreases in IL-1a production. The 1% Sample 4 treatment reduced IL-1A production to 13.4 pg/ml, which is significant with a p value of less than 0.05. The decrease in IL-1a production indicates that the truncated collagen has anti-inflammatory effects.

TABLE 9

IL-1a Assay	
Treatment	IL-1a pg/ml
Non-UVB Exposed	4.9 ± 2.1
Untreated	18.2 ± 1.4
1% Hydrocortisone	6.7 ± 0.6
5% Sample 4	18.1 ± 1.1
1% Sample 4	13.4 ± 0.9*

TABLE 9-continued

IL-1a Assay	
Treatment	IL-1a pg/ml
0.5% Sample 4	15.7 ± 0.7
0.1% Sample 4	13.9 ± 1.7*
5% Sample 3	15.8 ± 2.0

Denotes values that are significantly different from Untreated (p < 0.05)

Example 11: Urban Dust Protection by Truncated Collagen

[0319] A keratinocyte cell culture model was used to assess the ability of truncated collagens to exert a protective effect by promoting cell survival after exposure to urban dust.

[0320] Human epidermal keratinocytes were pretreated with the test materials and then exposed to urban dust. At the end of the treatment period changes in cell viability were then determined via an MTT assay.

[0321] Keratinocytes were seeded into the individual wells of a 96 well plate in 100 µl of medium and incubated overnight at 37±2° C. and 5±1% CO₂. On the following day the media was removed via aspiration to eliminate any non-adherent cells and replaced with 100 µl of fresh medium. The cells were grown until confluent, with a media change every 48 to 72 hours.

Pretreatment with Test Material Followed by Urban Dust Treatment

[0322] Test materials were prepared at 2× their final desired concentrations in cell culture media. Urban dust (NIST 1649B from Sigma Chemicals) was also prepared at 2× solutions. For the pretreatment, 50 µl of 2× test material was combined with 50 µl of culture media and the cells were incubated for 24 hours. At the end of the pretreatment period the test material containing culture media was removed and replaced with 50 µl of 2× urban dust and 50 µl of media. Another set of cells was treated with media alone (non-dust exposed) and used as a reference control to represent 100% cell viability. The cells were then incubated for 24 hours and then subjected to an MTT assay to determine changes in cell viability.

[0323] At the end of the treatment period, the cell culture medium was removed and the cells were washed with PBS. After the wash, 100 µl of cell culture media supplemented with 0.5 mg/ml MTT was added to each well and the cells were incubated for 30 minutes at 37±2° C. and 5±1% CO₂. After the incubation, the media/MTT solution was removed and the cells were washed again once with PBS and then 100 µl of isopropyl alcohol was added to the wells to extract the purple formazin crystals. The 96-well plate was then read at 540 nm using isopropyl alcohol as a blank.

[0324] The mean MTT absorbance value for the non-dust exposed cells was calculated and used to represent 100% value for cell number. The individual MTT values from the cells undergoing the various treatments was then divided by the mean value for the non-dust exposed cells and expressed as a percent to determine the change in cell number caused by each treatment.

[0325] The MTT results for the pretreatment with the test material then dust treatments are presented in Table 10. Table 10 shows that as the cells were treated with increasing amounts of collagen, cell viability increased upon pretreatment with truncated collagen and subsequent exposure to

urban dust. These results show that truncated collagen protects against the decline in cell viability associated with urban dust exposure.

TABLE 10

MTT Assay, Truncated Collagen Pretreatment		
Treatment	Viability (% Non-Dust Exposed) 4 mg/ml Urban Dust	Viability (% Non-Dust Exposed) 2 mg/ml Urban Dust
Non-Dust Exposed	100 ± 4.4*	100 ± 1.8
Untreated	59 ± 3.8	70 ± 0.9
0.1% Collagen	61 ± 4.7	72 ± 3.5
0.5% Collagen	59 ± 2.2	73 ± 2.2
1% Collagen	58 ± 0.5	74 ± 2.6
5% Collagen	66 ± 1.0	83 ± 6.5*

*Denotes values that are significantly different from untreated group (p < 0.05)

Example 12: Truncated *Chondrosia reniformis* (Kidney Sponge) Collagen

[0326] Truncated *Chondrosia reniformis* (Kidney Sponge) Fibrillar Collagen 1 with DsbA Secretion and FLAG Tag

[0327] The amino acid sequence of truncated *Chondrosia reniformis* fibrillar collagen 1 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 102. The DsbA secretion tag is encoded by nucleotides 1-57 of SEQ ID NO: 103 and the amino acid sequences are amino acids 1-19 of SEQ ID NO: 102. The fibrillary collagen nucleotide sequences are nucleotides 58-792 of SEQ ID NO: 103 and the amino acid sequences are amino acids 20-264 of SEQ ID NO: 102. The FLAG nucleotide sequences are nucleotides 793-819 of SEQ ID NO: 103 and the amino acid sequences are amino acids 265-273 of SEQ ID NO: 102.

(SEQ ID NO: 102)

MKKIWLALAGLVLAFSASAPVRRGPKGSRGDPGDGGAAGPKGPEGV
 DGLIGEPGQPGPIGAEGSSGLEFLGDKSGKARGGPGNRGRPGQDG
 VPGQDRAGEKGEGETGDRGQQLRGRKVGDPGLVGLGAQGPQGSQ
 GLVGGPPIGPEPGSGGEPDQGRGPEGPQGSQPGVRGGRGERGTPGA
 VGPKGPPGKNGADGPRGLPGASGPPGSPGNQGPESRGADGNNGFPG
 DDGENGLVGIPEGEPKARGTRGELGKTGDYKDDDDK

[0328] The nucleic acid sequence of truncated *Chondrosia reniformis* fibrillar collagen 1 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 103.

(SEQ ID NO: 103)

ATGAAAAAGATTGGCTGGCGCTGGCTGGTTTATGTTTGGCTTTAG
 CGCATCGCGCGCGTGGTCTGCTGGTCCGAAAGGTAGCCCTGGTG
 ATCCTGGTGATGGTGGTGCAGCAGGTCCTAAAGGTCCGGAAGGTGTT
 GATGGTCTGATTGGTGAACCGGGTCAGCCTGGTCCGATTGGCGCAGA
 AGGTAGCAGCGGTCTGGAAGGTTTTCTGGGTGATAAAGGTAGCAAAG
 GTGCACGTGGTGGTCCGGGTAATCGCGGTGCTCTGGTCCAGGATGGT
 GTTCCGGGTCAAGATGGTCTGCCGGTGAAGAGGTGAAGGTGGTGA

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AACCGGTGATCGCGGTGACGAGGGTCTGCGTGGTAAAGTTGGTGATC
 CAGGCTCTGGTGGGTGATCTGGGTGCACAGGGTCCGACAGGGTAGCCAA
 GGTCTGGTGGTCCGCTGGTATTCGGGTGAACCTGGTAGCGGTGG
 CGAACCGGGTATCAGGGTCTCGCGGTCCAGAAGGTCTCAGGGTT
 CACCGGGTGTTCGCGGTGGTCTGGTGAACCTGGTACACCGGGTGCA
 GTTGACCGAAAGGTCCGCCAGGTAAAAATGGTGCAGATGGTCCCGG
 TGGTCTGCCTGGTGAAGCGGTCTCCGGGTAGTCTGGTAAACAGG
 GTCTGAAGGTTCTCGTGGTCCGATGGTAATAATGGTTTTCCAGGT
 GATGATGGTGAAGTGGCTGGTGGTATCCCTGGCGAACCAGGTCC
 AAAAGGCGCACGCGGTACACGCGGTGAACTGGGTAAAACCGGTGACT
 ACAAGACGACGACGACAAAa

[0329] The polynucleotide of SEQ ID NO: 103 was subcloned into vector pET28a, expressed host *E. coli* cells and the truncated *Chondrosia reniformis* fibrillar collagen 1 was purified as described herein. The purified fibrillary collagen produced a clear band on SDS-PAGE and an anti-FLAG western was observed at around 40 kilodaltons. There were no existing bands that appear at that location on the gel in the absence of expression of this protein.

Truncated *Chondrosia reniformis* (Kidney Sponge) Fibrillar Collagen 2 with DsbA Secretion and FLAG Tag

[0330] The amino acid sequence of truncated *Chondrosia reniformis* fibrillar collagen 2 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 104. The DsbA secretion tag is encoded by nucleotides 1-57 of SEQ ID NO: 105 and the amino acid sequences are amino acids 1-19 of SEQ ID NO: 104. The fibrillary collagen nucleotide sequences are nucleotides 58-1323 of SEQ ID NO: 105 and the amino acid sequences are amino acids 20-441 of SEQ ID NO: 104. The FLAG nucleotide sequences are nucleotides 1324-1350 of SEQ ID NO: 105 and the amino acid sequences are amino acids 442-450 of SEQ ID NO: 105.

(SEQ ID NO: 104)

MKKIWLALAGLVLAFSASAGRGGPAGLQGAAGNPGDPDRGQAGEIG
 LPGTGQRGQGGSRGDDGIGGQSGTDGDPGNDGVAGIRGARGEPGAT
 GPEGAAGQKGRDRGRFGEQGRPGNDGPPGRRGRVGNLGETGAEGDEGT
 RGYTGDRGPEGAIGISVGTGNPQPQGIKPPGDTGHPGRQGPSQPQG
 PPGIPGTDGLTIHNLIKPPSQFFDATSSSDPLTDAVVESILKSFQYA
 ELEIDLTKKPDGTMKYPATSCDDLHKDYPQLPSGNYLDPNGGCKND
 AFETYCEFNNVSKMCLTPKIPLLPMGTYYKYYVNSEGYSPNDFGLN
 LRFFEYGSVTQLKFLQTKATRVQTIRVLCKNYDPLHKQPVFIMGN
 DETVMDEPRMEENQCQYFNGLSAHVELELSSNDPSYLPYEMRLYL
 RKTNEELGIELGDLCFEYGDYKDDDDK

[0331] The nucleic acid sequence of truncated *Chondrosia reniformis* fibrillar collagen 2 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 105.

(SEQ ID NO: 105)
 ATGAAAAAGATTGGCTGGCGCTGGCTGGTTAGTTTTAGCGTTTAG
 CGCATCGGCGGGTCTGGCGGTCCGGCAGGTCTGCAGGGTCTGCAG
 GTAATCCTGGCGACCTGGCGATCGTGGTCAGGCAGGCGAAATTGGT
 CTGCCAGGCACCGAAGGTGAGCGTGGTCAAGTGGTTCACGTGGTGA
 TGACGGTATTGGTGGTCTAGAGCGGCACCGATGGCGATCCGGGTAACG
 ATGGTGTGACAGTATTCTGGTGCACGCGGAGAACCTGGTGCCACC
 GGACCTGAAGGTGCAGCCGGTCAAGAGGTGATCGTGGCCGTTTTGG
 CGAACAGGGTCTCGCGGAAATGATGGTCCACCGGGTCGCGTGGCC
 GTGTGGCAATCTGGGTGAAACAGGTGCCGAAAGTGTGAAGGCACC
 CGTGGTTATACAGGTGACCGTGGACCGGAAGCGCAATTGGTATTAG
 CGGTGTGACCGGTAATCCGGGTCCACAGGGCATTAAAGGCCCTCCGG
 GTGATACGGGTCTCCGGTCTGCAGGACCGAGCGGTCCGCAAGGA
 CCACCGGGTATCCAGGTACAGATGGCCTGACCATTATAATCTGAT
 TAAACCGCCTAGCCAGTTTTTTGATGCAACCAGCAGCAGCGATCCGC
 TGACCGATGCAGTTGTTGAAAGCATTCTGAAATCTTTTCAGTATGCC
 GAGCTGGAAATTGACCTGACCAAAAAACCGGATGGCACCATGAAATA
 TCCGGCAATTAGCTGTGATGATCTGCACAAAGATTATCCGAGCTGC
 CGAGCGGTAATTATACCTGGATCCGAATGGTGGTTGTAATAATGAT
 GCCTTTGAAACCTATTGCGAGTTCAACAATAGCGTGAATAATGTGTCT
 GACCCGAAAAATCCGACACTGCTGCCGATGGGCACCTATAAATACT
 ATGTTAATAGCAGGGTTACTACAGCCGAATGATTTTTGGTCTGAAT
 CTGCGCTTTTTGAGTATTATGGTAGCGTTACCCAGCTGAAATTTCT
 GCAGACCAAAGCAACCCGTGTACCAGACCATTCGTGTTCTGTGTA
 AAACTATGATCCGCTGCATAAACAGCCGGTTTTTATTGGTATGAAT
 GACGAAACCGTTATGGATGAACCGGTATGGAAGAAAATCAGTGCCA
 GTATTTAACGGTCTGAGCGCACATGTTGAACTGGAACGTAGCAGCA
 ATGATCCGAGCTATCTGCCGATTTATGAAATGCGTCTGTATCTGGGT
 CGTAAACCAATGAAGAACTGGGCATTGAACTGGCGATCTGTGTTT
 TGAATATGGTGACTACAAAGACGACGACGACAAAaa

[0332] The polynucleotide of SEQ ID NO: 105 was sub-cloned into vector pET28a, expressed host *E. coli* cells and the truncated *Chondrosia reniformis* fibrillar collagen 2 was purified as described herein. The purified fibrillary collagen produced a clear band on SDS-PAGE and an anti-FLAG western was observed at around 55 kilodaltons. There were no existing bands that appear at that location on the gel in the absence of expression of this protein.

Truncated *Chondrosia reniformis* (Kidney Sponge) Non-Fibrillar Collagen 1 with DsbA Secretion and FLAG Tag

[0333] The amino acid sequence of truncated *Chondrosia reniformis* non-fibrillar collagen 1 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 106. The DsbA secretion tag is encoded by nucleotides 1-57 of SEQ ID NO: 107 and the amino acid sequences are amino acids 1-19 of SEQ ID NO: 106. The non-fibrillar collagen nucleotide

sequences are nucleotides 58-831 of SEQ ID NO: 107 and the amino acid sequences are amino acids 20-277 of SEQ ID NO: 106. The FLAG nucleotide sequences are nucleotides 832-858 of SEQ ID NO: 107 and the amino acid sequences are amino acids 278-286 of SEQ ID NO: 106.

(SEQ ID NO: 106)
 MKKIWLALAGLVLAFSASAETSSKVALMTVLVVITGALIIIEGTSIT
 RGSTHVNRLRKRQTSNEDNCEAVKVLPRDRGREGPPPGPPAGRDRG
 RDAVCSNQTTGLGAKGDRPPGTPGPFPEVGRPGPPGADGIPGPQGE
 RGAVPGGKPGPRGEVGTGADGADGATGATGVQVQPDGAKGKAGSAG
 TAGLKGEKGDTCIPDSNSTLGMPTGAGGSKGQKGESGIVGPKGER
 GEIGTPGHGPRGADGEPGHKGVPRAGAQQDRGDPGDGLTGDYDK
 DDDK

[0334] The nucleic acid sequence of truncated *Chondrosia reniformis* non-fibrillar collagen 1 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 107.

(SEQ ID NO: 107)
 ATGAAAAAGATTGGCTGGCGCTGGCTGGTTAGTTTTAGCGTTTAG
 CGCATCGGCGGAAAAAACAGCAGCAAAGTTGCACTGATGACCGTTT
 TGTTGTTATTACCAGGTGCACTGATTATTGAAGGCACAGCATTACC
 CGTGGTAGCACCCATGTTAATCGTGGTCTGCGTAAACGTGACACCAG
 CGAAGATAATTGTGAAGCAGTTAAAGTTGGTCTGCCAGGTCTGTGATG
 GTCGTGAAGGTCCCTCCGGTCCCGCTGGTCCGGCTGGCAGAGATGGC
 CGTGTGCAAGTTTGTAGCAATCAGACCACCGGTCTGGTGCAAAAGG
 TGATCGTGGTCCGCCAGGTACACCGGGTTTTCCGGGTGAAGTTGGCC
 GTCCGGTCCACCGGTGAGATGTTATCCGGTCTCAGGGTGAA
 CGTGGTGCAGTTGGTCTGGTGGTAAACCTGGTCCCGCTGGTGAAGT
 GGGCACCCCTGGTCCGATGGCGCAGATGGTGAACCGGTGCGACCG
 GTGTTCAAGGTCTGATGGTGCACAAAGGCGAAAAAGGTGAAGCGGC
 ACCGCAAGTCTGAAAGGTGAGAAAGCGATACCTGTATTCCGGATAG
 CAATAGCACCCCTGGGTATGCCTGGTACACAGGTGCCGGTGGTAGCA
 AAGGCCAGAAAGGTGAAAGTGGTATTGTTGGTCCGAAAGGCGAACGC
 GGTGAAATTGGCACACCGGGTCTCCTGGTTTTCTGGTGGCGGATGG
 TGAACAGGTGATGAAAGGTGTTCCGGGTGTCGGGTGCGCAGGGTG
 ATCGCGGTGATCCGGGTGATGATGGTCTGACCGGTGACTACAAAGAC
 GACGACGACAAAaa

[0335] The polynucleotide of SEQ ID NO: 107 was sub-cloned into vector pET28a, expressed host *E. coli* cells and the truncated *Chondrosia reniformis* non-fibrillar collagen 1 was purified as described herein. The purified non-fibrillar collagen produced a clear band on SDS-PAGE and an anti-FLAG western was observed at around 30 kilodaltons. There were no existing bands that appear at that location on the gel in the absence of expression of this protein. Truncated *Chondrosia reniformis* (Kidney Sponge) Non-Fibrillar Collagen 2 with DsbA Secretion and FLAG Tag

[0336] The amino acid sequence of truncated *Chondrosia reniformis* non-fibrillar collagen 2 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 108. The DsbA secretion tag is encoded by nucleotides 1-57 of SEQ ID NO: 109 and the amino acid sequences are amino acids 1-19 of SEQ ID NO: 108. The non-fibrillar collagen nucleotide sequences are nucleotides 58-1509 of SEQ ID NO: 109 and the amino acid sequences are amino acids 20-503 of SEQ ID NO: 108. The FLAG nucleotide sequences are nucleotides 1510-1536 of SEQ ID NO: 109 and the amino acid sequences are amino acids 504-512 of SEQ ID NO: 108.

(SEQ ID NO: 108)
 MKKIWLALAGLVLAFSASAGPFPAGADGAPGQKQKELGAVGPQGT
 LSGPSGPTGPPGPKGVRGAPGSSGAKGDAGNPGDDGPVGPQGV
 GSPGQKGETGRVGRHGDINGTPGEDGATGFPDPGAKGKGTSGT
 AGLKGEKGDTCIPDNSLGLMPGTPGAGWSKQKGESGIVGPKGEK
 EIGTPGPPGFRGADGEPGQGRGEPGRAGAQRGERGAPGNNGRDFP
 GADGAPGQKQKELGALGHPGFSGPSGSPGTPPPGPKGVRGAQGR
 RGSPPDVGPIGAPGPPGADGVPGLTGVQQRDGPKGESASSGAVY
 RVRTTTPSGADVVSRAAGAKYDHSGGTSDHHCLPNNPQYLS
 EDNTN ALGAQLYGVVEYIHDRSSPYNSLDQSDMPCVVCNAN
 GRSQLMLVPAR YTCPTGWSREYYGYMSEKAKNREGRKTTIC
 MDFSABEAVPGSGANT NPSPGIMMRANGLACPPYQSNPLT
 CAVCTKGDYKDDDDK

[0337] The nucleic acid sequence of truncated *Chondrosia reniformis* non-fibrillar collagen 2 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 109.

(SEQ ID NO: 109)
 ATGAAAAAGATTGGCTGGCGCTGGCTGGTTTAGTTTTAGCGTT
 TAG CGCATCGGCGGGTTTTCTCGCGCTCCGGGTGCCGACGGT
 GCTCCGGTGTCAAAAAAGGTGAACTGGGTGCCGTGGGTCCG
 CAGGGCACTCCGGGTCTGAGTGGTCTAGTGGTCCGACCGG
 TCCACAGGTCCAAAAAGGCGT GCGTGGTGACCCGGTAGC
 AGCGGACCAAGGTGATGACGGTAACC CTGGTGATGACGG
 TCCGGTCCGACAGGGCGTTCAGGTGTGAT GGTAGCCCTG
 GCAAAAGGGTGAACCGGTCTGTGGTCTCGTGC TCATGAT
 GGTATTAATGGCACCCAGGTGAAGATGGTGCACAGGCT
 TTCCAGGTCCGGATGGCGCAAAGGGTGAGAAGGGCACCA
 GCGGTACA GCTGGCTGAAGGGCGAAAAGGGCGATACATG
 CATCCCGGATCAAA TTCAACACTGGGCATGCCAGGTACG
 CCTGGCGCAGGTTGGAGTAAAG GACAAAAAGGCGAATC
 AGGCATTTGTGGGACCTAAAGCGGAGAAGGGT GAGATT
 GGTACTCCGGGACCGCCAGGCTTTTCGGGTGCAGACGGC
 GA ACCGGTTCAGCGTGGCGAACCTGGTCTGTGACGGCC
 ACAGGTGAAC GCGGAGCCCTGGTAATAATGGACGTGAT
 GGCTTTCTGGTGTATCCA GGTGCAGATGGCGCACCTG
 GGCAGAAAGGCGAACCTGGGAGCAATTGG TCATCCGGG
 ATTTAGCGGTCCGTGACGGTCCGACGGACCGCACAGGCT

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CTCCTGGACCGAAAGGTGTACGTGGCGCACAGGGTCGCTGGCGAT
 CGTGGCAGTCCAGGTGATGTGGGTCCGATTGGTGACCTGGTCTCC
 AGGTGCGGACGGCGTGCCTGGTTTAAACAGGTGTGCAGGGTCCG
 CAGC GTCCAAAAGGTGAATCAGCAAGCAGCGGTGCAGTTTATG
 TTGTTGTTG GGTCTGACCACTGTCTTAGCGGAGCAGATGTGTT
 TATAGCGGTCCG CAGCGCGGTGCAAAATATGATCATTGAGT
 GGCACCTCAGATCATC ATTGTCTGCCGAATAATCCGAGTAT
 CTGAGCGAAGATGATACCAAT GCACTGGGTGCACAGCTGT
 ATGGTGTGGAATGAAATTCGTGATCG TAGCAGCCCGTATA
 ATAGCCTGGATCAGAGCGATATGCCGTGTGTTG TTTGTA
 ATGCAAATGGTCGTAGCCAGCTGCTGATGGTTCCGGCACGT
 TATACATGCCCCACCGGTGGAGCCGTGAATATTATGGTTAT
 ATGAT GAGCGAAGGCAAAAGCCAAAAATCGCGAAGGTCTG
 TAAAACCACCATTT GTATGGATTTTAGCGCAGAAGCAGTT
 CCTGGTAGCGGTGCAAAATACC AATCCGAGTCCGGGTAT
 TATGATGCGTGCAAATGTAATGGTCTGGC ATGTCCGCCT
 TATCAGAGCAATACACCGCTGACCTGTGCCGTTTGTGA
 CCAAAGGTGACTACAAGACGACGACGACAAATaa

[0338] The polynucleotide of SEQ ID NO: 109 was subcloned into vector pET28a, expressed host *E. coli* cells and the truncated *Chondrosia reniformis* non-fibrillar collagen 2 was purified as described herein. The purified fibrillar collagen produced a clear band on SDS-PAGE and an anti-FLAG western was observed at around 60 kilodaltons. There were no existing bands that appear at that location on the gel in the absence of expression of this protein.

Example 13: Truncated *Rhincodon typus* (Whale Shark) Collagen

[0339] Truncated *Rhincodon typus* (Whale Shark) Collagen Type 1 Alpha 1 Truncation 1 with DsbA Secretion and FLAG Tag

[0340] The amino acid sequence of truncated *Rhincodon typus* collagen type 1 truncation 1 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 110. The DsbA secretion tag is encoded by nucleotides 1-57 of SEQ ID NO: 111 and the amino acid sequences are amino acids 1-19 of SEQ ID NO: 110. The collagen nucleotide sequences are nucleotides 58-630 of SEQ ID NO: 111 and the amino acid sequences are amino acids 20-210 of SEQ ID NO: 110. The FLAG nucleotide sequences are nucleotides 631-657 of SEQ ID NO: 111 and the amino acid sequences are amino acids 211-219 of SEQ ID NO: 110.

(SEQ ID NO: 110)
 MKKIWLALAGLVLAFSASAGPAGKGPSGDI GRPGESGSPGARGHSG
 QPRTGIAGNQGLPGTAGEEGRTPPPGAPLRGQAGMMFPKGA
 A GLPGKPGDRGNVGLAGPRGAPKDGVEVGAQPPGVAGPTGPR
 GETGL AGSVGFQGMPPSGAAGEPGKPGNQGLRGDAGSPGMI
 GPRGERGLPG ERGASGAQGLLGRGTS GAPGLGDKDDDDK

[0341] The nucleic acid sequence of truncated *Rhincodon typus* collagen type 1 truncation 1 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 111.

(SEQ ID NO: 111)

ATGAAAAAGATTTGGCTGGCGCTGGCTGGTTAGTTTTAGCGTTTATG
 CGCATCGGCGGGTCCGGCAGGCGCAAAAGGTCGAGCGGTGATATTG
 GTCGTCGGGTGAAAAGCGGTAGTCCGGGTGCACGTGGTCATAGCGGT
 CAGCCTGGTCTGACCGGTATTGCAGGTAATCAGGGTCTGCCTGGTAC
 AGCCGGTGAAGAAGGTCGCACCGGTCCGCCAGGTCTGCAGGTCTGC
 GTGGTCAGGCAGGTATGATGGTTTTCCGGGTCCGAAAGGTGCAGCG
 GGTCTGCCAGGCAAACCGGGTATCGTGGTAATGTTGGTCTGGCTGG
 TCCGCGTGGTGACCCGGTAAAGATGGTGAAGTTGGTGACAGGGTC
 CTCCGGGTGTTGCAGGTCCGACCGGTCTCGTGGTGAACCGGTCTG
 GCAGGTAGCGTTGGTTTTCAGGGTATGCCAGGTCCGTCAGGTGCAGC
 AGGCGAACCTGGTAAACCGGGTAAACAGGGCCTGCGTGGTATGCCG
 GTTCCACCGGTATGATTGGTCCACGCGGTGAACGTGGCTGCCTGGC
 GAACGTGGTGAAGCGGTGCACAAGGTCTGCTGGTCCACGTGGCAC
 CTCAGGCGCACCGGTCTGGGTGACTACAAAGACGACGACGACAAAT
 aa

[0342] The polynucleotide of SEQ ID NO: 111 was subcloned into vector pET28a, expressed host *E. coli* cells and the truncated *Rhincodon typus* collagen type 1 truncation 1 was purified as described herein. The purified collagen produced a clear band on SDS-PAGE and an anti-FLAG western was observed at around 25 kilodaltons. There were no existing bands that appear at that location on the gel in the absence of expression of this protein.

Truncated *Rhincodon typus* (Whale Shark) Collagen Type 6 Alpha 1 Truncation 2 with DsbA Secretion and FLAG Tag

[0343] The amino acid sequence of truncated *Rhincodon typus* collagen type 6 truncation 2 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 112. The DsbA secretion tag is encoded by nucleotides 1-57 of SEQ ID NO: 113 and the amino acid sequences are amino acids 1-19 of SEQ ID NO: 112. The collagen nucleotide sequences are nucleotides 58-684 of SEQ ID NO: 113 and the amino acid sequences are amino acids 20-228 of SEQ ID NO: 112. The FLAG nucleotide sequences are nucleotides 685-711 of SEQ ID NO: 113 and the amino acid sequences are amino acids 229-237 of SEQ ID NO: 112.

(SEQ ID NO: 112)

MKKIWLALAGLVLAFAASAAQGIPIGASAGKEGGKGDPLGSPGKPGPD
 GLRGFAGARGLPGAAGPPGLKGAEGPMGAPGLTGSTGERPMPGPA
 IGLPGRPGGPPGPPVGEKGDPGDKLPGPAGDDGVQGMGLPGPIG
 SQGPPGDYGDKGELGKPKQKSKGDKGESGPPPIGIQGPVIGHPGPI
 GSDGSPGLRGLYLMRQKQKDDGIRGLPGSAGPVGLQGLPGGDYKDD
 DK

[0344] The nucleic acid sequence of truncated *Rhincodon typus* collagen type 6 truncation 2 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 113.

(SEQ ID NO: 113)

ATGAAAAAGATTTGGCTGGCGCTGGCTGGTTAGTTTTAGCGTTTATG
 CGCATCGGCGAGGGTATTCCGGGTAGCGCAGGTAAGAAGGTTGGTA
 AAGGCGATCCGGGTCCGGTGGTTCCACCGGTAAACCGGGTCTGAT
 GGCTGCGTGGTTTTGCGGTGCACGTGGTCTGCCTGGTGCAGCAGG
 TCCGCTGGTCTGAAAGGTGCCGAAGGTCCGATGGGTCTCCGGGT
 TGACCGGTAGCACCGGTGAACCGGTCCGAATGGTCCGGCAGGCGCA
 ATTGGTCTGCCAGGTCTGCTGGTGGTCCGGTCTCTCTGGTCCGGT
 TGGTGAAAAAGGTGATCCTGGTGATAAAGGCTGCCTGGTCTGCCG
 GTGATGATGGTGTTCAGGGTCCCATGGGCTTACCGGGTCCGATTGGT
 AGCCAGGGTCTCCGGGTGATTATGGCGATAAAGGTGAACTGGGTAA
 ACCTGGCCAGAAAGGTAGCAAAGGTGACAAAGGCGAAAGCGGTCCGC
 CAGGTCCGATCGGCATTAGGGTCTATTGGTCAATCCAGGTCCAATT
 GGTTCAGATGGCTCACCGGGACTGCGTGGCTATCTGGGTATGCGTGG
 ACAGAAAGGTGATGACGGTATTCGTGGCTGCCAGGTAGTCAGGTC
 CGGTGGTCTGCAGGGACTGCCTGGTGGTACTACAAAGACGACGAC
 GACAAAtaa

[0345] The polynucleotide of SEQ ID NO: 113 was subcloned into vector pET28a, expressed host *E. coli* cells and the truncated *Rhincodon typus* collagen type 6 truncation 2 was purified as described herein. The purified collagen produced a clear band on SDS-PAGE and an anti-FLAG western was observed at around 35 kilodaltons. There were no existing bands that appear at that location on the gel in the absence of expression of this protein.

Truncated *Rhincodon typus* (Whale Shark) Collagen Type 6 Alpha 1 Truncation 3 with DsbA Secretion and FLAG Tag

[0346] The amino acid sequence of truncated *Rhincodon typus* collagen type 6 alpha 1 truncation 3 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 114. The DsbA secretion tag is encoded by nucleotides 1-57 of SEQ ID NO: 115 and the amino acid sequences are amino acids 1-19 of SEQ ID NO: 114. The collagen nucleotide sequences are nucleotides 58-735 of SEQ ID NO: 115 and the amino acid sequences are amino acids 20-245 of SEQ ID NO: 114. The FLAG nucleotide sequences are nucleotides 736-762 of SEQ ID NO: 115 and the amino acid sequences are amino acids 246-254 of SEQ ID NO: 114.

(SEQ ID NO: 114)

MKKIWLALAGLVLAFAASAKGETGEAGDPGTPGEPGIAGPKGDVGDG
 GDAGPPGAAGPAGVKGPPGEDGAKGDVGPAGFPDGPPTGEPVPGM
 DGGVGEKGSGLDPGLTGPARGASGEPGPPSPGKRGPPGAPGREG
 LKGSKSGSPQEGPVGRTGPIGPQSGPNVGPGLRGIPTGEGQLL
 GPPGQAGPPGMPGMPGLRGAQGLKGDKGHVGLIIGLIPPGEMGE
 KGDQGLPGIQDYGKDDDDK

[0347] The nucleic acid sequence of truncated *Rhincodon typus* collagen type 6 alpha 1 truncation 3 with DsbA secretion and FLAG tag is disclosed in SEQ ID NO: 115.

(SEQ ID NO: 115)
 ATGAAAAAGATTGGCTGGCGCTGGCTGGTTTAGTTTAGCGTTTAG
 CGCATCGGCGAAAGGTGAAACCGGTGAAGCGGGTGATCCGGGTACAC
 CGGGTGAACCTGGTATTGCAGGTCCGAAAGGTGATGTTGGTGATAAA
 GGTGACGCAGGTCCGCCTGGTGCAGCAGGTCCGGCAGGCGTTAAAGG
 TCCTCCGGGTGAAGATGGTGCAAAAGGCGACGTTGGTCTGCAGGTT
 TTCCTGGCGATCCGGGTCCGACTGGTGAACCGGGTGTGCCAGGTATG
 GATGGTGGTGTGGGTGAAAAAGGTAGCCTGGGTGATCCTGGTCTGAC
 CGGTCCGCGTGGCGCAAGTGGTGAACCAAGTCCACCGGGTAGTCCGG
 GTAAACGTGGTCTCTCGGACCGGCTGGTCCGGAAGGTCGTGAAGGT

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 CTGAAAGGTAGCAAAGGTTACCCGGGTCAAGAAGGTCCGGTTGGTCCG
 TACCCGGTCCGATTGGTCCGCGAGGGCTCACCCGGTAATGTTGGTCTTA
 AAGGTCTCGTGGTATTCCGGGTCTACAGCGCAACAGGGTCTGCTG
 GGTCGCCAGGCCAAGCAGGTCTCCAGGTCTATGGGTCCACCTGG
 TATGCTTGGCTGCGTGGTGCACAGGGCTGAAAGCGGATAAAGGCC
 ATGTTGGTCTGATTGGCCTGATTGGTCCACCAGGTGAAATGGGAGAA
 AAAGCGATCAGGGCCTGCCTGGTATTGAGGGTACTACAAGACGA
 CGACGACAAAtaa

[0348] The polynucleotide of SEQ ID NO: 115 was subcloned into vector pET28a, expressed host *E. coli* cells and the truncated *Rhincodon typus* collagen type 1 truncation 1 was purified as described herein. The purified collagen produced a clear band on SDS-PAGE and an anti-FLAG western was observed at around 25 kilodaltons. There were no existing bands that appear at that location on the gel in the absence of expression of this protein.

SEQUENCE LISTING

<160> NUMBER OF SEQ ID NOS: 153

<210> SEQ ID NO 1

<211> LENGTH: 429

<212> TYPE: PRT

<213> ORGANISM: Podocoryna carnea

<400> SEQUENCE: 1

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 Glu Lys Gly Glu Gln Gly Arg Thr Gly Ala Ala Gly Lys Gln Gly Ser
 20 25 30
 Pro Gly Ala Asp Gly Ala Arg Gly Pro Leu Gly Ser Ile Gly Gln Gln
 35 40 45
 Gly Ala Arg Gly Glu Pro Gly Asp Pro Gly Ser Pro Gly Leu Arg Gly
 50 55 60
 Asp Thr Gly Leu Ala Gly Val Lys Gly Val Ala Gly Pro Ser Gly Arg
 65 70 75 80
 Pro Gly Gln Pro Gly Ala Asn Gly Leu Pro Gly Val Asn Gly Arg Gly
 85 90 95
 Gly Leu Arg Gly Lys Pro Gly Ala Lys Gly Ile Ala Gly Ser Asp Gly
 100 105 110
 Glu Ala Gly Glu Ser Gly Ala Pro Gly Gln Ser Gly Pro Thr Gly Pro
 115 120 125
 Arg Gly Gln Arg Gly Pro Ser Gly Glu Asp Gly Asn Pro Gly Leu Gln
 130 135 140
 Gly Leu Pro Gly Ser Asp Gly Glu Pro Gly Glu Glu Gly Gln Pro Gly
 145 150 155 160
 Arg Ser Gly Gln Pro Gly Gln Gln Gly Pro Arg Gly Ser Pro Gly Glu
 165 170 175
 Val Gly Pro Arg Gly Ser Lys Gly Pro Ser Gly Asp Arg Gly Asp Arg
 180 185 190
 Gly Glu Arg Gly Val Pro Gly Gln Thr Gly Ser Ala Gly Asn Val Gly
 195 200 205

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Glu	Asp	Gly	Glu	Gln	Gly	Gly	Lys	Gly	Val	Asp	Gly	Ala	Ser	Gly	Pro
210						215					220				
Ser	Gly	Ala	Leu	Gly	Ala	Arg	Gly	Pro	Pro	Gly	Ser	Arg	Gly	Asp	Thr
225					230					235					240
Gly	Ala	Val	Gly	Pro	Pro	Gly	Pro	Thr	Gly	Arg	Ser	Gly	Leu	Pro	Gly
				245					250					255	
Asn	Ala	Gly	Gln	Lys	Gly	Pro	Ser	Gly	Glu	Pro	Gly	Ser	Pro	Gly	Lys
			260					265					270		
Ala	Gly	Ser	Ala	Gly	Glu	Gln	Gly	Pro	Pro	Gly	Lys	Asp	Gly	Ser	Asn
		275					280					285			
Gly	Glu	Pro	Gly	Ser	Pro	Gly	Lys	Glu	Gly	Glu	Arg	Gly	Leu	Ala	Gly
	290						295				300				
Pro	Pro	Gly	Pro	Asp	Gly	Arg	Arg	Gly	Glu	Thr	Gly	Ser	Pro	Gly	Ile
305					310					315					320
Ala	Gly	Ala	Leu	Gly	Lys	Pro	Gly	Leu	Glu	Gly	Pro	Lys	Gly	Tyr	Pro
				325					330					335	
Gly	Leu	Arg	Gly	Arg	Asp	Gly	Thr	Asn	Gly	Lys	Arg	Gly	Glu	Gln	Gly
			340					345					350		
Glu	Thr	Gly	Pro	Asp	Gly	Val	Arg	Gly	Ile	Pro	Gly	Asn	Asp	Gly	Gln
		355					360					365			
Ser	Gly	Lys	Pro	Gly	Ile	Asp	Gly	Ile	Asp	Gly	Thr	Asn	Gly	Gln	Pro
	370					375					380				
Gly	Glu	Ala	Gly	Tyr	Gln	Gly	Gly	Arg	Gly	Thr	Arg	Gly	Gln	Leu	Gly
385					390					395					400
Glu	Thr	Gly	Asp	Val	Gly	Gln	Asn	Gly	Asp	Arg	Gly	Ala	Pro	Gly	Pro
				405					410					415	
Asp	Gly	Ser	Lys	Gly	Ser	Ala	Gly	Arg	Pro	Gly	Leu	Arg			
			420					425							

<210> SEQ ID NO 2
 <211> LENGTH: 1289
 <212> TYPE: DNA
 <213> ORGANISM: Podocoryna carnea

<400> SEQUENCE: 2

ggaccacaag	gtgtttagg	agctgatggc	aaagatggaa	caccgggaga	gaaaggtgag	60
caaggacgaa	ccggagctgc	aggaaaacag	ggaagccctg	gagcagatgg	agcaagaggc	120
cctcttgat	caattggaca	acaagtgct	cgtggagaac	ctggtgatcc	aggatctccc	180
ggcttaagag	gagatactgg	attggctgga	gtcaaaggag	tagcaggacc	atctggtcga	240
cctggacaac	ccggtgcaaa	tggattacct	ggtgtgaatg	gcagaggcgg	tttgagaggc	300
aaacctggtg	ctaaaggaat	tgctggcagt	gatggagaag	cgggagaatc	tggcgcacct	360
ggacagtcog	gacctaccgg	tccacgtggt	caacgaggac	caagtggatg	ggatggtaat	420
cctggattac	agggattgcc	tggttctgat	ggagagcccc	gagaggaagg	acaacctgga	480
agatctggtc	aaccaggaca	gcaaggacca	cgtggttccc	ctggagaggt	aggaccaaga	540
ggatctaaag	gtccatcagg	agatcgtggt	gacaggggag	agagaggtgt	tcctggacaa	600
acaggttcgg	ctgaaatgt	aggagaagat	ggagagcaag	gaggcaaagg	tgtcgatgga	660
gcgagtggac	caagtggagc	tcttggtgct	cgtggtcccc	caggaagtag	aggtgacacc	720
ggggcagtgg	gacctccogg	acctactggg	cgatctggtt	tacctggaaa	cgcaggacaa	780

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aagggaccaa	gtggtgaacc	aggtagtcca	ggaaaagcag	gatcagctgg	tgaacagggt	840
cctcctggta	aagacggatc	aatggtgaa	cctggatctc	ctggcaaaga	gggtgaacgt	900
ggtcttgctg	gtccaccagg	tccagatggc	agacgtggtg	aaacgggatc	tccaggtatc	960
gctggtgctc	ttggtaaacc	aggtttggaa	ggacctaaag	gttatccagg	attaagagga	1020
agagatggaa	ccaatggcaa	acgaggagaa	caaggagaaa	ctggtcctga	tggagtcaga	1080
ggtattcctg	gaaatgatgg	acaatctggc	aaaccaggta	ttgatgggat	tgacggaaca	1140
aatggtcaac	caggtgaggc	tggataccaa	ggtggtagag	gtacacgtgg	tcagttaggt	1200
gaaactggtg	atgtcggaca	gaatggagat	cgaggagctc	ctggtcctga	tggatctaaa	1260
ggttctgctg	gtagaccagg	acttctggtg				1289

<210> SEQ ID NO 3

<211> LENGTH: 1425

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 3

atgaaaaaga	tttggtggc	gctggctggt	ttagttag	cgtttagcgc	atcggcggcg	60
cagtatgaag	atcaccatca	ccaccaccac	catcaccact	ctggctcgag	cctggtgccg	120
cgcggcagcc	atatgggtcc	gcaggggtgt	gttgggtgag	atggtaaaga	cggtaccccg	180
ggtgaaaaag	gagaacaggg	acgtacaggt	gcagcaggtg	aacagggcag	cccgggtgcc	240
gatggtgccc	gtggcccgc	gggtagcatt	ggtcagcagg	gtgcaagagg	cgaaccgggc	300
gatccgggta	gtccgggct	gcgtggtgat	acgggtctgg	ccggtgtaa	aggcggtgca	360
ggtccttcag	gtcgtccagg	tcaaccgggt	gcaaatggtc	tgccgggtgt	taatggtcgt	420
ggcggctctg	gtggcaaac	gggagcaaaa	ggtattgcag	gtagcagtg	agaagccggt	480
gaaagcggtg	ccccgggtca	gagtgggtcc	accggtccgc	gcggtcagcg	tggctccgtct	540
ggtgaagatg	gcaatccggg	tctgcaggg	ctgcctggta	gtgatggcga	accaggtgaa	600
gaaggtcagc	cgggtcgttc	aggccagccg	ggccagcagg	gcccgcgtgg	tagcccgggc	660
gaagttggcc	cgcggggtag	taaaggtcct	agtggcgatc	gcggtgatcg	tgggtgaaacg	720
ggtgttcctg	gtcagaccgg	tagcgcaggt	aatgttggcg	aagatggtga	acaggggtggc	780
aaaggtgttg	atggtgcaag	cggtccgagc	ggtgcactgg	gtgcacgtgg	tcctccgggc	840
agccgtggtg	acaccgggtg	agttggtccg	cctggcccga	ccggccgtag	tggcttaccg	900
ggtaatgcag	gtcagaaaag	tccgtcaggt	gaacctggca	gccctggtaa	agcaggtagt	960
gccggtgagc	aggggtccgc	gggcaaaagat	ggtagtaatg	gtgagccggg	tagccctggc	1020
aaagaaggtg	aacgtggtct	ggcaggaccg	ccgggtcctg	atggtcgccg	cggtgaaacg	1080
ggttcaccgg	gtattgcggg	tgccctgggt	aaaccaggtc	tggaaggtcc	gaaaggttat	1140
cctggtctgc	gcggtcgtga	tggtaccaat	ggcaaacgtg	gcgaacaggg	cgaaacccggt	1200
ccagatggtg	ttcgtggtat	tccgggtaac	gatggtcaga	gcggtaaacc	gggcattgat	1260
ggtattgatg	gcaccaatgg	tcagcctggc	gaagcaggtt	atcaggggtg	tcgcggtacc	1320
cgtggtcagc	tgggtgaaac	aggtgatggt	ggtcagaatg	gtgatcgccg	cgcaccgggt	1380

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 ccggatggta gcaaaggtag cgccggtcgt ccgggtttac gttaa 1425

<210> SEQ ID NO 4
 <211> LENGTH: 1425
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polynucleotide

<400> SEQUENCE: 4

atgaaaaaga tttggctggc gctggctggt ttagttttag cgtttagcgc atcgcgcgcg 60
 cagtatgaag atcaccatca ccaccaccac catcaccact ctggctcgag cctggtgccg 120
 cgcgcgagcc atatgggtcc gcagggtggt gttggtgcag atggtaaaga cggtaccccc 180
 ggtgaaaaag gtgaacaggg tcgtaccggt gcagcaggt aacagggcag cccgggtgcc 240
 gatggtgccc gtggcccgtc gggtagcatt ggtcagcagg gtgcacgtgg cgaaccgggc 300
 gatccgggta gcccgggcct gcgtggtgat acgggtctcg ccggtgttaa aggcgttgca 360
 ggtccttctg gtcgtccagg tcaaccgggt gcaaatggtc tgccgggtgt taatggtcgt 420
 ggcggtctgc gtggcaaac gggtgcaaaa ggtattgcag gtagcgatgg cgaagccggt 480
 gaaagcggtg ccccggttca gagcggtcgc accggtcgcg gcggtcagcg tggtcctctc 540
 ggtgaagatg gcaatccggg tctgcagggt ctgcctggta gcgatggcga accaggtgaa 600
 gaaggtcagc cgggtcgttc tggccagcgc gcccagcagg gcccgctgg tagcccgggc 660
 gaagttggcc cgccgggttc taaaggtcct agcggcgcgc gcggtgatcg tggatgaacgc 720
 ggtgttccctg gtcagaccgg tagcgcaggt aatgttggcg aagatggtga acaggggtggc 780
 aaagtggtg atggtgcaag cggtcgcgagc ggtgcactgg gtgcacgtgg tcctccgggc 840
 agccgtggtg acaccggtgc agttggtccg cctggcccga ccggccgtag cggcctgccc 900
 ggtaatgcag gtcagaaaagg tccgtctggt gaacctggca gccctggtaa agcaggttagc 960
 gccggtgagc agggtcgcc gggcaaatg ggtagcaatg gtgagccggg tagccctggc 1020
 aaagaaggtg aacgtgtctc ggcaggtcgc ccgggtcctg atggtcgcgc cggatgaaac 1080
 ggttctccgg gtattgccgg tgccctgggt aaaccaggtc tggaaggtcc gaaaggttat 1140
 cctggtctgc gcggtcgtga tggtagcaat ggcaaacgtg gcgaacaggg cgaaacccggt 1200
 ccagatggtg ttcgtggtat tccgggtaac gatggtcaga gcggtaaacc gggcattgat 1260
 ggtattgatg gcaccaatgg tcagcctggc gaagcaggtt atcaggggtg tcgcggtacc 1320
 cgtggtcagc tgggtgaaac cggtgatggt ggtcagaatg gtgatcgcgc cgcaccgggt 1380
 ccggatggta gcaaaggtag cgccggtcgt ccgggtctgc gttaa 1425

<210> SEQ ID NO 5
 <211> LENGTH: 474
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polypeptide

<400> SEQUENCE: 5

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
 1 5 10 15
 Ala Ser Ala Ala Gln Tyr Glu Asp His His His His His His His

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20			25			30									
His	Ser	Gly	Ser	Ser	Leu	Val	Pro	Arg	Gly	Ser	His	Met	Gly	Pro	Gln
	35						40					45			
Gly	Val	Val	Gly	Ala	Asp	Gly	Lys	Asp	Gly	Thr	Pro	Gly	Glu	Lys	Gly
	50						55				60				
Glu	Gln	Gly	Arg	Thr	Gly	Ala	Ala	Gly	Lys	Gln	Gly	Ser	Pro	Gly	Ala
	65				70					75					80
Asp	Gly	Ala	Arg	Gly	Pro	Leu	Gly	Ser	Ile	Gly	Gln	Gln	Gly	Ala	Arg
				85						90					95
Gly	Glu	Pro	Gly	Asp	Pro	Gly	Ser	Pro	Gly	Leu	Arg	Gly	Asp	Thr	Gly
		100						105						110	
Leu	Ala	Gly	Val	Lys	Gly	Val	Ala	Gly	Pro	Ser	Gly	Arg	Pro	Gly	Gln
		115						120						125	
Pro	Gly	Ala	Asn	Gly	Leu	Pro	Gly	Val	Asn	Gly	Arg	Gly	Gly	Leu	Arg
	130						135				140				
Gly	Lys	Pro	Gly	Ala	Lys	Gly	Ile	Ala	Gly	Ser	Asp	Gly	Glu	Ala	Gly
	145				150						155				160
Glu	Ser	Gly	Ala	Pro	Gly	Gln	Ser	Gly	Pro	Thr	Gly	Pro	Arg	Gly	Gln
				165						170					175
Arg	Gly	Pro	Ser	Gly	Glu	Asp	Gly	Asn	Pro	Gly	Leu	Gln	Gly	Leu	Pro
		180						185							190
Gly	Ser	Asp	Gly	Glu	Pro	Gly	Glu	Glu	Gly	Gln	Pro	Gly	Arg	Ser	Gly
		195					200						205		
Gln	Pro	Gly	Gln	Gln	Gly	Pro	Arg	Gly	Ser	Pro	Gly	Glu	Val	Gly	Pro
	210						215				220				
Arg	Gly	Ser	Lys	Gly	Pro	Ser	Gly	Asp	Arg	Gly	Asp	Arg	Gly	Glu	Arg
	225				230						235				240
Gly	Val	Pro	Gly	Gln	Thr	Gly	Ser	Ala	Gly	Asn	Val	Gly	Glu	Asp	Gly
				245						250					255
Glu	Gln	Gly	Gly	Lys	Gly	Val	Asp	Gly	Ala	Ser	Gly	Pro	Ser	Gly	Ala
		260						265						270	
Leu	Gly	Ala	Arg	Gly	Pro	Pro	Gly	Ser	Arg	Gly	Asp	Thr	Gly	Ala	Val
		275						280						285	
Gly	Pro	Pro	Gly	Pro	Thr	Gly	Arg	Ser	Gly	Leu	Pro	Gly	Asn	Ala	Gly
	290						295				300				
Gln	Lys	Gly	Pro	Ser	Gly	Glu	Pro	Gly	Ser	Pro	Gly	Lys	Ala	Gly	Ser
	305				310						315				320
Ala	Gly	Glu	Gln	Gly	Pro	Pro	Gly	Lys	Asp	Gly	Ser	Asn	Gly	Glu	Pro
				325						330					335
Gly	Ser	Pro	Gly	Lys	Glu	Gly	Glu	Arg	Gly	Leu	Ala	Gly	Pro	Pro	Gly
		340						345						350	
Pro	Asp	Gly	Arg	Arg	Gly	Glu	Thr	Gly	Ser	Pro	Gly	Ile	Ala	Gly	Ala
		355						360						365	
Leu	Gly	Lys	Pro	Gly	Leu	Glu	Gly	Pro	Lys	Gly	Tyr	Pro	Gly	Leu	Arg
	370						375				380				
Gly	Arg	Asp	Gly	Thr	Asn	Gly	Lys	Arg	Gly	Glu	Gln	Gly	Glu	Thr	Gly
	385				390						395				400
Pro	Asp	Gly	Val	Arg	Gly	Ile	Pro	Gly	Asn	Asp	Gly	Gln	Ser	Gly	Lys
				405						410					415
Pro	Gly	Ile	Asp	Gly	Ile	Asp	Gly	Thr	Asn	Gly	Gln	Pro	Gly	Glu	Ala
				420				425							430

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Gly Tyr Gln Gly Gly Arg Gly Thr Arg Gly Gln Leu Gly Glu Thr Gly
 435 440 445

Asp Val Gly Gln Asn Gly Asp Arg Gly Ala Pro Gly Pro Asp Gly Ser
 450 455 460

Lys Gly Ser Ala Gly Arg Pro Gly Leu Arg
 465 470

<210> SEQ ID NO 6
 <211> LENGTH: 1362
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 6

```

atgaaaaaga tttggctggc gctggctggt ttagttagcgc atcggcggcg      60
cagtatgaag atggtccgca ggggtgtggt ggtgcagatg gtaaagacgg taccocgggt      120
gaaaaaggag aacagggacg tacaggtgca gcaggtaaac agggcagccc gggtgccgat      180
ggtgcccctg gcccgcctgg tagcattggt cagcagggtg caagaggcga accgggcat      240
ccgggtagtc cgggcctgcg tggtgatacg ggtctggcgc gtgttaaagg cgttgccaggt      300
ccttcaggtc gtccaggtea accgggtgca aatggtctgc cgggtgttaa tggctgtggc      360
ggtctgctg gcaaaccggg agcaaaaggt attgcagta gcgatggaga agccggtgaa      420
agcggtgccc cgggtcagag tggctccgacc ggtccgcgcg gtcagcgtgg tccgtctggt      480
gaagatggca atccgggtct gcagggtctg cctggtatg atggcgaacc aggtgaagaa      540
ggtcagccgg gtcgttcagg ccagccgggc cagcagggcc cgcgtggtag cccgggcaaa      600
gttgcccgcg ggggtagtaa aggtcctagt ggcgatcgcg gtgatcgtgg tgaacgcggt      660
gttccctggtc agaccggtag cgcaggtaat gttggcgaag atggtgaaca ggggtggcaaa      720
ggtgttgatg gtgcaagcgg tccgagcggg gcaactgggtg cacgtggtcc tccgggcagc      780
cgtggtgaca ccggtgcagt tggctccgct ggcccgaacc gccgtagtgg cttaccgggt      840
aatgcaggtc agaaaaggtcc gtcaggtgaa cctggcagcc ctggtaaagc aggtagtgcc      900
ggtgagcagg gtccgccggg caaagatggt agtaatggtg agccgggtag ccctggcaaa      960
gaaggtgaac gtggtctggc aggaccgccg ggtcctgatg gtcgccgcgg tgaacgggt      1020
tcaccgggta ttgccggtgc cctgggtaaa ccaggtctgg aaggtccgaa aggttatcct      1080
ggtctgctg gtcgtgatgg taccaatggc aaactggcg aacagggcga aaccggtcca      1140
gatggtgttc gtggtattcc gggtaacgat ggtcagagcg gtaaaccggg cattgatggt      1200
attgatggca ccaatggtca gcctggcgaa gcaggttacc aggggtggtc cggtagccgt      1260
ggtcagctgg gtgaacagg tgatgttggc cagaatggtg atcgcggcgc accgggtccg      1320
gatgtagca aaggtagcgc cggtcgtccg ggtttacgtt aa      1362
    
```

<210> SEQ ID NO 7
 <211> LENGTH: 1362
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

-continued

<400> SEQUENCE: 7

```

atgaaaaaga tttggctggc gctggctggt ttagtttag cgttttagcgc atcggcggcg      60
cagtatgaag atggtccgca ggggtgtggt ggtgcagatg gtaaagacgg taccocgggt      120
gaaaaaggtg aacagggctg taccggtgca gcaggtaac agggcagccc gggtgccgat      180
ggtgcccgtg gcccgctggg tagcattggt cagcagggtg cacgtggcga accgggcgat      240
ccgggtagcc cgggcctgcg tggtgatacg ggtctggcgg gtgttaaagg cgttgccaggt      300
ccttctggtc gtccaggta accgggtgca aatggtctgc cgggtgttaa tggctcgtggc      360
ggctctgcgtg gcaaacgggg tgcaaaaggt attgcaggtg gcgatggcga agccggtgaa      420
agcggtgccc cgggtcagag cgttccgacc ggtccgcgcg gtcagcgtgg tccgtctggt      480
gaagatggca atccgggtct gcagggctg cctggttagcg atggcgaacc aggtgaagaa      540
ggtcagccgg gtcgttctgg ccagccgggc cagcagggcc cgcgtggtag cccgggcgaa      600
gttgccccgc gcggttctaa aggtcctagc ggcgatcgcg gtgatcgtgg tgaacgcggt      660
gttctcgtgc agaccggtag cgcaggtaat gttggcgaag atggtgaaca gggtgccaaa      720
gggtgtgatg gtgcaagcgg tccgagcggg gcaactgggtg cacgtggtcc tccgggcagc      780
cgtggtgaca ccgggtgcagt tggtcgcgct ggcccagacc gccgtagcgg cctgccgggt      840
aatgcaggtc agaaaggtcc gtctggtgaa cctggcagcc ctggtaaagc aggtagcggc      900
ggtagagcagg gtccgccggg caaagatggt agcaatggtg agccgggtag ccctggcaaa      960
gaaggtgaac gtggtctggc aggtccgccc ggtcctgatg gtcgccgcgg tgaacgggt      1020
tctccgggta ttgccggtgc cctgggtaaa ccaggtctgg aaggtccgaa aggttatcct      1080
ggctctgcgcg gtcgtgatgg taccaatggc aaacgtggcg aacagggcga aaccgggtcca      1140
gatggtgttc gtggtattcc gggtaacgat ggtcagagcg gtaaaccggg cattgatggt      1200
attgatggca ccaatggtca gcctggcgaa gcaggttatc aggggtggtcg cggtagccgt      1260
ggtcagctgg gtgaaccggg tgatgttggg cagaatggtg atcgccgcgc accgggtccg      1320
gatggtagca aaggtagcgc cggctcgtccg ggtctgcggt aa                          1362
    
```

<210> SEQ ID NO 8

<211> LENGTH: 453

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 8

```

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
 1           5           10           15
Ala Ser Ala Ala Gln Tyr Glu Asp Gly Pro Gln Gly Val Val Gly Ala
 20           25           30
Asp Gly Lys Asp Gly Thr Pro Gly Glu Lys Gly Glu Gln Gly Arg Thr
 35           40           45
Gly Ala Ala Gly Lys Gln Gly Ser Pro Gly Ala Asp Gly Ala Arg Gly
 50           55           60
Pro Leu Gly Ser Ile Gly Gln Gln Gly Ala Arg Gly Glu Pro Gly Asp
 65           70           75           80
Pro Gly Ser Pro Gly Leu Arg Gly Asp Thr Gly Leu Ala Gly Val Lys
 85           90           95
    
```

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Gly Val Ala Gly Pro Ser Gly Arg Pro Gly Gln Pro Gly Ala Asn Gly
 100 105 110

Leu Pro Gly Val Asn Gly Arg Gly Gly Leu Arg Gly Lys Pro Gly Ala
 115 120 125

Lys Gly Ile Ala Gly Ser Asp Gly Glu Ala Gly Glu Ser Gly Ala Pro
 130 135 140

Gly Gln Ser Gly Pro Thr Gly Pro Arg Gly Gln Arg Gly Pro Ser Gly
 145 150 155 160

Glu Asp Gly Asn Pro Gly Leu Gln Gly Leu Pro Gly Ser Asp Gly Glu
 165 170 175

Pro Gly Glu Glu Gly Gln Pro Gly Arg Ser Gly Gln Pro Gly Gln Gln
 180 185 190

Gly Pro Arg Gly Ser Pro Gly Glu Val Gly Pro Arg Gly Ser Lys Gly
 195 200 205

Pro Ser Gly Asp Arg Gly Asp Arg Gly Glu Arg Gly Val Pro Gly Gln
 210 215 220

Thr Gly Ser Ala Gly Asn Val Gly Glu Asp Gly Glu Gln Gly Gly Lys
 225 230 235 240

Gly Val Asp Gly Ala Ser Gly Pro Ser Gly Ala Leu Gly Ala Arg Gly
 245 250 255

Pro Pro Gly Ser Arg Gly Asp Thr Gly Ala Val Gly Pro Pro Gly Pro
 260 265 270

Thr Gly Arg Ser Gly Leu Pro Gly Asn Ala Gly Gln Lys Gly Pro Ser
 275 280 285

Gly Glu Pro Gly Ser Pro Gly Lys Ala Gly Ser Ala Gly Glu Gln Gly
 290 295 300

Pro Pro Gly Lys Asp Gly Ser Asn Gly Glu Pro Gly Ser Pro Gly Lys
 305 310 315 320

Glu Gly Glu Arg Gly Leu Ala Gly Pro Pro Gly Pro Asp Gly Arg Arg
 325 330 335

Gly Glu Thr Gly Ser Pro Gly Ile Ala Gly Ala Leu Gly Lys Pro Gly
 340 345 350

Leu Glu Gly Pro Lys Gly Tyr Pro Gly Leu Arg Gly Arg Asp Gly Thr
 355 360 365

Asn Gly Lys Arg Gly Glu Gln Gly Glu Thr Gly Pro Asp Gly Val Arg
 370 375 380

Gly Ile Pro Gly Asn Asp Gly Gln Ser Gly Lys Pro Gly Ile Asp Gly
 385 390 395 400

Ile Asp Gly Thr Asn Gly Gln Pro Gly Glu Ala Gly Tyr Gln Gly Gly
 405 410 415

Arg Gly Thr Arg Gly Gln Leu Gly Glu Thr Gly Asp Val Gly Gln Asn
 420 425 430

Gly Asp Arg Gly Ala Pro Gly Pro Asp Gly Ser Lys Gly Ser Ala Gly
 435 440 445

Arg Pro Gly Leu Arg
 450

<210> SEQ ID NO 9
 <211> LENGTH: 825
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:

-continued

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 9

```

atgaaaaaga tttggctggc gctggctggt ttagttttag cgtttagcgc atcgggcgcg      60
cagtatgaag atcaccatca ccaccaccac catcaccact ctggctcgag cctggtgccg      120
cgcggcagcc atatgggtcc gcagggtggt gttggtgcag atggtaaaga cggtagcccc      180
ggtgaaaaag gagaacaggg acgtacaggt gcagcagga aacagggcag cccgggtgcc      240
gatggtgccc gtggcccgcg gggtagcatt ggtcagcagg gtgcaagagg cgaaccgggc      300
gatccgggta gtcggggcct gcgtggtgat acgggtctgg ccggtgttaa aggcggtgca      360
ggtccttcag gtcgtccagg tcaaccgggt gcaaatggtc tgccgggtgt taatggtcgt      420
ggcggctctg aacgtggtct ggcaggaccg ccgggtcctg atggtcgccg cggtgaaacg      480
ggttcaccgg gtattgccgg tgccctgggt aaaccaggtc tggaaggtcc gaaaggttat      540
cctggtctgc gcggtctgta tggtagcaat ggcaaacctg gcgaacaggg cgaaccgggt      600
ccagatggtg ttcgtggtat tccgggtaac gatggtcaga gcggtaaacc gggcattgat      660
ggtattgatg gcaccaatgg tcagcctggc gaagcaggtt atcagggtgg tcgcggtacc      720
cgtggtcagc tgggtgaaac aggtgatggt ggtcagaatg gtgatcgccg cgcaccgggt      780
ccggatggta gcaaaggtag cgcgggtcgt ccgggtttac gttaa                               825
    
```

<210> SEQ ID NO 10

<211> LENGTH: 274

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 10

```

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
 1           5           10           15
Ala Ser Ala Ala Gln Tyr Glu Asp His His His His His His His His
 20           25           30
His Ser Gly Ser Ser Leu Val Pro Arg Gly Ser His Met Gly Pro Gln
 35           40           45
Gly Val Val Gly Ala Asp Gly Lys Asp Gly Thr Pro Gly Glu Lys Gly
 50           55           60
Glu Gln Gly Arg Thr Gly Ala Ala Gly Lys Gln Gly Ser Pro Gly Ala
 65           70           75           80
Asp Gly Ala Arg Gly Pro Leu Gly Ser Ile Gly Gln Gln Gly Ala Arg
 85           90           95
Gly Glu Pro Gly Asp Pro Gly Ser Pro Gly Leu Arg Gly Asp Thr Gly
100          105          110
Leu Ala Gly Val Lys Gly Val Ala Gly Pro Ser Gly Arg Pro Gly Gln
115          120          125
Pro Gly Ala Asn Gly Leu Pro Gly Val Asn Gly Arg Gly Gly Leu Glu
130          135          140
Arg Gly Leu Ala Gly Pro Pro Gly Pro Asp Gly Arg Arg Gly Glu Thr
145          150          155          160
Gly Ser Pro Gly Ile Ala Gly Ala Leu Gly Lys Pro Gly Leu Glu Gly
165          170          175
    
```

-continued

Pro Lys Gly Tyr Pro Gly Leu Arg Gly Arg Asp Gly Thr Asn Gly Lys
 180 185 190

Arg Gly Glu Gln Gly Glu Thr Gly Pro Asp Gly Val Arg Gly Ile Pro
 195 200 205

Gly Asn Asp Gly Gln Ser Gly Lys Pro Gly Ile Asp Gly Ile Asp Gly
 210 215 220

Thr Asn Gly Gln Pro Gly Glu Ala Gly Tyr Gln Gly Gly Arg Gly Thr
 225 230 235 240

Arg Gly Gln Leu Gly Glu Thr Gly Asp Val Gly Gln Asn Gly Asp Arg
 245 250 255

Gly Ala Pro Gly Pro Asp Gly Ser Lys Gly Ser Ala Gly Arg Pro Gly
 260 265 270

Leu Arg

<210> SEQ ID NO 11
 <211> LENGTH: 642
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 11

atgaaaaaga tttggctggc gctggctggt ttagtttag cgtttagcgc atcgcgggcg 60
 cagtatgaag atggtccgca ggggtgtggt ggtgcagatg gtaaagacgg taccocgggt 120
 aatgcaggtc agaaaggctc gtcaggtgaa cctggcagcc ctggtaaagc aggtagtgcc 180
 ggtgagcagg gtccgccggg caaagatggt agtaatggtg agccgggtag ccctggcaaa 240
 gaaggtgaac gtggtctggc aggaccgccc ggtcctgatg gtcgccgccc tgaaacgggt 300
 tcaccgggta ttgccgtgac cctgggtaaa ccaggtctgg aaggtccgaa aggttatcct 360
 ggtctgcccgc gtcgtgatgg taccaatggc aaacgtggcg aacagggcga aaccgggtcca 420
 gatggtgttc gtggtattcc gggtaacgat ggtcagagcg gtaaaccggg cattgatggt 480
 attgatggca ccaatggtca gcctggcgaa gcaggttatc agggtggtcg cggtaccctg 540
 ggtcagctgg gtgaaacagg tgatgttggc cagaatggtg atcgccggcg accgggtccg 600
 gatggtagca aaggtagcgc cggctgctcc ggtttacggt aa 642

<210> SEQ ID NO 12
 <211> LENGTH: 213
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 12

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
 1 5 10 15

Ala Ser Ala Ala Gln Tyr Glu Asp Gly Pro Gln Gly Val Val Gly Ala
 20 25 30

Asp Gly Lys Asp Gly Thr Pro Gly Asn Ala Gly Gln Lys Gly Pro Ser
 35 40 45

Gly Glu Pro Gly Ser Pro Gly Lys Ala Gly Ser Ala Gly Glu Gln Gly
 50 55 60

-continued

Pro Pro Gly Lys Asp Gly Ser Asn Gly Glu Pro Gly Ser Pro Gly Lys
65 70 75 80

Glu Gly Glu Arg Gly Leu Ala Gly Pro Pro Gly Pro Asp Gly Arg Arg
85 90 95

Gly Glu Thr Gly Ser Pro Gly Ile Ala Gly Ala Leu Gly Lys Pro Gly
100 105 110

Leu Glu Gly Pro Lys Gly Tyr Pro Gly Leu Arg Gly Arg Asp Gly Thr
115 120 125

Asn Gly Lys Arg Gly Glu Gln Gly Glu Thr Gly Pro Asp Gly Val Arg
130 135 140

Gly Ile Pro Gly Asn Asp Gly Gln Ser Gly Lys Pro Gly Ile Asp Gly
145 150 155 160

Ile Asp Gly Thr Asn Gly Gln Pro Gly Glu Ala Gly Tyr Gln Gly Gly
165 170 175

Arg Gly Thr Arg Gly Gln Leu Gly Glu Thr Gly Asp Val Gly Gln Asn
180 185 190

Gly Asp Arg Gly Ala Pro Gly Pro Asp Gly Ser Lys Gly Ser Ala Gly
195 200 205

Arg Pro Gly Leu Arg
210

<210> SEQ ID NO 13
<211> LENGTH: 696
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
polynucleotide

<400> SEQUENCE: 13

```

atgaaaaaga tttggctggc gctggctggt ttagttttag cgtttagcgc atcggcggcg      60
cagtatgaag atggtgaaaa aggtgaaaag ggcgagaaaag gtgagaaaag cgaaggggt      120
gaaaaaggtc cgcagggtgt tgttggtgca gatggtaaag acggtacccc gggtaaatgca      180
ggtcagaaaag gtcctcagg tgaacctggc agccctggta aagcaggtag tgccggtag      240
cagggtccgc cgggcaaaga tgtagtaat ggtgagccgg gtagccctgg caaagaagg      300
gaacgtggtc tggcaggacc gccgggtcct gatggtcgcc gcggtgaaac gggttcaccg      360
ggtattgccc gtgccctggg taaaccagg ctggaaggtc cgaaggtta tcctggtctg      420
cgcggtcgtg atggtaccaa tggcaaacgt ggcgaacagg gcgaaaccgg tccagatggt      480
gttcgtggta ttcgggtaa cgatggtcag agcggtaaac cgggcattga tggtattgat      540
ggcaccaatg gtcagcctgg cgaagcagg tatcagggtg gtcgcggtac ccgtggtcag      600
ctgggtgaaa caggatgatg tggtcagaat ggtgatcgcg gcgcaccggg tccggatggt      660
agcaaaggta gcgccggtcg tccgggttta cgtaa      696

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<210> SEQ ID NO 14
<211> LENGTH: 231
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
polypeptide

<400> SEQUENCE: 14

-continued

Met	Lys	Lys	Ile	Trp	Leu	Ala	Leu	Ala	Gly	Leu	Val	Leu	Ala	Phe	Ser
1			5						10					15	
Ala	Ser	Ala	Ala	Gln	Tyr	Glu	Asp	Gly	Glu	Lys	Gly	Glu	Lys	Gly	Glu
			20					25					30		
Lys	Gly	Glu	Lys	Gly	Glu	Lys	Gly	Glu	Lys	Gly	Pro	Gln	Gly	Val	Val
		35					40					45			
Gly	Ala	Asp	Gly	Lys	Asp	Gly	Thr	Pro	Gly	Asn	Ala	Gly	Gln	Lys	Gly
	50					55					60				
Pro	Ser	Gly	Glu	Pro	Gly	Ser	Pro	Gly	Lys	Ala	Gly	Ser	Ala	Gly	Glu
65					70					75					80
Gln	Gly	Pro	Pro	Gly	Lys	Asp	Gly	Ser	Asn	Gly	Glu	Pro	Gly	Ser	Pro
				85					90					95	
Gly	Lys	Glu	Gly	Glu	Arg	Gly	Leu	Ala	Gly	Pro	Pro	Gly	Pro	Asp	Gly
			100					105					110		
Arg	Arg	Gly	Glu	Thr	Gly	Ser	Pro	Gly	Ile	Ala	Gly	Ala	Leu	Gly	Lys
		115					120				125				
Pro	Gly	Leu	Glu	Gly	Pro	Lys	Gly	Tyr	Pro	Gly	Leu	Arg	Gly	Arg	Asp
	130					135					140				
Gly	Thr	Asn	Gly	Lys	Arg	Gly	Glu	Gln	Gly	Glu	Thr	Gly	Pro	Asp	Gly
145					150					155					160
Val	Arg	Gly	Ile	Pro	Gly	Asn	Asp	Gly	Gln	Ser	Gly	Lys	Pro	Gly	Ile
				165					170					175	
Asp	Gly	Ile	Asp	Gly	Thr	Asn	Gly	Gln	Pro	Gly	Glu	Ala	Gly	Tyr	Gln
		180						185					190		
Gly	Gly	Arg	Gly	Thr	Arg	Gly	Gln	Leu	Gly	Glu	Thr	Gly	Asp	Val	Gly
		195					200					205			
Gln	Asn	Gly	Asp	Arg	Gly	Ala	Pro	Gly	Pro	Asp	Gly	Ser	Lys	Gly	Ser
	210					215					220				
Ala	Gly	Arg	Pro	Gly	Leu	Arg									
225					230										

<210> SEQ ID NO 15
 <211> LENGTH: 696
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 15

```

atgaaaaaga tttggctggc gctggctggt ttagtttag cgttagcgc atcgcgcg 60
cagtatgaag atggtgataa aggtgataa ggcgacaaag gtgacaaagg cgataagggt 120
gataaaggtc cgcagggtgt tgttggtgca gatggtaaag acggtacccc gggtaatgca 180
ggtcagaaag gtccgtcagg tgaacctggc agccctggta aagcaggtag tgccggtgag 240
cagggtcctc cgggcaaaga tgtagtaat ggtgagccgg tagccctgg caaagaaggt 300
gaacgtggtc tggcaggacc gccgggtcct gatggtcgcc gcggtgaaac gggttcaccg 360
ggtattgccc gtgccctggg taaaccaggt ctggaaggtc cgaaaggta tccgtgtctg 420
cgcggtcgtg atggtaccaa tggcaaactg gccgaacagg gcgaaaccgg tccagatggt 480
gttcgtggta ttccgggtaa cgatggtcag agcggtaaac cgggcattga tggattgat 540
ggcaccaatg gtcagcctgg cgaagcaggt taccaggggt gtcgcggtac ccgtggtcag 600
    
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-continued

ctgggtgaaa caggtgatgt tggtcagaat ggtgatcgcg gcgaccggg tccggatggt 660
 agcaaaggta ggcgggtcg tccgggttta cgtaa 696

<210> SEQ ID NO 16
 <211> LENGTH: 231
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 16

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
 1 5 10 15
 Ala Ser Ala Ala Gln Tyr Glu Asp Gly Asp Lys Gly Asp Lys Gly Asp
 20 25 30
 Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Pro Gln Gly Val Val
 35 40 45
 Gly Ala Asp Gly Lys Asp Gly Thr Pro Gly Asn Ala Gly Gln Lys Gly
 50 55 60
 Pro Ser Gly Glu Pro Gly Ser Pro Gly Lys Ala Gly Ser Ala Gly Glu
 65 70 75 80
 Gln Gly Pro Pro Gly Lys Asp Gly Ser Asn Gly Glu Pro Gly Ser Pro
 85 90 95
 Gly Lys Glu Gly Glu Arg Gly Leu Ala Gly Pro Pro Gly Pro Asp Gly
 100 105 110
 Arg Arg Gly Glu Thr Gly Ser Pro Gly Ile Ala Gly Ala Leu Gly Lys
 115 120 125
 Pro Gly Leu Glu Gly Pro Lys Gly Tyr Pro Gly Leu Arg Gly Arg Asp
 130 135 140
 Gly Thr Asn Gly Lys Arg Gly Glu Gln Gly Glu Thr Gly Pro Asp Gly
 145 150 155 160
 Val Arg Gly Ile Pro Gly Asn Asp Gly Gln Ser Gly Lys Pro Gly Ile
 165 170 175
 Asp Gly Ile Asp Gly Thr Asn Gly Gln Pro Gly Glu Ala Gly Tyr Gln
 180 185 190
 Gly Gly Arg Gly Thr Arg Gly Gln Leu Gly Glu Thr Gly Asp Val Gly
 195 200 205
 Gln Asn Gly Asp Arg Gly Ala Pro Gly Pro Asp Gly Ser Lys Gly Ser
 210 215 220
 Ala Gly Arg Pro Gly Leu Arg
 225 230

<210> SEQ ID NO 17
 <211> LENGTH: 2232
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 17

atgaaaaaga tttggctggc gctggctggt ttagtttag cgtttagcgc atcggcggcg 60
 cagtatgaag atcaccatca ccaccaccac catcaccact ctggctcgag cctgggtgccc 120

-continued

cgcggcagcc	atatgtctgg	ctcgcagcgt	aaaggtgaag	aactgttcac	cggtgttgtt	180
cggatcctgg	ttgaactgga	tggtgatggt	aacggccaca	aattctctgt	tcgtggtgaa	240
ggtgaaggtg	atgcaaccaa	cggtaaactg	accctgaaat	tcatctgcac	taccggtaaa	300
ctgccgggtc	catggccgac	tctggtgact	accctgacct	atggtgttca	gtgtttttct	360
cgttaccocg	atcacatgaa	gcagcatgat	ttcttcaaat	ctgcaatgcc	ggaaggttat	420
gtacaggagc	gcaccatttc	tttcaaagac	gatggcacct	acaaaacocg	tgcagaggtt	480
aaatttgaag	gtgatactct	ggtgaaccgt	attgaaactga	aaggcattga	tttcaaagag	540
gacggcaaca	tcctgggcca	caaactggaa	tataacttca	actcccataa	cgtttacatc	600
accgcagaca	aacagaagaa	cggtatcaaa	gctaacttca	aaattcgcca	taacgttgaa	660
gacggtagcg	tacagctggc	ggaccactac	cagcagaaca	ctccgatcgg	tgatggtccg	720
gttctgctgc	cggataacca	ctacctgtcc	accagctcta	aactgtccaa	agaccggaac	780
gaaaagcgcg	accacatggt	gctgctggag	ttcgttactg	cagcaggtat	cacgcacggc	840
atggatgaac	tctacaaatc	tggcgcgcgc	ggcggctccg	aggggtgtgt	tggtgcagat	900
ggtaaagacg	gtaccocggg	taatgcaggt	cagaaaggtc	cgtcaggtga	acctggcagc	960
cctggtaaag	caggtagtgc	cggtagcagc	ggtccgcocg	gcaaagatgg	tagtaatggt	1020
gagccgggta	gccctggcaa	agaaggtgaa	cgtggtctgg	caggaccgcc	gggtcctgat	1080
ggtcgcgcgc	gtgaaacggg	ttcaccgggt	attgocggtg	ccctgggtaa	accaggtctg	1140
gaaggtccga	aaggttatcc	tggtctgcgc	ggtcgtgatg	gtaccaatgg	caaacgtggc	1200
gaacagggcg	aaaccggctc	agatggtggt	cgtggtattc	cgggtaacga	tggtcagagc	1260
ggtaaaccgg	gcattgatgg	tattgatggc	accaatggtc	agcctggcga	agcaggttat	1320
cagggtggtc	gcggtaccoc	tggtcagctg	ggtgaaacag	gtgatgttgg	tcagaatggt	1380
gatcgcggcg	caccgggtcc	ggatggtagc	aaaggtagcg	ccggtcgtcc	gggtttacgt	1440
caccagaaa	cgctggtgaa	agtaaaagat	gctgaagatc	agttgggtgc	acgagtggtt	1500
tacatcgaac	tggatctcaa	cagcggtaag	atccttgaga	gttttcgccc	cgaagaacgt	1560
tttccaatga	tgagcacttt	taaagttctg	ctatgtggcg	cggattatc	ccgtattgac	1620
gccgggcaag	agcaactcgg	tcgcccata	cactattctc	agaatgactt	ggttgagtac	1680
tcaccagtca	cagaaaagca	tcttacggat	ggcatgacag	taagagaatt	atgcagtgct	1740
gccataacca	tgagtgataa	cactgcggcc	aacttacttc	tgacaacgat	cggaggaccg	1800
aaggagctaa	ccgctttttt	gcacaacatg	ggggatcatg	taactcgcct	tgatcgttgg	1860
gaaccgggag	tgaatgaagc	cataccaaac	gacgagcgtg	acaccacgat	gcctgtagca	1920
atggcaacaa	cgttgcgcaa	actattaact	ggcgaactac	ttactctagc	ttcccggcaa	1980
caattaatag	actggatgga	ggcggataaa	ggtgcaggac	cacttctgcg	ctcggccctt	2040
cggctggct	ggtttattgc	tgataaatct	ggagcgggtg	agcgtgggtc	tcgcggtatc	2100
attgcagcac	tggggccaga	tggtaaagcc	tcccgtatcg	tagttatcta	cacgacgggg	2160
agtcaggcaa	ctatggatga	acgaaataga	cagatcgctg	agataggtgc	ctcactgatt	2220
aagcattggt	aa					2232

<210> SEQ ID NO 18

<211> LENGTH: 743

<212> TYPE: PRT

-continued

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 18

```

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
1           5           10           15
Ala Ser Ala Ala Gln Tyr Glu Asp His His His His His His His
20           25           30
His Ser Gly Ser Ser Leu Val Pro Arg Gly Ser His Met Ser Gly Ser
35           40           45
Ser Ser Lys Gly Glu Glu Leu Phe Thr Gly Val Val Pro Ile Leu Val
50           55           60
Glu Leu Asp Gly Asp Val Asn Gly His Lys Phe Ser Val Arg Gly Glu
65           70           75           80
Gly Glu Gly Asp Ala Thr Asn Gly Lys Leu Thr Leu Lys Phe Ile Cys
85           90           95
Thr Thr Gly Lys Leu Pro Val Pro Trp Pro Thr Leu Val Thr Thr Leu
100          105          110
Thr Tyr Gly Val Gln Cys Phe Ser Arg Tyr Pro Asp His Met Lys Gln
115          120          125
His Asp Phe Phe Lys Ser Ala Met Pro Glu Gly Tyr Val Gln Glu Arg
130          135          140
Thr Ile Ser Phe Lys Asp Asp Gly Thr Tyr Lys Thr Arg Ala Glu Val
145          150          155          160
Lys Phe Glu Gly Asp Thr Leu Val Asn Arg Ile Glu Leu Lys Gly Ile
165          170          175
Asp Phe Lys Glu Asp Gly Asn Ile Leu Gly His Lys Leu Glu Tyr Asn
180          185          190
Phe Asn Ser His Asn Val Tyr Ile Thr Ala Asp Lys Gln Lys Asn Gly
195          200          205
Ile Lys Ala Asn Phe Lys Ile Arg His Asn Val Glu Asp Gly Ser Val
210          215          220
Gln Leu Ala Asp His Tyr Gln Gln Asn Thr Pro Ile Gly Asp Gly Pro
225          230          235          240
Val Leu Leu Pro Asp Asn His Tyr Leu Ser Thr Gln Ser Lys Leu Ser
245          250          255
Lys Asp Pro Asn Glu Lys Arg Asp His Met Val Leu Leu Glu Phe Val
260          265          270
Thr Ala Ala Gly Ile Thr His Gly Met Asp Glu Leu Tyr Lys Ser Gly
275          280          285
Ala Pro Gly Gly Pro Gln Gly Val Val Gly Ala Asp Gly Lys Asp Gly
290          295          300
Thr Pro Gly Asn Ala Gly Gln Lys Gly Pro Ser Gly Glu Pro Gly Ser
305          310          315          320
Pro Gly Lys Ala Gly Ser Ala Gly Glu Gln Gly Pro Pro Gly Lys Asp
325          330          335
Gly Ser Asn Gly Glu Pro Gly Ser Pro Gly Lys Glu Gly Glu Arg Gly
340          345          350
Leu Ala Gly Pro Pro Gly Pro Asp Gly Arg Arg Gly Glu Thr Gly Ser
355          360          365

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Pro Gly Ile Ala Gly Ala Leu Gly Lys Pro Gly Leu Glu Gly Pro Lys
 370 375 380
 Gly Tyr Pro Gly Leu Arg Gly Arg Asp Gly Thr Asn Gly Lys Arg Gly
 385 390 395 400
 Glu Gln Gly Glu Thr Gly Pro Asp Gly Val Arg Gly Ile Pro Gly Asn
 405 410 415
 Asp Gly Gln Ser Gly Lys Pro Gly Ile Asp Gly Ile Asp Gly Thr Asn
 420 425 430
 Gly Gln Pro Gly Glu Ala Gly Tyr Gln Gly Gly Arg Gly Thr Arg Gly
 435 440 445
 Gln Leu Gly Glu Thr Gly Asp Val Gly Gln Asn Gly Asp Arg Gly Ala
 450 455 460
 Pro Gly Pro Asp Gly Ser Lys Gly Ser Ala Gly Arg Pro Gly Leu Arg
 465 470 475 480
 His Pro Glu Thr Leu Val Lys Val Lys Asp Ala Glu Asp Gln Leu Gly
 485 490 495
 Ala Arg Val Gly Tyr Ile Glu Leu Asp Leu Asn Ser Gly Lys Ile Leu
 500 505 510
 Glu Ser Phe Arg Pro Glu Glu Arg Phe Pro Met Met Ser Thr Phe Lys
 515 520 525
 Val Leu Leu Cys Gly Ala Val Leu Ser Arg Ile Asp Ala Gly Gln Glu
 530 535 540
 Gln Leu Gly Arg Arg Ile His Tyr Ser Gln Asn Asp Leu Val Glu Tyr
 545 550 555 560
 Ser Pro Val Thr Glu Lys His Leu Thr Asp Gly Met Thr Val Arg Glu
 565 570 575
 Leu Cys Ser Ala Ala Ile Thr Met Ser Asp Asn Thr Ala Ala Asn Leu
 580 585 590
 Leu Leu Thr Thr Ile Gly Gly Pro Lys Glu Leu Thr Ala Phe Leu His
 595 600 605
 Asn Met Gly Asp His Val Thr Arg Leu Asp Arg Trp Glu Pro Glu Leu
 610 615 620
 Asn Glu Ala Ile Pro Asn Asp Glu Arg Asp Thr Thr Met Pro Val Ala
 625 630 635 640
 Met Ala Thr Thr Leu Arg Lys Leu Leu Thr Gly Glu Leu Leu Thr Leu
 645 650 655
 Ala Ser Arg Gln Gln Leu Ile Asp Trp Met Glu Ala Asp Lys Val Ala
 660 665 670
 Gly Pro Leu Leu Arg Ser Ala Leu Pro Ala Gly Trp Phe Ile Ala Asp
 675 680 685
 Lys Ser Gly Ala Gly Glu Arg Gly Ser Arg Gly Ile Ile Ala Ala Leu
 690 695 700
 Gly Pro Asp Gly Lys Pro Ser Arg Ile Val Val Ile Tyr Thr Thr Gly
 705 710 715 720
 Ser Gln Ala Thr Met Asp Glu Arg Asn Arg Gln Ile Ala Glu Ile Gly
 725 730 735
 Ala Ser Leu Ile Lys His Trp
 740

<210> SEQ ID NO 19

<211> LENGTH: 2232

<212> TYPE: DNA

-continued

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 19

```

atgaaaaaga tttggctggc gctggctggt ttagtttag cgtttagcgc atcgcgcgcg    60
cagtatgaag atcacatca ccaccaccac catcaccact ctggctcgag cctggtgccg    120
cgcgcgagcc atatgtctgg ctcgagcagt aaaggtgaag aactgttcac cggtgttgtt    180
ccgatcctgg ttgaactgga tggtagtgtt aacggccaca aattctctgt tcgtggtgaa    240
ggtgaaggtg atgcaaccaa cggtaaactg accctgaaat tcatctgcac taccggtaaa    300
ctgccggttc catggcgcac tctggtgact accctgacct atggtgttca gtgtttttct    360
cgttaccggy atcacatgaa gcagcatgat ttcttcaaat ctgcaatgcc ggaaggttat    420
gtacaggagc gcaccatttc tttcaaagac gatggcacct acaaaaccgg tgcagaggtt    480
aaatttgaag gtgatactct ggtgaaccgt attgaactga aaggcattga tttcaaagag    540
gacggcaaca tcctgggcca caaactggaa tataacttca actcccataa cgtttacatc    600
accgcagaca aacagaagaa cggtatcaaa gctaacttca aaattcgcca taacggtgaa    660
gacggtagcg tacagctggc ggaccactac cagcagaaca ctccgatcgg tgatggtccg    720
gttctgctgc cggataacca ctacctgtcc acccagteta aactgtccaa agaccggaac    780
gaaaagcgcg accacatggt gctgctggag ttcgttactg cagcaggtat cacgcaocggc    840
atggatgaac tctacaaatc tggcgcgcgg ggcggtcocg agggtgttgt tggtgagat    900
ggtaaagaag gtacccccggg taatgcaggt cagaaaggtc cgtcagggtga acctggcagc    960
cctggtaaag caggtagtgc cggtagcag ggtccgcccgg gcaaagatgg tagtaatggt    1020
gagccgggta gccctggcaa agaaggtgaa cgtggtctgg caggaccgcc gggtcctgat    1080
ggtcgcgcgc gtgaaacggg ttcaccgggt attgcccgtg ccctgggtaa accaggtctg    1140
gaaggtccga aaggttatcc tggctctgcgc ggtcgtgatg gtaccaatgg caaacgtggc    1200
gaacagggcg aaaccggtcc agatggtgtt cgtggtatcc cgggtaacga tggtcagagc    1260
ggtaaacccg gcattgatgg tattgatggc accaatggtc agcctggcga agcaggttat    1320
caggtggttc gcggtaccgg tggtcagctg ggtgaaacag gtgatgttgg tcagaatggt    1380
gatcgcggcg caccgggtcc ggatggtagc aaaggtagcg ccggtcgtcc gggtttacgt    1440
caccagaaa cgctgggtgaa agtaaaagat gctgaagatc agttgggtgc acgagtgggt    1500
tacatcgaac tggatctcaa cagcggtaag atccttgaga gttttcgccc cgaagaacgt    1560
tttccaatga tgagcacttt taaagttctg ctatgtggcg cggattatcc ccgtattgac    1620
gccgggcaag agcaactcgg tcgcccata cactattctc agaatgactt ggttagtagc    1680
tcaccagtc cagaaaagca tcttacggat ggcatgacag taagagaatt atgcagtgct    1740
gccataacca tgagtataa cactgcggcc aacttacttc tgacaacgat cggaggaccg    1800
aaggagctaa ccgctttttt gcacaacatg ggggatcatg taactcgcc tgcctgttgg    1860
gaaccgggag tgaatgaagc cataccaaac gacgagcgtg acaccacgat gcctgtagca    1920
atggcaacaa cgttgcgcaa actattaact ggccaactac ttactctage ttcccgcaa    1980
caattaatag actggatgga ggcggataaa gttgcaggac cacttctgcg ctccggcctt    2040
ccggtggtct ggtttattgc tgataaatct ggagccgggt agcgtgggtc tcgcggtatc    2100

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attgcagcac tggggccaga tggtaagccc tcccgtatcg tagttatcta cagcagggg 2160
agtcaggcaa ctatggatga acgaaataga cagatcgctg agataggtgc ctcaactgatt 2220
aagcattggt aa 2232

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<210> SEQ ID NO 20
<211> LENGTH: 743
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
      polypeptide

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<400> SEQUENCE: 20

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```

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
 1          5          10          15
Ala Ser Ala Ala Gln Tyr Glu Asp His His His His His His His His
 20          25          30
His Ser Gly Ser Ser Leu Val Pro Arg Gly Ser His Met Ser Gly Ser
 35          40          45
Ser Ser Lys Gly Glu Glu Leu Phe Thr Gly Val Val Pro Ile Leu Val
 50          55          60
Glu Leu Asp Gly Asp Val Asn Gly His Lys Phe Ser Val Arg Gly Glu
 65          70          75          80
Gly Glu Gly Asp Ala Thr Asn Gly Lys Leu Thr Leu Lys Phe Ile Cys
 85          90          95
Thr Thr Gly Lys Leu Pro Val Pro Trp Pro Thr Leu Val Thr Thr Leu
100          105          110
Thr Tyr Gly Val Gln Cys Phe Ser Arg Tyr Pro Asp His Met Lys Gln
115          120          125
His Asp Phe Phe Lys Ser Ala Met Pro Glu Gly Tyr Val Gln Glu Arg
130          135          140
Thr Ile Ser Phe Lys Asp Asp Gly Thr Tyr Lys Thr Arg Ala Glu Val
145          150          155          160
Lys Phe Glu Gly Asp Thr Leu Val Asn Arg Ile Glu Leu Lys Gly Ile
165          170          175
Asp Phe Lys Glu Asp Gly Asn Ile Leu Gly His Lys Leu Glu Tyr Asn
180          185          190
Phe Asn Ser His Asn Val Tyr Ile Thr Ala Asp Lys Gln Lys Asn Gly
195          200          205
Ile Lys Ala Asn Phe Lys Ile Arg His Asn Val Glu Asp Gly Ser Val
210          215          220
Gln Leu Ala Asp His Tyr Gln Gln Asn Thr Pro Ile Gly Asp Gly Pro
225          230          235          240
Val Leu Leu Pro Asp Asn His Tyr Leu Ser Thr Gln Ser Lys Leu Ser
245          250          255
Lys Asp Pro Asn Glu Lys Arg Asp His Met Val Leu Leu Glu Phe Val
260          265          270
Thr Ala Ala Gly Ile Thr His Gly Met Asp Glu Leu Tyr Lys Ser Gly
275          280          285
Ala Pro Gly Gly Pro Gln Gly Val Val Gly Ala Asp Gly Lys Asp Gly
290          295          300
Thr Pro Gly Asn Ala Gly Gln Lys Gly Pro Ser Gly Glu Pro Gly Ser

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305				310						315				320	
Pro	Gly	Lys	Ala	Gly	Ser	Ala	Gly	Glu	Gln	Gly	Pro	Pro	Gly	Lys	Asp
				325					330					335	
Gly	Ser	Asn	Gly	Glu	Pro	Gly	Ser	Pro	Gly	Lys	Glu	Gly	Glu	Arg	Gly
			340					345					350		
Leu	Ala	Gly	Pro	Pro	Gly	Pro	Asp	Gly	Arg	Arg	Gly	Glu	Thr	Gly	Ser
		355					360					365			
Pro	Gly	Ile	Ala	Gly	Ala	Leu	Gly	Lys	Pro	Gly	Leu	Glu	Gly	Pro	Lys
	370					375					380				
Gly	Tyr	Pro	Gly	Leu	Arg	Gly	Arg	Asp	Gly	Thr	Asn	Gly	Lys	Arg	Gly
	385				390					395					400
Glu	Gln	Gly	Glu	Thr	Gly	Pro	Asp	Gly	Val	Arg	Gly	Ile	Pro	Gly	Asn
				405					410						415
Asp	Gly	Gln	Ser	Gly	Lys	Pro	Gly	Ile	Asp	Gly	Ile	Asp	Gly	Thr	Asn
			420					425					430		
Gly	Gln	Pro	Gly	Glu	Ala	Gly	Tyr	Gln	Gly	Gly	Arg	Gly	Thr	Arg	Gly
		435					440					445			
Gln	Leu	Gly	Glu	Thr	Gly	Asp	Val	Gly	Gln	Asn	Gly	Asp	Arg	Gly	Ala
	450					455					460				
Pro	Gly	Pro	Asp	Gly	Ser	Lys	Gly	Ser	Ala	Gly	Arg	Pro	Gly	Leu	Arg
	465				470				475						480
His	Pro	Glu	Thr	Leu	Val	Lys	Val	Lys	Asp	Ala	Glu	Asp	Gln	Leu	Gly
			485						490						495
Ala	Arg	Val	Gly	Tyr	Ile	Glu	Leu	Asp	Leu	Asn	Ser	Gly	Lys	Ile	Leu
			500					505					510		
Glu	Ser	Phe	Arg	Pro	Glu	Glu	Arg	Phe	Pro	Met	Met	Ser	Thr	Phe	Lys
		515					520					525			
Val	Leu	Leu	Cys	Gly	Ala	Val	Leu	Ser	Arg	Ile	Asp	Ala	Gly	Gln	Glu
	530					535					540				
Gln	Leu	Gly	Arg	Arg	Ile	His	Tyr	Ser	Gln	Asn	Asp	Leu	Val	Glu	Tyr
	545				550					555					560
Ser	Pro	Val	Thr	Glu	Lys	His	Leu	Thr	Asp	Gly	Met	Thr	Val	Arg	Glu
			565						570						575
Leu	Cys	Ser	Ala	Ala	Ile	Thr	Met	Ser	Asp	Asn	Thr	Ala	Ala	Asn	Leu
		580						585							590
Leu	Leu	Thr	Thr	Ile	Gly	Gly	Pro	Lys	Glu	Leu	Thr	Ala	Phe	Leu	His
		595					600					605			
Asn	Met	Gly	Asp	His	Val	Thr	Arg	Leu	Asp	Arg	Trp	Glu	Pro	Glu	Leu
	610					615					620				
Asn	Glu	Ala	Ile	Pro	Asn	Asp	Glu	Arg	Asp	Thr	Thr	Met	Pro	Val	Ala
	625				630					635					640
Met	Ala	Thr	Thr	Leu	Arg	Lys	Leu	Leu	Thr	Gly	Glu	Leu	Leu	Thr	Leu
			645						650						655
Ala	Ser	Arg	Gln	Gln	Leu	Ile	Asp	Trp	Met	Glu	Ala	Asp	Lys	Val	Ala
			660					665					670		
Gly	Pro	Leu	Leu	Arg	Ser	Ala	Leu	Pro	Ala	Gly	Trp	Phe	Ile	Ala	Asp
		675					680					685			
Lys	Ser	Gly	Ala	Gly	Glu	Arg	Gly	Ser	Arg	Gly	Ile	Ile	Ala	Ala	Leu
	690					695					700				
Gly	Pro	Asp	Gly	Lys	Pro	Ser	Arg	Ile	Val	Val	Ile	Tyr	Thr	Thr	Gly
	705				710					715					720

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Ser Gln Ala Thr Met Asp Glu Arg Asn Arg Gln Ile Ala Glu Ile Gly
 725 730 735

Ala Ser Leu Ile Lys His Trp
 740

<210> SEQ ID NO 21
 <211> LENGTH: 786
 <212> TYPE: PRT
 <213> ORGANISM: Homo sapiens

<400> SEQUENCE: 21

Met Ala Gly Leu Thr Ala Ala Ala Pro Arg Pro Gly Val Leu Leu Leu
 1 5 10 15

Leu Leu Ser Ile Leu His Pro Ser Arg Pro Gly Gly Val Pro Gly Ala
 20 25 30

Ile Pro Gly Gly Val Pro Gly Gly Val Phe Tyr Pro Gly Ala Gly Leu
 35 40 45

Gly Ala Leu Gly Gly Gly Ala Leu Gly Pro Gly Gly Lys Pro Leu Lys
 50 55 60

Pro Val Pro Gly Gly Leu Ala Gly Ala Gly Leu Gly Ala Gly Leu Gly
 65 70 75 80

Ala Phe Pro Ala Val Thr Phe Pro Gly Ala Leu Val Pro Gly Gly Val
 85 90 95

Ala Asp Ala Ala Ala Tyr Lys Ala Ala Lys Ala Gly Ala Gly Leu
 100 105 110

Gly Gly Val Pro Gly Val Gly Gly Leu Gly Val Ser Ala Gly Ala Val
 115 120 125

Val Pro Gln Pro Gly Ala Gly Val Lys Pro Gly Lys Val Pro Gly Val
 130 135 140

Gly Leu Pro Gly Val Tyr Pro Gly Gly Val Leu Pro Gly Ala Arg Phe
 145 150 155 160

Pro Gly Val Gly Val Leu Pro Gly Val Pro Thr Gly Ala Gly Val Lys
 165 170 175

Pro Lys Ala Pro Gly Val Gly Gly Ala Phe Ala Gly Ile Pro Gly Val
 180 185 190

Gly Pro Phe Gly Gly Pro Gln Pro Gly Val Pro Leu Gly Tyr Pro Ile
 195 200 205

Lys Ala Pro Lys Leu Pro Gly Gly Tyr Gly Leu Pro Tyr Thr Thr Gly
 210 215 220

Lys Leu Pro Tyr Gly Tyr Gly Pro Gly Gly Val Ala Gly Ala Ala Gly
 225 230 235 240

Lys Ala Gly Tyr Pro Thr Gly Thr Gly Val Gly Pro Gln Ala Ala Ala
 245 250 255

Ala Ala Ala Ala Lys Ala Ala Ala Lys Phe Gly Ala Gly Ala Ala Gly
 260 265 270

Val Leu Pro Gly Val Gly Gly Ala Gly Val Pro Gly Val Pro Gly Ala
 275 280 285

Ile Pro Gly Ile Gly Gly Ile Ala Gly Val Gly Thr Pro Ala Ala Ala
 290 295 300

Ala Ala Ala Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala Ala
 305 310 315 320

Gly Leu Val Pro Gly Gly Pro Gly Phe Gly Pro Gly Val Val Gly Val

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325					330					335					
Pro	Gly	Ala	Gly	Val	Pro	Gly	Val	Gly	Val	Pro	Gly	Ala	Gly	Ile	Pro
			340					345						350	
Val	Val	Pro	Gly	Ala	Gly	Ile	Pro	Gly	Ala	Ala	Val	Pro	Gly	Val	Val
		355					360					365			
Ser	Pro	Glu	Ala	Ala	Ala	Lys	Ala	Ala	Ala	Lys	Ala	Ala	Lys	Tyr	Gly
	370					375					380				
Ala	Arg	Pro	Gly	Val	Gly	Val	Gly	Gly	Ile	Pro	Thr	Tyr	Gly	Val	Gly
	385			390					395					400	
Ala	Gly	Gly	Phe	Pro	Gly	Phe	Gly	Val	Gly	Val	Gly	Gly	Ile	Pro	Gly
			405						410					415	
Val	Ala	Gly	Val	Pro	Gly	Val	Gly	Gly	Val	Pro	Gly	Val	Gly	Gly	Val
		420					425						430		
Pro	Gly	Val	Gly	Ile	Ser	Pro	Glu	Ala	Gln	Ala	Ala	Ala	Ala	Ala	Lys
		435					440						445		
Ala	Ala	Lys	Tyr	Gly	Ala	Ala	Gly	Ala	Gly	Val	Leu	Gly	Gly	Leu	Val
	450			455							460				
Pro	Gly	Pro	Gln	Ala	Ala	Val	Pro	Gly	Val	Pro	Gly	Thr	Gly	Gly	Val
	465			470					475					480	
Pro	Gly	Val	Gly	Thr	Pro	Ala	Ala	Ala	Ala	Ala	Lys	Ala	Ala	Ala	Lys
			485					490						495	
Ala	Ala	Gln	Phe	Gly	Leu	Val	Pro	Gly	Val	Gly	Val	Ala	Pro	Gly	Val
		500					505					510			
Gly	Val	Ala	Pro	Gly	Val	Gly	Val	Ala	Pro	Gly	Val	Gly	Leu	Ala	Pro
		515					520					525			
Gly	Val	Gly	Val	Ala	Pro	Gly	Val	Gly	Val	Ala	Pro	Gly	Val	Gly	Val
	530			535					540						
Ala	Pro	Gly	Ile	Gly	Pro	Gly	Gly	Val	Ala	Ala	Ala	Ala	Lys	Ser	Ala
	545			550					555						560
Ala	Lys	Val	Ala	Ala	Lys	Ala	Gln	Leu	Arg	Ala	Ala	Ala	Gly	Leu	Gly
			565					570						575	
Ala	Gly	Ile	Pro	Gly	Leu	Gly	Val	Gly	Val	Pro	Gly	Leu	Gly		
		580					585					590			
Val	Gly	Ala	Gly	Val	Pro	Gly	Leu	Gly	Val	Gly	Ala	Gly	Val	Pro	Gly
		595					600					605			
Phe	Gly	Ala	Gly	Ala	Asp	Glu	Gly	Val	Arg	Arg	Ser	Leu	Ser	Pro	Glu
	610			615					620						
Leu	Arg	Glu	Gly	Asp	Pro	Ser	Ser	Ser	Gln	His	Leu	Pro	Ser	Thr	Pro
	625			630					635					640	
Ser	Ser	Pro	Arg	Val	Pro	Gly	Ala	Leu	Ala	Ala	Ala	Lys	Ala	Ala	Lys
			645					650						655	
Tyr	Gly	Ala	Ala	Val	Pro	Gly	Val	Leu	Gly	Gly	Leu	Gly	Ala	Leu	Gly
		660						665					670		
Gly	Val	Gly	Ile	Pro	Gly	Gly	Val	Val	Gly	Ala	Gly	Pro	Ala	Ala	Ala
		675					680					685			
Ala	Ala	Ala	Ala	Lys	Ala	Ala	Ala	Lys	Ala	Ala	Gln	Phe	Gly	Leu	Val
				690			695					700			
Gly	Ala	Ala	Gly	Leu	Gly	Gly	Leu	Gly	Val	Gly	Gly	Leu	Gly	Val	Pro
	705			710					715					720	
Gly	Val	Gly	Gly	Leu	Gly	Gly	Ile	Pro	Pro	Ala	Ala	Ala	Ala	Lys	Ala
			725					730						735	

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Ala Lys Tyr Gly Ala Ala Gly Leu Gly Gly Val Leu Gly Gly Ala Gly
 740 745 750

Gln Phe Pro Leu Gly Gly Val Ala Ala Arg Pro Gly Phe Gly Leu Ser
 755 760 765

Pro Ile Phe Pro Gly Gly Ala Cys Leu Gly Lys Ala Cys Gly Arg Lys
 770 775 780

Arg Lys
 785

<210> SEQ ID NO 22
 <211> LENGTH: 2361
 <212> TYPE: DNA
 <213> ORGANISM: Homo sapiens

<400> SEQUENCE: 22

```

atggcggggtc tgacggcggc gggccccggg cccggagtcc tctgtctct gctgtccatc    60
ctccaccctc ctgcggctgg aggggtccct ggggccattc ctggtggagt tcctggagga    120
gtctttttatc caggggctgg tctcggagcc cttggaggag gagecgtggg gcctggaggg    180
aaacctctta agccagttcc cggagggett gcgggtgctg gccttggggc agggctcggc    240
gccttccccg cagttacctt tccgggggct ctggtgcctg gtggagtggc tgacgtgct    300
gcagcctata aagctgctaa ggtgggcgtt gggcttgggt gtgtcccagg agttggtggc    360
ttaggagtgct ctgcaggtgc ggtggttccct cagcctggag ccggagtgaa gcctgggaaa    420
gtgccgggtg tggggctgcc aggtgtatac ccaggtggcg tgctcccagg agtcgggttc    480
cccgtgtggt ggggtctccc tggagtcccc actggagcag gagttaagcc caaggetcca    540
ggtgtagggtg gagcttttgc tggaatccca ggagttggac cctttggggg accgcaacct    600
ggagtcccac tggggtatcc catcaaggcc cccaagctgc ctggtggcta tggactgccc    660
tacaccacag ggaactgcc ctatggctat gggcccggag gagtggctgg tgcagcgggc    720
aaggctgggtt acccaacagg gacaggggtt gggccccagg cagcagcagc agcggcagct    780
aaagcagcag caaagtctcg tctcggagca gccggagtcc tccctggtgt tggaggggct    840
ggtgttccctg gcgtgcctgg ggcattccct ggaattggag gcatcgcagg cgttgggact    900
ccagctgcag ctgcagctgc agcagcagcc gctaaggcag ccaagtatgg agctgctgca    960
ggcttagtgc ctggtgggcc aggccttggc ccgggagtag ttggtgtccc aggagctggc    1020
gttccagggtg ttggtgtccc aggagctggg attccagttg tcccagggtc tgggatccca    1080
ggtgctgcgg ttccaggggt tgtgtcacca gaagcagctg ctaaggcagc tgcaaaaggca    1140
gccccaaatcg gggccaggcc cggagtcgga gttggaggca ttccactta cggggttgga    1200
gctgggggctt tccccggtt tgggtctgga gtcggaggta tccctggagt cgcaggtgtc    1260
cctggtgtcg gaggtgttcc cggagtcgga ggtgtcccgg gagtggcat tcccccgaa    1320
gctcaggcag cagctgccgc caaggctgcc aagtacggtg ctgcaggagc aggagtgctg    1380
ggtgggctag tgccaggctc ccaggcggca gtcccagggt tgcccggcac gggaggagtg    1440
ccaggagtgg ggaacccagc agctgcagct gctaagcag ccgcccaggc cgcccagttt    1500
gggttagttc ctggtgtcgg cgtggctcct ggagttggcg tggtccctgg tgtcgggtgtg    1560
gctcctggag ttggcttggc tctcggagtt ggcgtggctc ctggagttgg tgtggctcct    1620
ggcgttggcg tggctcccgg cattggccct ggtggagttg cagctgcagc aaaatccgct    1680
    
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gccaaggtgg ctgccaagc ccagctccga gctgcagctg ggcttgggtg tggcatccct 1740
ggacttgag ttggtgtcgg cgtccctgga cttggagttg gtgctggtgt tcttgactt 1800
ggagttggtg ctggtgttcc tggcttcggg gcaggtgcag atgagggagt taggcggagc 1860
ctgtccctg agctcagga aggagatccc tctctctctc agcacctccc cagcaccccc 1920
tcatcaccca gggtagctgg agccctggct gccgctaaag cagccaaata tggagcagca 1980
gtgctggggg tccttgagg gctcggggct ctcggtggag taggcatccc aggcggtgtg 2040
gtgggagcgg gaccgcgcgc cgcctgtgcc gcagccaaag ctgctgcca agcgcgccag 2100
tttgccctag tgggagcgc tgggctcggg ggactcggag tcggagggtt tggagttcca 2160
ggtgttgggg gccttgagg tatacctcca gctgcagcgc ctaaagcagc taaatacggc 2220
gctgctggcc ttggaggtgt cctagggggg gccgggcagt tcccacttg aggagtggca 2280
gcaagacctg gcttcgatt gtctccatt ttcccagggt gggcctgctt ggggaaagct 2340
tgtggccgga agagaaaatg a 2361

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<210> SEQ ID NO 23

<211> LENGTH: 2418

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 23

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atgaaaaaga tttggctggc gctggctggt ttagttagc cgttagcgc atcggcggcg 60
cagtatgaag atcaccatca ccaccaccac catcaccact ctggctcag cctggtgccc 120
cgcggcagcc atatgggtgg cgtaccagc gcaattcctg ggggtgtccc aggcggtgtt 180
ttttatccgg gcgcgggtct tggcgcactg ggtggcgggt cactgggccc gggcggcaaa 240
ccgctgaaac cggtagcagg tggttagca ggcgcggct taggcagcagg tctgggagca 300
tttcggcag ttaaccttcc aggggcactg gttcctggag gtgtggcga tgcagccgcg 360
gcataaaag ccgctaaagc cggtgccggg ttaggagcgc tcccaggtgt cggtgccctg 420
ggtgttagcg ccggtgcagt tgttccgcag ccgggagcag gggttaaac tggtaaagt 480
ccgggagtag gtctgccagg cgtttatcct ggtggtgttt tgccgggtgc cgttttccg 540
ggcgttgggt ttcttccagg cgtgccgacc ggagccgggt taaaccgaa agcccccggt 600
gttgagggtg catttgcagg catcccggga gttggcccgt ttggtggtcc gcaacctggg 660
gttccgtag gttatccgat taaagcaccg aaactgcccg gcggttatgg tctgccgtac 720
acaaccggtg aactgccgta tggttatggc ccgggtggag ttgcgggtgc agcaggtaaa 780
gcgggttatc ctaccggaac cgggttaggt ccgagggcgc ctgctgccgc cgcgcgcaaa 840
gcagcgggta aatttggcgc cggagcagc ggtgttctgc ctggagttgg tgggtgcggc 900
gtgccagggg tacctggtgc aattccgggt attggtggta ttgcccgggt cggcaccgcc 960
gccgcggcag ctgcccagc ggcggctgcc aaagctgcta aatacgggtc cgcggcgggt 1020
ctggtgccag gagtccggg ttttggctcc ggagtgggtg gcgtgcctgg cgcagcgctt 1080
cctggtgtgg gcgttccagg tgcagggatt cctgtgtgct ctggtgccgg tattcccgcc 1140
gcggccggtc cgggggtggt tagcccggaa gccgcagcga aggctgcggc aaaggcagca 1200

```

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aagtatggcg cacgcccagg agtcggcgtg ggtggtatcc cgacctatgg ggtgggcgca 1260
gggggttttc ctggtttcgg cgtaggtgta ggaggatac cgggcgtggc cgggtgtacca 1320
gggggttggtg gcgtccctgg tgttgccggt gtgccaggtg ttggtatttc accggaagca 1380
caggcagcag ccgcagctaa ggcagcgaaa tatggtgccg ccggcgcagg agtttttaggt 1440
gggctggttc cgggcccgca ggcagctgtg ccgggggttc caggcacccg tggtgtccct 1500
ggagtcggta cgccggctgc agcggcagcc aaagcggctg cgaaagcagc acagtttggc 1560
ttagtaccgg gtgtgggagt tgccccggc gttggcgttg ctccaggggt ggggtgttget 1620
cctggcgtcg gtctggctcc tggagtgggc gtagcaccgg gtgtgggggt ggccccgggt 1680
gttggggttg caccgggtat cgtccgggc ggtgtcgcag cagcagctaa aagcgcggcg 1740
aaagttgcgg ccaaagccca actgcgcgcc gccgcgggcc tcggtgcagg tattccgggg 1800
ctgggtgtcg gagttggagt cccgggtttg ggcgtgggcg cgggagttcc gggactggga 1860
gtgggtgccg gagttcctgg ctttggtgca ggcgcagatg aaggtgttcg tcgtagcctg 1920
agtccggaac tgcgtgaagg tgatccgagt agcagccagc atctgccgag caccgccagc 1980
agcccgcgtg ttccgggtgc attagctgca gcaaaagccg ccaagtatgg tgcagccgtg 2040
ccgggcgtct taggtgtct gggcgcctcg ggtggtgtag gcattccggg aggtgttgtg 2100
ggtgcaggac cggccgcgcg agctgcggcc gccaaagcag ctgcaaaagc ggcccagttt 2160
ggtttagtgg gcgcccagc tttaggcggg ttaggtgtgg gtggactggg tgtacctggc 2220
gtaggcggtc tgggtggaat tccgccgca cgggcgcgca aagcggcaaa atatggcgcg 2280
gcaggcctgg gcggcgtgct gggtggggca ggtcagtttc cgctgggcgg ggttgccgca 2340
cgccgggat ttggtctgag cccgattttc cctggcggcg catgtctggg taaagcatgt 2400
ggtcgtaaac gtaaataa 2418
    
```

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<210> SEQ ID NO 24
<211> LENGTH: 805
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
    polypeptide
    
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<400> SEQUENCE: 24

```

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
1           5           10           15
Ala Ser Ala Ala Gln Tyr Glu Asp His His His His His His His His
20          25          30
His Ser Gly Ser Ser Leu Val Pro Arg Gly Ser His Met Gly Gly Val
35          40          45
Pro Gly Ala Ile Pro Gly Gly Val Pro Gly Gly Val Phe Tyr Pro Gly
50          55          60
Ala Gly Leu Gly Ala Leu Gly Gly Gly Ala Leu Gly Pro Gly Gly Lys
65          70          75          80
Pro Leu Lys Pro Val Pro Gly Gly Leu Ala Gly Ala Gly Leu Gly Ala
85          90          95
Gly Leu Gly Ala Phe Pro Ala Val Thr Phe Pro Gly Ala Leu Val Pro
100         105         110
Gly Gly Val Ala Asp Ala Ala Ala Tyr Lys Ala Ala Lys Ala Gly
115         120         125
    
```

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Ala Gly Leu Gly Gly Val Pro Gly Val Gly Gly Leu Gly Val Ser Ala
130 135 140

Gly Ala Val Val Pro Gln Pro Gly Ala Gly Val Lys Pro Gly Lys Val
145 150 155 160

Pro Gly Val Gly Leu Pro Gly Val Tyr Pro Gly Gly Val Leu Pro Gly
165 170 175

Ala Arg Phe Pro Gly Val Gly Val Leu Pro Gly Val Pro Thr Gly Ala
180 185 190

Gly Val Lys Pro Lys Ala Pro Gly Val Gly Gly Ala Phe Ala Gly Ile
195 200 205

Pro Gly Val Gly Pro Phe Gly Gly Pro Gln Pro Gly Val Pro Leu Gly
210 215 220

Tyr Pro Ile Lys Ala Pro Lys Leu Pro Gly Gly Tyr Gly Leu Pro Tyr
225 230 235 240

Thr Thr Gly Lys Leu Pro Tyr Gly Tyr Gly Pro Gly Gly Val Ala Gly
245 250 255

Ala Ala Gly Lys Ala Gly Tyr Pro Thr Gly Thr Gly Val Gly Pro Gln
260 265 270

Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala Ala Lys Phe Gly Ala Gly
275 280 285

Ala Ala Gly Val Leu Pro Gly Val Gly Gly Ala Gly Val Pro Gly Val
290 295 300

Pro Gly Ala Ile Pro Gly Ile Gly Gly Ile Ala Gly Val Gly Thr Pro
305 310 315 320

Ala Ala Ala Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala Lys Tyr Gly
325 330 335

Ala Ala Ala Gly Leu Val Pro Gly Gly Pro Gly Phe Gly Pro Gly Val
340 345 350

Val Gly Val Pro Gly Ala Gly Val Pro Gly Val Gly Val Pro Gly Ala
355 360 365

Gly Ile Pro Val Val Pro Gly Ala Gly Ile Pro Gly Ala Ala Val Pro
370 375 380

Gly Val Val Ser Pro Glu Ala Ala Ala Lys Ala Ala Ala Lys Ala Ala
385 390 395 400

Lys Tyr Gly Ala Arg Pro Gly Val Gly Val Gly Gly Ile Pro Thr Tyr
405 410 415

Gly Val Gly Ala Gly Gly Phe Pro Gly Phe Gly Val Gly Val Gly Gly
420 425 430

Ile Pro Gly Val Ala Gly Val Pro Gly Val Gly Gly Val Pro Gly Val
435 440 445

Gly Gly Val Pro Gly Val Gly Ile Ser Pro Glu Ala Gln Ala Ala Ala
450 455 460

Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala Gly Ala Gly Val Leu Gly
465 470 475 480

Gly Leu Val Pro Gly Pro Gln Ala Ala Val Pro Gly Val Pro Gly Thr
485 490 495

Gly Gly Val Pro Gly Val Gly Thr Pro Ala Ala Ala Ala Ala Lys Ala
500 505 510

Ala Ala Lys Ala Ala Gln Phe Gly Leu Val Pro Gly Val Gly Val Ala
515 520 525

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Pro Gly Val Gly Val Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly
 530 535 540

Leu Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Val Ala Pro Gly
 545 550 555 560

Val Gly Val Ala Pro Gly Ile Gly Pro Gly Gly Val Ala Ala Ala Ala
 565 570 575

Lys Ser Ala Ala Lys Val Ala Ala Lys Ala Gln Leu Arg Ala Ala Ala
 580 585 590

Gly Leu Gly Ala Gly Ile Pro Gly Leu Gly Val Gly Val Gly Val Pro
 595 600 605

Gly Leu Gly Val Gly Ala Gly Val Pro Gly Leu Gly Val Gly Ala Gly
 610 615 620

Val Pro Gly Phe Gly Ala Gly Ala Asp Glu Gly Val Arg Arg Ser Leu
 625 630 635 640

Ser Pro Glu Leu Arg Glu Gly Asp Pro Ser Ser Ser Gln His Leu Pro
 645 650 655

Ser Thr Pro Ser Ser Pro Arg Val Pro Gly Ala Leu Ala Ala Ala Lys
 660 665 670

Ala Ala Lys Tyr Gly Ala Ala Val Pro Gly Val Leu Gly Gly Leu Gly
 675 680 685

Ala Leu Gly Gly Val Gly Ile Pro Gly Gly Val Val Gly Ala Gly Pro
 690 695 700

Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala Ala Lys Ala Ala Gln Phe
 705 710 715 720

Gly Leu Val Gly Ala Ala Gly Leu Gly Gly Leu Gly Val Gly Gly Leu
 725 730 735

Gly Val Pro Gly Val Gly Gly Leu Gly Gly Ile Pro Pro Ala Ala Ala
 740 745 750

Ala Lys Ala Ala Lys Tyr Gly Ala Ala Gly Leu Gly Gly Val Leu Gly
 755 760 765

Gly Ala Gly Gln Phe Pro Leu Gly Gly Val Ala Ala Arg Pro Gly Phe
 770 775 780

Gly Leu Ser Pro Ile Phe Pro Gly Gly Ala Cys Leu Gly Lys Ala Cys
 785 790 795 800

Gly Arg Lys Arg Lys
 805

<210> SEQ ID NO 25
 <211> LENGTH: 2358
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polynucleotide

<400> SEQUENCE: 25

atgaaaaaga tttggctggc gctggctggt ttagtttag cgtttagcgc atcgcgggcg 60

cagtatgaag atatgggtgg cgtaccaggc gcaattcctg ggggtgtccc aggcggtggt 120

ttttatccgg gcgccggtct tggcgcactg ggtggcggtg cactgggccc gggcgcaaaa 180

ccgctgaaac cggtagcagg tggttttagca ggcgcggct taggcgcagg tctgggagca 240

tttccggcag ttacctttcc aggggcactg gttcctggag gtgtggccga tgcagccgcg 300

gcataaaag ccgctaaagc cgggtcgggt ttaggaggcg tcccaggtgt cgggtggcctg 360

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gggtgtagcg cgggtgcagt tgttccgcag cggggagcag gggttaaacc tggtaaagtg 420
ccgggagtag gtctgccagg cgtttatcct ggtggtgttt tgccgggtgc cgtttttccg 480
ggcgttggtg ttcttccagg cgtgccgacc ggagccggtg ttaaaccgaa agcccccggt 540
gttgaggtg catttgcagg catcccggga gttggcccgt ttggtggtcc gcaacctggg 600
gttccgtag gttatccgat taaagcaccg aaactgcccg gcggttatgg tctgccgtac 660
acaaccggtg aactgccgta tggttatggc cgggtgggag ttgcccgtgc agcaggtaaa 720
gcccgttata ctaccggaac cgtgttaggt ccgcaggccg ctgctgcgcg cgcgcgcaaaa 780
gcagcggcta aatttggcgc cggagcagcg ggtgttctgc ctggagtggg tgggtcgggc 840
gtgccagggg tacctggtgc aattccgggt attggtggtg ttgcccgtgt cggcaacccc 900
gcccggcag ctgcccagc ggcggctgcc aaagctgcta aatacgggtg cgcggcgggt 960
ctggtgccag gaggtccggg ttttggctcg ggagtgggtg gcgtgcccgg cgcaggcgtt 1020
cctggtgtgg gcgttccagg tgcagggatt cctgttctgc ctggtgccgg tattcccggc 1080
gcccgcgttc cgggggtggt tagcccggaa gccgcagcga aggctgcgcg aaaggcagca 1140
aagtatggcg cacgcccagg agtcggcgtg ggtggtatcc cgacctatgg ggtgggcgca 1200
gggggttttc ctggtttcgg cgtaggtgta ggaggtatac cgggcgtggc cgggtgtacca 1260
ggggttggtg gcgtcccctg tgttggcggg gtgccaggtg ttggtatttc accggaagca 1320
caggcagcag ccgcagctaa ggcagcgaaa tatggtgccg ccggcgcagg agttttaggt 1380
gggctggttc cgggcccgcg gccagctgtg cgggggggtc caggcacccg tgggtgtccct 1440
ggagtcggta cgcggcgtgc agcggcagcc aaagcggctg cgaaagcagc acagtttggc 1500
ttagtaccgg gtgtgggagt tgccccggc gttggcgttg ctccaggggt ggggtgttgc 1560
cctggcgtcg gtctggtctc tggagtgggc gtagcaccgg gtgtgggggt ggccccgggt 1620
gttgggggtg caccgggtat cgttccgggc ggtgtcgcag cagcagctaa aagcgcggcg 1680
aaagttgcgg ccaaagccca actgcgcgcc gccgcgggcc tcggtgcagg tattccgggg 1740
ctgggtgtcg gagttggagt cccgggtttg ggcgtggcgc cgggagttcc gggactggga 1800
gtgggtgccc gagttcctgg ctttgggtgca ggcgcagatg aaggtgttcg tcgtagcctg 1860
agtccggaac tgcgtgaagg tgatccgagt agcagccagc atctgccgag caccocgagc 1920
agcccgcgtg ttccgggtgc attagctgca gcaaaagccg ccaagtatgg tgcagccgtg 1980
ccgggcgtct taggtggtct gggcgcctcg ggtggtgtag gcattccggg aggtgttgtg 2040
ggtgcaggac cggccgcgcg agctgcggcc gccaaagcag ctgcaaaagc ggcccagttt 2100
ggtttagtgg gcgcccagc tttaggcggg ttaggtgtgg gtggactggg tgtacctggc 2160
gtaggcggtc tgggtggaat tccgccgcga gcggccgcga aagcggcaaa atatggcgcg 2220
gcagccctgg gcggcgtgct ggggtgggca ggtcagtttc cgctgggccc ggttgcgcga 2280
cgtccgggat ttggtctgag cccgattttc cctggcggcg catgtctggg taaagcatgt 2340
ggtcgtaaac gtaaataa 2358

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<210> SEQ ID NO 26

<211> LENGTH: 785

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic

-continued

polypeptide

<400> SEQUENCE: 26

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
 1 5 10 15

Ala Ser Ala Ala Gln Tyr Glu Asp Met Gly Gly Val Pro Gly Ala Ile
 20 25 30

Pro Gly Gly Val Pro Gly Gly Val Phe Tyr Pro Gly Ala Gly Leu Gly
 35 40 45

Ala Leu Gly Gly Gly Ala Leu Gly Pro Gly Gly Lys Pro Leu Lys Pro
 50 55 60

Val Pro Gly Gly Leu Ala Gly Ala Gly Leu Gly Ala Gly Leu Gly Ala
 65 70 75 80

Phe Pro Ala Val Thr Phe Pro Gly Ala Leu Val Pro Gly Gly Val Ala
 85 90 95

Asp Ala Ala Ala Ala Tyr Lys Ala Ala Lys Ala Gly Ala Gly Leu Gly
 100 105 110

Gly Val Pro Gly Val Gly Gly Leu Gly Val Ser Ala Gly Ala Val Val
 115 120 125

Pro Gln Pro Gly Ala Gly Val Lys Pro Gly Lys Val Pro Gly Val Gly
 130 135 140

Leu Pro Gly Val Tyr Pro Gly Gly Val Leu Pro Gly Ala Arg Phe Pro
 145 150 155 160

Gly Val Gly Val Leu Pro Gly Val Pro Thr Gly Ala Gly Val Lys Pro
 165 170 175

Lys Ala Pro Gly Val Gly Gly Ala Phe Ala Gly Ile Pro Gly Val Gly
 180 185 190

Pro Phe Gly Gly Pro Gln Pro Gly Val Pro Leu Gly Tyr Pro Ile Lys
 195 200 205

Ala Pro Lys Leu Pro Gly Gly Tyr Gly Leu Pro Tyr Thr Thr Gly Lys
 210 215 220

Leu Pro Tyr Gly Tyr Gly Pro Gly Gly Val Ala Gly Ala Ala Gly Lys
 225 230 235 240

Ala Gly Tyr Pro Thr Gly Thr Gly Val Gly Pro Gln Ala Ala Ala Ala
 245 250 255

Ala Ala Ala Lys Ala Ala Ala Lys Phe Gly Ala Gly Ala Ala Gly Val
 260 265 270

Leu Pro Gly Val Gly Gly Ala Gly Val Pro Gly Val Pro Gly Ala Ile
 275 280 285

Pro Gly Ile Gly Gly Ile Ala Gly Val Gly Thr Pro Ala Ala Ala Ala
 290 295 300

Ala Ala Ala Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala Ala Gly
 305 310 315 320

Leu Val Pro Gly Gly Pro Gly Phe Gly Pro Gly Val Val Gly Val Pro
 325 330 335

Gly Ala Gly Val Pro Gly Val Gly Val Pro Gly Ala Gly Ile Pro Val
 340 345 350

Val Pro Gly Ala Gly Ile Pro Gly Ala Ala Val Pro Gly Val Val Ser
 355 360 365

Pro Glu Ala Ala Ala Lys Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala
 370 375 380

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Arg Pro Gly Val Gly Val Gly Gly Ile Pro Thr Tyr Gly Val Gly Ala
 385 390 395 400

Gly Gly Phe Pro Gly Phe Gly Val Gly Val Gly Gly Ile Pro Gly Val
 405 410 415

Ala Gly Val Pro Gly Val Gly Gly Val Pro Gly Val Gly Gly Val Pro
 420 425 430

Gly Val Gly Ile Ser Pro Glu Ala Gln Ala Ala Ala Ala Lys Ala
 435 440 445

Ala Lys Tyr Gly Ala Ala Gly Ala Gly Val Leu Gly Gly Leu Val Pro
 450 455 460

Gly Pro Gln Ala Ala Val Pro Gly Val Pro Gly Thr Gly Gly Val Pro
 465 470 475 480

Gly Val Gly Thr Pro Ala Ala Ala Ala Ala Lys Ala Ala Ala Lys Ala
 485 490 495

Ala Gln Phe Gly Leu Val Pro Gly Val Gly Val Ala Pro Gly Val Gly
 500 505 510

Val Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Leu Ala Pro Gly
 515 520 525

Val Gly Val Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Val Ala
 530 535 540

Pro Gly Ile Gly Pro Gly Gly Val Ala Ala Ala Ala Lys Ser Ala Ala
 545 550 555 560

Lys Val Ala Ala Lys Ala Gln Leu Arg Ala Ala Ala Gly Leu Gly Ala
 565 570 575

Gly Ile Pro Gly Leu Gly Val Gly Val Gly Val Pro Gly Leu Gly Val
 580 585 590

Gly Ala Gly Val Pro Gly Leu Gly Val Gly Ala Gly Val Pro Gly Phe
 595 600 605

Gly Ala Gly Ala Asp Glu Gly Val Arg Arg Ser Leu Ser Pro Glu Leu
 610 615 620

Arg Glu Gly Asp Pro Ser Ser Ser Gln His Leu Pro Ser Thr Pro Ser
 625 630 635 640

Ser Pro Arg Val Pro Gly Ala Leu Ala Ala Ala Lys Ala Ala Lys Tyr
 645 650 655

Gly Ala Ala Val Pro Gly Val Leu Gly Gly Leu Gly Ala Leu Gly Gly
 660 665 670

Val Gly Ile Pro Gly Gly Val Val Gly Ala Gly Pro Ala Ala Ala Ala
 675 680 685

Ala Ala Ala Lys Ala Ala Ala Lys Ala Ala Gln Phe Gly Leu Val Gly
 690 695 700

Ala Ala Gly Leu Gly Gly Leu Gly Val Gly Gly Leu Gly Val Pro Gly
 705 710 715 720

Val Gly Gly Leu Gly Gly Ile Pro Pro Ala Ala Ala Ala Lys Ala Ala
 725 730 735

Lys Tyr Gly Ala Ala Gly Leu Gly Gly Val Leu Gly Gly Ala Gly Gln
 740 745 750

Phe Pro Leu Gly Gly Val Ala Ala Arg Pro Gly Phe Gly Leu Ser Pro
 755 760 765

Ile Phe Pro Gly Gly Ala Cys Leu Gly Lys Ala Cys Gly Arg Lys Arg
 770 775 780

Lys

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785

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<210> SEQ ID NO 27
<211> LENGTH: 3945
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
        polynucleotide

<400> SEQUENCE: 27

atgaaaaaga tttggctggc gctggctggt ttagttttag cgtttagcgc atcgcgcgcg      60
cagtatgaag atcaccatca ccaccaccac catcaccact ctggctcgag cctggtgccg      120
cgcggcagcc atatgtctgg ctcgagcagt aaaggtgaag aactgttcac cggtgttgtt      180
ccgatcctgg ttgaactgga tggtagtgtt aacggccaca aattctctgt tctggtgtaa      240
ggtgaaggtg atgcaaccaa cggtaaacctg accctgaaat tcatctgcac taccggtaaa      300
ctgccggttc catggcgcac tctggtgact accctgacct atggtgttca gtgtttttct      360
cgttaccggy atcacatgaa gcagcatgat ttcttcaaat ctgcaatgcc ggaaggttat      420
gtacaggagc gcaccatttc tttcaaagac gatggcacct acaaaaccgc tgcagaggtt      480
aaatttgaag gtgatactct ggtgaaccgt attgaactga aaggcattga tttcaaagag      540
gacggcaaca tcctggggcca caaacggaa tataacttca actcccataa cgtttacatc      600
accgcagaca aacagaagaa cggtatcaaa gctaacttca aaattcgcca taacgttgaa      660
gacggtagcg tacagctggc ggaccactac cagcagaaca ctccgatcgg tgatggtccg      720
gttctgctgc cggataacca ctacctgtcc acccagteta aactgtccaa agaccggaac      780
gaaaagcgcg accacatggt gctgctggag ttcgttactg cagcaggtat cacgcacggc      840
atggtatgaac tctacaaatc tggcgcgcgc ggcggtggcg taccaggcgc aatctcctggg      900
ggtgtcccag cgggtgtttt ttatccgggc gccggtcttg gcgcactggg tggcggtgca      960
ctgggcccgg gcgcgaaacc gctgaaaccg gtaccaggtg gtttagcagg cgccgggetta      1020
ggcgcaggtc tgggagcatt tccggcagtt acctttccag gggcactggt tcctggaggt      1080
gtggccgatg cagccgcggc atataaagcc gctaaagccg gtgcgggttt aggaggcgtc      1140
ccaggtgtcg gtggcctggg tgttagcgcc ggtgcagttg ttccgcagcc gggagcaggg      1200
gttaaacctg gtaaaagtcc gggagtaggt ctgccagcgg tttatcctgg tgggtgtttg      1260
ccgggtgccc gttttccggg cgttgggtgtt cttccaggcg tgccgaccgg agccgggtgtt      1320
aaaccgaaag cccccggtgt tggaggtgca tttgcaggca tcccgggagt tggcccgttt      1380
ggtggtccgc aacctggggt tccggttaggt tatccgatta aagcaccgaa actgccccggc      1440
ggttatggtc tgccgtacac aaccggtaaa ctgccgatg gttatggccc ggggtggagtt      1500
gccccgtgag caggtaaagc gggttatcct accggaaccg gtgtaggtcc gcaggccgct      1560
gctgccgcgc ccgcaaaagc agcggctaaa tttggcgcgc gacgagcggg tgttctgcct      1620
ggagttggtg gtgcgggctg gccaggggta cctggtgcaa ttccgggtat tgggtgtatt      1680
gccggtgtcg gcaccccggc cgcggcagct gcggcagcgg cggctgcaaa agctgctaaa      1740
tacggtgccc cggcgggtct ggtgccagga ggtccgggtt ttggtccggg agtgggttggc      1800
gtgcctggcg caggcgttcc tgggtggggc gttccaggtg cagggattcc tgttgtgcct      1860
ggtgccggta ttccccgcgc ggcggttccg ggggtggtta gcccggaagc cgcagcgaag      1920

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gctgcggaag aggcagcaaa gtatggcgca cccccaggag tcggcgtggg tggatcccc 1980
acctatgggg tgggcgcagg gggttttcct ggtttcggcg taggtgtagg aggtataccg 2040
ggcgtggcgg gtgtaccagg ggttggtggc gtccctggcg ttggcgggtg gccaggtggt 2100
ggtattttcac cggaaacaca ggcagcagcc gcagetaagg cagcgaataa tggtgccgcc 2160
ggcgcaggag ttttaggtgg gctggttccg gccccgcagg cagctgtgcc gggggttcca 2220
ggcaccgggtg gtgtccctgg agtcggtagc ccggctgcag cggcagccaa agcggctgcg 2280
aaagcagcac agtttggtct agtaccgggt gtgggagttg cccccggcgt tggcgttget 2340
ccaggggtgg gtgttctcc tggcgtcggc ctggctcctg gagtggcgt agcaccgggt 2400
gtgggggtgg cccccgggtg tggggttgca ccgggtatcg gtccgggcgg tgtcgcagca 2460
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ggtgcaggta tccggggct ggggtgcgga gttggagtc cgggtttggg cgtgggcgcg 2580
ggagttccgg gactgggagt ggggtgcgga gttcctggct ttggtgcagg cgcagatgaa 2640
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ctgccagaca ccccagcag cccgcgtggt ccgggtgcat tagctgcagc aaaagccgcc 2760
aagtatggtg cagccgtgcc gggcgtctta ggtggtctgg gcgccctggg tgggttaggc 2820
attccgggag gtgttgggg tgcaggaccg gccgcgcag ctgcggcgcg caaagcagct 2880
gcaaaagcgg cccagtttg tttagtgggc gccgcagggt taggcggttt aggtgtgggt 2940
ggactgggtg tacctggcgt aggcggtctg ggtggaattc cgcgcgcagc ggccgcgaaa 3000
gcccgaaaat atggcgcggc aggcctgggc ggcgtgctgg gtggggcagg tcagtttccg 3060
ctgggcgggg ttgccgcag tccgggattt ggtctgagcc cgattttccc tggcggcgca 3120
tgtctgggta aagcatgtgg tcgtaaacgt aaaccccag aaacgctggt gaaagtaaaa 3180
gatgctgaag atcagttggg tgcacgagtg ggttacatcg aactggatct caacagcgg 3240
aagatccttg agagttttcg ccccgaagaa cgttttccaa tgatgagcac ttttaaagtt 3300
ctgctatgtg gcgcgggtatt atccctgatt gacgcgggc aagagcaact cggtcgccc 3360
atacactatt ctcagaatga cttggttag tactcaccag tcacagaaaa gcattcttac 3420
gatggcatga cagtaagaga attatgcagt gctgccataa ccatgagtg taactactgcg 3480
gccaacttac ttctgacaac gatcggagga ccgaaggagc taaccgcttt tttgcacaac 3540
atgggggatc atgtaactcg ccttgatcgt tgggaaccgg agctgaaatga agccatacca 3600
aacgacgagc gtgacaccac gatgcctgta gcaatggcaa caacgctgcg caaactatta 3660
actggcgaac tacttactct agcttcccgg caacaattaa tagactggat ggaggcggat 3720
aaagttgcag gaccacttct gcgctcggcc cttccggctg gctggtttat tgctgataaa 3780
tctggagcgg gtgagcgtgg gtctcggcgt atcattgcag cactggggcc agatggttaag 3840
ccctcccgta tcgtagtat ctacacgacg gggagtcagg caactatgga tgaacgaaat 3900
agacagatcg ctgagatagg tgcctcactg attaagcatt ggtaa 3945

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<210> SEQ ID NO 28

<211> LENGTH: 1314

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic

-continued

polypeptide

<400> SEQUENCE: 28

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
 1 5 10 15

Ala Ser Ala Ala Gln Tyr Glu Asp His His His His His His His His
 20 25 30

His Ser Gly Ser Ser Leu Val Pro Arg Gly Ser His Met Ser Gly Ser
 35 40 45

Ser Ser Lys Gly Glu Glu Leu Phe Thr Gly Val Val Pro Ile Leu Val
 50 55 60

Glu Leu Asp Gly Asp Val Asn Gly His Lys Phe Ser Val Arg Gly Glu
 65 70 75 80

Gly Glu Gly Asp Ala Thr Asn Gly Lys Leu Thr Leu Lys Phe Ile Cys
 85 90 95

Thr Thr Gly Lys Leu Pro Val Pro Trp Pro Thr Leu Val Thr Thr Leu
 100 105 110

Thr Tyr Gly Val Gln Cys Phe Ser Arg Tyr Pro Asp His Met Lys Gln
 115 120 125

His Asp Phe Phe Lys Ser Ala Met Pro Glu Gly Tyr Val Gln Glu Arg
 130 135 140

Thr Ile Ser Phe Lys Asp Asp Gly Thr Tyr Lys Thr Arg Ala Glu Val
 145 150 155 160

Lys Phe Glu Gly Asp Thr Leu Val Asn Arg Ile Glu Leu Lys Gly Ile
 165 170 175

Asp Phe Lys Glu Asp Gly Asn Ile Leu Gly His Lys Leu Glu Tyr Asn
 180 185 190

Phe Asn Ser His Asn Val Tyr Ile Thr Ala Asp Lys Gln Lys Asn Gly
 195 200 205

Ile Lys Ala Asn Phe Lys Ile Arg His Asn Val Glu Asp Gly Ser Val
 210 215 220

Gln Leu Ala Asp His Tyr Gln Gln Asn Thr Pro Ile Gly Asp Gly Pro
 225 230 235 240

Val Leu Leu Pro Asp Asn His Tyr Leu Ser Thr Gln Ser Lys Leu Ser
 245 250 255

Lys Asp Pro Asn Glu Lys Arg Asp His Met Val Leu Leu Glu Phe Val
 260 265 270

Thr Ala Ala Gly Ile Thr His Gly Met Asp Glu Leu Tyr Lys Ser Gly
 275 280 285

Ala Pro Gly Gly Gly Val Pro Gly Ala Ile Pro Gly Gly Val Pro Gly
 290 295 300

Gly Val Phe Tyr Pro Gly Ala Gly Leu Gly Ala Leu Gly Gly Gly Ala
 305 310 315 320

Leu Gly Pro Gly Gly Lys Pro Leu Lys Pro Val Pro Gly Gly Leu Ala
 325 330 335

Gly Ala Gly Leu Gly Ala Gly Leu Gly Ala Phe Pro Ala Val Thr Phe
 340 345 350

Pro Gly Ala Leu Val Pro Gly Gly Val Ala Asp Ala Ala Ala Tyr
 355 360 365

Lys Ala Ala Lys Ala Gly Ala Gly Leu Gly Gly Val Pro Gly Val Gly
 370 375 380

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Gly Leu Gly Val Ser Ala Gly Ala Val Val Pro Gln Pro Gly Ala Gly
 385 390 395 400
 Val Lys Pro Gly Lys Val Pro Gly Val Gly Leu Pro Gly Val Tyr Pro
 405 410 415
 Gly Gly Val Leu Pro Gly Ala Arg Phe Pro Gly Val Gly Val Leu Pro
 420 425 430
 Gly Val Pro Thr Gly Ala Gly Val Lys Pro Lys Ala Pro Gly Val Gly
 435 440 445
 Gly Ala Phe Ala Gly Ile Pro Gly Val Gly Pro Phe Gly Gly Pro Gln
 450 455 460
 Pro Gly Val Pro Leu Gly Tyr Pro Ile Lys Ala Pro Lys Leu Pro Gly
 465 470 475 480
 Gly Tyr Gly Leu Pro Tyr Thr Thr Gly Lys Leu Pro Tyr Gly Tyr Gly
 485 490 495
 Pro Gly Gly Val Ala Gly Ala Ala Gly Lys Ala Gly Tyr Pro Thr Gly
 500 505 510
 Thr Gly Val Gly Pro Gln Ala Ala Ala Ala Ala Ala Lys Ala Ala
 515 520 525
 Ala Lys Phe Gly Ala Gly Ala Ala Gly Val Leu Pro Gly Val Gly Gly
 530 535 540
 Ala Gly Val Pro Gly Val Pro Gly Ala Ile Pro Gly Ile Gly Gly Ile
 545 550 555 560
 Ala Gly Val Gly Thr Pro Ala Ala Ala Ala Ala Ala Ala Ala Ala
 565 570 575
 Lys Ala Ala Lys Tyr Gly Ala Ala Ala Gly Leu Val Pro Gly Gly Pro
 580 585 590
 Gly Phe Gly Pro Gly Val Val Gly Val Pro Gly Ala Gly Val Pro Gly
 595 600 605
 Val Gly Val Pro Gly Ala Gly Ile Pro Val Val Pro Gly Ala Gly Ile
 610 615 620
 Pro Gly Ala Ala Val Pro Gly Val Val Ser Pro Glu Ala Ala Ala Lys
 625 630 635 640
 Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala Arg Pro Gly Val Gly Val
 645 650 655
 Gly Gly Ile Pro Thr Tyr Gly Val Gly Ala Gly Gly Phe Pro Gly Phe
 660 665 670
 Gly Val Gly Val Gly Gly Ile Pro Gly Val Ala Gly Val Pro Gly Val
 675 680 685
 Gly Gly Val Pro Gly Val Gly Gly Val Pro Gly Val Gly Ile Ser Pro
 690 695 700
 Glu Ala Gln Ala Ala Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala
 705 710 715 720
 Gly Ala Gly Val Leu Gly Gly Leu Val Pro Gly Pro Gln Ala Ala Val
 725 730 735
 Pro Gly Val Pro Gly Thr Gly Gly Val Pro Gly Val Gly Thr Pro Ala
 740 745 750
 Ala Ala Ala Ala Lys Ala Ala Ala Lys Ala Ala Gln Phe Gly Leu Val
 755 760 765
 Pro Gly Val Gly Val Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly
 770 775 780
 Val Ala Pro Gly Val Gly Leu Ala Pro Gly Val Gly Val Ala Pro Gly

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785	790	795	800
Val Gly Val Ala Pro Gly	Val Gly Val Ala Pro Gly	Val Ala Pro Gly Ile Gly Pro Gly	
	805	810	815
Gly Val Ala Ala Ala Ala Lys Ser Ala Ala Lys Val Ala Ala Lys Ala		825	830
820			
Gln Leu Arg Ala Ala Ala Gly Leu Gly Ala Gly Ile Pro Gly Leu Gly		840	845
835			
Val Gly Val Gly Val Pro Gly Leu Gly Val Gly Ala Gly Val Pro Gly		855	860
850			
Leu Gly Val Gly Ala Gly Val Pro Gly Phe Gly Ala Gly Ala Asp Glu		870	875
865			880
Gly Val Arg Arg Ser Leu Ser Pro Glu Leu Arg Glu Gly Asp Pro Ser		885	890
885			895
Ser Ser Gln His Leu Pro Ser Thr Pro Ser Ser Pro Arg Val Pro Gly		900	905
900			910
Ala Leu Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala Val Pro Gly		920	925
915			
Val Leu Gly Gly Leu Gly Ala Leu Gly Gly Val Gly Ile Pro Gly Gly		935	940
930			
Val Val Gly Ala Gly Pro Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala		950	955
945			960
Ala Lys Ala Ala Gln Phe Gly Leu Val Gly Ala Ala Gly Leu Gly Gly		965	970
965			975
Leu Gly Val Gly Gly Leu Gly Val Pro Gly Val Gly Gly Leu Gly Gly		980	985
980			990
Ile Pro Pro Ala Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala Gly		995	1000
995			1005
Leu Gly Gly Val Leu Gly Gly Ala Gly Gln Phe Pro Leu Gly Gly		1010	1015
1010			1020
Val Ala Ala Arg Pro Gly Phe Gly Leu Ser Pro Ile Phe Pro Gly		1025	1030
1025			1035
Gly Ala Cys Leu Gly Lys Ala Cys Gly Arg Lys Arg Lys His Pro		1040	1045
1040			1050
Glu Thr Leu Val Lys Val Lys Asp Ala Glu Asp Gln Leu Gly Ala		1055	1060
1055			1065
Arg Val Gly Tyr Ile Glu Leu Asp Leu Asn Ser Gly Lys Ile Leu		1070	1075
1070			1080
Glu Ser Phe Arg Pro Glu Glu Arg Phe Pro Met Met Ser Thr Phe		1085	1090
1085			1095
Lys Val Leu Leu Cys Gly Ala Val Leu Ser Arg Ile Asp Ala Gly		1100	1105
1100			1110
Gln Glu Gln Leu Gly Arg Arg Ile His Tyr Ser Gln Asn Asp Leu		1115	1120
1115			1125
Val Glu Tyr Ser Pro Val Thr Glu Lys His Leu Thr Asp Gly Met		1130	1135
1130			1140
Thr Val Arg Glu Leu Cys Ser Ala Ala Ile Thr Met Ser Asp Asn		1145	1150
1145			1155
Thr Ala Ala Asn Leu Leu Leu Thr Thr Ile Gly Gly Pro Lys Glu		1160	1165
1160			1170
Leu Thr Ala Phe Leu His Asn Met Gly Asp His Val Thr Arg Leu		1175	1180
1175			1185

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Asp Arg Trp Glu Pro Glu Leu Asn Glu Ala Ile Pro Asn Asp Glu
 1190 1195 1200
 Arg Asp Thr Thr Met Pro Val Ala Met Ala Thr Thr Leu Arg Lys
 1205 1210 1215
 Leu Leu Thr Gly Glu Leu Leu Thr Leu Ala Ser Arg Gln Gln Leu
 1220 1225 1230
 Ile Asp Trp Met Glu Ala Asp Lys Val Ala Gly Pro Leu Leu Arg
 1235 1240 1245
 Ser Ala Leu Pro Ala Gly Trp Phe Ile Ala Asp Lys Ser Gly Ala
 1250 1255 1260
 Gly Glu Arg Gly Ser Arg Gly Ile Ile Ala Ala Leu Gly Pro Asp
 1265 1270 1275
 Gly Lys Pro Ser Arg Ile Val Val Ile Tyr Thr Thr Gly Ser Gln
 1280 1285 1290
 Ala Thr Met Asp Glu Arg Asn Arg Gln Ile Ala Glu Ile Gly Ala
 1295 1300 1305
 Ser Leu Ile Lys His Trp
 1310

<210> SEQ ID NO 29

<211> LENGTH: 760

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 29

Gly Gly Val Pro Gly Ala Ile Pro Gly Gly Val Pro Gly Gly Val Phe
 1 5 10 15
 Tyr Pro Gly Ala Gly Leu Gly Ala Leu Gly Gly Gly Ala Leu Gly Pro
 20 25 30
 Gly Gly Lys Pro Leu Lys Pro Val Pro Gly Gly Leu Ala Gly Ala Gly
 35 40 45
 Leu Gly Ala Gly Leu Gly Ala Phe Pro Ala Val Thr Phe Pro Gly Ala
 50 55 60
 Leu Val Pro Gly Gly Val Ala Asp Ala Ala Ala Tyr Lys Ala Ala
 65 70 75 80
 Lys Ala Gly Ala Gly Leu Gly Gly Val Pro Gly Val Gly Gly Leu Gly
 85 90 95
 Val Ser Ala Gly Ala Val Val Pro Gln Pro Gly Ala Gly Val Lys Pro
 100 105 110
 Gly Lys Val Pro Gly Val Gly Leu Pro Gly Val Tyr Pro Gly Gly Val
 115 120 125
 Leu Pro Gly Ala Arg Phe Pro Gly Val Gly Val Leu Pro Gly Val Pro
 130 135 140
 Thr Gly Ala Gly Val Lys Pro Lys Ala Pro Gly Val Gly Gly Ala Phe
 145 150 155 160
 Ala Gly Ile Pro Gly Val Gly Pro Phe Gly Gly Pro Gln Pro Gly Val
 165 170 175
 Pro Leu Gly Tyr Pro Ile Lys Ala Pro Lys Leu Pro Gly Gly Tyr Gly
 180 185 190
 Leu Pro Tyr Thr Thr Gly Lys Leu Pro Tyr Gly Tyr Gly Pro Gly Gly

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195			200			205									
Val	Ala	Gly	Ala	Ala	Gly	Lys	Ala	Gly	Tyr	Pro	Thr	Gly	Thr	Gly	Val
210						215					220				
Gly	Pro	Gln	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Lys	Ala	Ala	Ala	Lys	Phe
225					230					235					240
Gly	Ala	Gly	Ala	Ala	Gly	Val	Leu	Pro	Gly	Val	Gly	Gly	Ala	Gly	Val
				245						250					255
Pro	Gly	Val	Pro	Gly	Ala	Ile	Pro	Gly	Ile	Gly	Gly	Ile	Ala	Gly	Val
		260						265						270	
Gly	Thr	Pro	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Lys	Ala	Ala
		275						280						285	
Lys	Tyr	Gly	Ala	Ala	Ala	Gly	Leu	Val	Pro	Gly	Gly	Pro	Gly	Phe	Gly
290						295					300				
Pro	Gly	Val	Val	Gly	Val	Pro	Gly	Ala	Gly	Val	Pro	Gly	Val	Gly	Val
305					310					315					320
Pro	Gly	Ala	Gly	Ile	Pro	Val	Val	Pro	Gly	Ala	Gly	Ile	Pro	Gly	Ala
				325						330					335
Ala	Val	Pro	Gly	Val	Val	Ser	Pro	Glu	Ala	Ala	Ala	Lys	Ala	Ala	Ala
		340						345						350	
Lys	Ala	Ala	Lys	Tyr	Gly	Ala	Arg	Pro	Gly	Val	Gly	Val	Gly	Gly	Ile
355							360						365		
Pro	Thr	Tyr	Gly	Val	Gly	Ala	Gly	Gly	Phe	Pro	Gly	Phe	Gly	Val	Gly
370						375					380				
Val	Gly	Gly	Ile	Pro	Gly	Val	Ala	Gly	Val	Pro	Gly	Val	Gly	Gly	Val
385					390					395					400
Pro	Gly	Val	Gly	Gly	Val	Pro	Gly	Val	Gly	Ile	Ser	Pro	Glu	Ala	Gln
				405						410					415
Ala	Ala	Ala	Ala	Ala	Lys	Ala	Ala	Lys	Tyr	Gly	Ala	Ala	Gly	Ala	Gly
				420				425						430	
Val	Leu	Gly	Gly	Leu	Val	Pro	Gly	Pro	Gln	Ala	Ala	Val	Pro	Gly	Val
		435					440						445		
Pro	Gly	Thr	Gly	Gly	Val	Pro	Gly	Val	Gly	Thr	Pro	Ala	Ala	Ala	Ala
450						455					460				
Ala	Lys	Ala	Ala	Ala	Lys	Ala	Ala	Gln	Phe	Gly	Leu	Val	Pro	Gly	Val
465					470					475					480
Gly	Val	Ala	Pro	Gly	Val	Gly	Val	Ala	Pro	Gly	Val	Gly	Val	Ala	Pro
				485				490							495
Gly	Val	Gly	Leu	Ala	Pro	Gly	Val	Gly	Val	Ala	Pro	Gly	Val	Gly	Val
			500					505					510		
Ala	Pro	Gly	Val	Gly	Val	Ala	Pro	Gly	Ile	Gly	Pro	Gly	Gly	Val	Ala
		515					520						525		
Ala	Ala	Ala	Lys	Ser	Ala	Ala	Lys	Val	Ala	Ala	Lys	Ala	Gln	Leu	Arg
530							535				540				
Ala	Ala	Ala	Gly	Leu	Gly	Ala	Gly	Ile	Pro	Gly	Leu	Gly	Val	Gly	Val
545					550					555					560
Gly	Val	Pro	Gly	Leu	Gly	Val	Gly	Ala	Gly	Val	Pro	Gly	Leu	Gly	Val
				565						570					575
Gly	Ala	Gly	Val	Pro	Gly	Phe	Gly	Ala	Gly	Ala	Asp	Glu	Gly	Val	Arg
				580				585						590	
Arg	Ser	Leu	Ser	Pro	Glu	Leu	Arg	Glu	Gly	Asp	Pro	Ser	Ser	Ser	Gln
		595					600								

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His Leu Pro Ser Thr Pro Ser Ser Pro Arg Val Pro Gly Ala Leu Ala
 610 615 620

Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala Val Pro Gly Val Leu Gly
 625 630 635 640

Gly Leu Gly Ala Leu Gly Gly Val Gly Ile Pro Gly Gly Val Val Gly
 645 650 655

Ala Gly Pro Ala Ala Ala Ala Ala Ala Lys Ala Ala Ala Lys Ala
 660 665 670

Ala Gln Phe Gly Leu Val Gly Ala Ala Gly Leu Gly Gly Leu Gly Val
 675 680 685

Gly Gly Leu Gly Val Pro Gly Val Gly Gly Leu Gly Gly Ile Pro Pro
 690 695 700

Ala Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala Gly Leu Gly Gly
 705 710 715 720

Val Leu Gly Gly Ala Gly Gln Phe Pro Leu Gly Gly Val Ala Ala Arg
 725 730 735

Pro Gly Phe Gly Leu Ser Pro Ile Phe Pro Gly Gly Ala Cys Leu Gly
 740 745 750

Lys Ala Cys Gly Arg Lys Arg Lys
 755 760

<210> SEQ ID NO 30
 <211> LENGTH: 2115
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polynucleotide

<400> SEQUENCE: 30

ggtggcgtac caggcgcaat tcctgggggt gtcccaggcg gtgttttta tccgggccc 60

ggtcttgccg cactgggtgg cgggtcactg ggcccggcgg gcaaaccgct gaaaccggta 120

ccaggtggtt tagcaggcgc cggcttaggc gcaggtctgg gagcatttcc ggcagttacc 180

tttccagggg cactggttcc tggaggtgtg gccgatgcag ccgcggcata taaagccgct 240

aaagccggtg cgggtttagg aggcgtccca ggtgtcggtg gcctgggtgt tagcgcgggt 300

gcagttgttc cgcagccggg agcaggggtt aaacctggta aagtgccggg agtaggtctg 360

ccaggcgttt atcctggtgg tgttttgccg ggtgcccgtt tccgggcccg tggtgttctt 420

ccaggcgctg cgaccggagc cgggttataa ccgaaagccc ccggtgttgg aggtgcattt 480

gcaggcatcc cgggagttgg cccggttggg ggtccgcaac ctggggttcc gttaggttat 540

ccgattaaag caccgaaact gcccgcggtt tatggtctgc cgtacacaac cggtaaactg 600

ccgtatggtt atggcccggg tggagttgcg ggtgcagcag gtaaagcggg ttatccctacc 660

ggaaccggtg taggtccgca ggcgctgct gccgcgcgca caaaagcagc ggctaaattt 720

ggcgccggag cagcgggtgt tetgcctgga gttggtggtg cgggcgtgcc aggggtacct 780

ggtgcaattc cgggtattgg tggtattgcc ggtgtcggca ccccggcccg ggcagctgcg 840

gcagcggcgg ctgccaaagc tgctaaatac ggtgcccggg cgggtctggt gccaggaggt 900

ccgggttttg gtccgggagt ggttgccgtg cctggcgcag gcgttccctg tggtggcggt 960

ccaggtgcag ggattcctgt tgtgctggt gccggtatcc ccggcgcggc cgttccgggg 1020

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gtggttagcc cggaagccgc agcgaagcct gcggcaaagg cagcaaagta tggcgcacgc 1080
ccaggagtcg gcgtgggtgg tatcccagacc tatgggggtgg gcgcaggggg ttttcctggt 1140
ttcggcgtag gtgtaggagg tataccgggc gtggccggtg taccaggggt tggcggcgtc 1200
cctggtgttg gcggtgtgcc aggtgttggg atttcaccgg aagcacaggc agcagccgca 1260
gctaaggcag cgaatatggt tgcgcgcggc gcaggagttt taggtgggct ggttccgggc 1320
ccgcaggcag ctgtgccggg ggttccaggc accggtggtg tccctggagt cggtacgccg 1380
gctgcagcgg cagccaaagc ggctgcgaaa gcagcacagt ttggcttagt accgggtgtg 1440
ggagttgccc ccggcgttgg cgttgtctca ggggtgggtg ttgctcctgg cgtcggctctg 1500
gctcctggag tggcgttagc acccggtgtg ggggtggccc cgggtgttgg ggttgaccgc 1560
ggtatcggtc cgggcggtgt cgcagcagca gctaaaagcg cggcgaaagt tgcggccaaa 1620
gcccactgc gcgccgcgc gggcctcggg gcaggatttc cggggtggg tgtcggagtt 1680
ggagtcccgc gtttggcgt gggcgcggga gttccgggac tgggagtggtg tgcggagtt 1740
cctggcttgg gtgcaggcgc agatgaaggt gttcgtcgtg gcctgagttc ggaactgctg 1800
gaaggtgatc cgagtagcag ccagcatctg ccgagcacc ccgagcagccc gcgtgttccg 1860
ggtgcattag ctgcagcaaa agccgccaag tatggtgcag ccgtgccggg cgtcttaggt 1920
ggtctggcgc ccctgggtgg tgtaggcatt ccgggaggtg ttgtgggtgc aggaccggcc 1980
gccgcagctg cggccgcaaa agcagctgca aaagcggccc agtttggttt agtgggcgcc 2040
gcaggtttag gcggtttagg tgtgggtgga ctgggtgtac ctggcgtagg cggctcgggt 2100
ggaattccgc cctaa 2115

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<210> SEQ ID NO 31

<211> LENGTH: 704

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 31

```

Gly Gly Val Pro Gly Ala Ile Pro Gly Gly Val Pro Gly Gly Val Phe
 1          5          10         15
Tyr Pro Gly Ala Gly Leu Gly Ala Leu Gly Gly Gly Ala Leu Gly Pro
 20         25         30
Gly Gly Lys Pro Leu Lys Pro Val Pro Gly Gly Leu Ala Gly Ala Gly
 35         40         45
Leu Gly Ala Gly Leu Gly Ala Phe Pro Ala Val Thr Phe Pro Gly Ala
 50         55         60
Leu Val Pro Gly Gly Val Ala Asp Ala Ala Ala Tyr Lys Ala Ala
 65         70         75         80
Lys Ala Gly Ala Gly Leu Gly Gly Val Pro Gly Val Gly Gly Leu Gly
 85         90         95
Val Ser Ala Gly Ala Val Val Pro Gln Pro Gly Ala Gly Val Lys Pro
100        105        110
Gly Lys Val Pro Gly Val Gly Leu Pro Gly Val Tyr Pro Gly Gly Val
115        120        125
Leu Pro Gly Ala Arg Phe Pro Gly Val Gly Val Leu Pro Gly Val Pro
130        135        140

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Thr Gly Ala Gly Val Lys Pro Lys Ala Pro Gly Val Gly Gly Ala Phe
 145 150 155 160

Ala Gly Ile Pro Gly Val Gly Pro Phe Gly Gly Pro Gln Pro Gly Val
 165 170 175

Pro Leu Gly Tyr Pro Ile Lys Ala Pro Lys Leu Pro Gly Gly Tyr Gly
 180 185 190

Leu Pro Tyr Thr Thr Gly Lys Leu Pro Tyr Gly Tyr Gly Pro Gly Gly
 195 200 205

Val Ala Gly Ala Ala Gly Lys Ala Gly Tyr Pro Thr Gly Thr Gly Val
 210 215 220

Gly Pro Gln Ala Ala Ala Ala Ala Ala Lys Ala Ala Ala Lys Phe
 225 230 235 240

Gly Ala Gly Ala Ala Gly Val Leu Pro Gly Val Gly Gly Ala Gly Val
 245 250 255

Pro Gly Val Pro Gly Ala Ile Pro Gly Ile Gly Gly Ile Ala Gly Val
 260 265 270

Gly Thr Pro Ala Ala Ala Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala
 275 280 285

Lys Tyr Gly Ala Ala Ala Gly Leu Val Pro Gly Gly Pro Gly Phe Gly
 290 295 300

Pro Gly Val Val Gly Val Pro Gly Ala Gly Val Pro Gly Val Gly Val
 305 310 315 320

Pro Gly Ala Gly Ile Pro Val Val Pro Gly Ala Gly Ile Pro Gly Ala
 325 330 335

Ala Val Pro Gly Val Val Ser Pro Glu Ala Ala Ala Lys Ala Ala Ala
 340 345 350

Lys Ala Ala Lys Tyr Gly Ala Arg Pro Gly Val Gly Val Gly Gly Ile
 355 360 365

Pro Thr Tyr Gly Val Gly Ala Gly Gly Phe Pro Gly Phe Gly Val Gly
 370 375 380

Val Gly Gly Ile Pro Gly Val Ala Gly Val Pro Gly Val Gly Gly Val
 385 390 395 400

Pro Gly Val Gly Gly Val Pro Gly Val Gly Ile Ser Pro Glu Ala Gln
 405 410 415

Ala Ala Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala Gly Ala Gly
 420 425 430

Val Leu Gly Gly Leu Val Pro Gly Pro Gln Ala Ala Val Pro Gly Val
 435 440 445

Pro Gly Thr Gly Gly Val Pro Gly Val Gly Thr Pro Ala Ala Ala Ala
 450 455 460

Ala Lys Ala Ala Ala Lys Ala Ala Gln Phe Gly Leu Val Pro Gly Val
 465 470 475 480

Gly Val Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Val Ala Pro
 485 490 495

Gly Val Gly Leu Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Val
 500 505 510

Ala Pro Gly Val Gly Val Ala Pro Gly Ile Gly Pro Gly Gly Val Ala
 515 520 525

Ala Ala Ala Lys Ser Ala Ala Lys Val Ala Ala Lys Ala Gln Leu Arg
 530 535 540

Ala Ala Ala Gly Leu Gly Ala Gly Ile Pro Gly Leu Gly Val Gly Val

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545	550	555	560
Gly Val Pro Gly Leu Gly Val Gly Ala Gly Val Pro Gly Leu Gly Val	565	570	575
Gly Ala Gly Val Pro Gly Phe Gly Ala Gly Ala Asp Glu Gly Val Arg	580	585	590
Arg Ser Leu Ser Pro Glu Leu Arg Glu Gly Asp Pro Ser Ser Ser Gln	595	600	605
His Leu Pro Ser Thr Pro Ser Ser Pro Arg Val Pro Gly Ala Leu Ala	610	615	620
Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala Val Pro Gly Val Leu Gly	625	630	635
Gly Leu Gly Ala Leu Gly Gly Val Gly Ile Pro Gly Gly Val Val Gly	645	650	655
Ala Gly Pro Ala Ala Ala Ala Ala Ala Lys Ala Ala Lys Ala	660	665	670
Ala Gln Phe Gly Leu Val Gly Ala Ala Gly Leu Gly Gly Leu Gly Val	675	680	685
Gly Gly Leu Gly Val Pro Gly Val Gly Gly Leu Gly Gly Ile Pro Pro	690	695	700

<210> SEQ ID NO 32
 <211> LENGTH: 2115
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 32

```

ggtggcgtag caggcgcaat tctgggggt gtcccaggcg gtgttttta tccgggagcc 60
ggtcttgagg cactgggtgg cggtgactg ggcccgggag gcaaacgct gaaaccggt 120
ccaggtggtt tagcaggcgc cggcttaggc gcaggtctgg gagcatttc gccagttacc 180
ttccaggagg cactggttcc tggaggtgtg gccgatgcag ccgcggcata taaagccgct 240
aaagccggtg cgggtttagg aggcgtcca ggtgtcggtg gcctgggtgt tagcgccggt 300
gcagttgttc cgcagccggg agcaggggtt aaacctgta aagtgcggg agtaggtctg 360
ccaggcggtt atcctggtgg tgttttgccg ggtgcccgtt tccgggagc ttggtgttctt 420
ccaggcgtgc cgaccggagc cgggtgtaaa ccgaaagccc ccggtgttgg aggtgcattt 480
gcaggcatcc cgggagttgg cccggttggg ggtccgcaac ctggggttcc gttaggttat 540
ccgattaaag caccgaaact gcccgccggt tatggtctgc cgtacacaac cggtaaaactg 600
ccgtatggtt atggcccggg tggagttgag ggtgcagcag gtaaagcggg ttatcctacc 660
ggaaccggtg taggtccgca gcccgctgct gccgcgcgcg caaaagcagc ggctaaattt 720
ggcgccggag cagcgggtgt tctgectgga gttggtggtg cgggcgtgcc aggggtacct 780
ggtgcaattc cgggtattgg tggtattgcc ggtgtcggca ccccgggcgc gccagctgag 840
gcagcggcgg ctgccaaagc tgctaaatac ggtgccgcgg cgggtctggt gccaggaggt 900
ccgggttttg gtccgggagt ggttggcgtg cctggcgcag gcgttcctgg tgtgggagtt 960
ccaggtgcag ggattcctgt tgtgectggt gccggtattc ccggcgcggc cgttccgggg 1020
gtggttagcc cggaagccgc agcgaaggct gcggcaaggg cagcaagta tggcgcagc 1080
    
```

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```

ccaggagtgc gcgtgggtgg taccgacc tatgggggtg gcgcaggggg ttttctgtgt 1140
ttcggcgtag gtgtaggagg tataccgggc gtggccggtg taccaggggt tggtagcgctc 1200
cctgtgtttg gcggtgtgcc aggtgttggg atttcaccgg aagcacaggc agcagccgca 1260
gctaaggcag cgaatatg tgccgccggc gcaggagttt taggtgggct ggttccgggc 1320
ccgcaggcag ctgtgccggg ggttccaggc accggtggtg tccttgagtg cggtaacggc 1380
gtgcagcggc cagccaaagc ggtgcgaaa gcagcacagt ttggcttagt accgggtgtg 1440
ggagttgccc ccggcgttgg cgttgctcca ggggtgggtg ttgctcctgg cgtcggtctg 1500
gtcctcgtag tggcgttagc acccggtgtg ggggtggccc cgggtgttgg ggttgcaccg 1560
ggtatcggtc cgggctgtgt gcagcagca gctaaaagcg cggcgaaagt tgcggccaaa 1620
gcccactgc gcgccgcgc gggcctcggg gcaggtattc cggggtggg tgtcggagtt 1680
ggagtcaccg gtttggcgt gggcgcggga gttccgggac tgggagtggg tgccggagtt 1740
cctggctttg gtgcaggcgc agatgaaggt gttcgtcgta gcctgagtc ggaactcgt 1800
gaagtgatc cgagtagcag ccagcatctg ccgagcacc ccagcagccc gcgtgttccg 1860
ggtgcattag ctgcagaaa agccgccaag tatggtgcag ccgtgccggg cgtcttaggt 1920
ggtctgggcg ccctgggtgg ttaggcatt ccgggaggtg ttgtgggtgc aggaccggcc 1980
gccgcagctg cggccgcca agcagctgca aaagcggccc agtttggtt agtggcgccc 2040
gcaggtttag gcggttagg tgtgggtgga ctgggtgtac ctggcgtagg cgtctgggt 2100
ggaattccgc cctaa 2115
    
```

```

<210> SEQ ID NO 33
<211> LENGTH: 676
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
    polypeptide
    
```

```

<400> SEQUENCE: 33
Gly Leu Gly Gly Val Pro Gly Val Gly Gly Leu Gly Val Ser Ala Gly
1          5          10          15
Ala Val Val Pro Gln Pro Gly Ala Gly Val Lys Pro Gly Lys Val Pro
20         25         30
Gly Val Gly Leu Pro Gly Val Tyr Pro Gly Gly Val Leu Pro Gly Ala
35         40         45
Arg Phe Pro Gly Val Gly Val Leu Pro Gly Val Pro Thr Gly Ala Gly
50         55         60
Val Lys Pro Lys Ala Pro Gly Val Gly Gly Ala Phe Ala Gly Ile Pro
65         70         75         80
Gly Val Gly Pro Phe Gly Gly Pro Gln Pro Gly Val Pro Leu Gly Tyr
85         90         95
Pro Ile Lys Ala Pro Lys Leu Pro Gly Gly Tyr Gly Leu Pro Tyr Thr
100        105        110
Thr Gly Lys Leu Pro Tyr Gly Tyr Gly Pro Gly Gly Val Ala Gly Ala
115        120        125
Ala Gly Lys Ala Gly Tyr Pro Thr Gly Thr Gly Val Gly Pro Gln Ala
130        135        140
Ala Ala Ala Ala Ala Lys Ala Ala Ala Lys Phe Gly Ala Gly Ala
145        150        155        160
    
```

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Ala Gly Val Leu Pro Gly Val Gly Gly Ala Gly Val Pro Gly Val Pro
165 170 175

Gly Ala Ile Pro Gly Ile Gly Gly Ile Ala Gly Val Gly Thr Pro Ala
180 185 190

Ala Ala Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala
195 200 205

Ala Ala Gly Leu Val Pro Gly Gly Pro Gly Phe Gly Pro Gly Val Val
210 215 220

Gly Val Pro Gly Ala Gly Val Pro Gly Val Gly Val Pro Gly Ala Gly
225 230 235 240

Ile Pro Val Val Pro Gly Ala Gly Ile Pro Gly Ala Ala Val Pro Gly
245 250 255

Val Val Ser Pro Glu Ala Ala Ala Lys Ala Ala Ala Lys Ala Ala Lys
260 265 270

Tyr Gly Ala Arg Pro Gly Val Gly Val Gly Gly Ile Pro Thr Tyr Gly
275 280 285

Val Gly Ala Gly Gly Phe Pro Gly Phe Gly Val Gly Val Gly Gly Ile
290 295 300

Pro Gly Val Ala Gly Val Pro Gly Val Gly Gly Val Pro Gly Val Gly
305 310 315 320

Gly Val Pro Gly Val Gly Ile Ser Pro Glu Ala Gln Ala Ala Ala Ala
325 330 335

Ala Lys Ala Ala Lys Tyr Gly Ala Ala Gly Ala Gly Val Leu Gly Gly
340 345 350

Leu Val Pro Gly Pro Gln Ala Ala Val Pro Gly Val Pro Gly Thr Gly
355 360 365

Gly Val Pro Gly Val Gly Thr Pro Ala Ala Ala Ala Ala Lys Ala Ala
370 375 380

Ala Lys Ala Ala Gln Phe Gly Leu Val Pro Gly Val Gly Val Ala Pro
385 390 395 400

Gly Val Gly Val Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Leu
405 410 415

Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Val Ala Pro Gly Val
420 425 430

Gly Val Ala Pro Gly Ile Gly Pro Gly Gly Val Ala Ala Ala Ala Lys
435 440 445

Ser Ala Ala Lys Val Ala Ala Lys Ala Gln Leu Arg Ala Ala Ala Gly
450 455 460

Leu Gly Ala Gly Ile Pro Gly Leu Gly Val Gly Val Gly Val Pro Gly
465 470 475 480

Leu Gly Val Gly Ala Gly Val Pro Gly Leu Gly Val Gly Ala Gly Val
485 490 495

Pro Gly Phe Gly Ala Gly Ala Asp Glu Gly Val Arg Arg Ser Leu Ser
500 505 510

Pro Glu Leu Arg Glu Gly Asp Pro Ser Ser Ser Gln His Leu Pro Ser
515 520 525

Thr Pro Ser Ser Pro Arg Val Pro Gly Ala Leu Ala Ala Ala Lys Ala
530 535 540

Ala Lys Tyr Gly Ala Ala Val Pro Gly Val Leu Gly Gly Leu Gly Ala
545 550 555 560

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Leu	Gly	Gly	Val	Gly	Ile	Pro	Gly	Gly	Val	Val	Gly	Ala	Gly	Pro	Ala
				565					570					575	
Ala	Ala	Ala	Ala	Ala	Ala	Lys	Ala	Ala	Ala	Lys	Ala	Ala	Gln	Phe	Gly
				580					585					590	
Leu	Val	Gly	Ala	Ala	Gly	Leu	Gly	Gly	Leu	Gly	Val	Gly	Gly	Leu	Gly
		595					600					605			
Val	Pro	Gly	Val	Gly	Gly	Leu	Gly	Gly	Ile	Pro	Pro	Ala	Ala	Ala	Ala
	610					615					620				
Lys	Ala	Ala	Lys	Tyr	Gly	Ala	Ala	Gly	Leu	Gly	Gly	Val	Leu	Gly	Gly
	625				630					635					640
Ala	Gly	Gln	Phe	Pro	Leu	Gly	Gly	Val	Ala	Ala	Arg	Pro	Gly	Phe	Gly
				645					650						655
Leu	Ser	Pro	Ile	Phe	Pro	Gly	Gly	Ala	Cys	Leu	Gly	Lys	Ala	Cys	Gly
			660					665						670	
Arg	Lys	Arg	Lys												
			675												

<210> SEQ ID NO 34
 <211> LENGTH: 2031
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 34

```

ggtttaggag ggcgtcccagg tgcctgggag ctgggtggtta gcgccggtgc agttgttccg   60
cagccggggag cagggggttaa acctggtaaa gtgccggggag taggtctgccc aggcggttat   120
cctggtggtg ttttgccggg tgcccgtttt ccgggcggtg gtgttcttcc aggcggtgccc   180
accggagccg gtgttaaaccc gaaagcccc ggtgttgagg gtgcatttgc aggcattccc   240
ggagttggcc cgtttggtgg tccgcaacct ggggttcccg taggttatcc gattaaagca   300
ccgaaactgc ccggcgggta tggctctgccc tacacaaccg gtaaaactgcc gtatggttat   360
ggccccgggtg gagttgcccgg tgcagcaggt aaagcggggt atcctaccgg aaccggtgta   420
ggtcgccagg ccgctgctgc cgcgccgcca aaagcagcgg ctaaatttgg cgccggagca   480
gccccgtgtc tgcctggagt tgggtgtgccc ggcgtgcccagg gggtagctgg tgcaattccc   540
ggtagttggtg gtattgcccgg tgcctgcccacc ccggccgccc cagctgcccgc agccggcgct   600
gccaaaagctg ctaaatacgg tgcgccggcg ggtctggtgc caggagggtcc gggttttggt   660
ccgggagtggt ttggcgtgccc tggcgcaggc gttcctggtg tgggcgttcc aggtgcccgg   720
attcctggtg tgccctggtg ccgtattccc ggcgccggccc ttccgggggt ggtagccccc   780
gaagccgcag cgaaggctgc ggcaaaggca gcaaagtat ggcacacccc aggagtcggc   840
gtgggtggta tcccaccta tggggtgggc gcagggggtt ttctggttt cgccgtaggt   900
gtaggaggtat taccggcgct ggcgggtgta ccagggggtg gtggcgtccc tggtgttgcc   960
ggtagtcccag gtgttggtat ttaccgggaa gcacaggcag cagccgcagc taaggcagc   1020
aaatatggtg ccgccggcgc aggagtttta ggtgggctgg ttccgggccc gcaggcagct   1080
gtgccggggg ttccaggcac ccgtggtgtc cctggagtcg gtacccgggc tgcagcggca   1140
gccaaaaggc ctgcgaaaag agcacagttt ggccttagtac cgggtgtggg agttgcccc   1200
ggcgttgccg ttgctccagg ggtgggtgtt gctcctggcg tcggctctggc tctgaggtg   1260

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ggcgtagcac ccggtgtggg ggtggccccg ggtgttgggg ttgcaccggg tatcggtccg 1320
ggcgggtgtcg cagcagcagc taaaagcgcg gcgaaagtg cggccaaagc ccaactgcgc 1380
gccgcgcggg gcctcgggtgc aggtattccg gggctgggtg tcggagtgg agtccccggg 1440
ttgggcgtgg gcgcgggagt tccgggactg ggagtgggtg ccggagtcc tggctttggt 1500
gcaggcgcag atgaaggtgt tcgtcgtagc ctgagtccgg aactgcgtga aggtgatccg 1560
agtagcagcc agcatctgcc gagcaccocg agcagcccgc gtgttccggg tgcattagct 1620
gcagcaaaag ccgccaagta tggtgagcc gtgcggggcg tcttaggtgg tctgggcgcc 1680
ctgggtggtg taggcattcc gggaggtgtt gtgggtgcag gaccggccgc cgcagctgcg 1740
gccgcaaaag cagctgcaaa agcggcccag tttggttag tgggcgcgc aggttttagc 1800
ggtttaggtg tgggtggact ggggttacct ggcgtaggcg gtctgggtgg aattccgcc 1860
gcagcggccg cgaaagcggc aaaatatgac gcggcaggcc tgggcggcgt gctgggtggg 1920
gcaggtcagt ttccgctggg cggggttgcc gcacgtccgg gatttggctc gagcccgatt 1980
ttccctggcg gcgcatgtct gggtaaagca tgtggtcgta aacgtaaata a 2031
    
```

```

<210> SEQ ID NO 35
<211> LENGTH: 659
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
    polypeptide
    
```

```

<400> SEQUENCE: 35
Gly Gly Val Pro Gly Ala Ile Pro Gly Gly Val Pro Gly Gly Val Phe
1 5 10 15
Tyr Pro Gly Ala Gly Leu Gly Ala Leu Gly Gly Gly Ala Leu Gly Pro
20 25 30
Gly Gly Lys Pro Leu Lys Pro Val Pro Gly Gly Leu Ala Gly Ala Gly
35 40 45
Leu Gly Ala Gly Leu Gly Ala Phe Pro Ala Val Thr Phe Pro Gly Ala
50 55 60
Leu Val Pro Gly Gly Val Ala Asp Ala Ala Ala Tyr Lys Ala Ala
65 70 75 80
Lys Ala Gly Ala Gly Leu Gly Gly Val Pro Gly Val Gly Gly Leu Gly
85 90 95
Val Ser Ala Gly Ala Val Val Pro Gln Pro Gly Ala Gly Val Lys Pro
100 105 110
Gly Lys Val Pro Gly Val Gly Leu Pro Gly Val Tyr Pro Gly Gly Val
115 120 125
Leu Pro Gly Ala Arg Phe Pro Gly Val Gly Val Leu Pro Gly Val Pro
130 135 140
Thr Gly Ala Gly Val Lys Pro Lys Ala Pro Gly Val Gly Gly Ala Phe
145 150 155 160
Ala Gly Ile Pro Gly Val Gly Pro Phe Gly Gly Pro Gln Pro Gly Val
165 170 175
Pro Leu Gly Tyr Pro Ile Lys Ala Pro Lys Leu Pro Gly Gly Tyr Gly
180 185 190
Leu Pro Tyr Thr Thr Gly Lys Leu Pro Tyr Gly Tyr Gly Pro Gly Gly
195 200 205
    
```

-continued

Val Ala Gly Ala Ala Gly Lys Ala Gly Tyr Pro Thr Gly Thr Gly Val
 210 215 220
 Gly Pro Gln Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala Ala Lys Phe
 225 230 235 240
 Gly Ala Gly Ala Ala Gly Val Leu Pro Gly Val Gly Gly Ala Gly Val
 245 250 255
 Pro Gly Val Pro Gly Ala Ile Pro Gly Ile Gly Gly Ile Ala Gly Val
 260 265 270
 Gly Thr Pro Ala Ala Ala Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala
 275 280 285
 Lys Tyr Gly Ala Ala Ala Gly Leu Val Pro Gly Gly Pro Gly Phe Gly
 290 295 300
 Pro Gly Val Val Gly Val Pro Gly Ala Gly Val Pro Gly Val Gly Val
 305 310 315 320
 Pro Gly Ala Gly Ile Pro Val Val Pro Gly Ala Gly Ile Pro Gly Ala
 325 330 335
 Ala Val Pro Gly Val Val Ser Pro Glu Ala Ala Ala Lys Ala Ala Ala
 340 345 350
 Lys Ala Ala Lys Tyr Gly Ala Arg Pro Gly Val Gly Val Gly Gly Ile
 355 360 365
 Pro Thr Tyr Gly Val Gly Ala Gly Gly Phe Pro Gly Phe Gly Val Gly
 370 375 380
 Val Gly Gly Ile Pro Gly Val Ala Gly Val Pro Gly Val Gly Gly Val
 385 390 395 400
 Pro Gly Val Gly Gly Val Pro Gly Val Gly Ile Ser Pro Glu Ala Gln
 405 410 415
 Ala Ala Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala Gly Ala Gly
 420 425 430
 Val Leu Gly Gly Leu Val Pro Gly Pro Gln Ala Ala Val Pro Gly Val
 435 440 445
 Pro Gly Thr Gly Gly Val Pro Gly Val Gly Thr Pro Ala Ala Ala Ala
 450 455 460
 Ala Lys Ala Ala Ala Lys Ala Ala Gln Phe Gly Leu Val Pro Gly Val
 465 470 475 480
 Gly Val Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Val Ala Pro
 485 490 495
 Gly Val Gly Leu Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Val
 500 505 510
 Ala Pro Gly Val Gly Val Ala Pro Gly Ile Gly Pro Gly Gly Val Ala
 515 520 525
 Ala Ala Ala Lys Ser Ala Ala Lys Val Ala Ala Lys Ala Gln Leu Arg
 530 535 540
 Ala Ala Ala Gly Leu Gly Ala Gly Ile Pro Gly Leu Gly Val Gly Val
 545 550 555 560
 Gly Val Pro Gly Leu Gly Val Gly Ala Gly Val Pro Gly Leu Gly Val
 565 570 575
 Gly Ala Gly Val Pro Gly Phe Gly Ala Gly Ala Asp Glu Gly Val Arg
 580 585 590
 Arg Ser Leu Ser Pro Glu Leu Arg Glu Gly Asp Pro Ser Ser Ser Gln
 595 600 605

-continued

His Leu Pro Ser Thr Pro Ser Ser Pro Arg Val Pro Gly Ala Leu Ala
 610 615 620

Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala Val Pro Gly Val Leu Gly
 625 630 635 640

Gly Leu Gly Ala Leu Gly Gly Val Gly Ile Pro Gly Gly Val Val Gly
 645 650 655

Ala Gly Pro

<210> SEQ ID NO 36
 <211> LENGTH: 1980
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polynucleotide

<400> SEQUENCE: 36

ggtggcgtac caggcgcaat tctggggggt gtcccaggcg gtgtttttta tccggggcgc 60

ggtcttgccg cactgggtgg cggtgcaactg ggcccgggcg gcaaaccgct gaaaccggta 120

ccaggtgggt tagcaggcgc cggccttagc gcaggtctgg gagcatttcc gccagttacc 180

tttccagggg cactggttcc tggaggtgtg gccgatgcag ccgcggcata taaagccgct 240

aaagccggtg cgggtttagg aggcgtccca ggtgtcgggt gcctgggtgt tagcgcgggt 300

gcagttgttc cgcagccggg agcaggggtt aaacctggta aagtgccggg agtaggtctg 360

ccaggcggtt atcctgggtg tgttttgccg ggtgcccgtt ttccgggcgt tgggtttctt 420

ccaggcgtgc cgaccggagc cgggtgtaaa ccgaaagccc ccggtgttgg aggtgcattt 480

gcaggcatcc cgggagttgg cccgtttggg ggtccgcaac ctgggggttc gttaggttat 540

ccgattaagc caccgaaact gcccgccggt tatggtctgc cgtacacaac cggtaaaactg 600

ccgtatggtt atggcccggg tggagttgcg ggtgcagcag gtaaagcggg ttatcctacc 660

ggaaccggtg taggtccgca ggcgctgct gccgcgcccg caaaagcagc ggctaaattt 720

ggcgcgggag cagcgggtgt tctgcctgga gttggtggtg cgggcgtgcc aggggtacct 780

ggtgcaattc cgggtattgg tggattgccc ggtgtcggca ccccgccgc gccagctgcg 840

gcagcggcgg ctgccaaaag tgcataaac ggtgcccgcg cgggtctggt gccaggaggt 900

ccgggttttg gtcggggagt ggttgccgtg cctggcgcag gcgttccctg tgtgggcgtt 960

ccaggtgcag ggattcctgt tgtgcctggt gccggtattc ccggcgcggc cgttccgggg 1020

gtggttagcc cggaaagccg agcgaaggct gcggcaagg cagcaaaagta tggcgcaacg 1080

ccaggagtgc gcgtgggtgg tatcccagc tatgggggtg gcgcaggggg ttttccctgt 1140

ttcggcgtag gtgtaggagg tataccgggc gtggccgggt taccaggggt tgggtggcgtc 1200

cctggtgttg gcggtgtgcc aggtgttggg atttaccgg aagcacaggc agcagccgca 1260

gctaaggcag cgaatatagg tgccgcggc gcaggagttt taggtgggct ggttccgggc 1320

ccgcaggcag ctgtgccggg ggttccaggc accggtggtg tccctggagt cggtaacgcc 1380

gctgcagcgg cagccaaaag ggtgcgaaa gcagcacagt ttggcttagt accgggtgtg 1440

ggagttgccc ccggcgttgg cgttgctcca ggggtgggtg ttgctcctgg cgtcgggtctg 1500

gctcctggag tgggcgtagc acccggtgtg ggggtggccc cgggtgttgg ggttgcaacc 1560

ggtatcggtc cgggcggtgt cgcagcagca gctaaaagcg cggcgaaagt tccggccaaa 1620

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gcccaactgc gcgcccgcgc gggcctcggg gcaggtattc cggggctggg tgtcggagtt 1680
ggagtcocccg gtttgggctg gggcgcggga gttccgggac tgggagtggg tgccggagtt 1740
cctggccttg gtgcaggcgc agatgaaggt gttcgtcgta gcctgagtcc ggaactgcgt 1800
gaagtgatc cgagtagcag ccagcatctg ccgagcacc cagacagccc gcgtgttccg 1860
ggtgcattag ctgcagcaaa agccgccaag tatggtgcag ccgtgcccgg cgtcttaggt 1920
ggtctgggcg ccctgggtgg ttaggcatt ccgggaggtg ttgtgggtgc aggaccgtaa 1980
    
```

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<210> SEQ ID NO 37
<211> LENGTH: 622
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
                             polypeptide
    
```

<400> SEQUENCE: 37

```

Gly Gly Val Pro Gly Ala Ile Pro Gly Gly Val Pro Gly Gly Val Phe
1           5           10          15
Tyr Pro Gly Ala Gly Leu Gly Ala Leu Gly Gly Gly Ala Leu Gly Pro
20          25          30
Gly Gly Lys Pro Leu Lys Pro Val Pro Gly Gly Leu Ala Gly Ala Gly
35          40          45
Leu Gly Ala Gly Leu Gly Ala Phe Pro Ala Val Thr Phe Pro Gly Ala
50          55          60
Leu Val Pro Gly Gly Val Ala Asp Ala Ala Ala Tyr Lys Ala Ala
65          70          75          80
Lys Ala Gly Ala Gly Leu Gly Gly Val Pro Gly Val Gly Gly Leu Gly
85          90          95
Val Ser Ala Gly Ala Val Val Pro Gln Pro Gly Ala Gly Val Lys Pro
100         105        110
Gly Lys Val Pro Gly Val Gly Leu Pro Gly Val Tyr Pro Gly Gly Val
115        120        125
Leu Pro Gly Ala Arg Phe Pro Gly Val Gly Val Leu Pro Gly Val Pro
130        135        140
Thr Gly Ala Gly Val Lys Pro Lys Ala Pro Gly Val Gly Gly Ala Phe
145        150        155        160
Ala Gly Ile Pro Gly Val Gly Pro Phe Gly Gly Pro Gln Pro Gly Val
165        170        175
Pro Leu Gly Tyr Pro Ile Lys Ala Pro Lys Leu Pro Gly Gly Tyr Gly
180        185        190
Leu Pro Tyr Thr Thr Gly Lys Leu Pro Tyr Gly Tyr Gly Pro Gly Gly
195        200        205
Val Ala Gly Ala Ala Gly Lys Ala Gly Tyr Pro Thr Gly Thr Gly Val
210        215        220
Gly Pro Gln Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala Ala Lys Phe
225        230        235        240
Gly Ala Gly Ala Ala Gly Val Leu Pro Gly Val Gly Gly Ala Gly Val
245        250        255
Pro Gly Val Pro Gly Ala Ile Pro Gly Ile Gly Gly Ile Ala Gly Val
260        265        270
Gly Thr Pro Ala Ala Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala
275        280        285
    
```

-continued

Lys Tyr Gly Ala Ala Ala Gly Leu Val Pro Gly Gly Pro Gly Phe Gly
 290 295 300

Pro Gly Val Val Gly Val Pro Gly Ala Gly Val Pro Gly Val Gly Val
 305 310 315 320

Pro Gly Ala Gly Ile Pro Val Val Pro Gly Ala Gly Ile Pro Gly Ala
 325 330 335

Ala Val Pro Gly Val Val Ser Pro Glu Ala Ala Ala Lys Ala Ala Ala
 340 345 350

Lys Ala Ala Lys Tyr Gly Ala Arg Pro Gly Val Gly Val Gly Gly Ile
 355 360 365

Pro Thr Tyr Gly Val Gly Ala Gly Gly Phe Pro Gly Phe Gly Val Gly
 370 375 380

Val Gly Gly Ile Pro Gly Val Ala Gly Val Pro Gly Val Gly Gly Val
 385 390 395 400

Pro Gly Val Gly Gly Val Pro Gly Val Gly Ile Ser Pro Glu Ala Gln
 405 410 415

Ala Ala Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala Gly Ala Gly
 420 425 430

Val Leu Gly Gly Leu Val Pro Gly Pro Gln Ala Ala Val Pro Gly Val
 435 440 445

Pro Gly Thr Gly Gly Val Pro Gly Val Gly Thr Pro Ala Ala Ala Ala
 450 455 460

Ala Lys Ala Ala Ala Lys Ala Ala Gln Phe Gly Leu Val Pro Gly Val
 465 470 475 480

Gly Val Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Val Ala Pro
 485 490 495

Gly Val Gly Leu Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Val
 500 505 510

Ala Pro Gly Val Gly Val Ala Pro Gly Ile Gly Pro Gly Gly Val Ala
 515 520 525

Ala Ala Ala Lys Ser Ala Ala Lys Val Ala Ala Lys Ala Gln Leu Arg
 530 535 540

Ala Ala Ala Gly Leu Gly Ala Gly Ile Pro Gly Leu Gly Val Gly Val
 545 550 555 560

Gly Val Pro Gly Leu Gly Val Gly Ala Gly Val Pro Gly Leu Gly Val
 565 570 575

Gly Ala Gly Val Pro Gly Phe Gly Ala Gly Ala Asp Glu Gly Val Arg
 580 585 590

Arg Ser Leu Ser Pro Glu Leu Arg Glu Gly Asp Pro Ser Ser Ser Gln
 595 600 605

His Leu Pro Ser Thr Pro Ser Ser Pro Arg Val Pro Gly Ala
 610 615 620

<210> SEQ ID NO 38
 <211> LENGTH: 1869
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polynucleotide

<400> SEQUENCE: 38

ggtggcgtac caggcgcaat tctctgggggt gtcccaggcg gtgtttttta tccgggccc 60

-continued

ggtcttggcg cactgggtgg cggtgcaactg ggcccgggcg gcaaaccgct gaaaccggta	120
ccaggtgggt tagcaggcgc cggcttaggc gcaggtctgg gagcatttcc ggcagttacc	180
tttccagggg cactggttcc tggaggtgtg gccgatgcag ccgcggcata taaagccgct	240
aaagccggcg cgggtttagg aggcgtccca ggtgtcggtg gcctgggtgt tagcgcgggt	300
gcagttgttc cgcagccggg agcaggggtt aaacctggta aagtgccggg agtaggtctg	360
ccaggcggtt atcctgggtg tgttttgccg ggtgcccgtt ttccgggcgt tgggtttctt	420
ccaggcgtgc cgaccggagc cgggttataa cggaaagccc ccggtgttgg aggtgcattt	480
gcaggcatcc cgggagttgg cccgtttggt ggtccgcaac ctggggttcc gttaggttat	540
ccgattaaag caccgaaact gcccgcggtt tatggtctgc cgtacacaac cggtaaactg	600
ccgtatggtt atggcccggg tggagttgcg ggtgcagcag gtaaagcggg ttatcctacc	660
ggaaccggtg taggtccgca ggcgctgct gccgcgccc caaaagcagc ggctaaattt	720
ggcgcgggag cagcgggtgt tctgcctgga gttggtggtg cgggcgtgcc aggggtacct	780
ggtgcaattc cgggtattgg tggattgccc ggtgtcggca ccccgccgc ggcagctgcg	840
gcagcggcgg ctgccaaagc tgctaaatac ggtgcgcggg cgggtctggt gccaggaggt	900
ccgggttttg gtccgggagt ggttggcgtg cctggcgcag gcgttcctgg tgtgggcgtt	960
ccaggtgcag ggattcctgt tgtgcctggt gccggtatcc ccggcgcggc cgttccgggg	1020
gtggttagcc cggaaagcgc agcgaaggct gccgcaaagg cagcaaagta tggcgcacgc	1080
ccaggagtgc gcgtgggtgg tatcccagcc tatgggggtg gcgcaggggg ttttcctggt	1140
ttcggcgtag gtgtaggagg tataccgggc gtggcgggtg taccaggggt tgggtggcgtc	1200
cctggtgttg gcggtgtgccc aggtgttgggt atttcaccgg aagcacaggc agcagccgca	1260
gctaaggcag cgaatatatg tgcccgcggc gcaggagttt taggtgggct ggttccgggc	1320
ccgcaggcag ctgtgccggg ggttccaggc accggtgggtg tccctggagt cggtagcccg	1380
gctgcagcgg cagccaaagc ggtgcgaaa gcagcacagt ttggcttagt accgggtgtg	1440
ggagttgccc ccggcgttgg cgttgcctca ggggtgggtg ttgctcctgg cgtcggctctg	1500
gctcctggag tgggcgtagc acccggtgtg ggggtggccc cgggtgttgg ggttgcaccg	1560
ggtatcggtc cgggcgggtg cgcagcagca gctaaaagcg cggcgaaagt tgcggccaaa	1620
gcccactgccc gcgcccgcgc gggcctcggg gcaggatcc cggggctggg tgtcggagtt	1680
ggagttcccg gtttgggctg gggcgcggga gttccgggac tgggagtggtg tgcggagtt	1740
cctggctttg gtgcaggcgc agatgaagggt gttcgtcgta gcctgagttcc ggaactgctg	1800
gaaggtgatc cgagtagcag ccagcatctg ccgagcacc ccgagcagccc gcgtgttccg	1860
ggtgcataa	1869

<210> SEQ ID NO 39
 <211> LENGTH: 527
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide
 <400> SEQUENCE: 39

Gly Gly Val Pro Gly Ala Ile Pro Gly Gly Val Pro Gly Gly Val Phe
 1 5 10 15

-continued

Tyr	Pro	Gly	Ala	Gly	Leu	Gly	Ala	Leu	Gly	Gly	Gly	Ala	Leu	Gly	Pro
			20					25					30		
Gly	Gly	Lys	Pro	Leu	Lys	Pro	Val	Pro	Gly	Gly	Leu	Ala	Gly	Ala	Gly
		35					40					45			
Leu	Gly	Ala	Gly	Leu	Gly	Ala	Phe	Pro	Ala	Val	Thr	Phe	Pro	Gly	Ala
		50				55					60				
Leu	Val	Pro	Gly	Gly	Val	Ala	Asp	Ala	Ala	Ala	Ala	Tyr	Lys	Ala	Ala
		65			70					75					80
Lys	Ala	Gly	Ala	Gly	Leu	Gly	Gly	Val	Pro	Gly	Val	Gly	Gly	Leu	Gly
				85					90					95	
Val	Ser	Ala	Gly	Ala	Val	Val	Pro	Gln	Pro	Gly	Ala	Gly	Val	Lys	Pro
			100					105						110	
Gly	Lys	Val	Pro	Gly	Val	Gly	Leu	Pro	Gly	Val	Tyr	Pro	Gly	Gly	Val
			115				120					125			
Leu	Pro	Gly	Ala	Arg	Phe	Pro	Gly	Val	Gly	Val	Leu	Pro	Gly	Val	Pro
			130			135					140				
Thr	Gly	Ala	Gly	Val	Lys	Pro	Lys	Ala	Pro	Gly	Val	Gly	Gly	Ala	Phe
			145		150						155				160
Ala	Gly	Ile	Pro	Gly	Val	Gly	Pro	Phe	Gly	Gly	Pro	Gln	Pro	Gly	Val
				165				170						175	
Pro	Leu	Gly	Tyr	Pro	Ile	Lys	Ala	Pro	Lys	Leu	Pro	Gly	Gly	Tyr	Gly
			180					185						190	
Leu	Pro	Tyr	Thr	Thr	Gly	Lys	Leu	Pro	Tyr	Gly	Tyr	Gly	Pro	Gly	Gly
			195				200					205			
Val	Ala	Gly	Ala	Ala	Gly	Lys	Ala	Gly	Tyr	Pro	Thr	Gly	Thr	Gly	Val
			210			215					220				
Gly	Pro	Gln	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Lys	Ala	Ala	Ala	Lys	Phe
					230					235					240
Gly	Ala	Gly	Ala	Ala	Gly	Val	Leu	Pro	Gly	Val	Gly	Gly	Ala	Gly	Val
				245					250					255	
Pro	Gly	Val	Pro	Gly	Ala	Ile	Pro	Gly	Ile	Gly	Gly	Ile	Ala	Gly	Val
			260					265					270		
Gly	Thr	Pro	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Lys	Ala	Ala
			275			280							285		
Lys	Tyr	Gly	Ala	Ala	Ala	Gly	Leu	Val	Pro	Gly	Gly	Pro	Gly	Phe	Gly
			290			295						300			
Pro	Gly	Val	Val	Gly	Val	Pro	Gly	Ala	Gly	Val	Pro	Gly	Val	Gly	Val
					310					315					320
Pro	Gly	Ala	Gly	Ile	Pro	Val	Val	Pro	Gly	Ala	Gly	Ile	Pro	Gly	Ala
				325					330					335	
Ala	Val	Pro	Gly	Val	Val	Ser	Pro	Glu	Ala	Ala	Ala	Lys	Ala	Ala	Ala
				340				345					350		
Lys	Ala	Ala	Lys	Tyr	Gly	Ala	Arg	Pro	Gly	Val	Gly	Val	Gly	Gly	Ile
			355				360						365		
Pro	Thr	Tyr	Gly	Val	Gly	Ala	Gly	Gly	Phe	Pro	Gly	Phe	Gly	Val	Gly
			370				375					380			
Val	Gly	Gly	Ile	Pro	Gly	Val	Ala	Gly	Val	Pro	Gly	Val	Gly	Gly	Val
					390					395					400
Pro	Gly	Val	Gly	Gly	Val	Pro	Gly	Val	Gly	Ile	Ser	Pro	Glu	Ala	Gln
				405					410						415

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Ala Ala Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala Gly Ala Gly
 420 425 430

Val Leu Gly Gly Leu Val Pro Gly Pro Gln Ala Ala Val Pro Gly Val
 435 440 445

Pro Gly Thr Gly Gly Val Pro Gly Val Gly Thr Pro Ala Ala Ala Ala
 450 455 460

Ala Lys Ala Ala Ala Lys Ala Ala Gln Phe Gly Leu Val Pro Gly Val
 465 470 475 480

Gly Val Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Val Ala Pro
 485 490 495

Gly Val Gly Leu Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Val
 500 505 510

Ala Pro Gly Val Gly Val Ala Pro Gly Ile Gly Pro Gly Gly Val
 515 520 525

<210> SEQ ID NO 40
 <211> LENGTH: 1584
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 40

```

ggtggcgtac caggcgcaat tcctgggggt gtcccaggcg gtgtttttta tccgggcgcc      60
ggtcttggcg cactgggtgg cggtgcaact ggccccggcg gcaaaccgct gaaaccggta      120
ccaggtggtt tagcaggcgc cggcttaggc gcaggtctgg gagcatttcc ggcagttacc      180
tttccagggg cactggttcc tggaggtgtg gccgatgcag ccgcggcata taaagccgct      240
aaagccggtg cgggttttag aggcgtocca ggtgtcggtg gcctgggtgt tagcgcgggt      300
gcagttgttc cgcagccggg agcagggggt aaacctggta aagtgccggg agtaggtctg      360
ccaggcgttt atcctggtgg tgttttgccg ggtgcccggt ttccgggcgt tggtgttctt      420
ccaggcggtc cgaccggagc cgggtgtaaa ccgaaagccc ccggtgttgg aggtgcattt      480
gcaggcatcc cgggagttgg cccgtttggg ggtccgcaac ctggggttcc gttaggttat      540
ccgattaaag caccgaaact gcccgcggtt tatggtctgc cgtacacaac cggtaaactg      600
ccgtatggtt atggccccgg tggagttgcg ggtgcagcag gtaaagcggg ttatcctacc      660
ggaaccggtg taggtccgca ggcgctgct gccgcgccc caaaagcagc ggctaaattt      720
ggcgccggag cagcgggtgt tctgcctgga gttggtggtg cgggcgtgcc aggggtacct      780
ggtgcaattc cgggtattgg tggatttgcc ggtgtcggca ccccgccgc ggcagctgcg      840
gcagcggcgg ctgccaaagc tgctaatac ggtgcgcggc cgggtctggt gccaggaggt      900
ccgggttttg gtccgggagt ggttggcgtg cctggcgcag gcgttcctgg tgtgggcgtt      960
ccaggtgcag ggattcctgt tgtgcctggt gccggtattc ccggcgcggc cgttccgggg      1020
gtggttagcc cggaaagcgc agcgaaggct gcggcaaaag cagcaaaagta tggcgcacgc      1080
ccaggagtgc gcgtgggtgg tatcccagcc tatgggggtg gcgcaggggg ttttctctgt      1140
ttcggcgtag gtgtaggagg tataccgggc gtggccgggtg taccaggggt tgggtggcgtc      1200
cctggtgttg gcggtgtgcc aggtgttggg atttcaccgg aagcacaggc agcagccgca      1260
gctaaggcag cgaaaatatg tgccgcggc gcaggagttt taggtgggct ggttccgggc      1320
    
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ccgcaggeag ctgtgccggg ggttccagge accggtgggtg tccctggagt cggtagcgcg 1380
gtgcagcggg cagccaaagc ggctgcgaaa gcagcacagt ttggcttagt accgggtgtg 1440
ggagttgccc cggcgcttgg cgttgetcca ggggtgggtg ttgctcctgg cgtcggtctg 1500
gctcctggag tgggcgtage acccggtgtg ggggtggccc cgggtgttgg ggttgcaccg 1560
ggtatcggtc cgggcggtgt ctaa 1584

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<210> SEQ ID NO 41
<211> LENGTH: 515
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
      polypeptide

```

```

<400> SEQUENCE: 41

```

```

Gly Val Leu Pro Gly Val Gly Gly Ala Gly Val Pro Gly Val Pro Gly
 1          5          10          15
Ala Ile Pro Gly Ile Gly Gly Ile Ala Gly Val Gly Thr Pro Ala Ala
 20          25          30
Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala
 35          40          45
Ala Gly Leu Val Pro Gly Gly Pro Gly Phe Gly Pro Gly Val Val Gly
 50          55          60
Val Pro Gly Ala Gly Val Pro Gly Val Gly Val Pro Gly Ala Gly Ile
 65          70          75          80
Pro Val Val Pro Gly Ala Gly Ile Pro Gly Ala Ala Val Pro Gly Val
 85          90          95
Val Ser Pro Glu Ala Ala Ala Lys Ala Ala Ala Lys Ala Ala Lys Tyr
 100         105         110
Gly Ala Arg Pro Gly Val Gly Val Gly Gly Ile Pro Thr Tyr Gly Val
 115         120         125
Gly Ala Gly Gly Phe Pro Gly Phe Gly Val Gly Val Gly Gly Ile Pro
 130         135         140
Gly Val Ala Gly Val Pro Gly Val Gly Gly Val Pro Gly Val Gly Gly
 145         150         155         160
Val Pro Gly Val Gly Ile Ser Pro Glu Ala Gln Ala Ala Ala Ala Ala
 165         170         175
Lys Ala Ala Lys Tyr Gly Ala Ala Gly Ala Gly Val Leu Gly Gly Leu
 180         185         190
Val Pro Gly Pro Gln Ala Ala Val Pro Gly Val Pro Gly Thr Gly Gly
 195         200         205
Val Pro Gly Val Gly Thr Pro Ala Ala Ala Ala Ala Lys Ala Ala Ala
 210         215         220
Lys Ala Ala Gln Phe Gly Leu Val Pro Gly Val Gly Val Ala Pro Gly
 225         230         235         240
Val Gly Val Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Leu Ala
 245         250         255
Pro Gly Val Gly Val Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly
 260         265         270
Val Ala Pro Gly Ile Gly Pro Gly Gly Val Ala Ala Ala Ala Lys Ser
 275         280         285
Ala Ala Lys Val Ala Ala Lys Ala Gln Leu Arg Ala Ala Ala Gly Leu

```

-continued

290	295	300
Gly Ala Gly Ile Pro Gly Leu Gly Val Gly Val Gly Val Pro Gly Leu		
305	310	315 320
Gly Val Gly Ala Gly Val Pro Gly Leu Gly Val Gly Ala Gly Val Pro		
	325	330 335
Gly Phe Gly Ala Gly Ala Asp Glu Gly Val Arg Arg Ser Leu Ser Pro		
	340	345 350
Glu Leu Arg Glu Gly Asp Pro Ser Ser Ser Gln His Leu Pro Ser Thr		
	355	360 365
Pro Ser Ser Pro Arg Val Pro Gly Ala Leu Ala Ala Ala Lys Ala Ala		
	370	375 380
Lys Tyr Gly Ala Ala Val Pro Gly Val Leu Gly Gly Leu Gly Ala Leu		
	385	390 395 400
Gly Gly Val Gly Ile Pro Gly Gly Val Val Gly Ala Gly Pro Ala Ala		
	405	410 415
Ala Ala Ala Ala Ala Lys Ala Ala Ala Lys Ala Ala Gln Phe Gly Leu		
	420	425 430
Val Gly Ala Ala Gly Leu Gly Gly Leu Gly Val Gly Gly Leu Gly Val		
	435	440 445
Pro Gly Val Gly Gly Leu Gly Gly Ile Pro Pro Ala Ala Ala Ala Lys		
	450	455 460
Ala Ala Lys Tyr Gly Ala Ala Gly Leu Gly Gly Val Leu Gly Gly Ala		
	465	470 475 480
Gly Gln Phe Pro Leu Gly Gly Val Ala Ala Arg Pro Gly Phe Gly Leu		
	485	490 495
Ser Pro Ile Phe Pro Gly Gly Ala Cys Leu Gly Lys Ala Cys Gly Arg		
	500	505 510
Lys Arg Lys		
	515	

<210> SEQ ID NO 42
 <211> LENGTH: 1548
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 42

```

gggtgttctgc ctggagttgg tgggtgcgggc gtgccagggg tacctggtgc aattccgggt    60
attggtggta ttgccggtgt cggcaccocg gccgcggcag ctgcggcagc ggcggctgcc    120
aaagctgcta aatacggctg cgcggcgggt ctggtgccag gaggtccggg ttttggtccg    180
ggagtggttg gcgtgcctgg cgcaggcgtt cctggtgtgg gcgttccagg tgcagggatt    240
cctgttgtgc ctggtgccgg tattcccggc gcggccggtc cgggggtggt tagcccggaa    300
gccgcagcga aggctgcggc aaagccagca aagtatggcg cacgcccagg agtcggcgtg    360
ggtggtatcc cgacctatgg ggtgggcgca gggggttttc ctggtttcgg cgtaggtgta    420
ggaggtatac cgggcgtggc cgggtgtacca ggggttggtg gcgtccctgg tgttgccggt    480
gtgccaggtg ttggtatttc accggaagca caggcagcag ccgcagctaa ggcagcgaaa    540
tatggtgccg ccggcgcagg agtttttaggt gggctggttc cgggcccgca ggcagctgtg    600
cggggggttc caggcaccgg tgggtgcctt ggagtcggta cgccggtgc agcggcagcc    660
    
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aaagcggctg cgaagcagc acagtttggc ttagtaccgg gtgtgggagt tgccccggc 720
gttggcgttg ctccaggggt ggggtgtgct cctggcgctg gtctggtccc tggagtgggc 780
gtagcaccgg gtgtgggggt gggccccggg gtgggggttg caccgggtat cggtcggggc 840
ggtgtcgcag cagcagctaa aagcgcggcg aaagtgcgg ccaaagccca actgcgcgcc 900
gccgcggggc tcggtgcagg tattccgggg ctgggtgctg gagttggagt cccgggtttg 960
ggcgtggggc cgggagttcc gggactggga gtgggtgccc gagttcctgg ctttgggtgca 1020
ggcgcagatg aagggtgttc tcgtagcctg agtccggaac tgcgtgaagg tgatccgagt 1080
agcagccagc atctgccgag cccccgagc agcccgcgtg ttccgggtgc attagctgca 1140
gcaaaagccg ccaagtatgg tgcagccgtg ccgggcgtct taggtggtct gggcgcctcg 1200
ggtggtgtag gcattccggg aggtgtttgt ggtgcaggac cggcgcgcgc agctgcggcc 1260
gcaaagcag ctgcaaaagc ggcccagttt ggtttagtgg gcgcccagc tttaggcggg 1320
ttaggtgtgg gtggactggg tgtacctggc gtaggcggtc tgggtggaat tccgcccga 1380
gcccgcgcga aagcggcaaa atatggcgcg gcaggcctgg gcggcgtgct ggggtgggca 1440
ggtcagtttc cgctggggcg ggttgcgcga cgccgggat ttggtctgag cccgattttc 1500
cctggcggcg catgtctggg taaagcatgt ggtcgtaaac gtaaataa 1548

```

<210> SEQ ID NO 43

<211> LENGTH: 466

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 43

```

Gly Leu Val Pro Gly Gly Pro Gly Phe Gly Pro Gly Val Val Gly Val
1          5          10          15
Pro Gly Ala Gly Val Pro Gly Val Gly Val Pro Gly Ala Gly Ile Pro
20        25        30
Val Val Pro Gly Ala Gly Ile Pro Gly Ala Ala Val Pro Gly Val Val
35        40        45
Ser Pro Glu Ala Ala Ala Lys Ala Ala Ala Lys Ala Ala Lys Tyr Gly
50        55        60
Ala Arg Pro Gly Val Gly Val Gly Gly Ile Pro Thr Tyr Gly Val Gly
65        70        75        80
Ala Gly Gly Phe Pro Gly Phe Gly Val Gly Val Gly Gly Ile Pro Gly
85        90        95
Val Ala Gly Val Pro Gly Val Gly Gly Val Pro Gly Val Gly Gly Val
100       105       110
Pro Gly Val Gly Ile Ser Pro Glu Ala Gln Ala Ala Ala Ala Lys
115       120       125
Ala Ala Lys Tyr Gly Ala Ala Gly Ala Gly Val Leu Gly Gly Leu Val
130       135       140
Pro Gly Pro Gln Ala Ala Val Pro Gly Val Pro Gly Thr Gly Gly Val
145       150       155       160
Pro Gly Val Gly Thr Pro Ala Ala Ala Ala Ala Lys Ala Ala Ala Lys
165       170       175
Ala Ala Gln Phe Gly Leu Val Pro Gly Val Gly Val Ala Pro Gly Val

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	180		185		190																
Gly	Val	Ala	Pro	Gly	Val	Gly	Val	Ala	Pro	Gly	Val	Gly	Leu	Ala	Pro						
	195						200					205									
Gly	Val	Gly	Val	Ala	Pro	Gly	Val	Gly	Val	Ala	Pro	Gly	Val	Gly	Val						
	210						215				220										
Ala	Pro	Gly	Ile	Gly	Pro	Gly	Gly	Val	Ala	Ala	Ala	Ala	Lys	Ser	Ala						
	225				230					235					240						
Ala	Lys	Val	Ala	Ala	Lys	Ala	Gln	Leu	Arg	Ala	Ala	Ala	Gly	Leu	Gly						
			245						250					255							
Ala	Gly	Ile	Pro	Gly	Leu	Gly	Val	Gly	Val	Gly	Val	Pro	Gly	Leu	Gly						
	260						265						270								
Val	Gly	Ala	Gly	Val	Pro	Gly	Leu	Gly	Val	Gly	Ala	Gly	Val	Pro	Gly						
	275						280					285									
Phe	Gly	Ala	Gly	Ala	Asp	Glu	Gly	Val	Arg	Arg	Ser	Leu	Ser	Pro	Glu						
	290					295					300										
Leu	Arg	Glu	Gly	Asp	Pro	Ser	Ser	Ser	Gln	His	Leu	Pro	Ser	Thr	Pro						
	305				310					315					320						
Ser	Ser	Pro	Arg	Val	Pro	Gly	Ala	Leu	Ala	Ala	Ala	Lys	Ala	Ala	Lys						
				325					330						335						
Tyr	Gly	Ala	Ala	Val	Pro	Gly	Val	Leu	Gly	Gly	Leu	Gly	Ala	Leu	Gly						
	340							345					350								
Gly	Val	Gly	Ile	Pro	Gly	Gly	Val	Val	Gly	Ala	Gly	Pro	Ala	Ala	Ala						
	355						360					365									
Ala	Ala	Ala	Ala	Lys	Ala	Ala	Ala	Lys	Ala	Ala	Gln	Phe	Gly	Leu	Val						
	370				375						380										
Gly	Ala	Ala	Gly	Leu	Gly	Gly	Leu	Gly	Val	Gly	Gly	Leu	Gly	Val	Pro						
	385				390					395					400						
Gly	Val	Gly	Gly	Leu	Gly	Gly	Ile	Pro	Pro	Ala	Ala	Ala	Ala	Lys	Ala						
				405					410						415						
Ala	Lys	Tyr	Gly	Ala	Ala	Gly	Leu	Gly	Gly	Val	Leu	Gly	Gly	Ala	Gly						
	420							425					430								
Gln	Phe	Pro	Leu	Gly	Gly	Val	Ala	Ala	Arg	Pro	Gly	Phe	Gly	Leu	Ser						
	435					440						445									
Pro	Ile	Phe	Pro	Gly	Gly	Ala	Cys	Leu	Gly	Lys	Ala	Cys	Gly	Arg	Lys						
	450					455					460										
Arg	Lys																				
	465																				

<210> SEQ ID NO 44
 <211> LENGTH: 1401
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 44

ggtctggtgc caggaggctcc gggttttggt ccgggagtggt ttggcgtgcc tggcgcaggc	60
gttctctggtg tgggcgttcc aggtgcaggg attcctgttg tgctgtgtgc cggtattccc	120
ggcgcggcgc ttccgggggt ggttagcccc gaagccgcag cgaaggctgc ggcaaaggca	180
gcaaagtatg gcgcacgccc aggagtcggc gtgggtggta tccccaccta tggggtgggc	240
gcaggggggtt ttctgtgttt cggcgtaggt gtaggagta taccggcggt ggccggtgta	300

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ccaggggttg gtggcgtoce tgggtttggtc ggtgtgccag gtgttggtat ttcaccggaa 360
gcacagggcag cagccgcagc taaggcagcg aaatatggtg ccgcccggcg aggagtttta 420
ggtgggcttg ttccggggccc gcagggcagct gtgcccggggg ttccaggcac cggtggtgtc 480
cctggagtcg gtaagccggc tgcagcggca gccaaagcgg ctgcccgaagc agcacagttt 540
ggcttagtac cgggtgtggg agttgcccc gccgttggcg ttgctccagg ggtgggtgtt 600
gtcctggcg tgggtctggc tctggagtg ggcgttagcac ccggtgtggg ggtggccccg 660
ggtgttgggg ttgcccggg tatcggtcg gccggtgtcg cagcagcagc taaaagcgcg 720
gcgaaagtg cggccaaagc ccaactgcgc gccgcccggg gcctcgggtc aggtattccg 780
gggctgggtg tccgagttgg agtccccggg ttgggcgtgg gcgcccggag tccgggactg 840
ggagtggttg ccggagttcc tggctttggt gcagggcagc atgaaggtgt tegtctgtag 900
ctgagtcagg aactgcgtga aggtgatccg agtagcagcc agcatctgcc gagcaccgcc 960
agcagcccgc gtgttccggg tgcattagct gcagcaaaag ccgccaagta tgggtgcagcc 1020
gtgcccggcg tcttagtggt tctgggcgcc ctgggtgttg taggcattcc gggaggtgtt 1080
gtgggtgcag gaccggccgc cgcagctgcg gccgccaagc cagctgcaa agcggcccag 1140
tttggttag tgggcgcgc aggttttagc ggtttagtg tgggtggact ggtgtacct 1200
ggcgtaggcg gtctgggtg aattccgcc gcagcggcg cgaaagcggc aaaatatggc 1260
gcggcagccc tggcggcgt gctgggtgg gcaggtcagt ttccgctggg cggggttggc 1320
gcacgtccgg gatttggtct gagcccagtt ttccctggcg gcgcatgtct gggtaaagca 1380
tgtggtcgta aacgtaaata a 1401

```

<210> SEQ ID NO 45

<211> LENGTH: 460

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 45

```

Gly Gly Val Pro Gly Ala Ile Pro Gly Gly Val Pro Gly Gly Val Phe
1           5           10           15
Tyr Pro Gly Ala Gly Leu Gly Ala Leu Gly Gly Gly Ala Leu Gly Pro
20          25          30
Gly Gly Lys Pro Leu Lys Pro Val Pro Gly Gly Leu Ala Gly Ala Gly
35          40          45
Leu Gly Ala Gly Leu Gly Ala Phe Pro Ala Val Thr Phe Pro Gly Ala
50          55          60
Leu Val Pro Gly Gly Val Ala Asp Ala Ala Ala Tyr Lys Ala Ala
65          70          75          80
Lys Ala Gly Ala Gly Leu Gly Gly Val Pro Gly Val Gly Gly Leu Gly
85          90          95
Val Ser Ala Gly Ala Val Val Pro Gln Pro Gly Ala Gly Val Lys Pro
100         105         110
Gly Lys Val Pro Gly Val Gly Leu Pro Gly Val Tyr Pro Gly Gly Val
115         120         125
Leu Pro Gly Ala Arg Phe Pro Gly Val Gly Val Leu Pro Gly Val Pro
130         135         140

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Thr	Gly	Ala	Gly	Val	Lys	Pro	Lys	Ala	Pro	Gly	Val	Gly	Gly	Ala	Phe
145					150					155					160
Ala	Gly	Ile	Pro	Gly	Val	Gly	Pro	Phe	Gly	Gly	Pro	Gln	Pro	Gly	Val
				165					170					175	
Pro	Leu	Gly	Tyr	Pro	Ile	Lys	Ala	Pro	Lys	Leu	Pro	Gly	Gly	Tyr	Gly
		180						185						190	
Leu	Pro	Tyr	Thr	Thr	Gly	Lys	Leu	Pro	Tyr	Gly	Tyr	Gly	Pro	Gly	Gly
		195					200					205			
Val	Ala	Gly	Ala	Ala	Gly	Lys	Ala	Gly	Tyr	Pro	Thr	Gly	Thr	Gly	Val
	210					215					220				
Gly	Pro	Gln	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Lys	Ala	Ala	Ala	Lys	Phe
225					230					235					240
Gly	Ala	Gly	Ala	Ala	Gly	Val	Leu	Pro	Gly	Val	Gly	Gly	Ala	Gly	Val
				245					250					255	
Pro	Gly	Val	Pro	Gly	Ala	Ile	Pro	Gly	Ile	Gly	Gly	Ile	Ala	Gly	Val
		260						265					270		
Gly	Thr	Pro	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Lys	Ala	Ala
		275					280						285		
Lys	Tyr	Gly	Ala	Ala	Ala	Gly	Leu	Val	Pro	Gly	Gly	Pro	Gly	Phe	Gly
	290					295					300				
Pro	Gly	Val	Val	Gly	Val	Pro	Gly	Ala	Gly	Val	Pro	Gly	Val	Gly	Val
305					310					315					320
Pro	Gly	Ala	Gly	Ile	Pro	Val	Val	Pro	Gly	Ala	Gly	Ile	Pro	Gly	Ala
				325					330					335	
Ala	Val	Pro	Gly	Val	Val	Ser	Pro	Glu	Ala	Ala	Ala	Lys	Ala	Ala	Ala
			340					345					350		
Lys	Ala	Ala	Lys	Tyr	Gly	Ala	Arg	Pro	Gly	Val	Gly	Val	Gly	Gly	Ile
	355						360						365		
Pro	Thr	Tyr	Gly	Val	Gly	Ala	Gly	Gly	Phe	Pro	Gly	Phe	Gly	Val	Gly
	370					375					380				
Val	Gly	Gly	Ile	Pro	Gly	Val	Ala	Gly	Val	Pro	Gly	Val	Gly	Gly	Val
385					390					395					400
Pro	Gly	Val	Gly	Gly	Val	Pro	Gly	Val	Gly	Ile	Ser	Pro	Glu	Ala	Gln
				405					410					415	
Ala	Ala	Ala	Ala	Ala	Lys	Ala	Ala	Lys	Tyr	Gly	Ala	Ala	Gly	Ala	Gly
				420				425					430		
Val	Leu	Gly	Gly	Leu	Val	Pro	Gly	Pro	Gln	Ala	Ala	Val	Pro	Gly	Val
	435						440					445			
Pro	Gly	Thr	Gly	Gly	Val	Pro	Gly	Val	Gly	Thr	Pro				
	450						455				460				

<210> SEQ ID NO 46
 <211> LENGTH: 1383
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 46

ggtggcgctac caggcgcaat tcctgggggt gtcccaggcg gtgtttttta tccgggccc 60

ggtcttggcg cactgggtgg cggtgcaactg ggcccgggcg gcaaacegct gaaaccgta 120

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ccaggtggtt tagcaggcgc cggcttaggc gcaggtctgg ggcatttcc ggcagttacc 180
tttccagggg cactggttcc tggaggtgtg gccgatgcag ccgcggcata taaagccgct 240
aaagccggtg cgggtttagg aggcgtccca ggtgtcggtg gcctgggtgt tagcgcgggt 300
gcagttgttc cgcagccggg agcaggggtt aaacctggtg aagtgccggg agtaggtctg 360
ccaggcgttt atcctgggtg tgttttgccg ggtgcccgtt ttccgggcgt tgggtttctt 420
ccaggcggtc cgaccggagc cgggtttaa cggaaagccc ccggtgttgg aggtgcattt 480
gcaggcatcc cgggagttgg cccgtttggt ggtccgcaac ctggggttcc gttaggttat 540
ccgattaaag caccgaaact gcccgcggtt tatggtctgc cgtacacaac cggtaaactg 600
ccgtatggtt atgcccggg tggagttgcg ggtgcagcag gtaaagcggg ttatcctacc 660
ggaaccggtg taggtccgca ggcgctgct gccgcccgg caaaagcagc ggctaaattt 720
ggcgccggag cagcgggtgt tctgcctgga gttggtggtg cgggcgtgcc aggggtacct 780
ggtgcaattc cgggtattgg tggattgcc ggtgtcgca ccccgccgc ggcagctgcg 840
gcagcggcgg ctgccaaagc tgctaaatac ggtgcccggg cgggtctggt gccaggaggt 900
ccgggttttg gtccgggagt ggttggcgtg cctggcgcag gcgttcctgg tgtgggcgtt 960
ccaggtgcag ggattcctgt tgtgcctggt gccggtatcc ccggcgcggc cgttccgggg 1020
gtggttagcc cggaaagcgc agcgaaggct gcggcaaagg cagcaaagta tggcgcacgc 1080
ccaggagtgc gcgtgggtgg tatcccagcc tatgggggtg gcgcaggggg ttttcctggt 1140
ttcggcgtag gtgtaggagg tataccgggc gtggccggtg taccaggggt tgggtggcgtc 1200
cctggtgttg gcggtgtgcc aggtgttggg atttcaccgg aagcacaggc agcagccgca 1260
gctaaggcag cgaatatg tgccgcggc gcaggagttt taggtgggct ggttccgggc 1320
ccgcaggcag ctgtgccggg ggttccaggc accggtggtg tccctggagt cggtacgccg 1380
taa 1383

```

```

<210> SEQ ID NO 47
<211> LENGTH: 416
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
        polypeptide

```

<400> SEQUENCE: 47

```

Gly Gly Val Pro Gly Ala Ile Pro Gly Gly Val Pro Gly Gly Val Phe
 1             5             10             15
Tyr Pro Gly Ala Gly Leu Gly Ala Leu Gly Gly Gly Ala Leu Gly Pro
      20             25             30
Gly Gly Lys Pro Leu Lys Pro Val Pro Gly Gly Leu Ala Gly Ala Gly
      35             40             45
Leu Gly Ala Gly Leu Gly Ala Phe Pro Ala Val Thr Phe Pro Gly Ala
      50             55             60
Leu Val Pro Gly Gly Val Ala Asp Ala Ala Ala Ala Tyr Lys Ala Ala
      65             70             75             80
Lys Ala Gly Ala Gly Leu Gly Gly Val Pro Gly Val Gly Gly Leu Gly
      85             90             95
Val Ser Ala Gly Ala Val Val Pro Gln Pro Gly Ala Gly Val Lys Pro
      100            105            110

```

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Gly Lys Val Pro Gly Val Gly Leu Pro Gly Val Tyr Pro Gly Gly Val
 115 120 125

Leu Pro Gly Ala Arg Phe Pro Gly Val Gly Val Leu Pro Gly Val Pro
 130 135 140

Thr Gly Ala Gly Val Lys Pro Lys Ala Pro Gly Val Gly Gly Ala Phe
 145 150 155 160

Ala Gly Ile Pro Gly Val Gly Pro Phe Gly Gly Pro Gln Pro Gly Val
 165 170 175

Pro Leu Gly Tyr Pro Ile Lys Ala Pro Lys Leu Pro Gly Gly Tyr Gly
 180 185 190

Leu Pro Tyr Thr Thr Gly Lys Leu Pro Tyr Gly Tyr Gly Pro Gly Gly
 195 200 205

Val Ala Gly Ala Ala Gly Lys Ala Gly Tyr Pro Thr Gly Thr Gly Val
 210 215 220

Gly Pro Gln Ala Ala Ala Ala Ala Ala Lys Ala Ala Ala Lys Phe
 225 230 235 240

Gly Ala Gly Ala Ala Gly Val Leu Pro Gly Val Gly Gly Ala Gly Val
 245 250 255

Pro Gly Val Pro Gly Ala Ile Pro Gly Ile Gly Gly Ile Ala Gly Val
 260 265 270

Gly Thr Pro Ala Ala Ala Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala
 275 280 285

Lys Tyr Gly Ala Ala Ala Gly Leu Val Pro Gly Gly Pro Gly Phe Gly
 290 295 300

Pro Gly Val Val Gly Val Pro Gly Ala Gly Val Pro Gly Val Gly Val
 305 310 315 320

Pro Gly Ala Gly Ile Pro Val Val Pro Gly Ala Gly Ile Pro Gly Ala
 325 330 335

Ala Val Pro Gly Val Val Ser Pro Glu Ala Ala Ala Lys Ala Ala Ala
 340 345 350

Lys Ala Ala Lys Tyr Gly Ala Arg Pro Gly Val Gly Val Gly Gly Ile
 355 360 365

Pro Thr Tyr Gly Val Gly Ala Gly Gly Phe Pro Gly Phe Gly Val Gly
 370 375 380

Val Gly Gly Ile Pro Gly Val Ala Gly Val Pro Gly Val Gly Gly Val
 385 390 395 400

Pro Gly Val Gly Gly Val Pro Gly Val Gly Ile Ser Pro Glu Ala Gln
 405 410 415

<210> SEQ ID NO 48
 <211> LENGTH: 1251
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polynucleotide

<400> SEQUENCE: 48

ggtggcgctac caggcgcaat tcttgggggt gtcccaggcg gtgtttttta tccggggcgc 60

ggtcttggcg cactgggtgg cggtgcactg ggcccgggcg gcaaaccgct gaaaccggta 120

ccaggtggtt tagcaggcgc cggttaggc gcaggtctgg gagcatttcc ggcagttacc 180

tttccagggg cactggttcc tggaggtgtg gccgatgcag ccgcggcata taaagcgcct 240

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aaagccggtg cgggtttagg agcgctccca ggtgtcgggtg gcctgggtgt tagcgccggt 300
gcagttgttc cgcagccggg agcaggggtt aaacctggta aagtgccggg agtaggtctg 360
ccaggcgttt atcctggtgg tgttttgccg ggtgcccgtt ttecgggcgt tgggtttctt 420
ccaggcggtc cgaccggagc cggtgtaaa ccgaaagccc ccggtgttg aggtgcattt 480
gcaggcatcc cgggagttgg cccgtttggt ggtccgcaac ctggggttcc gttaggttat 540
ccgattaag caccgaaact gcccgcggt tatggtctgc cgtacacaac cggtaaactg 600
ccgtatggtt atggcccggg tggagtgcg ggtgcagcag gtaaagcggg ttatcctacc 660
ggaaccggtg taggtccgca ggcgctgct gccgcccgcg caaaagcagc ggctaaattt 720
ggcgcgggag cagcgggtgt tctgcctgga gttggtggtg cgggcgtgcc aggggtacct 780
ggtgcaattc cgggtattgg tggattgcc ggtgtcggca ccccgccgc ggcagctgcg 840
gcagcggcgg ctgccaaagc tgctaaatac ggtgcccggc cgggtctggt gccaggaggt 900
ccgggttttg gtccgggagt ggttggcgtg cctggcgcag gcgttcctgg tgtggcggtt 960
ccaggtgcag ggattcctgt tgtgcctggt gccggtattc ccggcgcggc cgttccgggg 1020
gtggttagcc cggaaagcgc agcgaaggct gcggcaaagg cagcaaagta tggcgcacgc 1080
ccaggagtgc gcgtgggtgg tatcccagacc tatgggggtg gcgcaggggg ttttccgtgt 1140
ttcggcgtag gtgtaggagg tataccgggc gtggccggtg taccaggggt tggtgcgctc 1200
cctggtgttg gcggtgtgcc aggtgttggg atttaccggg aagcacagta a 1251
    
```

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<210> SEQ ID NO 49
<211> LENGTH: 401
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
        polypeptide
    
```

<400> SEQUENCE: 49

```

Arg Pro Gly Val Gly Val Gly Gly Ile Pro Thr Tyr Gly Val Gly Ala
1          5          10          15
Gly Gly Phe Pro Gly Phe Gly Val Gly Val Gly Gly Ile Pro Gly Val
20          25          30
Ala Gly Val Pro Gly Val Gly Gly Val Pro Gly Val Gly Gly Val Pro
35          40          45
Gly Val Gly Ile Ser Pro Glu Ala Gln Ala Ala Ala Ala Lys Ala
50          55          60
Ala Lys Tyr Gly Ala Ala Gly Ala Gly Val Leu Gly Gly Leu Val Pro
65          70          75          80
Gly Pro Gln Ala Ala Val Pro Gly Val Pro Gly Thr Gly Gly Val Pro
85          90          95
Gly Val Gly Thr Pro Ala Ala Ala Ala Lys Ala Ala Ala Lys Ala
100         105         110
Ala Gln Phe Gly Leu Val Pro Gly Val Gly Val Ala Pro Gly Val Gly
115         120         125
Val Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Leu Ala Pro Gly
130         135         140
Val Gly Val Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Val Ala
145         150         155         160
Pro Gly Ile Gly Pro Gly Gly Val Ala Ala Ala Ala Lys Ser Ala Ala
    
```

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165					170					175						
Lys	Val	Ala	Ala	Lys	Ala	Gln	Leu	Arg	Ala	Ala	Ala	Ala	Gly	Leu	Gly	Ala
		180						185						190		
Gly	Ile	Pro	Gly	Leu	Gly	Val	Gly	Val	Gly	Val	Pro	Gly	Leu	Gly	Val	
		195					200					205				
Gly	Ala	Gly	Val	Pro	Gly	Leu	Gly	Val	Gly	Ala	Gly	Val	Pro	Gly	Phe	
	210					215					220					
Gly	Ala	Gly	Ala	Asp	Glu	Gly	Val	Arg	Arg	Ser	Leu	Ser	Pro	Glu	Leu	
	225					230					235				240	
Arg	Glu	Gly	Asp	Pro	Ser	Ser	Ser	Gln	His	Leu	Pro	Ser	Thr	Pro	Ser	
			245						250					255		
Ser	Pro	Arg	Val	Pro	Gly	Ala	Leu	Ala	Ala	Ala	Lys	Ala	Ala	Lys	Tyr	
			260					265						270		
Gly	Ala	Ala	Val	Pro	Gly	Val	Leu	Gly	Gly	Leu	Gly	Ala	Leu	Gly	Gly	
		275					280					285				
Val	Gly	Ile	Pro	Gly	Gly	Val	Val	Gly	Ala	Gly	Pro	Ala	Ala	Ala	Ala	
	290					295					300					
Ala	Ala	Ala	Lys	Ala	Ala	Ala	Lys	Ala	Ala	Gln	Phe	Gly	Leu	Val	Gly	
	305					310					315				320	
Ala	Ala	Gly	Leu	Gly	Gly	Leu	Gly	Val	Gly	Gly	Leu	Gly	Val	Pro	Gly	
			325					330						335		
Val	Gly	Gly	Leu	Gly	Gly	Ile	Pro	Pro	Ala	Ala	Ala	Ala	Lys	Ala	Ala	
			340					345						350		
Lys	Tyr	Gly	Ala	Ala	Gly	Leu	Gly	Gly	Val	Leu	Gly	Gly	Ala	Gly	Gln	
		355					360						365			
Phe	Pro	Leu	Gly	Gly	Val	Ala	Ala	Arg	Pro	Gly	Phe	Gly	Leu	Ser	Pro	
	370					375					380					
Ile	Phe	Pro	Gly	Gly	Ala	Cys	Leu	Gly	Lys	Ala	Cys	Gly	Arg	Lys	Arg	
	385					390					395				400	

Lys

<210> SEQ ID NO 50
 <211> LENGTH: 1206
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 50

```

cgcccaggag tcggcgtggg tggtatcccg acctatgggg tgggcgcagg gggttttcct    60
ggtttcggcg taggtgtagg aggtataccg ggcgtggccg gtgtaccagg ggttggtggc    120
gtccctggtg ttggcgtgtg gccaggtgtt ggtatttcac cggaagcaca ggcagcagcc    180
gcagctaagg cagcgaataa tggtgccgcc ggccgaggag ttttaggtgg gctggttccg    240
ggcccgcagg cagctgtgcc gggggttcca ggcaccggtg gtgtccctgg agtcggtacg    300
ccggctgcag cggcagccaa agcgggtgct aaagcagcac agtttgctt agtaccgggt    360
gtgggagttg cccccggcgt tgccgttget ccaggggtgg gtgttgetcc tggcgtcggt    420
ctggtcctcg gagtggcgt agcaccgggt gtgggggtgg ccccggtgtg tggggttgea    480
ccgggtatcg gtcggggcgg tgtcgcagca gcagctaaaa gcgcggcgaa agttgcggcc    540
aaagcccaac tgcgcgccgc cgcgggctc ggtgcagta ttccggggct ggggtgtcga    600

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gttgagatcc cgggtttggg cgtgggcgcg ggagttccgg gactgggagt gggtgccgga	660
gttctcggct ttggtgcagg cgcagatgaa ggtgttcgtc gtagectgag tccggaactg	720
cgtgaagggtg atccgagtag cagccagcat ctgccgagca ccccgagcag cccgcgtgtt	780
ccgggtgcat tagctgcagc aaaagccgcc aagtatggtg cagccgtgcc gggcgtotta	840
ggtggtctgg gcgccctggg tgggttaggc attccgggag gtgttggtgg tgcaggaccg	900
gccgccgag ctgcggcgcg caaagcagct gcaaaagcgg cccagtttgg tttagtgggc	960
gccgcaggtt taggcggttt aggtgtgggt ggactgggtg tacctggcgt aggcggtctg	1020
ggtggaattc cgcccgcagc ggccgcgaaa gcggcaaaat atggcgcggc aggcctgggc	1080
ggcgtgctgg gtggggcagg tcagtttccg ctgggcgggg ttgccgcacg tccgggattt	1140
ggtctgagcc cgattttccc tggcggcgcg tgctgggta aagcatgtgg tcgtaaacgt	1200
aaataa	1206

<210> SEQ ID NO 51
 <211> LENGTH: 371
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 51

Gly Gly Val Pro Gly Ala Ile Pro Gly Gly Val Pro Gly Gly Val Phe
 1 5 10 15
 Tyr Pro Gly Ala Gly Leu Gly Ala Leu Gly Gly Gly Ala Leu Gly Pro
 20 25 30
 Gly Gly Lys Pro Leu Lys Pro Val Pro Gly Gly Leu Ala Gly Ala Gly
 35 40 45
 Leu Gly Ala Gly Leu Gly Ala Phe Pro Ala Val Thr Phe Pro Gly Ala
 50 55 60
 Leu Val Pro Gly Gly Val Ala Asp Ala Ala Ala Tyr Lys Ala Ala
 65 70 75 80
 Lys Ala Gly Ala Gly Leu Gly Gly Val Pro Gly Val Gly Gly Leu Gly
 85 90 95
 Val Ser Ala Gly Ala Val Val Pro Gln Pro Gly Ala Gly Val Lys Pro
 100 105 110
 Gly Lys Val Pro Gly Val Gly Leu Pro Gly Val Tyr Pro Gly Gly Val
 115 120 125
 Leu Pro Gly Ala Arg Phe Pro Gly Val Gly Val Leu Pro Gly Val Pro
 130 135 140
 Thr Gly Ala Gly Val Lys Pro Lys Ala Pro Gly Val Gly Gly Ala Phe
 145 150 155 160
 Ala Gly Ile Pro Gly Val Gly Pro Phe Gly Gly Pro Gln Pro Gly Val
 165 170 175
 Pro Leu Gly Tyr Pro Ile Lys Ala Pro Lys Leu Pro Gly Gly Tyr Gly
 180 185 190
 Leu Pro Tyr Thr Thr Gly Lys Leu Pro Tyr Gly Tyr Gly Pro Gly Gly
 195 200 205
 Val Ala Gly Ala Ala Gly Lys Ala Gly Tyr Pro Thr Gly Thr Gly Val
 210 215 220

-continued

Gly Pro Gln Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala Ala Lys Phe
 225 230 235 240

Gly Ala Gly Ala Ala Gly Val Leu Pro Gly Val Gly Gly Ala Gly Val
 245 250 255

Pro Gly Val Pro Gly Ala Ile Pro Gly Ile Gly Gly Ile Ala Gly Val
 260 265 270

Gly Thr Pro Ala Ala Ala Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala
 275 280 285

Lys Tyr Gly Ala Ala Ala Gly Leu Val Pro Gly Gly Pro Gly Phe Gly
 290 295 300

Pro Gly Val Val Gly Val Pro Gly Ala Gly Val Pro Gly Val Gly Val
 305 310 315 320

Pro Gly Ala Gly Ile Pro Val Val Pro Gly Ala Gly Ile Pro Gly Ala
 325 330 335

Ala Val Pro Gly Val Val Ser Pro Glu Ala Ala Ala Lys Ala Ala Ala
 340 345 350

Lys Ala Ala Lys Tyr Gly Ala Arg Pro Gly Val Gly Val Gly Gly Ile
 355 360 365

Pro Thr Tyr
 370

<210> SEQ ID NO 52
 <211> LENGTH: 1116
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 52

ggtggcgctac caggcgcaat tcttgggggt gtcccaggcg gtgtttttta tccgggcgcc 60

ggtcttggcg cactgggttg cggtgcaact ggccccggcg gcaaaccgct gaaaccggta 120

ccaggtgggt tagcaggcgc cggcttaggc gcaggtctgg gagcatttcc ggcagttacc 180

tttccagggg cactggttcc tggaggtgtg gccgatgcag ccgcggcata taaagccgct 240

aaagccggtg cgggtttagg aggcgtccca ggtgtcggtg gcctgggtgt tagcggcggt 300

gcagttgttc cgcagccggg agcagggggt aaacctggta aagtgccggg agtaggtctg 360

ccaggcgttt atcctgtgtg tgttttgccg ggtgcccgtt tccggggcgt tgggtgttctt 420

ccaggcggtg cgaccgggag cgggtgttaa ccgaaagccc ccggtgttgg aggtgcattt 480

gcaggcatcc cgggagttgg cccgtttggt ggtccgcaac ctggggttcc gttaggttat 540

ccgattaaag caccgaaact gcccgcggtt tatggtctgc cgtacacaac cggtaaaactg 600

ccgtatggtt atggccccgg tggagttgcy ggtgcagcag gtaaagcggg ttatcctacc 660

ggaaccggtg taggtccgca ggccgctgct gccgccccg caaaagcagc ggctaaattt 720

ggcgccggag cagcgggtgt tctgcctgga gttggtggtg cgggcgtgcc aggggtacct 780

ggtgcaattc cgggtatttg tggatttggc ggtgtcggca ccccgccgc ggcagctgcy 840

gcagcggcgg ctgccaaagc tgctaaatac ggtgcccggc cgggtctggt gccaggaggt 900

ccgggttttg gtccgggagt ggttggcgtg cctggcgcag gcgttccctg tgtggcggtt 960

ccaggtgcag ggattcctgt tgtgcctggt gccggtattc ccggcgcggc cgttccgggg 1020

gtggttagcc cggaaagccg agcgaaggtt gccgcaaagg cagcaaagta tggcgcacgc 1080

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ccaggagtcg gcgtgggtgg tatcccgacc tatta

1116

<210> SEQ ID NO 53

<211> LENGTH: 345

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 53

Gly Gly Val Pro Gly Ala Ile Pro Gly Gly Val Pro Gly Gly Val Phe
 1 5 10 15
 Tyr Pro Gly Ala Gly Leu Gly Ala Leu Gly Gly Gly Ala Leu Gly Pro
 20 25 30
 Gly Gly Lys Pro Leu Lys Pro Val Pro Gly Gly Leu Ala Gly Ala Gly
 35 40 45
 Leu Gly Ala Gly Leu Gly Ala Phe Pro Ala Val Thr Phe Pro Gly Ala
 50 55 60
 Leu Val Pro Gly Gly Val Ala Asp Ala Ala Ala Tyr Lys Ala Ala
 65 70 75 80
 Lys Ala Gly Ala Gly Leu Gly Gly Val Pro Gly Val Gly Gly Leu Gly
 85 90 95
 Val Ser Ala Gly Ala Val Val Pro Gln Pro Gly Ala Gly Val Lys Pro
 100 105 110
 Gly Lys Val Pro Gly Val Gly Leu Pro Gly Val Tyr Pro Gly Gly Val
 115 120 125
 Leu Pro Gly Ala Arg Phe Pro Gly Val Gly Val Leu Pro Gly Val Pro
 130 135 140
 Thr Gly Ala Gly Val Lys Pro Lys Ala Pro Gly Val Gly Gly Ala Phe
 145 150 155 160
 Ala Gly Ile Pro Gly Val Gly Pro Phe Gly Gly Pro Gln Pro Gly Val
 165 170 175
 Pro Leu Gly Tyr Pro Ile Lys Ala Pro Lys Leu Pro Gly Gly Tyr Gly
 180 185 190
 Leu Pro Tyr Thr Thr Gly Lys Leu Pro Tyr Gly Tyr Gly Pro Gly Gly
 195 200 205
 Val Ala Gly Ala Ala Gly Lys Ala Gly Tyr Pro Thr Gly Thr Gly Val
 210 215 220
 Gly Pro Gln Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala Ala Lys Phe
 225 230 235 240
 Gly Ala Gly Ala Ala Gly Val Leu Pro Gly Val Gly Gly Ala Gly Val
 245 250 255
 Pro Gly Val Pro Gly Ala Ile Pro Gly Ile Gly Gly Ile Ala Gly Val
 260 265 270
 Gly Thr Pro Ala Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala
 275 280 285
 Lys Tyr Gly Ala Ala Ala Gly Leu Val Pro Gly Gly Pro Gly Phe Gly
 290 295 300
 Pro Gly Val Val Gly Val Pro Gly Ala Gly Val Pro Gly Val Gly Val
 305 310 315 320
 Pro Gly Ala Gly Ile Pro Val Val Pro Gly Ala Gly Ile Pro Gly Ala
 325 330 335

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Ala Val Pro Gly Val Val Ser Pro Glu
 340 345

<210> SEQ ID NO 54
 <211> LENGTH: 1038
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polynucleotide

<400> SEQUENCE: 54

```

ggtggcgtag caggcgcaat tctggggggt gtcccaggcg gtgtttttta tccgggagcc    60
ggtcttgagc cactgggtgg cgggtgactg ggcccgggag gaaaccgct gaaaccggta    120
ccaggtggtt tagcaggcgc cggcttaggc gcaggtctgg gagcatttcc ggcagttacc    180
tttccagggg cactggttcc tggaggtgtg gccgatgcag ccgcggcata taaagccgct    240
aaagccgggt cgggttttag aggcgtccca ggtgtcgggt gcctgggtgt tagcgcgggt    300
gcagttgttc cgcagccggg agcaggggtt aaacctggta aagtgcgggg agtaggtctg    360
ccaggcggtt atcctggtgg tgttttgccg ggtgcccgtt tccggggcgt tgggtttctt    420
ccaggcgtgc cgaccggagc cgggtgtaaa ccgaaagccc ccggtgttgg aggtgcattt    480
gcaggcattc cgggagttgg cccggttggg ggtccgcaac ctggggttcc gttaggttat    540
ccgattaaag caccgaaact gcccgcggtt tatggtctgc cgtacacaac cggtaaactg    600
ccgtatggtt atggcccggg tggagttgag ggtgcagcag gtaaagcggg ttatcctacc    660
ggaaccgggt taggtccgca ggcgctgctt gccgcgccc caaaagcagc ggctaaattt    720
ggcgcgggag cagcgggtgt tctgcctgga gttggtggtg cgggcgtgcc aggggtacct    780
ggtgcaattc cgggtattgg tggatttgcg ggtgtcggca ccccggcgcg gccagctgag    840
gcagcggcgg ctgcccgaag tgcataatac ggtgcccggc cgggtctggt gccaggaggt    900
ccgggttttg gtccgggagt ggttggcgtg cctggcgcag gcgttcctgg tgtgggcgtt    960
ccaggtgcag ggattcctgt tgtgctggtt gccggtatcc ccggcggcgc cgttccgggg    1020
gtggttagcc cggaataa                                     1038
  
```

<210> SEQ ID NO 55
 <211> LENGTH: 336
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polypeptide

<400> SEQUENCE: 55

```

Lys Tyr Gly Ala Ala Gly Ala Gly Val Leu Gly Gly Leu Val Pro Gly
1           5           10           15
Pro Gln Ala Ala Val Pro Gly Val Pro Gly Thr Gly Gly Val Pro Gly
20           25           30
Val Gly Thr Pro Ala Ala Ala Ala Lys Ala Ala Ala Lys Ala Ala
35           40           45
Gln Phe Gly Leu Val Pro Gly Val Gly Val Ala Pro Gly Val Gly Val
50           55           60
Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Leu Ala Pro Gly Val
65           70           75           80
  
```

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Gly Val Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Val Ala Pro
85 90 95

Gly Ile Gly Pro Gly Gly Val Ala Ala Ala Ala Lys Ser Ala Ala Lys
100 105 110

Val Ala Ala Lys Ala Gln Leu Arg Ala Ala Ala Gly Leu Gly Ala Gly
115 120 125

Ile Pro Gly Leu Gly Val Gly Val Gly Val Pro Gly Leu Gly Val Gly
130 135 140

Ala Gly Val Pro Gly Leu Gly Val Gly Ala Gly Val Pro Gly Phe Gly
145 150 155 160

Ala Gly Ala Asp Glu Gly Val Arg Arg Ser Leu Ser Pro Glu Leu Arg
165 170 175

Glu Gly Asp Pro Ser Ser Ser Gln His Leu Pro Ser Thr Pro Ser Ser
180 185 190

Pro Arg Val Pro Gly Ala Leu Ala Ala Ala Lys Ala Ala Lys Tyr Gly
195 200 205

Ala Ala Val Pro Gly Val Leu Gly Gly Leu Gly Ala Leu Gly Gly Val
210 215 220

Gly Ile Pro Gly Gly Val Val Gly Ala Gly Pro Ala Ala Ala Ala Ala
225 230 235 240

Ala Ala Lys Ala Ala Ala Lys Ala Ala Gln Phe Gly Leu Val Gly Ala
245 250 255

Ala Gly Leu Gly Gly Leu Gly Val Gly Gly Leu Gly Val Pro Gly Val
260 265 270

Gly Gly Leu Gly Gly Ile Pro Pro Ala Ala Ala Ala Lys Ala Ala Lys
275 280 285

Tyr Gly Ala Ala Gly Leu Gly Gly Val Leu Gly Gly Ala Gly Gln Phe
290 295 300

Pro Leu Gly Gly Val Ala Ala Arg Pro Gly Phe Gly Leu Ser Pro Ile
305 310 315 320

Phe Pro Gly Gly Ala Cys Leu Gly Lys Ala Cys Gly Arg Lys Arg Lys
325 330 335

<210> SEQ ID NO 56
<211> LENGTH: 1011
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
polynucleotide

<400> SEQUENCE: 56

aaatatggtg cgcgccggcg aggagtttta ggtgggctgg ttccgggccc gcaggcagct 60

gtgccggggg ttccaggcac cgttggtgtc cctggagtcg gtacgccggc tgcagcggca 120

gccaaaagcg ctgcgaaagc agcacagttt ggcttagtac cgggtgtggg agttgcccc 180

ggcgttggcg ttgctccagg ggtgggtgtt gctcctggcg tcggtctggc tctggagtg 240

ggcgtagcac ccggtgtggg ggtggccccg ggtgttgggg ttgcaccggg tatcggtccg 300

ggcgtgtcgc cagcagcagc taaaagcgcg gcgaaagttg cggccaaagc ccaactgcgc 360

gccgccggcg gcctcggtgc aggtattccg gggctgggtg tcggagtgg agtccccggg 420

ttgggcgtgg gcgcgggagt tccgggactg ggagtgggtg ccggagttec tggctttggt 480

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gcagggcgcag atgaaggtgt tcgtcgtagc ctgagtcagg aactgcgtga aggtgatccg 540
agtagcagcc agcatctgcc gagcaccgcc agcagccccc gtgttccggg tgcattagct 600
gcagcaaaaag ccgccaagta tgggtgcagcc gtgccggggc tcttaggtgg tctggggccc 660
ctgggtgggtg taggcattcc gggaggtgtt gtgggtgcag gaccggccgc cgcagctgcg 720
gccgcaaaag cagctgcaaa agcggccccc tttggtttag tggggcggcc aggttttagc 780
ggtttaggtg tgggtggact ggggtgtacct ggcgtaggcg gtctgggtgg aattccgccc 840
gcagcggccg cgaaagcggc aaaatatggc gcggcaggcc tggggcggcg gctgggtggg 900
gcaggtcagt ttccgctggg cgggggttgc gcacgtccgg gatttggctc gagcccgatt 960
ttccctggcg gcgcattgct gggtaaagca tgtggtcgta aacgtaaata a 1011
    
```

```

<210> SEQ ID NO 57
<211> LENGTH: 288
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
    polypeptide
    
```

<400> SEQUENCE: 57

```

Gln Phe Gly Leu Val Pro Gly Val Gly Val Ala Pro Gly Val Gly Val
1           5           10          15
Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Leu Ala Pro Gly Val
20          25          30
Gly Val Ala Pro Gly Val Gly Val Ala Pro Gly Val Gly Val Ala Pro
35          40          45
Gly Ile Gly Pro Gly Gly Val Ala Ala Ala Ala Lys Ser Ala Ala Lys
50          55          60
Val Ala Ala Lys Ala Gln Leu Arg Ala Ala Ala Gly Leu Gly Ala Gly
65          70          75          80
Ile Pro Gly Leu Gly Val Gly Val Gly Val Pro Gly Leu Gly Val Gly
85          90          95
Ala Gly Val Pro Gly Leu Gly Val Gly Ala Gly Val Pro Gly Phe Gly
100         105        110
Ala Gly Ala Asp Glu Gly Val Arg Arg Ser Leu Ser Pro Glu Leu Arg
115        120        125
Glu Gly Asp Pro Ser Ser Ser Gln His Leu Pro Ser Thr Pro Ser Ser
130        135        140
Pro Arg Val Pro Gly Ala Leu Ala Ala Ala Lys Ala Ala Lys Tyr Gly
145        150        155        160
Ala Ala Val Pro Gly Val Leu Gly Gly Leu Gly Ala Leu Gly Gly Val
165        170        175
Gly Ile Pro Gly Gly Val Val Gly Ala Gly Pro Ala Ala Ala Ala
180        185        190
Ala Ala Lys Ala Ala Ala Lys Ala Ala Gln Phe Gly Leu Val Gly Ala
195        200        205
Ala Gly Leu Gly Gly Leu Gly Val Gly Gly Leu Gly Val Pro Gly Val
210        215        220
Gly Gly Leu Gly Gly Ile Pro Pro Ala Ala Ala Ala Lys Ala Ala Lys
225        230        235        240
Tyr Gly Ala Ala Gly Leu Gly Gly Val Leu Gly Gly Ala Gly Gln Phe
245        250        255
    
```

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Pro Leu Gly Gly Val Ala Ala Arg Pro Gly Phe Gly Leu Ser Pro Ile
 260 265 270

Phe Pro Gly Gly Ala Cys Leu Gly Lys Ala Cys Gly Arg Lys Arg Lys
 275 280 285

<210> SEQ ID NO 58
 <211> LENGTH: 867
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polynucleotide

<400> SEQUENCE: 58

cagtttggt tagtaccggg tgtgggagtt gccccggcg ttggcgttgc tccaggggtg 60
 ggtgttgctc ctggcgctcg tctggctcct ggagtggcg tagcaccggg tgtgggggtg 120
 gccccgggtg ttggggttgc accgggtatc ggccggggcg gtgtgcgagc agcagctaaa 180
 agcgcggcga aagttgctggc caaagcccaa ctgctggcgg ccgctgggct cggtgcaggt 240
 attccggggc tgggtgtcgg agttggagtc ccgggtttgg gcgtggggcg gggagttccg 300
 ggactgggag tgggtgcccg agttcctggc tttggtgcag gcgcagatga aggtgttcgt 360
 cgtagcctga gtccggaact gcgtgaaggt gatccgagta gcagccagca tctgcccgagc 420
 accccgagca gcccgctgt tccgggtgca ttagctgcag caaaagccgc caagtatggt 480
 gcagccgtgc cgggcgtctt aggtgtctg ggcgcccgg gtggtgtagg cattccggga 540
 ggtgttgggt gtgcaggacc ggccgcccga gctgcggccg ccaaagcagc tgcaaaagcg 600
 gccagtttg gtttagtggg ccgcccaggt ttaggcggtt taggtgtggg tggactgggt 660
 gtacctggcg taggcggtct ggttgaatt ccgcccgcag cggccgcgaa agcggcaaaa 720
 tatggcgcgg caggcctggg ccgctgctg ggtggggcag gtcagtttcc gctgggcccg 780
 gttccgcac gtccgggatt tggcttgagc ccgattttcc ctggcggcgc atgtctgggt 840
 aaagcatgtg gtcgtaaacg taaataa 867

<210> SEQ ID NO 59
 <211> LENGTH: 275
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polypeptide

<400> SEQUENCE: 59

Gly Gly Val Pro Gly Ala Ile Pro Gly Gly Val Pro Gly Gly Val Phe
 1 5 10 15

Tyr Pro Gly Ala Gly Leu Gly Ala Leu Gly Gly Gly Ala Leu Gly Pro
 20 25 30

Gly Gly Lys Pro Leu Lys Pro Val Pro Gly Gly Leu Ala Gly Ala Gly
 35 40 45

Leu Gly Ala Gly Leu Gly Ala Phe Pro Ala Val Thr Phe Pro Gly Ala
 50 55 60

Leu Val Pro Gly Gly Val Ala Asp Ala Ala Ala Tyr Lys Ala Ala
 65 70 75 80

Lys Ala Gly Ala Gly Leu Gly Gly Val Pro Gly Val Gly Gly Leu Gly
 85 90 95

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Val	Ser	Ala	Gly	Ala	Val	Val	Pro	Gln	Pro	Gly	Ala	Gly	Val	Lys	Pro
			100					105						110	
Gly	Lys	Val	Pro	Gly	Val	Gly	Leu	Pro	Gly	Val	Tyr	Pro	Gly	Gly	Val
		115					120					125			
Leu	Pro	Gly	Ala	Arg	Phe	Pro	Gly	Val	Gly	Val	Leu	Pro	Gly	Val	Pro
		130				135					140				
Thr	Gly	Ala	Gly	Val	Lys	Pro	Lys	Ala	Pro	Gly	Val	Gly	Gly	Ala	Phe
145					150					155					160
Ala	Gly	Ile	Pro	Gly	Val	Gly	Pro	Phe	Gly	Gly	Pro	Gln	Pro	Gly	Val
				165					170					175	
Pro	Leu	Gly	Tyr	Pro	Ile	Lys	Ala	Pro	Lys	Leu	Pro	Gly	Gly	Tyr	Gly
			180					185						190	
Leu	Pro	Tyr	Thr	Thr	Gly	Lys	Leu	Pro	Tyr	Gly	Tyr	Gly	Pro	Gly	Gly
		195					200					205			
Val	Ala	Gly	Ala	Ala	Gly	Lys	Ala	Gly	Tyr	Pro	Thr	Gly	Thr	Gly	Val
	210					215					220				
Gly	Pro	Gln	Ala	Ala	Ala	Ala	Ala	Ala	Lys	Ala	Ala	Ala	Lys	Phe	
225					230				235					240	
Gly	Ala	Gly	Ala	Ala	Gly	Val	Leu	Pro	Gly	Val	Gly	Gly	Ala	Gly	Val
				245					250					255	
Pro	Gly	Val	Pro	Gly	Ala	Ile	Pro	Gly	Ile	Gly	Gly	Ile	Ala	Gly	Val
		260						265						270	
Gly	Thr	Pro													
		275													

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<210> SEQ ID NO 60
<211> LENGTH: 828
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
        polynucleotide

<400> SEQUENCE: 60
ggtggcgtac caggcgcaat tcctgggggt gtcccaggcg gtgttttta tccgggcgcc      60
ggtcttggcg cactgggtgg cggtgcaact ggcccgggcg gcaaacgct gaaaccggta      120
ccaggtggtt tagcaggcgc cggcttaggc gcaggtctgg gagcatttc ggcagttacc      180
tttccagggg cactggttcc tggaggtgtg gccgatgcag ccgcggcata taaagccgct      240
aaagccggtg cgggtttagg aggcgtccca ggtgtcggtg gcctgggtgt tagcgcgggt      300
gcagttgttc cgcagccggg agcaggggtt aaacctggta aagtgccggg agtaggtctg      360
ccaggcgttt atcctggtgg tgttttgccg ggtgcccgtt ttccgggcgt tggtgttctt      420
ccaggcgctg cgaccgggag cgggtgtaaa ccgaaagccc ccggtgttgg aggtgcattt      480
gcaggcatcc cgggagttgg cccggttggg ggtccgcaac ctggggttcc gttaggttat      540
ccgattaaag caccgaaact gcccgccggt tatggtctgc cgtacacaac cggtaaactg      600
ccgtatggtt atggcccggg tggagttgcg ggtgcagcag gtaaagcggg ttatcctacc      660
ggaaccggtg taggtccgca ggccgctgct gccgcgcccg caaaagcagc ggctaaattt      720
ggcgccggag cagcgggtgt tctgcctgga gttggtggtg cgggcgtgcc aggggtacct      780
ggtgcaattc cgggtattgg tggtattgcc ggtgtcggca ccccgtaa      828
    
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<210> SEQ ID NO 61
 <211> LENGTH: 227
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 61

Gly Gly Val Pro Gly Ala Ile Pro Gly Gly Val Pro Gly Gly Val Phe
 1 5 10 15
 Tyr Pro Gly Ala Gly Leu Gly Ala Leu Gly Gly Gly Ala Leu Gly Pro
 20 25 30
 Gly Gly Lys Pro Leu Lys Pro Val Pro Gly Gly Leu Ala Gly Ala Gly
 35 40 45
 Leu Gly Ala Gly Leu Gly Ala Phe Pro Ala Val Thr Phe Pro Gly Ala
 50 55 60
 Leu Val Pro Gly Gly Val Ala Asp Ala Ala Ala Tyr Lys Ala Ala
 65 70 75 80
 Lys Ala Gly Ala Gly Leu Gly Gly Val Pro Gly Val Gly Gly Leu Gly
 85 90 95
 Val Ser Ala Gly Ala Val Val Pro Gln Pro Gly Ala Gly Val Lys Pro
 100 105 110
 Gly Lys Val Pro Gly Val Gly Leu Pro Gly Val Tyr Pro Gly Gly Val
 115 120 125
 Leu Pro Gly Ala Arg Phe Pro Gly Val Gly Val Leu Pro Gly Val Pro
 130 135 140
 Thr Gly Ala Gly Val Lys Pro Lys Ala Pro Gly Val Gly Gly Ala Phe
 145 150 155 160
 Ala Gly Ile Pro Gly Val Gly Pro Phe Gly Gly Pro Gln Pro Gly Val
 165 170 175
 Pro Leu Gly Tyr Pro Ile Lys Ala Pro Lys Leu Pro Gly Gly Tyr Gly
 180 185 190
 Leu Pro Tyr Thr Thr Gly Lys Leu Pro Tyr Gly Tyr Gly Pro Gly Gly
 195 200 205
 Val Ala Gly Ala Ala Gly Lys Ala Gly Tyr Pro Thr Gly Thr Gly Val
 210 215 220
 Gly Pro Gln
 225

<210> SEQ ID NO 62
 <211> LENGTH: 684
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 62

ggtggcgctac caggcgcaat tcctgggggt gtcccaggcg gtgtttttta tccgggcgcc 60
 ggtcttgccg cactgggtgg cggtgcactg ggcccgggcg gcaaaccgct gaaaccggta 120
 ccagtggtt tagcaggcgc cggcttaggc gcaggtctgg gagcatttcc ggcagttacc 180
 tttccagggg cactggttcc tggaggtgtg gccgatgcag ccgcggcata taaagccgct 240
 aaagccggtg cgggtttagg aggcgtocca ggtgtcggtg gcctgggtgt tagcggcggc 300

-continued

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gcagttgttc cgcagccggg agcaggggtt aaacctggta aagtgccggg agtaggtctg   360
ccaggcggtt atcctggtgg tgttttgcg ggtgcccgtt ttccgggcgt tgggtttctt   420
ccaggcggtc cgaccgggagc cgggtgttaa cgaagagccc ccggtgttgg aggtgcattt   480
gcaggcatcc cgggagtgg cccgtttggt ggtccgcaac ctggggttcc gttaggttat   540
ccgattaaag caccgaaact gcccgcggt tatggtctgc cgtacacaac cggtaaactg   600
ccgtatggtt atggccggg tggagtgcg ggtgcagcag gtaaagcggg ttatcctacc   660
ggaaccggtg taggtccgca gtaa                                     684

```

```

<210> SEQ ID NO 63
<211> LENGTH: 219
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
                             polypeptide

```

```

<400> SEQUENCE: 63

```

```

Gln Leu Arg Ala Ala Ala Gly Leu Gly Ala Gly Ile Pro Gly Leu Gly
 1           5           10          15
Val Gly Val Gly Val Pro Gly Leu Gly Val Gly Ala Gly Val Pro Gly
 20          25          30
Leu Gly Val Gly Ala Gly Val Pro Gly Phe Gly Ala Gly Ala Asp Glu
 35          40          45
Gly Val Arg Arg Ser Leu Ser Pro Glu Leu Arg Glu Gly Asp Pro Ser
 50          55          60
Ser Ser Gln His Leu Pro Ser Thr Pro Ser Ser Pro Arg Val Pro Gly
 65          70          75          80
Ala Leu Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala Val Pro Gly
 85          90          95
Val Leu Gly Gly Leu Gly Ala Leu Gly Gly Val Gly Ile Pro Gly Gly
100         105         110
Val Val Gly Ala Gly Pro Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala
115         120         125
Ala Lys Ala Ala Gln Phe Gly Leu Val Gly Ala Ala Gly Leu Gly Gly
130         135         140
Leu Gly Val Gly Gly Leu Gly Val Pro Gly Val Gly Gly Leu Gly Gly
145         150         155         160
Ile Pro Pro Ala Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala Gly
165         170         175
Leu Gly Gly Val Leu Gly Gly Ala Gly Gln Phe Pro Leu Gly Gly Val
180         185         190
Ala Ala Arg Pro Gly Phe Gly Leu Ser Pro Ile Phe Pro Gly Gly Ala
195         200         205
Cys Leu Gly Lys Ala Cys Gly Arg Lys Arg Lys
210         215

```

```

<210> SEQ ID NO 64
<211> LENGTH: 660
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
                             polynucleotide

```

-continued

<400> SEQUENCE: 64

```

caactgcgcg cgcgccggg cctcggtgca ggtattccgg ggctgggtgt cggagttgga    60
gtcccggggt tggcgtggg cgcgggagtt ccgggactgg gagtgggtgc cggagttcct    120
ggctttggtg caggcgcaga tgaagtggtt cgtcgtagcc tgagtccgga actgcgtgaa    180
ggtgatccga gtagcagcca gcatctgccg agcaccccga gcagcccgcg tgttccgggt    240
gcattagctg cagcaaaagc cgccaagtat ggtgcagccg tgcccggcgt cttaggtggt    300
ctgggcgccc tgggtggtgt aggcattccg ggaggtgttg tgggtgcagg aceggccgcc    360
gcagctgcgg ccgcaaaagc agctgcaaaa gcggcccagt ttggtttagt gggcgccgca    420
ggtttaggcg gtttagtggt ggtgggactg ggtgtacctg gcgtaggcgg tctgggtgga    480
attccgcccc cagcggccgc gaaagcggca aaatatggcg cggcaggcct gggcgcgctg    540
ctgggtgggg caggtcagtt tccgctgggc ggggttgcg cactccggg atttggctctg    600
agcccgattt tccctggcgg cgcattgtctg ggtaaagcat gtggtcgtaa acgtaaataa    660

```

<210> SEQ ID NO 65

<211> LENGTH: 126

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 65

```

Val Pro Gly Val Leu Gly Gly Leu Gly Ala Leu Gly Gly Val Gly Ile
1           5           10          15
Pro Gly Gly Val Val Gly Ala Gly Pro Ala Ala Ala Ala Ala Ala Ala
20          25          30
Lys Ala Ala Ala Lys Ala Ala Gln Phe Gly Leu Val Gly Ala Ala Gly
35          40          45
Leu Gly Gly Leu Gly Val Gly Gly Leu Gly Val Pro Gly Val Gly Gly
50          55          60
Leu Gly Gly Ile Pro Pro Ala Ala Ala Ala Lys Ala Ala Lys Tyr Gly
65          70          75          80
Ala Ala Gly Leu Gly Gly Val Leu Gly Gly Ala Gly Gln Phe Pro Leu
85          90          95
Gly Gly Val Ala Ala Arg Pro Gly Phe Gly Leu Ser Pro Ile Phe Pro
100         105         110
Gly Gly Ala Cys Leu Gly Lys Ala Cys Gly Arg Lys Arg Lys
115         120         125

```

<210> SEQ ID NO 66

<211> LENGTH: 381

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 66

```

gtgcggggcg tcttagtggt tctgggcgcc ctgggtggtg taggcattcc gggaggtgtt    60
gtgggtgcag gaccggccgc cgcagctgcg gccgcaaaag cagctgcaaa agcggcccag    120
tttggttagg tgggcgcccc aggttttaggc ggtttaggtg tgggtggact ggtgtacct    180

```

-continued

ggcgtaggcg gtctgggtgg aattccgccc gcagcggcgg cgaaagcggc aaaatatggc 240
 gcggcaggcc tgggcccggcgt gctgggtggg gcaggtcagt ttccgctggg cgggggttgc 300
 gcacgtccgg gatttggtct gagcccgatt ttccctggcg gcgcatgtct gggtaaagca 360
 tgtggtcgta aacgtaaata a 381

<210> SEQ ID NO 67
 <211> LENGTH: 87
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 67

Gln Phe Gly Leu Val Gly Ala Ala Gly Leu Gly Gly Leu Gly Val Gly
 1 5 10 15
 Gly Leu Gly Val Pro Gly Val Gly Gly Leu Gly Gly Ile Pro Pro Ala
 20 25 30
 Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala Gly Leu Gly Gly Val
 35 40 45
 Leu Gly Gly Ala Gly Gln Phe Pro Leu Gly Gly Val Ala Ala Arg Pro
 50 55 60
 Gly Phe Gly Leu Ser Pro Ile Phe Pro Gly Gly Ala Cys Leu Gly Lys
 65 70 75 80
 Ala Cys Gly Arg Lys Arg Lys
 85

<210> SEQ ID NO 68
 <211> LENGTH: 264
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 68

cagtttggtt tagtggggcg cgcaggttta ggcggtttag gtgtgggtgg actgggtgta 60
 cctggcgtag gcggtctggg tgaattccg cccgcagcgg ccgcaaaagc ggcaaaatat 120
 ggcgcgccag gcctggggcgg cgtgctgggt ggggcaggtc agtttccgct gggcgggggt 180
 gccgcacgtc cgggatttgg tctgagcccg attttccctg gcggcgcgatg tctgggtaaa 240
 gcatgtggtc gtaaacgtaa ataa 264

<210> SEQ ID NO 69
 <211> LENGTH: 72
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 69

Gly Gly Val Pro Gly Ala Ile Pro Gly Gly Val Pro Gly Gly Val Phe
 1 5 10 15
 Tyr Pro Gly Ala Gly Leu Gly Ala Leu Gly Gly Gly Ala Leu Gly Pro
 20 25 30

-continued

Gly Gly Lys Pro Leu Lys Pro Val Pro Gly Gly Leu Ala Gly Ala Gly
 35 40 45
 Leu Gly Ala Gly Leu Gly Ala Phe Pro Ala Val Thr Phe Pro Gly Ala
 50 55 60
 Leu Val Pro Gly Gly Val Ala Asp
 65 70

<210> SEQ ID NO 70
 <211> LENGTH: 219
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 70
 ggtggcgctac caggcgcaat tcctgggggt gtcccaggcg gtgtttttaa tccgggcgcc 60
 ggtcttgagg cactgggtgg cggtgcaact ggcccggggc gcaaaccgct gaaaccggta 120
 ccaggtgggt tagcaggcgc cggcttaggc gcaggtctgg gagcatttcc ggcagttacc 180
 tttccagggg cactggttcc tggaggtgtg gccgattaa 219

<210> SEQ ID NO 71
 <211> LENGTH: 44
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 71
 Gly Leu Gly Gly Val Leu Gly Gly Ala Gly Gln Phe Pro Leu Gly Gly
 1 5 10 15
 Val Ala Ala Arg Pro Gly Phe Gly Leu Ser Pro Ile Phe Pro Gly Gly
 20 25 30
 Ala Cys Leu Gly Lys Ala Cys Gly Arg Lys Arg Lys
 35 40

<210> SEQ ID NO 72
 <211> LENGTH: 135
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 72
 ggccctgggcg gcgtgctggg tggggcaggt cagtttccgc tgggcggggg tgccgcacgt 60
 ccgggatttg gtctgagccc gattttccct ggccggcgcg gtctgggtaa agcatgtggt 120
 cgtaaacgta aataa 135

<210> SEQ ID NO 73
 <211> LENGTH: 636
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 73

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atgaaaaaga tttggctggc gctggctggt ttagttttag cgtttagcgc atcggcgcg 60
cagtatgaag atgcaggttt tccgggtctg cctggtcagg caggcgaacc gggtcgcat 120
ggtaaagatg gtctgatggg tagtccgggt tttaaagggt aagcaggttc accgggtgca 180
cctggtcagg atggcaccgg tgggaaccg ggtattccgg gatttccggg taatcgtggc 240
ctgatgggtc agaaggtga aattgggtccg cctggtcagg agggtaaaaa aggcgcaccg 300
ggtatgccag gactgatggg ttcaaaggc agtccgggtc agccaggcac accgggttca 360
aaagtagca aaggcgaacc tggattcag ggtatgctg gtgcaagcgg tctgaaaggc 420
gagccagggt ccaccgggtc tccgggtgaa ccaggttata tgggtctgcc aggtatccaa 480
ggcaaaaaag gtgataaagg taatcagggc gaaaaaggca ttcagggccca gaaaggcgaa 540
aatggccgtc agggatttcc aggcagcag ggcattccagg gtcattcatgg tgcaaaaggt 600
gaactgggtg aaaagggcga accaggtggt cgtaa 636

```

<210> SEQ ID NO 74

<211> LENGTH: 211

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 74

```

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
1           5           10          15
Ala Ser Ala Ala Gln Tyr Glu Asp Ala Gly Phe Pro Gly Leu Pro Gly
20          25          30
Pro Ala Gly Glu Pro Gly Arg His Gly Lys Asp Gly Leu Met Gly Ser
35          40          45
Pro Gly Phe Lys Gly Glu Ala Gly Ser Pro Gly Ala Pro Gly Gln Asp
50          55          60
Gly Thr Arg Gly Glu Pro Gly Ile Pro Gly Phe Pro Gly Asn Arg Gly
65          70          75          80
Leu Met Gly Gln Lys Gly Glu Ile Gly Pro Pro Gly Gln Gln Gly Lys
85          90          95
Lys Gly Ala Pro Gly Met Pro Gly Leu Met Gly Ser Asn Gly Ser Pro
100         105         110
Gly Gln Pro Gly Thr Pro Gly Ser Lys Gly Ser Lys Gly Glu Pro Gly
115         120         125
Ile Gln Gly Met Pro Gly Ala Ser Gly Leu Lys Gly Glu Pro Gly Ala
130         135         140
Thr Gly Ser Pro Gly Glu Pro Gly Tyr Met Gly Leu Pro Gly Ile Gln
145         150         155         160
Gly Lys Lys Gly Asp Lys Gly Asn Gln Gly Glu Lys Gly Ile Gln Gly
165         170         175
Gln Lys Gly Glu Asn Gly Arg Gln Gly Ile Pro Gly Gln Gln Gly Ile
180         185         190
Gln Gly His His Gly Ala Lys Gly Glu Arg Gly Glu Lys Gly Glu Pro
195         200         205
Gly Val Arg
210

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<210> SEQ ID NO 75
<211> LENGTH: 565
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
      polynucleotide

<400> SEQUENCE: 75

tgcaggtttt cccgggtctgc ctgggtccggc aggcgaaccg ggtegtcatg gtaaagatgg      60
tctgatgggt agtccggggt ttaaagggtga agcaggttca cccgggtgcac ctggtcagga      120
tggcaccctt ggtgaaccgg gtattccggg atttccgggt aatcgtggcc tgatgggtca      180
gaaagggtgaa attggtccgc ctgggtcagca gggtaaaaaa ggcgcaccgg gtatgccagg      240
actgatgggt tcaaatggca gtccgggtca gccaggcaca cccgggttcaa aaggtagcaa      300
aggcgaacct ggtattcagg gtatgcctgg tgcaagcggg ctgaaaggcg agccagggtgc      360
caccggttct cccgggtgaac caggttatat ggggtctgcca ggtatccaag gcaaaaaagg      420
tgataaagggt aatcagggcg aaaaaggcat tcagggccag aaaggcgaaa atggccgtca      480
gggtattcca ggccagcagg gcattccagg tcatcatggt gcaaaagggtg aacgtggtga      540
aaaggcgcaa ccagggtgttc gttaa                                         565

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<210> SEQ ID NO 76
<211> LENGTH: 187
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
      polypeptide

<400> SEQUENCE: 76

Ala Gly Phe Pro Gly Leu Pro Gly Pro Ala Gly Glu Pro Gly Arg His
 1          5          10          15
Gly Lys Asp Gly Leu Met Gly Ser Pro Gly Phe Lys Gly Glu Ala Gly
 20          25          30
Ser Pro Gly Ala Pro Gly Gln Asp Gly Thr Arg Gly Glu Pro Gly Ile
 35          40          45
Pro Gly Phe Pro Gly Asn Arg Gly Leu Met Gly Gln Lys Gly Glu Ile
 50          55          60
Gly Pro Pro Gly Gln Gln Gly Lys Lys Gly Ala Pro Gly Met Pro Gly
 65          70          75          80
Leu Met Gly Ser Asn Gly Ser Pro Gly Gln Pro Gly Thr Pro Gly Ser
 85          90          95
Lys Gly Ser Lys Gly Glu Pro Gly Ile Gln Gly Met Pro Gly Ala Ser
100          105          110
Gly Leu Lys Gly Glu Pro Gly Ala Thr Gly Ser Pro Gly Glu Pro Gly
115          120          125
Tyr Met Gly Leu Pro Gly Ile Gln Gly Lys Lys Gly Asp Lys Gly Asn
130          135          140
Gln Gly Glu Lys Gly Ile Gln Gly Gln Lys Gly Glu Asn Gly Arg Gln
145          150          155          160
Gly Ile Pro Gly Gln Gln Gly Ile Gln Gly His His Gly Ala Lys Gly
165          170          175
Glu Arg Gly Glu Lys Gly Glu Pro Gly Val Arg
180          185

```

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<210> SEQ ID NO 77
 <211> LENGTH: 639
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 77

```

atgaaaaaga tttggctggc gctggctggt ttagtttag cgttagcgc atcggcggcg    60
cagtatgaag atatgggtcc gctcggtagc cgtggtgcaa gtgtccggc aggcgctcgt    120
ggtcggaatg gtgatgcagg tcgtccgggt gaaccgggtc tgatgggtcc tcgtggtctg    180
cctggttcac cgggtaatat tggctctgca ggtaaagaag gtccggttgg tctgccaggt    240
attgatggcc gtccgggtcc gattggtcca gccggtgcac gtggtgaacc tggcaatatt    300
ggttttccgg gtccctaaagg tccgaccggt gatccgggta aaaatggtga taaaggtcat    360
gcaggtctgg caggcgcacg cggtgcacct ggtccggatg gtaataatgg tgcacagggt    420
ccaccgggtc cgcagggtgt tcaaggtggt aaaggcgaac aggtcctgc cggtcctccg    480
ggttttcagg gactgcctgg tccgagcggc cctgcccggg aagttggtaa acctggtgaa    540
cgcggtctgc atggtgaatt tggcctgcct gggcctgcag gtcccggtgg cgaacgtggt    600
ccgccagggtg aaagcgggtgc agcagggtccg acaggttaa    639
    
```

<210> SEQ ID NO 78
 <211> LENGTH: 212
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 78

```

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
1           5           10          15
Ala Ser Ala Ala Gln Tyr Glu Asp Met Gly Pro Pro Gly Ser Arg Gly
20          25          30
Ala Ser Gly Pro Ala Gly Val Arg Gly Pro Asn Gly Asp Ala Gly Arg
35          40          45
Pro Gly Glu Pro Gly Leu Met Gly Pro Arg Gly Leu Pro Gly Ser Pro
50          55          60
Gly Asn Ile Gly Pro Ala Gly Lys Glu Gly Pro Val Gly Leu Pro Gly
65          70          75          80
Ile Asp Gly Arg Pro Gly Pro Ile Gly Pro Ala Gly Ala Arg Gly Glu
85          90          95
Pro Gly Asn Ile Gly Phe Pro Gly Pro Lys Gly Pro Thr Gly Asp Pro
100         105         110
Gly Lys Asn Gly Asp Lys Gly His Ala Gly Leu Ala Gly Ala Arg Gly
115         120         125
Ala Pro Gly Pro Asp Gly Asn Asn Gly Ala Gln Gly Pro Pro Gly Pro
130         135         140
Gln Gly Val Gln Gly Gly Lys Gly Glu Gln Gly Pro Ala Gly Pro Pro
145         150         155         160
Gly Phe Gln Gly Leu Pro Gly Pro Ser Gly Pro Ala Gly Glu Val Gly
    
```


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165 170 175

Lys Pro Gly Glu Arg Gly Leu His Gly Glu Phe Gly Leu Pro Gly Pro
 180 185 190

Ala Gly Pro Arg Gly Glu Arg Gly Pro Pro Gly Glu Ser Gly Ala Ala
 195 200 205

Gly Pro Thr Gly
 210

<210> SEQ ID NO 79
 <211> LENGTH: 567
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 79

atgggtccgc ctggtagccg tgggtcaagt ggtccggcag gcgttcgtgg tccgaatggt 60

gatgcaggtc gtccgggtga accgggtctg atgggtcctc gtggtctgcc tggttcaccg 120

ggtaaatattg gtccctgcagg taaagaaggt ccggttggtc tgccaggat tgatggccgt 180

ccgggtccga ttggtccagc cgggtcacgt ggtgaacctg gcaatattgg ttttccgggt 240

cctaaaggtc cgaccggtga tccgggtaaa aatggtgata aaggtcatgc aggtctggca 300

ggcgcacgcg gtgcacctgg tccggatggt aataatggtg cacagggtcc accgggtccg 360

cagggtgttc aagggtgtaa aggcgaacag ggtcctgccg gtccctccggg ttttcagga 420

ctgcctggtc cgagcgggtcc tgcgggtgaa gttgtaaac ctggtgaacg cggtctgcat 480

ggtgaatttg gcctgcctgg gcctgcaggt ccgcgtggcg aacgtgggtcc gccaggtgaa 540

agcggtgccg caggtccgac aggttaa 567

<210> SEQ ID NO 80
 <211> LENGTH: 188
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 80

Met Gly Pro Pro Gly Ser Arg Gly Ala Ser Gly Pro Ala Gly Val Arg
 1 5 10 15

Gly Pro Asn Gly Asp Ala Gly Arg Pro Gly Glu Pro Gly Leu Met Gly
 20 25 30

Pro Arg Gly Leu Pro Gly Ser Pro Gly Asn Ile Gly Pro Ala Gly Lys
 35 40 45

Glu Gly Pro Val Gly Leu Pro Gly Ile Asp Gly Arg Pro Gly Pro Ile
 50 55 60

Gly Pro Ala Gly Ala Arg Gly Glu Pro Gly Asn Ile Gly Phe Pro Gly
 65 70 75 80

Pro Lys Gly Pro Thr Gly Asp Pro Gly Lys Asn Gly Asp Lys Gly His
 85 90 95

Ala Gly Leu Ala Gly Ala Arg Gly Ala Pro Gly Pro Asp Gly Asn Asn
 100 105 110

Gly Ala Gln Gly Pro Pro Gly Pro Gln Gly Val Gln Gly Gly Lys Gly
 115 120 125

-continued

Glu Gln Gly Pro Ala Gly Pro Pro Gly Phe Gln Gly Leu Pro Gly Pro
 130 135 140

Ser Gly Pro Ala Gly Glu Val Gly Lys Pro Gly Glu Arg Gly Leu His
 145 150 155 160

Gly Glu Phe Gly Leu Pro Gly Pro Ala Gly Pro Arg Gly Glu Arg Gly
 165 170 175

Pro Pro Gly Glu Ser Gly Ala Ala Gly Pro Thr Gly
 180 185

<210> SEQ ID NO 81
 <211> LENGTH: 612
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polynucleotide

<400> SEQUENCE: 81

atgaaaaaga tttggctggc gctggctggt ttagttagcgc atcggcggcg 60
 cagtatgaag atggttttca ggtcctgcc ggtgaaccgg gtgaacctgg tcagacaggt 120
 ccggcaggcg cacgtggtcc tgcaggtcct cctggtaaag ccggtgaaga tggatcaccg 180
 ggtaaacccg gtcgtcctgg tgaacgtggt gttgttggtc cgcagggtgc ccgtggtttt 240
 ccgggtactc cgggtctgcc aggttttaaa ggtattcgtg gtcataatgg tctggatggt 300
 ctgaaaggtc agcctggtgc accgggtggt aaagggtgaac cagggtgctcc gggtgaaaat 360
 ggcacaccgg gtcagaccgg tgcgcgtggt ctgcctggcg aacgcggtcg tgttggtgca 420
 cctggtccag ccgggtgcac cggtagtgat ggtagcgttg gtccggttgg tccagcgggt 480
 ccgattggtg gcgcaggctc accgggtttt ccaggcgcac cgggtccgaa aggtgaaatt 540
 ggtgcagttg gtaatgcagg ccctgccggg ccagcaggac cgcgtggtga agttggcctg 600
 cctggtctgt aa 612

<210> SEQ ID NO 82
 <211> LENGTH: 203
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polypeptide

<400> SEQUENCE: 82

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
 1 5 10 15

Ala Ser Ala Ala Gln Tyr Glu Asp Gly Phe Gln Gly Pro Ala Gly Glu
 20 25 30

Pro Gly Glu Pro Gly Gln Thr Gly Pro Ala Gly Ala Arg Gly Pro Ala
 35 40 45

Gly Pro Pro Gly Lys Ala Gly Glu Asp Gly His Pro Gly Lys Pro Gly
 50 55 60

Arg Pro Gly Glu Arg Gly Val Val Gly Pro Gln Gly Ala Arg Gly Phe
 65 70 75 80

Pro Gly Thr Pro Gly Leu Pro Gly Phe Lys Gly Ile Arg Gly His Asn
 85 90 95

Gly Leu Asp Gly Leu Lys Gly Gln Pro Gly Ala Pro Gly Val Lys Gly

-continued

100	105	110
Glu Pro Gly Ala Pro Gly Glu Asn Gly Thr Pro Gly Gln Thr Gly Ala		
115	120	125
Arg Gly Leu Pro Gly Glu Arg Gly Arg Val Gly Ala Pro Gly Pro Ala		
130	135	140
Gly Ala Arg Gly Ser Asp Gly Ser Val Gly Pro Val Gly Pro Ala Gly		
145	150	155
Pro Ile Gly Ser Ala Gly Pro Pro Gly Phe Pro Gly Ala Pro Gly Pro		
165	170	175
Lys Gly Glu Ile Gly Ala Val Gly Asn Ala Gly Pro Ala Gly Pro Ala		
180	185	190
Gly Pro Arg Gly Glu Val Gly Leu Pro Gly Leu		
195	200	

<210> SEQ ID NO 83
 <211> LENGTH: 540
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 83

```

ggttttcagg gtctctgccg tgaaccgggt gaacctggtc agacagggtcc ggcaggcgca      60
cgtggtcctg caggtcctcc tggtaaagcc ggtgaagatg gtcacccggg taaaccgggt      120
cgtctcggtg aacgtggtgt tgttggtccg cagggtgccc gtggttttcc gggtactccg      180
ggctctgccag gttttaaagg tattcgtggt cataaatggtc tggatggtct gaaaggtcag      240
cctggtgcac cgggtgtaa agtgaaacca ggtgctccgg gtgaaaatgg cacaccgggt      300
cagaccggtg cgcgtggtct gctcggcgaa cgcggtcgtg ttggtgcacc tggteccagcc      360
ggtgcaacgq gtagtgatgg tagcgttggc cgggttggtc cagcgggtcc gattggttagc      420
gcaggtccac cgggttttcc aggcgcaccg ggtccgaaag gtgaaattgg tgcagttggt      480
aatgcaggcc ctgccggtcc agcaggaccg cgtggtgaag ttggcctgcc tggctctgtaa      540
    
```

<210> SEQ ID NO 84
 <211> LENGTH: 179
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 84

Gly Phe Gln Gly Pro Ala Gly Glu Pro Gly Glu Pro Gly Gln Thr Gly		
1	5	10
Pro Ala Gly Ala Arg Gly Pro Ala Gly Pro Pro Gly Lys Ala Gly Glu		
20	25	30
Asp Gly His Pro Gly Lys Pro Gly Arg Pro Gly Glu Arg Gly Val Val		
35	40	45
Gly Pro Gln Gly Ala Arg Gly Phe Pro Gly Thr Pro Gly Leu Pro Gly		
50	55	60
Phe Lys Gly Ile Arg Gly His Asn Gly Leu Asp Gly Leu Lys Gly Gln		
65	70	75
Pro Gly Ala Pro Gly Val Lys Gly Glu Pro Gly Ala Pro Gly Glu Asn		

-continued

	85		90		95	
Gly Thr Pro	Gly Gln Thr	Gly Ala Arg	Gly Leu Pro	Gly Glu Arg	Gly	
	100		105		110	
Arg Val Gly	Ala Pro Gly	Pro Ala Gly	Ala Arg Gly	Ser Asp Gly	Ser	
	115		120		125	
Val Gly Pro	Val Gly Pro	Ala Gly Pro	Ile Gly Ser	Ala Gly Pro	Pro	
	130		135		140	
Gly Phe Pro	Gly Ala Pro	Gly Pro Lys	Gly Glu Ile	Gly Ala Val	Gly	
	145		150		155	160
Asn Ala Gly	Pro Ala Gly	Pro Ala Gly	Pro Arg Gly	Glu Val Gly	Leu	
	165		170		175	
Pro Gly Leu						

<210> SEQ ID NO 85
 <211> LENGTH: 689
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 85

```

gtccgcaggg tgttgttggg gcagatggta aagacggtac cccgggtgaa aaaggagaac    60
agggacgtac aggtgcagca ggtaaacagg gcagcccggg tgccgatggt gccctgggcc    120
cgctgggtag catttggtcag caggggtgcaa gaggcgaacc gggcgatccg ggtagtccgg    180
gcctgcgtgg tgatacgggt ctggcccgtg ttaaaggcgt tgcaggtcct tcaggtcgtc    240
caggtcaacc gggtgcaaat ggtctgccgg gtgttaatgg tcgtggcggg ctggaacgtg    300
gtctggcagg accgccgggt cctgatggtc gccgcgggtg aacgggttca ccgggtattg    360
ccggtgccct gggtaaacca ggtctggaag gtccgaaagg ttatcctggt ctgocgggtc    420
gtgatggtac caatggcaaa cgtggcgaac agggcgaaac cggtcocgat ggtgttcgtg    480
gtattccggg taacgatggt cagagcggta aaccgggcat tgatggtatt gatggcacca    540
atggtcagcc tggcgaagca ggttatcagg gtggtcgcgg taccogtggg cagctgggtg    600
aaacaggtga tgttggtcag aatggtgatc gcgggcacc gggtccggat ggtagcaaa    660
gtagcgcggg tcgtccgggt ttacgttaa    689
    
```

<210> SEQ ID NO 86
 <211> LENGTH: 229
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 86

Gly Pro Gln	Gly Val Val	Gly Ala Asp	Gly Lys Asp	Gly Thr Pro	Gly
1	5		10		15
Glu Lys Gly	Glu Gln Gly	Arg Thr Gly	Ala Ala Gly	Lys Gln Gly	Ser
	20		25		30
Pro Gly Ala	Asp Gly Ala	Arg Gly Pro	Leu Gly Ser	Ile Gly Gln	Gln
	35		40		45
Gly Ala Arg	Gly Glu Pro	Gly Asp Pro	Gly Ser Pro	Gly Leu Arg	Gly
	50		55		60

-continued

Asp Thr Gly Leu Ala Gly Val Lys Gly Val Ala Gly Pro Ser Gly Arg
 65 70 75 80

Pro Gly Gln Pro Gly Ala Asn Gly Leu Pro Gly Val Asn Gly Arg Gly
 85 90 95

Gly Leu Glu Arg Gly Leu Ala Gly Pro Pro Gly Pro Asp Gly Arg Arg
 100 105 110

Gly Glu Thr Gly Ser Pro Gly Ile Ala Gly Ala Leu Gly Lys Pro Gly
 115 120 125

Leu Glu Gly Pro Lys Gly Tyr Pro Gly Leu Arg Gly Arg Asp Gly Thr
 130 135 140

Asn Gly Lys Arg Gly Glu Gln Gly Glu Thr Gly Pro Asp Gly Val Arg
 145 150 155 160

Gly Ile Pro Gly Asn Asp Gly Gln Ser Gly Lys Pro Gly Ile Asp Gly
 165 170 175

Ile Asp Gly Thr Asn Gly Gln Pro Gly Glu Ala Gly Tyr Gln Gly Gly
 180 185 190

Arg Gly Thr Arg Gly Gln Leu Gly Glu Thr Gly Asp Val Gly Gln Asn
 195 200 205

Gly Asp Arg Gly Ala Pro Gly Pro Asp Gly Ser Lys Gly Ser Ala Gly
 210 215 220

Arg Pro Gly Leu Arg
 225

<210> SEQ ID NO 87
 <211> LENGTH: 2283
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polynucleotide

<400> SEQUENCE: 87

ggtggcgtac caggcgcaat tcctgggggt gtcccaggcg gtgttttta tccgggccc 60

ggtcttgccg cactgggtgg cggtgccactg ggcccgggcg gcaaaccgct gaaaccggta 120

ccagtggtt tagcaggcgc cggcttaggc gcaggtctgg gagcatttc gccagttacc 180

tttccagggg cactggttcc tggagtggtg gccgatgcag ccgcggcata taaagccgct 240

aaagccggtg cgggttagg aggcgtccca ggtgtcggtg gcctgggtgt tagcgcgggt 300

gcagttgttc cgcagccggg agcaggggtt aaacctggta aagtgccggg agtaggtctg 360

ccaggcgttt atcctggtgg tgtttgccc ggtgcccgtt ttccgggctg tggtgttctt 420

ccaggcgtgc cgaccggagc cgggttataa ccgaaagccc ccggtgttgg aggtgcattt 480

gcaggcatcc cgggagttgg cccgtttggt ggtccgcaac ctggggttcc gttaggttat 540

ccgattaaag caccgaaact gcccgccggt tatggtctgc cgtacacaac cggtaaactg 600

ccgtatggtt atggcccggg tggagttgcg ggtgcagcag gtaaagcggg ttatcctacc 660

ggaaccggtg taggtccgca ggccgctgct gccgcgccg caaaagcagc ggctaaattt 720

ggcgccggag cagcgggtgt tctgcctgga gttggtggtg cgggcgtgcc aggggtacct 780

ggtgcaattc cgggtattgg tggtattgcc ggtgtcggca ccccgccgc ggcagctgcg 840

gcagcggcgg ctgccaaagc tgctaaatac ggtgcccgcg cgggtctggt gccaggaggt 900

ccgggttttg gtccgggagt ggttggcgtg cctggcgcag gcgttccctg tgtgggcgtt 960

-continued

ccaggtgcag	ggattcctgt	tgtgectggt	gccggtattc	ccggcgcggc	cgttccgggg	1020
gtggttagcc	cggaagccgc	agcgaaggct	gcggcaaagg	cagcaaagta	tggcgcacgc	1080
ccaggagtgc	gcgtgggtgg	tatcccagacc	tatgggggtg	gcgcaggggg	ttttcctggt	1140
ttcggcgtag	gtgtaggagg	tataccgggc	gtggccggtg	taccaggggt	tggtagcgtc	1200
cctggtgttg	gcggtgtgcc	aggtgttgg	atttcaccgg	aagcacaggc	agcagccgca	1260
gctaaggcag	cgaaatatgg	tgccgcggc	gcaggagttt	taggtgggct	ggttccgggg	1320
ccgcaggcag	ctgtgcccgg	ggttccaggc	accggtggtg	tcctggaggt	cggtacgccc	1380
gctgcagcgg	cagccaaagc	ggctgcaaaa	gcagcacagt	ttggcttagt	accgggtgtg	1440
ggagttgccc	ccggcgttgg	cgttgcctca	ggggtgggtg	ttgctcctgg	cgtcggctctg	1500
gctcctggag	tgggcgtagc	acccggtgtg	ggggtggccc	cggtgttgg	ggttgcaccg	1560
ggtatcggtc	cgggcggtgt	gcgagcagca	gctaaaagcg	cggcgaaagt	tcgggcaaaa	1620
gcccaactgc	gcgccgcgcg	gggcctcgg	gcaggtattc	cggggctggg	tgtcggagtt	1680
ggagtccecg	gtttggcgt	gggcgcggga	gttccgggac	tgggagtggt	tgccggagtt	1740
cctggctttg	gtgcaggcgc	agatgaaggt	gttcgctgta	gcctgagttc	ggaactgcgt	1800
gaaggtgatc	cgagtagcag	ccagcatctg	ccgagcacc	cgagcagccc	gcgtgttccc	1860
ggtgcattag	ctgcagcaaa	agccgccaag	tatggtgcag	ccgtgccggg	cgtcttaggt	1920
ggtctggggc	ccctgggtgg	tgtaggaatt	ccgggaggtg	ttgtgggtgc	aggaccggcc	1980
gccgcagctg	cgccgcgcaa	agcagctgca	aaagcggccc	agtttggttt	agtgggcgcc	2040
gcaggtttag	gcggtttagg	tgtgggtgga	ctgggtgtac	ctggcgtagg	cggtctgggt	2100
ggaattccgc	ccgcagcggc	cgcaaaagcg	gcaaaatatg	gcgcggcagg	cctgggcggc	2160
gtgctgggtg	gggcaggtca	gtttccgctg	ggcgggggtg	ccgcacgtcc	gggatttgg	2220
ctgagcccga	ttttccctgg	cggcgcagtgt	ctgggtaaaag	catgtgggtc	taaactgaaa	2280
taa						2283

<210> SEQ ID NO 88
 <211> LENGTH: 760
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 88

Gly Gly Val Pro Gly Ala Ile Pro Gly Gly Val Pro Gly Gly Val Phe
 1 5 10 15

Tyr Pro Gly Ala Gly Leu Gly Ala Leu Gly Gly Gly Ala Leu Gly Pro
 20 25 30

Gly Gly Lys Pro Leu Lys Pro Val Pro Gly Gly Leu Ala Gly Ala Gly
 35 40 45

Leu Gly Ala Gly Leu Gly Ala Phe Pro Ala Val Thr Phe Pro Gly Ala
 50 55 60

Leu Val Pro Gly Gly Val Ala Asp Ala Ala Ala Tyr Lys Ala Ala
 65 70 75 80

Lys Ala Gly Ala Gly Leu Gly Gly Val Pro Gly Val Gly Gly Leu Gly
 85 90 95

-continued

Val	Ser	Ala	Gly	Ala	Val	Val	Pro	Gln	Pro	Gly	Ala	Gly	Val	Lys	Pro
			100					105					110		
Gly	Lys	Val	Pro	Gly	Val	Gly	Leu	Pro	Gly	Val	Tyr	Pro	Gly	Gly	Val
		115					120					125			
Leu	Pro	Gly	Ala	Arg	Phe	Pro	Gly	Val	Gly	Val	Leu	Pro	Gly	Val	Pro
	130					135					140				
Thr	Gly	Ala	Gly	Val	Lys	Pro	Lys	Ala	Pro	Gly	Val	Gly	Gly	Ala	Phe
	145				150					155					160
Ala	Gly	Ile	Pro	Gly	Val	Gly	Pro	Phe	Gly	Gly	Pro	Gln	Pro	Gly	Val
				165					170					175	
Pro	Leu	Gly	Tyr	Pro	Ile	Lys	Ala	Pro	Lys	Leu	Pro	Gly	Gly	Tyr	Gly
			180					185						190	
Leu	Pro	Tyr	Thr	Thr	Gly	Lys	Leu	Pro	Tyr	Gly	Tyr	Gly	Pro	Gly	Gly
		195					200					205			
Val	Ala	Gly	Ala	Ala	Gly	Lys	Ala	Gly	Tyr	Pro	Thr	Gly	Thr	Gly	Val
	210					215					220				
Gly	Pro	Gln	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Lys	Ala	Ala	Ala	Lys	Phe
	225				230					235					240
Gly	Ala	Gly	Ala	Ala	Gly	Val	Leu	Pro	Gly	Val	Gly	Gly	Ala	Gly	Val
				245					250					255	
Pro	Gly	Val	Pro	Gly	Ala	Ile	Pro	Gly	Ile	Gly	Gly	Ile	Ala	Gly	Val
			260					265					270		
Gly	Thr	Pro	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Ala	Lys	Ala	Ala
		275					280					285			
Lys	Tyr	Gly	Ala	Ala	Ala	Gly	Leu	Val	Pro	Gly	Gly	Pro	Gly	Phe	Gly
	290					295					300				
Pro	Gly	Val	Val	Gly	Val	Pro	Gly	Ala	Gly	Val	Pro	Gly	Val	Gly	Val
	305				310					315					320
Pro	Gly	Ala	Gly	Ile	Pro	Val	Val	Pro	Gly	Ala	Gly	Ile	Pro	Gly	Ala
				325					330					335	
Ala	Val	Pro	Gly	Val	Val	Ser	Pro	Glu	Ala	Ala	Ala	Lys	Ala	Ala	Ala
			340					345					350		
Lys	Ala	Ala	Lys	Tyr	Gly	Ala	Arg	Pro	Gly	Val	Gly	Val	Gly	Gly	Ile
		355					360					365			
Pro	Thr	Tyr	Gly	Val	Gly	Ala	Gly	Gly	Phe	Pro	Gly	Phe	Gly	Val	Gly
	370					375					380				
Val	Gly	Gly	Ile	Pro	Gly	Val	Ala	Gly	Val	Pro	Gly	Val	Gly	Gly	Val
	385				390					395					400
Pro	Gly	Val	Gly	Gly	Val	Pro	Gly	Val	Gly	Ile	Ser	Pro	Glu	Ala	Gln
				405					410					415	
Ala	Ala	Ala	Ala	Ala	Lys	Ala	Ala	Lys	Tyr	Gly	Ala	Ala	Gly	Ala	Gly
				420				425					430		
Val	Leu	Gly	Gly	Leu	Val	Pro	Gly	Pro	Gln	Ala	Ala	Val	Pro	Gly	Val
		435					440					445			
Pro	Gly	Thr	Gly	Gly	Val	Pro	Gly	Val	Gly	Thr	Pro	Ala	Ala	Ala	Ala
	450					455					460				
Ala	Lys	Ala	Ala	Ala	Lys	Ala	Ala	Gln	Phe	Gly	Leu	Val	Pro	Gly	Val
	465				470					475					480
Gly	Val	Ala	Pro	Gly	Val	Gly	Val	Ala	Pro	Gly	Val	Gly	Val	Ala	Pro
				485					490					495	
Gly	Val	Gly	Leu	Ala	Pro	Gly	Val	Gly	Val	Ala	Pro	Gly	Val	Gly	Val

-continued

500	505	510
Ala Pro Gly Val Gly Val	Ala Pro Gly Ile Gly	Pro Gly Gly Val Ala
515	520	525
Ala Ala Ala Lys Ser Ala	Ala Lys Val Ala Ala	Lys Ala Gln Leu Arg
530	535	540
Ala Ala Ala Gly Leu Gly	Ala Gly Ile Pro Gly	Leu Gly Val Gly Val
545	550	555
Gly Val Pro Gly Leu Gly	Val Gly Ala Gly Val	Pro Gly Leu Gly Val
565	570	575
Gly Ala Gly Val Pro Gly	Phe Gly Ala Gly Ala	Asp Glu Gly Val Arg
580	585	590
Arg Ser Leu Ser Pro Glu	Leu Arg Glu Gly Asp	Pro Ser Ser Ser Gln
595	600	605
His Leu Pro Ser Thr Pro	Ser Ser Pro Arg Val	Pro Gly Ala Leu Ala
610	615	620
Ala Ala Lys Ala Ala Lys	Tyr Gly Ala Ala Val	Pro Gly Val Leu Gly
625	630	635
Gly Leu Gly Ala Leu Gly	Gly Val Gly Ile Pro	Gly Gly Val Val Gly
645	650	655
Ala Gly Pro Ala Ala Ala	Ala Ala Ala Lys Ala	Ala Ala Lys Ala
660	665	670
Ala Gln Phe Gly Leu Val	Gly Ala Ala Gly Leu	Gly Gly Leu Gly Val
675	680	685
Gly Gly Leu Gly Val Pro	Gly Val Gly Gly Leu	Gly Gly Ile Pro Pro
690	695	700
Ala Ala Ala Ala Lys Ala	Ala Ala Lys Tyr Gly	Ala Ala Gly Leu Gly Gly
705	710	715
Val Leu Gly Gly Ala Gly	Gln Phe Pro Leu Gly	Gly Val Ala Ala Arg
725	730	735
Pro Gly Phe Gly Leu Ser	Pro Ile Phe Pro Gly	Gly Ala Cys Leu Gly
740	745	750
Lys Ala Cys Gly Arg Lys	Arg Lys	
755	760	

<210> SEQ ID NO 89
 <211> LENGTH: 429
 <212> TYPE: PRT
 <213> ORGANISM: Podocoryna carnea

<400> SEQUENCE: 89

Gly Pro Gln Gly Val Val	Gly Ala Asp Gly Lys Asp Gly Thr Pro Gly
1	5 10 15
Glu Lys Gly Glu Gln Gly	Arg Thr Gly Ala Ala Gly Lys Gln Gly Ser
20	25 30
Pro Gly Ala Asp Gly Ala	Arg Gly Pro Leu Gly Ser Ile Gly Gln Gln
35	40 45
Gly Ala Arg Gly Glu Pro	Gly Asp Pro Gly Ser Pro Gly Leu Arg Gly
50	55 60
Asp Thr Gly Leu Ala Gly	Val Lys Gly Val Ala Gly Pro Ser Gly Arg
65	70 75 80
Pro Gly Gln Pro Gly Ala	Asn Gly Leu Pro Gly Val Asn Gly Arg Gly
85	90 95

-continued

Gly Leu Arg Gly Lys Pro Gly Ala Lys Gly Ile Ala Gly Ser Asp Gly
 100 105 110

Glu Ala Gly Glu Ser Gly Ala Pro Gly Gln Ser Gly Pro Thr Gly Pro
 115 120 125

Arg Gly Gln Arg Gly Pro Ser Gly Glu Asp Gly Asn Pro Gly Leu Gln
 130 135 140

Gly Leu Pro Gly Ser Asp Gly Glu Pro Gly Glu Glu Gly Gln Pro Gly
 145 150 155 160

Arg Ser Gly Gln Pro Gly Gln Gln Gly Pro Arg Gly Ser Pro Gly Glu
 165 170 175

Val Gly Pro Arg Gly Ser Lys Gly Pro Ser Gly Asp Arg Gly Asp Arg
 180 185 190

Gly Glu Arg Gly Val Pro Gly Gln Thr Gly Ser Ala Gly Asn Val Gly
 195 200 205

Glu Asp Gly Glu Gln Gly Gly Lys Gly Val Asp Gly Ala Ser Gly Pro
 210 215 220

Ser Gly Ala Leu Gly Ala Arg Gly Pro Pro Gly Ser Arg Gly Asp Thr
 225 230 235 240

Gly Ala Val Gly Pro Pro Gly Pro Thr Gly Arg Ser Gly Leu Pro Gly
 245 250 255

Asn Ala Gly Gln Lys Gly Pro Ser Gly Glu Pro Gly Ser Pro Gly Lys
 260 265 270

Ala Gly Ser Ala Gly Glu Gln Gly Pro Pro Gly Lys Asp Gly Ser Asn
 275 280 285

Gly Glu Pro Gly Ser Pro Gly Lys Glu Gly Glu Arg Gly Leu Ala Gly
 290 295 300

Pro Pro Gly Pro Asp Gly Arg Arg Gly Glu Thr Gly Ser Pro Gly Ile
 305 310 315 320

Ala Gly Ala Leu Gly Lys Pro Gly Leu Glu Gly Pro Lys Gly Tyr Pro
 325 330 335

Gly Leu Arg Gly Arg Asp Gly Thr Asn Gly Lys Arg Gly Glu Gln Gly
 340 345 350

Glu Thr Gly Pro Asp Gly Val Arg Gly Ile Pro Gly Asn Asp Gly Gln
 355 360 365

Ser Gly Lys Pro Gly Ile Asp Gly Ile Asp Gly Thr Asn Gly Gln Pro
 370 375 380

Gly Glu Ala Gly Tyr Gln Gly Gly Arg Gly Thr Arg Gly Gln Leu Gly
 385 390 395 400

Glu Thr Gly Asp Val Gly Gln Asn Gly Asp Arg Gly Ala Pro Gly Pro
 405 410 415

Asp Gly Ser Lys Gly Ser Ala Gly Arg Pro Gly Leu Arg
 420 425

<210> SEQ ID NO 90
 <211> LENGTH: 570
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polynucleotide

<400> SEQUENCE: 90

ggtccgcagg gtgttgttgg tgcagatggt aaagacggta ccccggttaa tgcaggtcag 60

-continued

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aaaggtccgt caggtgaacc tggcagccct ggtaaagcag gtagtgccgg tgagcagggt 120
ccgccgggca aagatggtag taatggtag cgggtagcc ctggcaaaga aggtgaacgt 180
ggtctggcag gaccgccggg tctgatggt cgcgcgggtg aaacgggttc accgggtatt 240
gccggtgccc tgggtaaacc aggtctgga ggtccgaaag gttatcctgg tctgcgcggt 300
cgtgatggta ccaatggcaa acgtggcgaa cagggcgaaa cccgtccaga tgggttctcg 360
ggtattccgg gtaacgatgg tcagagcgg aaaccgggca ttgatggat tgatggcacc 420
aatggtcagc ctggcgaagc aggttatcag ggtggtcgcg gtaccctgg tcagctgggt 480
gaaacaggtg atgttggtca gaatggtgat cgcggcgcac cgggtccgga tggtagcaaa 540
ggtagcgcg gtcgtccggg tttacgtaa 570

```

```

<210> SEQ ID NO 91
<211> LENGTH: 189
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
        polypeptide

```

```

<400> SEQUENCE: 91

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Gly Pro Gln Gly Val Val Gly Ala Asp Gly Lys Asp Gly Thr Pro Gly
1          5          10          15
Asn Ala Gly Gln Lys Gly Pro Ser Gly Glu Pro Gly Ser Pro Gly Lys
20         25         30
Ala Gly Ser Ala Gly Glu Gln Gly Pro Pro Gly Lys Asp Gly Ser Asn
35         40         45
Gly Glu Pro Gly Ser Pro Gly Lys Glu Gly Glu Arg Gly Leu Ala Gly
50         55         60
Pro Pro Gly Pro Asp Gly Arg Arg Gly Glu Thr Gly Ser Pro Gly Ile
65         70         75         80
Ala Gly Ala Leu Gly Lys Pro Gly Leu Glu Gly Pro Lys Gly Tyr Pro
85         90         95
Gly Leu Arg Gly Arg Asp Gly Thr Asn Gly Lys Arg Gly Glu Gln Gly
100        105        110
Glu Thr Gly Pro Asp Gly Val Arg Gly Ile Pro Gly Asn Asp Gly Gln
115        120        125
Ser Gly Lys Pro Gly Ile Asp Gly Ile Asp Gly Thr Asn Gly Gln Pro
130        135        140
Gly Glu Ala Gly Tyr Gln Gly Gly Arg Gly Thr Arg Gly Gln Leu Gly
145        150        155        160
Glu Thr Gly Asp Val Gly Gln Asn Gly Asp Arg Gly Ala Pro Gly Pro
165        170        175
Asp Gly Ser Lys Gly Ser Ala Gly Arg Pro Gly Leu Arg
180        185

```

```

<210> SEQ ID NO 92
<211> LENGTH: 228
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
        polypeptide

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<400> SEQUENCE: 92

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Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
 1 5 10 15

Ala Ser Ala Gly Asp Gln Gly Pro Val Gly Arg Thr Gly Glu Val Gly
 20 25 30

Ala Val Gly Pro Pro Gly Phe Ala Gly Glu Lys Gly Pro Ser Gly Glu
 35 40 45

Ala Gly Thr Ala Gly Pro Pro Gly Thr Pro Gly Pro Gln Gly Leu Leu
 50 55 60

Gly Ala Pro Gly Ile Leu Gly Leu Pro Gly Ser Arg Gly Glu Arg Gly
 65 70 75 80

Leu Pro Gly Val Ala Gly Ala Val Gly Glu Pro Gly Pro Leu Gly Ile
 85 90 95

Ala Gly Pro Pro Gly Ala Arg Gly Pro Pro Gly Ala Val Gly Ser Pro
 100 105 110

Gly Val Asn Gly Ala Pro Gly Glu Ala Gly Arg Asp Gly Asn Pro Gly
 115 120 125

Asn Asp Gly Pro Pro Gly Arg Asp Gly Gln Pro Gly His Lys Gly Glu
 130 135 140

Arg Gly Tyr Pro Gly Asn Ile Gly Pro Val Gly Ala Ala Gly Ala Pro
 145 150 155 160

Gly Pro His Gly Pro Val Gly Pro Ala Gly Lys His Gly Asn Arg Gly
 165 170 175

Glu Thr Gly Pro Ser Gly Pro Val Gly Pro Ala Gly Ala Val Gly Pro
 180 185 190

Arg Gly Pro Ser Gly Pro Gln Gly Ile Arg Gly Asp Lys Gly Glu Pro
 195 200 205

Gly Glu Lys Gly Pro Arg Gly Leu Pro Gly Leu Gly Asp Tyr Lys Asp
 210 215 220

Asp Asp Asp Lys
 225

<210> SEQ ID NO 93
 <211> LENGTH: 687
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polynucleotide

<400> SEQUENCE: 93

```

atgaaaaaga tttggctggc gctggctggt ttagtttag cgtttagcgc atcgcggggt    60
gatcagggtc cggttggtcg taccggtgaa gttggtgcag tcgggccgcc gggttttgcg    120
ggtgaaaaag gcccgtcagg tgaagcaggc accgctggcc ctctggcac gcctggccca    180
cagggtttac tgggcgcacc tggaattctg ggaactgccg gcagccgtgg agaacgcggt    240
ttaccagggt ttgccgtg cgttggtgaa cctggtccac tgggcattgc agggccgcct    300
ggcgcacggg gaccgcctgg tgetgttgg agtccgggtg tgaatggtgc tccgggtgaa    360
gccggtcgtg acgtaatcc gggaaatgac ggcccgccag gccgcgatgg tcagccgggt    420
cataaagggt agcgtggtta cccaggtaat attggtccag tcggtgccgc cggtgccg    480
ggtcctcatg gccctgtcgg tccagccggt aaacatggtg atcgcggtga gacaggtccg    540
tcaggaccag tgggcccctgc tggcgcagtc ggtccgcgcg ggccgagtgg ccctcagggt    600
    
```

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 attcgtggcg ataaagggga accgggcgaa aaagggccgc ggggtctgcc aggctgggt 660

gactacaaag acgacgacga caaataa 687

<210> SEQ ID NO 94

<211> LENGTH: 228

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 94

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
1 5 10 15Ala Ser Ala Lys Gly His Asn Gly Leu Gln Gly Leu Pro Gly Ile Ala
20 25 30Gly His His Gly Asp Gln Gly Ala Pro Gly Ser Val Gly Pro Ala Gly
35 40 45Pro Arg Gly Pro Ala Gly Pro Ser Gly Pro Ala Gly Lys Asp Gly Arg
50 55 60Thr Gly His Pro Gly Thr Val Gly Pro Ala Gly Ile Arg Gly Pro Gln
65 70 75 80Gly His Gln Gly Pro Ala Gly Pro Pro Gly Pro Pro Gly Pro Pro Gly
85 90 95Pro Pro Gly Val Ser Gly Gly Gly Tyr Asp Phe Gly Tyr Asp Gly Asp
100 105 110Phe Tyr Arg Ala Asp Gln Pro Arg Ser Ala Pro Ser Leu Arg Pro Lys
115 120 125Asp Tyr Glu Val Asp Ala Thr Leu Lys Ser Leu Asn Asn Gln Ile Glu
130 135 140Thr Leu Leu Thr Pro Glu Gly Ser Arg Lys Asn Pro Ala Arg Thr Cys
145 150 155 160Arg Asp Leu Arg Leu Ser His Pro Glu Trp Ser Ser Gly Tyr Tyr Trp
165 170 175Ile Asp Pro Asn Gln Gly Cys Thr Met Asp Ala Ile Lys Val Tyr Cys
180 185 190Asp Phe Ser Thr Gly Glu Thr Cys Ile Arg Ala Gln Pro Glu Asn Ile
195 200 205Pro Ala Lys Asn Trp Tyr Arg Ser Ser Lys Asp Gly Asp Tyr Lys Asp
210 215 220Asp Asp Asp Lys
225

<210> SEQ ID NO 95

<211> LENGTH: 687

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 95

atgaaaaaga tttggctggc gctggctgggt ttagtttttag cgtttagcgc atcggcgaaa 60

ggtcacaatg gactgcaagg cctgccaggt attgcaggtc atcatggtga tcaaggtgcc 120

ccgggaagcg ttggtccggc ggggccgaga ggcctgcgg gaccttcagg tccggcaggc 180

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aaagatggtc ggacaggcca tccgggcacc gttggccctg caggaattcg tggaccgcag   240
ggtcacacagg gacctgctgg tccgccaggt cccccgggcc ctccgggacc accgggtgtt   300
agtgggtggg gttatgattt tggctatgat ggtgattttt atcgtgcaga tcagccgcgt   360
agcgcaccga gcctgcgtcc taaagattat gaagttgatg caaccctgaa aagcctgaat   420
aatcagattg aaacactgct gacaccggaa ggtagccgta aaaatccggc ccgtacctgt   480
cgtgatctgc gtctgagcca cccggaatgg agcagcggtt attattggat tgatccgaat   540
caaggttgta ccatggatgc aattaaagtt tattgtgatt ttagcacagg tgaaacatgt   600
atccgtgcac agccggaaaa tattccggcc aaaaattggt atcgtagtag caaagatggt   660
gactacaaag acgacgacga caaataa                                     687

```

<210> SEQ ID NO 96

<211> LENGTH: 262

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 96

```

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
 1                               5                               10                               15
Ala Ser Ala Tyr Glu Val Asp Ala Thr Leu Lys Ser Leu Asn Asn Gln
 20                               25                               30
Ile Glu Thr Leu Leu Thr Pro Glu Gly Ser Arg Lys Asn Pro Ala Arg
 35                               40                               45
Thr Cys Arg Asp Leu Arg Leu Ser His Pro Glu Trp Ser Ser Gly Tyr
 50                               55                               60
Tyr Trp Ile Asp Pro Asn Gln Gly Cys Thr Met Asp Ala Ile Lys Val
 65                               70                               75                               80
Tyr Cys Asp Phe Ser Thr Gly Glu Thr Cys Ile Arg Ala Gln Pro Glu
 85                               90                               95
Asn Ile Pro Ala Lys Asn Trp Tyr Arg Ser Ser Lys Asp Lys Lys His
 100                              105                              110
Val Trp Leu Gly Glu Thr Ile Asn Ala Gly Ser Gln Phe Glu Tyr Asn
 115                              120                              125
Val Glu Gly Val Thr Ser Lys Glu Met Ala Thr Gln Leu Ala Phe Met
 130                              135                              140
Arg Leu Leu Ala Asn Tyr Ala Ser Gln Asn Ile Thr Tyr His Cys Lys
 145                              150                              155                              160
Asn Ser Ile Ala Tyr Met Asp Glu Glu Thr Gly Asn Leu Lys Lys Ala
 165                              170                              175
Val Ile Leu Gln Gly Ser Asn Asp Val Glu Leu Val Ala Glu Gly Asn
 180                              185                              190
Ser Arg Phe Thr Tyr Thr Val Leu Val Asp Gly Cys Ser Lys Lys Thr
 195                              200                              205
Asn Glu Trp Gly Lys Thr Ile Ile Glu Tyr Lys Thr Asn Lys Pro Ser
 210                              215                              220
Arg Leu Pro Phe Leu Asp Ile Ala Pro Leu Asp Ile Gly Gly Ala Asp
 225                              230                              235                              240
Gln Glu Phe Phe Val Asp Ile Gly Pro Val Cys Phe Lys Gly Asp Tyr

```

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	245	250	255	
Lys Asp Asp Asp Asp Lys				
260				
<210> SEQ ID NO 97 <211> LENGTH: 788 <212> TYPE: DNA <213> ORGANISM: Artificial Sequence <220> FEATURE: <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide				
<400> SEQUENCE: 97				
tgaaaaagat ttggctggcg ctggctggtt tagtttttagc gtttagcgca tcggcgatg				60
aagttgatgc aacctgaaa agcctgaata atcagattga aacctgctg acaccggaag				120
gtagccgtaa aaatccggcc cgtacctgtc gtgatctgcg tctgagccac ccggaatgga				180
gcagcggtta ttattggatt gatccgaatc aaggttgtag catggatgca attaaagttt				240
attgtgattt tagcacaggt gaaacatgta tccgtgcaca gccggaaaat attccggcca				300
aaaattggta tcgtagtagc aaagataaaa aacatgtgtg gctgggtgaa accattaatg				360
caggtagcca gtttgaatac aatggtgaag gtgttaccag caaagaaatg gcaacacagc				420
tggcatttat gcgctctgtg gcaaattatg caagccagaa tattacatat cattgtaaaa				480
atagcattgc atatatggat gaagaaaccg gtaatctgaa aaaagcagtt attctgcagg				540
gtagcaatga tgttgaactg gttgccgaag gtaatagccg tttacatat accgttctgg				600
ttgatggtg tagcaaaaa accaatgaat ggggtaaaac catcattgaa tataaaacca				660
acaaaccgag ccgtctgccg tttctggata tcgctccgct ggatattggt ggtgccgatc				720
aggaattttt tgcgatatc ggtcctgtgt gttttaaagg tgactacaaa gacgacgacg				780
acaaataa				788
<210> SEQ ID NO 98 <211> LENGTH: 228 <212> TYPE: PRT <213> ORGANISM: Artificial Sequence <220> FEATURE: <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide				
<400> SEQUENCE: 98				
Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser				
1	5	10	15	
Ala Ser Ala Gly Gly Val Pro Gly Ala Ile Pro Gly Gly Val Pro Gly				
20	25	30		
Gly Val Phe Tyr Pro Gly Ala Gly Leu Gly Ala Leu Gly Gly Gly Ala				
35	40	45		
Leu Gly Pro Gly Gly Lys Pro Leu Lys Pro Val Pro Gly Gly Leu Ala				
50	55	60		
Gly Ala Gly Leu Gly Ala Gly Leu Gly Ala Phe Pro Ala Val Thr Phe				
65	70	75	80	
Pro Gly Ala Leu Val Pro Gly Gly Val Ala Asp Ala Ala Ala Ala Tyr				
85	90	95		
Lys Ala Ala Lys Ala Gly Ala Gly Leu Gly Gly Val Pro Gly Val Gly				
100	105	110		

-continued

Gly Leu Gly Val Ser Ala Gly Ala Val Val Pro Gln Pro Gly Ala Gly
 115 120 125

Val Lys Pro Gly Lys Val Pro Gly Val Gly Leu Pro Gly Val Tyr Pro
 130 135 140

Gly Gly Val Leu Pro Gly Ala Arg Phe Pro Gly Val Gly Val Leu Pro
 145 150 155 160

Gly Val Pro Thr Gly Ala Gly Val Lys Pro Lys Ala Pro Gly Val Gly
 165 170 175

Gly Ala Phe Ala Gly Ile Pro Gly Val Gly Pro Phe Gly Gly Pro Gln
 180 185 190

Pro Gly Val Pro Leu Gly Tyr Pro Ile Lys Ala Pro Lys Leu Pro Gly
 195 200 205

Gly Tyr Gly Leu Pro Tyr Thr Thr Gly Lys Leu Gly Asp Tyr Lys Asp
 210 215 220

Asp Asp Asp Lys
 225

<210> SEQ ID NO 99
 <211> LENGTH: 687
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polynucleotide

<400> SEQUENCE: 99

atgaaaaaga tttggctggc gctggctggt ttagttagcgc atcggcgggt 60
 ggcgtaccag ggcgaattcc tgggggtgtc ccaggcgggtg ttttttatcc gggcgccggt 120
 cttggcgcac tgggtggcgg tgcactgggc ccgggcccga aaccgctgaa accggtacca 180
 ggtggttag caggcgcgg cttaggcgcga ggtctgggag catttccggc agttaacctt 240
 ccaggggcac tggttcctgg aggtgtggcc gatgcagccg cggcatataa agccgctaaa 300
 gccggtgcgg gtttaggagg cgtcccagggt gtcggtggcc tgggtgtag cgccggtgca 360
 gttgttccgc agccgggagc aggggttaaa cctggtaaaag tgccgggaggt aggtctgcca 420
 ggcgtttatc ctggtggtgt tttgccgggt gcccgtttcc cgggcggttg tgttcttcca 480
 ggcgtgccga ccggagccgg tgtaaacccg aaagcccccg gtgttgagg tgcatttgca 540
 ggcaccccgg gagttggccc gtttggtggt ccgcaacctg gggttccggt aggttatccg 600
 attaaagcac cgaactgcc cggcggttat ggtctgccgt acacaaccgg taaactgggt 660
 gactacaag acgacgacga caataa 687

<210> SEQ ID NO 100
 <211> LENGTH: 228
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polypeptide

<400> SEQUENCE: 100

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
 1 5 10 15

Ala Ser Ala Pro Tyr Gly Tyr Gly Pro Gly Gly Val Ala Gly Ala Ala
 20 25 30

-continued

Gly Lys Ala Gly Tyr Pro Thr Gly Thr Gly Val Gly Pro Gln Ala Ala
 35 40 45

Ala Ala Ala Ala Ala Lys Ala Ala Ala Lys Phe Gly Ala Gly Ala Ala
 50 55 60

Gly Val Leu Pro Gly Val Gly Gly Ala Gly Val Pro Gly Val Pro Gly
 65 70 75 80

Ala Ile Pro Gly Ile Gly Gly Ile Ala Gly Val Gly Thr Pro Ala Ala
 85 90 95

Ala Ala Ala Ala Ala Ala Ala Ala Lys Ala Ala Lys Tyr Gly Ala Ala
 100 105 110

Ala Gly Leu Val Pro Gly Gly Pro Gly Phe Gly Pro Gly Val Val Gly
 115 120 125

Val Pro Gly Ala Gly Val Pro Gly Val Gly Val Pro Gly Ala Gly Ile
 130 135 140

Pro Val Val Pro Gly Ala Gly Ile Pro Gly Ala Ala Val Pro Gly Val
 145 150 155 160

Val Ser Pro Glu Ala Ala Ala Lys Ala Ala Ala Lys Ala Ala Lys Tyr
 165 170 175

Gly Ala Arg Pro Gly Val Gly Val Gly Gly Ile Pro Thr Tyr Gly Val
 180 185 190

Gly Ala Gly Gly Phe Pro Gly Phe Gly Val Gly Val Gly Gly Ile Pro
 195 200 205

Gly Val Ala Gly Val Pro Gly Val Gly Gly Val Gly Asp Tyr Lys Asp
 210 215 220

Asp Asp Asp Lys
 225

<210> SEQ ID NO 101
 <211> LENGTH: 687
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 101

```

atgaaaaaga tttggctggc gctggctggt ttagtttag cgtttagcgc atcggcgccc 60
tatggttatg gcccggtggt agttgctggg gcagcaggta aagcgggtta tctaccgga 120
accggtgtag gtccgcaggc cgctgctgcc gccgcgcaa aagcagcggc taaattggc 180
gccggagcag cgggtgttct gectggagtt ggtggtgccc gcgtgccagg ggtacctggt 240
gcaattccgg gtattggtgg tattgcccgt gtcggcacc cggccgcggc agctgcggca 300
gcggcggctg ccaaagctgc taaatacggg gccgcggcgg gtctggtgcc aggaggtccg 360
ggttttggtc cgggagtggt tggcgtgcct ggcgcaggcg ttctggtgt gggcggtcca 420
ggtgcaggga ttctgtgtgt gectggtgcc ggtattcccg gcgcggccgt tccgggggtg 480
gttagccccg aagccgcagc gaaggctgcg gcaaaggcag caaagtatgg cgcacgccc 540
ggagtcggcg tgggtggtat cccgacctat ggggtggcgg cagggggttt tctggtttc 600
ggcgtaggtg taggaggtat accggcgctg gccggtgtac caggggttgg tggcgtcgg 660
gactacaaag acgacgacga caaataa 687
    
```

<210> SEQ ID NO 102

-continued

<211> LENGTH: 273
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 102

```

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
 1          5          10          15
Ala Ser Ala Pro Val Gly Arg Arg Gly Pro Lys Gly Ser Arg Gly Asp
 20          25          30
Pro Gly Asp Gly Gly Ala Ala Gly Pro Lys Gly Pro Glu Gly Val Asp
 35          40          45
Gly Leu Ile Gly Glu Pro Gly Gln Pro Gly Pro Ile Gly Ala Glu Gly
 50          55          60
Ser Ser Gly Leu Glu Gly Phe Leu Gly Asp Lys Gly Ser Lys Gly Ala
 65          70          75          80
Arg Gly Gly Pro Gly Asn Arg Gly Arg Pro Gly Gln Asp Gly Val Pro
 85          90          95
Gly Gln Asp Gly Arg Ala Gly Glu Lys Gly Glu Gly Gly Glu Thr Gly
 100         105         110
Asp Arg Gly Gln Gln Gly Leu Arg Gly Lys Val Gly Asp Pro Gly Leu
 115         120         125
Val Gly Asp Leu Gly Ala Gln Gly Pro Gln Gly Ser Gln Gly Leu Val
 130         135         140
Gly Pro Pro Gly Ile Pro Gly Glu Pro Gly Ser Gly Gly Glu Pro Gly
 145         150         155         160
Asp Gln Gly Pro Arg Gly Pro Glu Gly Pro Gln Gly Ser Pro Gly Val
 165         170         175
Arg Gly Gly Arg Gly Glu Arg Gly Thr Pro Gly Ala Val Gly Pro Lys
 180         185         190
Gly Pro Pro Gly Lys Asn Gly Ala Asp Gly Pro Arg Gly Leu Pro Gly
 195         200         205
Ala Ser Gly Pro Pro Gly Ser Pro Gly Asn Gln Gly Pro Glu Gly Ser
 210         215         220
Arg Gly Ala Asp Gly Asn Asn Gly Phe Pro Gly Asp Asp Gly Glu Asn
 225         230         235         240
Gly Leu Val Gly Ile Pro Gly Glu Pro Gly Pro Lys Gly Ala Arg Gly
 245         250         255
Thr Arg Gly Glu Leu Gly Lys Thr Gly Asp Tyr Lys Asp Asp Asp Asp
 260         265         270
    
```

Lys

<210> SEQ ID NO 103
 <211> LENGTH: 822
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 103

```

atgaaaaaga tttggctggc gctggctggt ttagttagcgc atcggcgccc      60
gttggtcgtc gtggtccgaa aggtagccgt ggtgatcctg gtgatggtgg tgcagcaggt      120
    
```

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cctaaaggtc cggaaagggtg tgatggctcg attggtgaac cgggtcagcc tggcccgatt 180
ggcgcagaag gtagcagcgg tctggaaggt tttctgggtg ataaaggtag caaaggtgca 240
cgtggtggtc cgggtaatcg cggtcgctct ggtcaggatg gtgttccggg tcaagatggt 300
cgtgccggtg aaaaagggtga aggtgggtgaa accgggtgac gcggtcagca gggctctgctg 360
ggtaaagttg gtgatccagg tctggtgggt gatctgggtg cacaggggtcc gcagggtagc 420
caaggtctgg ttggtccgcc tggatttccg ggtgaacctg gtagcggtag cgaaccgggt 480
gatcagggtc ctcgcggtcc agaaggtcct cagggttcac cgggtgttcg cgggtgctg 540
ggtgaacgtg gtacaccggg tgcagttgga ccgaaaggtc cgccaggtaa aaatggtgca 600
gatggtccgc gtggtctgcc tggtgcaagc ggtcctccgg gtagtccctg taaccagggt 660
cctgaaggtt ctcgtggtgc cgatgtaaat aatggttttc caggatgata tggtgaaaat 720
ggcctggttg gtatccctgg cgaaccagggt ccaaaaggcg cacgcggtac acgcggtgaa 780
ctgggtaaaa ccggtgacta caaagacgac gacgacaaat aa 822

```

<210> SEQ ID NO 104

<211> LENGTH: 450

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 104

```

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
1           5           10          15
Ala Ser Ala Gly Arg Gly Gly Pro Ala Gly Leu Gln Gly Ala Ala Gly
20          25          30
Asn Pro Gly Asp Pro Gly Asp Arg Gly Gln Ala Gly Glu Ile Gly Leu
35          40          45
Pro Gly Thr Glu Gly Gln Arg Gly Gln Gly Gly Ser Arg Gly Asp Asp
50          55          60
Gly Ile Gly Gly Gln Ser Gly Thr Asp Gly Asp Pro Gly Asn Asp Gly
65          70          75          80
Val Ala Gly Ile Arg Gly Ala Arg Gly Glu Pro Gly Ala Thr Gly Pro
85          90          95
Glu Gly Ala Ala Gly Gln Lys Gly Asp Arg Gly Arg Phe Gly Glu Gln
100         105         110
Gly Arg Pro Gly Asn Asp Gly Pro Pro Gly Arg Arg Gly Arg Val Gly
115         120         125
Asn Leu Gly Glu Thr Gly Ala Glu Gly Asp Glu Gly Thr Arg Gly Tyr
130         135         140
Thr Gly Asp Arg Gly Pro Glu Gly Ala Ile Gly Ile Ser Gly Val Thr
145         150         155         160
Gly Asn Pro Gly Pro Gln Gly Ile Lys Gly Pro Pro Gly Asp Thr Gly
165         170         175
His Pro Gly Arg Gln Gly Pro Ser Gly Pro Gln Gly Pro Pro Gly Ile
180         185         190
Pro Gly Thr Asp Gly Leu Thr Ile His Asn Leu Ile Lys Pro Pro Ser
195         200         205
Gln Phe Phe Asp Ala Thr Ser Ser Ser Asp Pro Leu Thr Asp Ala Val

```

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210			215			220									
Val	Glu	Ser	Ile	Leu	Lys	Ser	Phe	Gln	Tyr	Ala	Glu	Leu	Glu	Ile	Asp
225					230					235					240
Leu	Thr	Lys	Lys	Pro	Asp	Gly	Thr	Met	Lys	Tyr	Pro	Ala	Ile	Ser	Cys
				245					250					255	
Asp	Asp	Leu	His	Lys	Asp	Tyr	Pro	Gln	Leu	Pro	Ser	Gly	Asn	Tyr	Thr
		260						265					270		
Leu	Asp	Pro	Asn	Gly	Gly	Cys	Lys	Asn	Asp	Ala	Phe	Glu	Thr	Tyr	Cys
		275					280					285			
Glu	Phe	Asn	Asn	Ser	Val	Lys	Met	Cys	Leu	Thr	Pro	Lys	Ile	Pro	Thr
290						295					300				
Leu	Leu	Pro	Met	Gly	Thr	Tyr	Lys	Tyr	Tyr	Val	Asn	Ser	Glu	Gly	Tyr
305					310					315					320
Tyr	Ser	Pro	Asn	Asp	Phe	Gly	Leu	Asn	Leu	Arg	Phe	Phe	Glu	Tyr	Tyr
			325						330					335	
Gly	Ser	Val	Thr	Gln	Leu	Lys	Phe	Leu	Gln	Thr	Lys	Ala	Thr	Arg	Val
		340						345						350	
Thr	Gln	Thr	Ile	Arg	Val	Leu	Cys	Lys	Asn	Tyr	Asp	Pro	Leu	His	Lys
		355					360					365			
Gln	Pro	Val	Phe	Ile	Gly	Met	Asn	Asp	Glu	Thr	Val	Met	Asp	Glu	Pro
370						375					380				
Arg	Met	Glu	Glu	Asn	Gln	Cys	Gln	Tyr	Phe	Asn	Gly	Leu	Ser	Ala	His
385				390						395					400
Val	Glu	Leu	Glu	Leu	Ser	Ser	Asn	Asp	Pro	Ser	Tyr	Leu	Pro	Ile	Tyr
				405					410					415	
Glu	Met	Arg	Leu	Tyr	Leu	Gly	Arg	Lys	Thr	Asn	Glu	Glu	Leu	Gly	Ile
		420						425						430	
Glu	Leu	Gly	Asp	Leu	Cys	Phe	Glu	Tyr	Gly	Asp	Tyr	Lys	Asp	Asp	Asp
		435					440					445			
Asp	Lys														
	450														

<210> SEQ ID NO 105
 <211> LENGTH: 1353
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 105
 atgaaaaaga tttggctggc gctggctggc ttagttagcgc atcggcgggt 60
 cgtggcggtc cggcaggctc gcagggtgct gcaggaatc ctggcgacc tggcgatcgt 120
 ggtcaggcag gcgaaattgg tctgccaggc accgaagtc agcgtggcca aggtggttca 180
 cgtggtgatg acggtattgg tggtcagagc ggcaccgatg gcgatccggg taacgatggt 240
 gttgcaggta ttcgtggtgc acgcgagaaa cctggtgcca ccggacctga aggtgcagcc 300
 ggtcagaaaag gtgatcgtgg ccgttttggc gaacagggtc gtccgggaaa tgatggtcca 360
 ccgggtcgcc gtggccgtgt gggcaatctg ggtgaaacag gtgccgaagg tgatgaagcc 420
 acccgtggtt atacaggatg ccgtggaccg gaaggcgcaa ttggtattag cgtgtgacc 480
 ggtaatccgg gtccacaggg cattaaagcc cctccgggtg atacgggtca tccgggtcgt 540

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cagggaccga gcggtccgca aggaccaccg ggtattccag gtacagatgg cctgaccatt 600
cataatctga ttaaacccgc tagccagttt tttgatgcaa ccagcagcag cgatccgctg 660
accgatgcag ttgttgaaag cattctgaaa tcttttcagt atgccgagct ggaattgac 720
ctgacaaaaa aaccggatgg caccatgaaa tatccggcaa ttagctgtga tgatctgcac 780
aaagattatc cgcagctgcc gagcggtaat tataccctgg atccgaatgg tggttgtaaa 840
aatgatgcct ttgaaaccta ttgcgagttc aacaatagcg tgaaaatgtg tctgaccccg 900
aaaattccga cactgctgcc gatgggcacc tataaatact atgttaatag cgagggttac 960
tacagcccga atgattttgg tctgaatctg cgcttttttg agtattatgg tagcgttacc 1020
cagctgaaat ttctgcagac caaagcaacc cgtgttacc agaccattcg tgttctgtgt 1080
aaaaactatg atccgctgca taaacagccg gtttttattg gtatgaatga cgaaaccggt 1140
atggatgaac cgcgatgga agaaaatcag tgccagtatt ttaacggctc gagcgacat 1200
gttgaactgg aactgagcag caatgatccg agctatctgc cgatttatga aatgcgtctg 1260
tatctgggtc gtaaaaccaa tgaagaactg ggcattgaac tgggcgatct gtgttttgaa 1320
tatggtgact acaaagacga cgacgacaaa taa 1353
    
```

```

<210> SEQ ID NO 106
<211> LENGTH: 286
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
        polypeptide
    
```

<400> SEQUENCE: 106

```

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
1           5           10          15
Ala Ser Ala Glu Lys Thr Ser Ser Lys Val Ala Leu Met Thr Val Leu
20          25          30
Val Val Ile Thr Gly Ala Leu Ile Ile Glu Gly Thr Ser Ile Thr Arg
35          40          45
Gly Ser Thr His Val Asn Arg Gly Leu Arg Lys Arg Gln Thr Ser Glu
50          55          60
Asp Asn Cys Glu Ala Val Lys Val Gly Leu Pro Gly Arg Asp Gly Arg
65          70          75          80
Glu Gly Pro Pro Gly Pro Pro Gly Pro Ala Gly Arg Asp Gly Arg Asp
85          90          95
Ala Val Cys Ser Asn Gln Thr Thr Gly Leu Gly Ala Lys Gly Asp Arg
100         105         110
Gly Pro Pro Gly Thr Pro Gly Phe Pro Gly Glu Val Gly Arg Pro Gly
115         120         125
Pro Pro Gly Ala Asp Gly Ile Pro Gly Pro Gln Gly Glu Arg Gly Ala
130         135         140
Val Gly Pro Gly Gly Lys Pro Gly Pro Arg Gly Glu Val Gly Thr Pro
145         150         155         160
Gly Ala Asp Gly Ala Asp Gly Ala Thr Gly Ala Thr Gly Val Gln Gly
165         170         175
Pro Asp Gly Ala Lys Gly Glu Lys Gly Ala Ser Gly Thr Ala Gly Leu
180         185         190
Lys Gly Glu Lys Gly Asp Thr Cys Ile Pro Asp Ser Asn Ser Thr Leu
    
```

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	195					200						205			
Gly	Met	Pro	Gly	Thr	Pro	Gly	Ala	Gly	Gly	Ser	Lys	Gly	Gln	Lys	Gly
	210					215					220				
Glu	Ser	Gly	Ile	Val	Gly	Pro	Lys	Gly	Glu	Arg	Gly	Glu	Ile	Gly	Thr
225					230					235					240
Pro	Gly	His	Pro	Gly	Phe	Arg	Gly	Ala	Asp	Gly	Glu	Pro	Gly	His	Lys
				245					250					255	
Gly	Val	Pro	Gly	Arg	Ala	Gly	Ala	Gln	Gly	Asp	Arg	Gly	Asp	Pro	Gly
	260						265						270		
Asp	Asp	Gly	Leu	Thr	Gly	Asp	Tyr	Lys	Asp	Asp	Asp	Asp	Lys		
	275						280						285		

<210> SEQ ID NO 107
 <211> LENGTH: 861
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 107

```

atgaaaaaga tttggctggc gctggctggt ttagtttag cgtttagcgc atcgcgcgaa    60
aaaaccagca gcaaagtgc actgatgacc gttctggtt ttattaccgg tgcactgatt    120
attgaaggca ccagcattac ccgtggtagc acccatgta atcgtggtct gcgtaaacgt    180
cagaccagcg aagataattg tgaagcagtt aaagtggctc tgccaggctc tgatggtcgt    240
gaaggtcctc cgggtccgcc tggctccgct ggcagagatg gccgtgatgc agttttagc    300
aatcagacca cgggtctggg tgcaaaaggt gatcgtggtc cgccaggtag accgggtttt    360
cgggtggaag ttggccgtcc ggtgccaccg ggtgcagatg gtattccggg tcctcagggt    420
gaacgtggtg cagttggtcc tggtggtaaa cctggtccgc gtggtgaagt gggcaccct    480
ggtgccgatg ggcagatgg tgcaaccggt gcgaccggtg ttcagggtcc tgatggtgcc    540
aaaggcgaaa aaggtgcaag cggcaccgca ggtctgaaag gtgagaaagg cgatacctgt    600
attccggata gcaatagcac cctgggtatg cctggtacac caggtgccgg tggtagcaaa    660
ggccagaaaag gtgaaagtgg tattgttggc cggaaaggcg aacgcggtga aattggcaca    720
ccgggtcacc ctggttttcg tggtgccgat ggtgaaccag gtcataaagg tgttccgggt    780
cgtgccggtg cgcagggtga tcgcggtgat ccgggtgatg atggtctgac cggtgactac    840
aaagacgacg acgacaaata a                                861
    
```

<210> SEQ ID NO 108
 <211> LENGTH: 512
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 108

Met	Lys	Lys	Ile	Trp	Leu	Ala	Leu	Ala	Gly	Leu	Val	Leu	Ala	Phe	Ser
1			5						10					15	
Ala	Ser	Ala	Gly	Phe	Pro	Gly	Ala	Pro	Gly	Ala	Asp	Gly	Ala	Pro	Gly
		20					25				30				
Gln	Lys	Gly	Glu	Leu	Gly	Ala	Val	Gly	Pro	Gln	Gly	Thr	Pro	Gly	Leu

-continued

	35					40						45					
Ser	Gly	Pro	Ser	Gly	Pro	Thr	Gly	Pro	Pro	Gly	Pro	Lys	Gly	Val	Arg		
	50					55					60						
Gly	Ala	Pro	Gly	Ser	Ser	Gly	Ala	Lys	Gly	Asp	Ala	Gly	Asn	Pro	Gly		
	65				70					75					80		
Asp	Asp	Gly	Pro	Val	Gly	Pro	Gln	Gly	Val	Pro	Gly	Val	Asp	Gly	Ser		
				85					90					95			
Pro	Gly	Gln	Lys	Gly	Glu	Thr	Gly	Arg	Val	Gly	Pro	Arg	Gly	His	Asp		
			100						105					110			
Gly	Ile	Asn	Gly	Thr	Pro	Gly	Glu	Asp	Gly	Ala	Thr	Gly	Phe	Pro	Gly		
			115					120					125				
Pro	Asp	Gly	Ala	Lys	Gly	Glu	Lys	Gly	Thr	Ser	Gly	Thr	Ala	Gly	Leu		
	130					135						140					
Lys	Gly	Glu	Lys	Gly	Asp	Thr	Cys	Ile	Pro	Asp	Ser	Asn	Ser	Thr	Leu		
	145				150					155					160		
Gly	Met	Pro	Gly	Thr	Pro	Gly	Ala	Gly	Trp	Ser	Lys	Gly	Gln	Lys	Gly		
				165					170					175			
Glu	Ser	Gly	Ile	Val	Gly	Pro	Lys	Gly	Glu	Lys	Gly	Glu	Ile	Gly	Thr		
			180						185					190			
Pro	Gly	Pro	Pro	Gly	Phe	Arg	Gly	Ala	Asp	Gly	Glu	Pro	Gly	Gln	Arg		
		195					200						205				
Gly	Glu	Pro	Gly	Arg	Ala	Gly	Ala	Gln	Gly	Glu	Arg	Gly	Ala	Pro	Gly		
	210					215						220					
Asn	Asn	Gly	Arg	Asp	Gly	Phe	Pro	Gly	Asp	Pro	Gly	Ala	Asp	Gly	Ala		
	225				230					235					240		
Pro	Gly	Gln	Lys	Gly	Glu	Leu	Gly	Ala	Ile	Gly	His	Pro	Gly	Phe	Ser		
				245					250					255			
Gly	Pro	Ser	Gly	Pro	Ser	Gly	Pro	Thr	Gly	Pro	Pro	Gly	Pro	Lys	Gly		
			260						265					270			
Val	Arg	Gly	Ala	Gln	Gly	Arg	Pro	Gly	Asp	Arg	Gly	Ser	Pro	Gly	Asp		
		275					280						285				
Val	Gly	Pro	Ile	Gly	Ala	Pro	Gly	Pro	Pro	Gly	Ala	Asp	Gly	Val	Pro		
	290					295						300					
Gly	Leu	Thr	Gly	Val	Gln	Gly	Arg	Asp	Gly	Pro	Lys	Gly	Glu	Ser	Ala		
	305				310					315					320		
Ser	Ser	Gly	Ala	Val	Tyr	Val	Arg	Trp	Gly	Arg	Thr	Thr	Cys	Pro	Ser		
				325					330					335			
Gly	Ala	Asp	Val	Val	Tyr	Ser	Gly	Arg	Ala	Ala	Gly	Ala	Lys	Tyr	Asp		
			340						345					350			
His	Ser	Gly	Gly	Thr	Ser	Asp	His	His	Cys	Leu	Pro	Asn	Asn	Pro	Gln		
		355					360							365			
Tyr	Leu	Ser	Glu	Asp	Asp	Thr	Asn	Ala	Leu	Gly	Ala	Gln	Leu	Tyr	Gly		
	370					375						380					
Val	Glu	Tyr	Glu	Ile	Arg	Asp	Arg	Ser	Ser	Pro	Tyr	Asn	Ser	Leu	Asp		
	385				390					395					400		
Gln	Ser	Asp	Met	Pro	Cys	Val	Val	Cys	Asn	Ala	Asn	Gly	Arg	Ser	Gln		
			405						410					415			
Leu	Leu	Met	Val	Pro	Ala	Arg	Tyr	Thr	Cys	Pro	Thr	Gly	Trp	Ser	Arg		
			420						425					430			
Glu	Tyr	Tyr	Gly	Tyr	Met	Met	Ser	Glu	Gly	Lys	Ala	Lys	Asn	Arg	Glu		
			435					440						445			

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Gly Arg Lys Thr Thr Ile Cys Met Asp Phe Ser Ala Glu Ala Val Pro
 450 455 460

Gly Ser Gly Ala Asn Thr Asn Pro Ser Pro Gly Ile Met Met Arg Ala
 465 470 475 480

Asn Cys Asn Gly Leu Ala Cys Pro Pro Tyr Gln Ser Asn Thr Pro Leu
 485 490 495

Thr Cys Ala Val Cys Thr Lys Gly Asp Tyr Lys Asp Asp Asp Asp Lys
 500 505 510

<210> SEQ ID NO 109

<211> LENGTH: 1539

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 109

```

atgaaaaaga tttggctggc gctggctggt ttagttagcgc atcggcgggt    60
tttctcggcg ctccgggtgc cgacgggtgct cgggtcaaa aaggtgaact gggtgccgtg    120
ggtcgcgagg gcaactccggg tctgagtggt cctagtggtc cgaccgggtcc accaggtcca    180
aaaggcgtgc gtggtgcacc gggtagcagc ggagccaaag gtgatgcagg taaccctggt    240
gatgacggtc cggttggtcc acagggcggt ccaggtggtg atggtagccc tggccaaaag    300
ggtgaaaccg gtcgtgtggg tctcgtggtt catgatggta ttaatggcac cccaggtgaa    360
gatggtgcga caggctttcc aggtccggat ggcgcaaagg gtgagaaggg caccagcggt    420
acagctggcc tgaagggcga aaagggcgat acatgcaccc cggattcaaa ttcaacactg    480
ggcatgccag gtacgcctgg cgcaggttgg agtaaaaggac aaaaaggcga atcaggcatt    540
gtgggacctg aagcgagaa gggtagagatt ggtactccgg gaccgccagg ctttcgcggt    600
gcagacggcg aaccgggtca gcgtggcga cctggtcgtg caggcgcaca aggtgaacgc    660
ggagcccctg gtaataatgg acgtgatggc tttcctgggt atccaggtgc agatggcgca    720
cctggccaga aaggcgaact gggagcaatt ggtcatccgg gatttagcgg tccgtcaggt    780
ccgagcggac cgacaggtcc tctcggaccg aaaggtgtac gtggcgcaca gggtcgtcct    840
ggcgatcgtg gcagtcagg tgatgtgggt ccgattgggt cacctgggtcc tccaggtgcg    900
gacggcgtgc ctggtttaac aggtgtgcag ggtcgcgacg gtcctaaagg tgaatcagca    960
agcagcggtg cagtttatgt tcgttggggt cgtaccacct gtcctagcgg agcagatggt   1020
gtttatagcg gtcgcgcagc cgtgcaaaa tatgatcatt caggtggcac ctcagatcat   1080
cattgtctgc cgaataatcc gcagtatctg agcgaagatg ataccaatgc actgggtgca   1140
cagctgtatg gtgtggaata tgaattcgt gatcgtagca gcccgataa tagcctggat   1200
cagagcgata tgccgtgtgt tgtttgtaat gcaaatggtc gtagccagct gctgatggtt   1260
ccggcacggt atacatgccc gaccggttgg agccgtgaat attatggta tatgatgagc   1320
gaaggcaaa ccaaaaatcg cgaaggtcgt aaaaccacca tttgtatgga ttttagcgca   1380
gaagcagttc ctggtagcgg tgcaaatacc aatccgagtc cgggtattat gatgcgtgca   1440
aattgtaatg gtctggcatg tccgccttat cagagcaata caccgctgac ctgtgccgtt   1500
tgtaccaaag gtgactacaa agacgacgac gacaaataa   1539

```

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<210> SEQ ID NO 110
 <211> LENGTH: 219
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 110

```

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
1           5           10           15
Ala Ser Ala Gly Pro Ala Gly Ala Lys Gly Pro Ser Gly Asp Ile Gly
20           25           30
Arg Pro Gly Glu Ser Gly Ser Pro Gly Ala Arg Gly His Ser Gly Gln
35           40           45
Pro Gly Arg Thr Gly Ile Ala Gly Asn Gln Gly Leu Pro Gly Thr Ala
50           55           60
Gly Glu Glu Gly Arg Thr Gly Pro Pro Gly Pro Ala Gly Leu Arg Gly
65           70           75           80
Gln Ala Gly Met Met Gly Phe Pro Gly Pro Lys Gly Ala Ala Gly Leu
85           90           95
Pro Gly Lys Pro Gly Asp Arg Gly Asn Val Gly Leu Ala Gly Pro Arg
100          105          110
Gly Ala Pro Gly Lys Asp Gly Glu Val Gly Ala Gln Gly Pro Pro Gly
115          120          125
Val Ala Gly Pro Thr Gly Pro Arg Gly Glu Thr Gly Leu Ala Gly Ser
130          135          140
Val Gly Phe Gln Gly Met Pro Gly Pro Ser Gly Ala Ala Gly Glu Pro
145          150          155          160
Gly Lys Pro Gly Asn Gln Gly Leu Arg Gly Asp Ala Gly Ser Pro Gly
165          170          175
Met Ile Gly Pro Arg Gly Glu Arg Gly Leu Pro Gly Glu Arg Gly Ala
180          185          190
Ser Gly Ala Gln Gly Leu Leu Gly Pro Arg Gly Thr Ser Gly Ala Pro
195          200          205
Gly Leu Gly Asp Tyr Lys Asp Asp Asp Asp Lys
210          215

```

<210> SEQ ID NO 111
 <211> LENGTH: 660
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 111

```

atgaaaaaga tttggctggc gctggctggt ttagtttag cgtttagcgc atcggcgggt    60
ccggcaggcg caaaaggctc gagcggtgat attggtcgtc cgggtgaaag cggtagtccg    120
gggtgcacgtg gtcatagcgg tcagcctggt cgtaccggta ttgcaggtaa tcagggtctg    180
cctggtacag ccggtgaaga aggtcgcacc ggtccgccag gtccctgcagg tctgcgtggt    240
caggcaggta tgatgggttt tccgggtccg aaaggtgcag cgggtctgcc aggcaaaccc    300
ggtgatcgtg gtaatgttgg tctggctggt ccgctggttg caccgggtaa agatggtgaa    360

```


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```

gttgggtgcac agggctectcc ggggtgttgea ggtccgaccg gtectcgtgg tgaaacccggt 420
ctggcaggta gcgcttggttt tcagggtatg ccagggtccgt cagggtgcagc aggcgaaacct 480
ggtaaacccgg gtaaccaggg cctgcgtggt gatgcccgtt caccgggtat gattggtcca 540
cgcggtgaac gtggcctgcc tggcgaacgt ggtgcaagcg gtgcacaagg tctgctgggt 600
ccacgtggca cctcaggcgc accaggtctg ggtgactaca aagacgacga cgacaaataa 660

```

```

<210> SEQ ID NO 112
<211> LENGTH: 237
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
        polypeptide

```

```

<400> SEQUENCE: 112

```

```

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
 1                               5                               10                               15
Ala Ser Ala Gln Gly Ile Pro Gly Ser Ala Gly Lys Glu Gly Gly Lys
 20                               25                               30
Gly Asp Pro Gly Pro Leu Gly Ser Pro Gly Lys Pro Gly Pro Asp Gly
 35                               40                               45
Leu Arg Gly Phe Ala Gly Ala Arg Gly Leu Pro Gly Ala Ala Gly Pro
 50                               55                               60
Pro Gly Leu Lys Gly Ala Glu Gly Pro Met Gly Ala Pro Gly Leu Thr
 65                               70                               75                               80
Gly Ser Thr Gly Glu Arg Gly Pro Asn Gly Pro Ala Gly Ala Ile Gly
 85                               90                               95
Leu Pro Gly Arg Pro Gly Gly Pro Gly Pro Pro Gly Pro Val Gly Glu
 100                              105                              110
Lys Gly Asp Pro Gly Asp Lys Gly Leu Pro Gly Pro Ala Gly Asp Asp
 115                              120                              125
Gly Val Gln Gly Ala Met Gly Leu Pro Gly Pro Ile Gly Ser Gln Gly
 130                              135                              140
Pro Pro Gly Asp Tyr Gly Asp Lys Gly Glu Leu Gly Lys Pro Gly Gln
 145                              150                              155                              160
Lys Gly Ser Lys Gly Asp Lys Gly Glu Ser Gly Pro Pro Gly Pro Ile
 165                              170                              175
Gly Ile Gln Gly Pro Ile Gly His Pro Gly Pro Ile Gly Ser Asp Gly
 180                              185                              190
Ser Pro Gly Leu Arg Gly Tyr Leu Gly Met Arg Gly Gln Lys Gly Asp
 195                              200                              205
Asp Gly Ile Arg Gly Leu Pro Gly Ser Ala Gly Pro Val Gly Leu Gln
 210                              215                              220
Gly Leu Pro Gly Gly Asp Tyr Lys Asp Asp Asp Asp Lys
 225                              230                              235

```

```

<210> SEQ ID NO 113
<211> LENGTH: 714
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
        polynucleotide

```

```

<400> SEQUENCE: 113

```

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atgaaaaaga tttggctggc gctggctggt ttagtttag cgtttagcgc atcggcgag	60
ggtattccgg gtagcgcagg taaagaaggt ggtaaaggcg atccgggtcc gctgggttca	120
ccgggtaaac cgggtcctga tggctctgct ggttttgccg gtgcacgtgg tctgcctggt	180
gcagcaggtc cgctgtgtct gaaaggtgcc gaaggtccga tgggtgctcc gggctgacc	240
ggtagcacgg gtgaacggcg tccgaatggt ccggcaggcg caattggtct gccaggtcgt	300
cctggtggtc cgggtcctcc tgggtccggt ggtgaaaaag gtgatcctgg tgataaaggc	360
ctgcctggtc ctgccgtga tgatggtgt cagggtgcca tgggcttacc gggctcgatt	420
ggtagccagg gtcctccggg tgattatggc gataaagggt aactgggtaa acctggccag	480
aaaggtagca aaggtgacaa aggcgaaagc ggtccgccag gtccgatcgg cattcagggt	540
cctattggtc atccaggctc aattggttca gatggctcac cgggactcgg tggctatctg	600
ggtatgcgtg gacagaaaagg tgatgacggt attcgtggcc tgccaggtag tgcaggtccg	660
gtgggtctgc agggactgcc tgggtgtgac tacaagacg acgacgacaa ataa	714

<210> SEQ ID NO 114
 <211> LENGTH: 254
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<400> SEQUENCE: 114

Met Lys Lys Ile Trp Leu Ala Leu Ala Gly Leu Val Leu Ala Phe Ser
 1 5 10 15
 Ala Ser Ala Lys Gly Glu Thr Gly Glu Ala Gly Asp Pro Gly Thr Pro
 20 25 30
 Gly Glu Pro Gly Ile Ala Gly Pro Lys Gly Asp Val Gly Asp Lys Gly
 35 40 45
 Asp Ala Gly Pro Pro Gly Ala Ala Gly Pro Ala Gly Val Lys Gly Pro
 50 55 60
 Pro Gly Glu Asp Gly Ala Lys Gly Asp Val Gly Pro Ala Gly Phe Pro
 65 70 75 80
 Gly Asp Pro Gly Pro Thr Gly Glu Pro Gly Val Pro Gly Met Asp Gly
 85 90 95
 Gly Val Gly Glu Lys Gly Ser Leu Gly Asp Pro Gly Leu Thr Gly Pro
 100 105 110
 Arg Gly Ala Ser Gly Glu Pro Gly Pro Pro Gly Ser Pro Gly Lys Arg
 115 120 125
 Gly Pro Pro Gly Pro Ala Gly Pro Glu Gly Arg Glu Gly Leu Lys Gly
 130 135 140
 Ser Lys Gly Ser Pro Gly Gln Glu Gly Pro Val Gly Arg Thr Gly Pro
 145 150 155 160
 Ile Gly Pro Gln Gly Ser Pro Gly Asn Val Gly Pro Lys Gly Leu Arg
 165 170 175
 Gly Ile Pro Gly Pro Thr Gly Glu Gln Gly Leu Leu Gly Pro Pro Gly
 180 185 190
 Gln Ala Gly Pro Pro Gly Pro Met Gly Pro Pro Gly Met Pro Gly Leu
 195 200 205
 Arg Gly Ala Gln Gly Leu Lys Gly Asp Lys Gly His Val Gly Leu Ile

-continued

210	215	220	
Gly Leu Ile Gly Pro	Pro Gly Glu Met Gly Glu	Lys Gly Asp Gln Gly	
225	230	235	240
Leu Pro Gly Ile Gln Gly Asp Tyr Lys Asp Asp Asp Lys			
	245	250	

<210> SEQ ID NO 115
 <211> LENGTH: 765
 <212> TYPE: DNA
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polynucleotide

<400> SEQUENCE: 115

```

atgaaaaaga tttggctggc gctggctggg ttagtttttag cgtttagcgc atcggcgaaa    60
ggtgaaaccg gtgaagcggg tgatccgggt acaccgggtg aacctgggat tgcaggtccg    120
aaaggtgatg ttggtgataa aggtgacgca ggtccgcctg gtgcagcagg tccggcaggc    180
gttaaaggtc ctccgggtga agatggtgca aaaggcgacg ttggtcctgc aggttttctc    240
ggcgatccgg gtccgactgg tgaaccgggt gtgccaggta tggatggtgg tgtgggtgaa    300
aaaggtagcc tgggtgatcc tggctctgacc ggtccgcctg gcgcaagtgg tgaaccaggt    360
ccaccgggta gtccgggtaa acgtggtcct cctggaccgg ctggtccgga aggtcgtgaa    420
ggtctgaaag gtagcaaagg ttcaccgggt caagaaggtc cggttggtcg taccggtccg    480
attggtccgc agggctcacc gggtaatggt ggtcctaaag gtctgcgtgg tattccgggt    540
cctacaggcg aacagggtct gctgggtccg ccaggccaag caggtcctcc aggtcctatg    600
ggtccacctg gtatgcctgg cctgcgtggt gccaggggcc tgaaggcga taaaggccat    660
gttggcttga ttggcctgat tggtcacca ggtgaaatgg gagaaaaagg cgatcagggc    720
ctgctgggta ttcagggtga ctacaaagac gacgacgaca aataa                    765
    
```

<210> SEQ ID NO 116
 <211> LENGTH: 30
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic His tag
 <220> FEATURE:
 <221> NAME/KEY: MISC_FEATURE
 <222> LOCATION: (1)..(30)
 <223> OTHER INFORMATION: This sequence may encompass 2-30 residues

<400> SEQUENCE: 116

```

His His His His His His His His His His His His His His His His
1          5          10         15

His His His His His His His His His His His His His His His His
20         25         30
    
```

<210> SEQ ID NO 117
 <211> LENGTH: 20
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic His tag
 <220> FEATURE:
 <221> NAME/KEY: MISC_FEATURE

-continued

<222> LOCATION: (1)..(20)
<223> OTHER INFORMATION: This sequence may encompass 2-20 residues

<400> SEQUENCE: 117

His His His His His His His His His His His His His His His His
1 5 10 15

His His His His
 20

<210> SEQ ID NO 118
<211> LENGTH: 15
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
His tag
<220> FEATURE:
<221> NAME/KEY: MISC_FEATURE
<222> LOCATION: (1)..(15)
<223> OTHER INFORMATION: This sequence may encompass 5-15 residues

<400> SEQUENCE: 118

His His His His His His His His His His His His His His His His
1 5 10 15

<210> SEQ ID NO 119
<211> LENGTH: 18
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
His tag
<220> FEATURE:
<221> NAME/KEY: MISC_FEATURE
<222> LOCATION: (1)..(18)
<223> OTHER INFORMATION: This sequence may encompass 5-18 residues

<400> SEQUENCE: 119

His His His His His His His His His His His His His His His His
1 5 10 15

His His

<210> SEQ ID NO 120
<211> LENGTH: 16
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
His tag
<220> FEATURE:
<221> NAME/KEY: MISC_FEATURE
<222> LOCATION: (1)..(16)
<223> OTHER INFORMATION: This sequence may encompass 5-16 residues

<400> SEQUENCE: 120

His His His His His His His His His His His His His His His His
1 5 10 15

<210> SEQ ID NO 121
<211> LENGTH: 14
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
His tag
<220> FEATURE:

-continued

<221> NAME/KEY: MISC_FEATURE
 <222> LOCATION: (1)..(14)
 <223> OTHER INFORMATION: This sequence may encompass 5-14 residues

<400> SEQUENCE: 121

His His His His His His His His His His His His His His His
 1 5 10

<210> SEQ ID NO 122
 <211> LENGTH: 13
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 His tag
 <220> FEATURE:
 <221> NAME/KEY: MISC_FEATURE
 <222> LOCATION: (1)..(13)
 <223> OTHER INFORMATION: This sequence may encompass 5-13 residues

<400> SEQUENCE: 122

His His His His His His His His His His His His His His His
 1 5 10

<210> SEQ ID NO 123
 <211> LENGTH: 12
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 His tag
 <220> FEATURE:
 <221> NAME/KEY: MISC_FEATURE
 <222> LOCATION: (1)..(12)
 <223> OTHER INFORMATION: This sequence may encompass 5-12 residues

<400> SEQUENCE: 123

His His His His His His His His His His His His His His His
 1 5 10

<210> SEQ ID NO 124
 <211> LENGTH: 11
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 His tag
 <220> FEATURE:
 <221> NAME/KEY: MISC_FEATURE
 <222> LOCATION: (1)..(11)
 <223> OTHER INFORMATION: This sequence may encompass 5-11 residues

<400> SEQUENCE: 124

His His His His His His His His His His His His His His His
 1 5 10

<210> SEQ ID NO 125
 <211> LENGTH: 10
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 His tag
 <220> FEATURE:
 <221> NAME/KEY: MISC_FEATURE
 <222> LOCATION: (1)..(10)
 <223> OTHER INFORMATION: This sequence may encompass 5-10 residues

-continued

<400> SEQUENCE: 125

His His His His His His His His His His His
1 5 10

<210> SEQ ID NO 126

<211> LENGTH: 12

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
His tag

<220> FEATURE:

<221> NAME/KEY: MISC_FEATURE

<222> LOCATION: (1)..(12)

<223> OTHER INFORMATION: This sequence may encompass 6-12 residues

<400> SEQUENCE: 126

His His His His His His His His His His His His
1 5 10

<210> SEQ ID NO 127

<211> LENGTH: 11

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
His tag

<220> FEATURE:

<221> NAME/KEY: MISC_FEATURE

<222> LOCATION: (1)..(11)

<223> OTHER INFORMATION: This sequence may encompass 6-11 residues

<400> SEQUENCE: 127

His His His His His His His His His His His
1 5 10

<210> SEQ ID NO 128

<211> LENGTH: 10

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
His tag

<220> FEATURE:

<221> NAME/KEY: MISC_FEATURE

<222> LOCATION: (1)..(10)

<223> OTHER INFORMATION: This sequence may encompass 7-10 residues

<400> SEQUENCE: 128

His His His His His His His His His His His
1 5 10

<210> SEQ ID NO 129

<211> LENGTH: 9

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
9xHis tag

<400> SEQUENCE: 129

His His His His His His His His His His
1 5

<210> SEQ ID NO 130

<211> LENGTH: 150

-continued

<212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide
 <220> FEATURE:
 <221> NAME/KEY: MISC_FEATURE
 <222> LOCATION: (1)..(150)
 <223> OTHER INFORMATION: This sequence may encompass 2-50 "Gly Glu Lys" repeating units

<400> SEQUENCE: 130

```
Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly
1          5          10         15
Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu
20        25        30
Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys
35        40        45
Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly
50        55        60
Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu
65        70        75        80
Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys
85        90        95
Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly
100       105      110
Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu
115      120      125
Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys
130      135      140
Gly Glu Lys Gly Glu Lys
145          150
```

<210> SEQ ID NO 131
 <211> LENGTH: 150
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide
 <220> FEATURE:
 <221> NAME/KEY: MISC_FEATURE
 <222> LOCATION: (1)..(150)
 <223> OTHER INFORMATION: This sequence may encompass 2-50 "Gly Asp Lys" repeating units

<400> SEQUENCE: 131

```
Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly
1          5          10         15
Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp
20        25        30
Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys
35        40        45
Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly
50        55        60
Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp
65        70        75        80
Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys
85        90        95
```

-continued

Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly
 100 105 110
 Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp
 115 120 125
 Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys
 130 135 140
 Gly Asp Lys Gly Asp Lys
 145 150

<210> SEQ ID NO 132
 <211> LENGTH: 120
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polypeptide
 <220> FEATURE:
 <221> NAME/KEY: MISC_FEATURE
 <222> LOCATION: (1)..(120)
 <223> OTHER INFORMATION: This sequence may encompass 2-40 "Gly Glu Lys"
 repeating units

<400> SEQUENCE: 132

Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly
 1 5 10 15
 Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu
 20 25 30
 Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys
 35 40 45
 Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly
 50 55 60
 Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu
 65 70 75 80
 Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys
 85 90 95
 Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly
 100 105 110
 Glu Lys Gly Glu Lys Gly Glu Lys
 115 120

<210> SEQ ID NO 133
 <211> LENGTH: 120
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
 polypeptide
 <220> FEATURE:
 <221> NAME/KEY: MISC_FEATURE
 <222> LOCATION: (1)..(120)
 <223> OTHER INFORMATION: This sequence may encompass 2-40 "Gly Asp Lys"
 repeating units

<400> SEQUENCE: 133

Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly
 1 5 10 15
 Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp
 20 25 30
 Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys

-continued

35	40	45
Gly Asp Lys Gly Asp Lys	Gly Asp Lys Gly Asp Lys	Lys Gly Asp Lys Gly
50	55	60
Asp Lys Gly Asp Lys Gly	Asp Lys Gly Asp Lys	Gly Asp Lys Gly Asp
65	70	75
Lys Gly Asp Lys Gly Asp	Lys Gly Asp Lys Gly Asp	Lys Gly Asp Lys Gly Asp
85	90	95
Gly Asp Lys Gly Asp Lys	Gly Asp Lys Gly Asp Lys	Gly Asp Lys Gly Asp
100	105	110
Asp Lys Gly Asp Lys Gly	Asp Lys	
115	120	

<210> SEQ ID NO 134
 <211> LENGTH: 90
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide
 <220> FEATURE:
 <221> NAME/KEY: MISC_FEATURE
 <222> LOCATION: (1)..(90)
 <223> OTHER INFORMATION: This sequence may encompass 2-30 "Gly Glu Lys" repeating units

<400> SEQUENCE: 134

Gly Glu Lys	Gly Glu Lys	Gly Glu Lys
1	5	10
Gly Glu Lys	Gly Glu Lys	Gly Glu Lys
20	25	30
Lys Gly Glu Lys	Gly Glu Lys	Gly Glu Lys
35	40	45
Gly Glu Lys	Gly Glu Lys	Gly Glu Lys
50	55	60
Gly Glu Lys	Gly Glu Lys	Gly Glu Lys
65	70	75
Lys Gly Glu Lys	Gly Glu Lys	Gly Glu Lys
85	90	

<210> SEQ ID NO 135
 <211> LENGTH: 90
 <212> TYPE: PRT
 <213> ORGANISM: Artificial Sequence
 <220> FEATURE:
 <223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide
 <220> FEATURE:
 <221> NAME/KEY: MISC_FEATURE
 <222> LOCATION: (1)..(90)
 <223> OTHER INFORMATION: This sequence may encompass 2-30 "Gly Asp Lys" repeating units

<400> SEQUENCE: 135

Gly Asp Lys	Gly Asp Lys	Gly Asp Lys
1	5	10
Asp Lys Gly Asp Lys	Gly Asp Lys Gly Asp Lys	Gly Asp Lys Gly Asp
20	25	30
Lys Gly Asp Lys	Gly Asp Lys	Gly Asp Lys
35	40	45

-continued

```

Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly
 50                    55                    60

Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp
65                    70                    75                    80

Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys
      85                    90

```

```

<210> SEQ ID NO 136
<211> LENGTH: 60
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
      polypeptide
<220> FEATURE:
<221> NAME/KEY: MISC_FEATURE
<222> LOCATION: (1)..(60)
<223> OTHER INFORMATION: This sequence may encompass 2-20 "Gly Glu Lys"
      repeating units

```

```

<400> SEQUENCE: 136

```

```

Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly
 1                    5                    10                    15

Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu
      20                    25                    30

Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys
      35                    40                    45

Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys
      50                    55                    60

```

```

<210> SEQ ID NO 137
<211> LENGTH: 60
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
      polypeptide
<220> FEATURE:
<221> NAME/KEY: MISC_FEATURE
<222> LOCATION: (1)..(60)
<223> OTHER INFORMATION: This sequence may encompass 2-20 "Gly Asp Lys"
      repeating units

```

```

<400> SEQUENCE: 137

```

```

Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly
 1                    5                    10                    15

Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp
      20                    25                    30

Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys
      35                    40                    45

Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys
      50                    55                    60

```

```

<210> SEQ ID NO 138
<211> LENGTH: 45
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
      polypeptide
<220> FEATURE:
<221> NAME/KEY: MISC_FEATURE
<222> LOCATION: (1)..(45)

```

-continued

<223> OTHER INFORMATION: This sequence may encompass 2-15 "Gly Glu Lys" repeating units

<400> SEQUENCE: 138

```
Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly
1           5           10          15
Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu
20          25          30
Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys
35          40          45
```

<210> SEQ ID NO 139

<211> LENGTH: 45

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<220> FEATURE:

<221> NAME/KEY: MISC_FEATURE

<222> LOCATION: (1)..(45)

<223> OTHER INFORMATION: This sequence may encompass 2-15 "Gly Asp Lys" repeating units

<400> SEQUENCE: 139

```
Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly
1           5           10          15
Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp
20          25          30
Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys
35          40          45
```

<210> SEQ ID NO 140

<211> LENGTH: 30

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<220> FEATURE:

<221> NAME/KEY: MISC_FEATURE

<222> LOCATION: (1)..(30)

<223> OTHER INFORMATION: This sequence may encompass 2-10 "Gly Glu Lys" repeating units

<400> SEQUENCE: 140

```
Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly
1           5           10          15
Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys
20          25          30
```

<210> SEQ ID NO 141

<211> LENGTH: 30

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic polypeptide

<220> FEATURE:

<221> NAME/KEY: MISC_FEATURE

<222> LOCATION: (1)..(30)

<223> OTHER INFORMATION: This sequence may encompass 2-10 "Gly Asp Lys" repeating units

<400> SEQUENCE: 141

-continued

Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly
1 5 10 15

Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys
20 25 30

<210> SEQ ID NO 142
<211> LENGTH: 27
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic peptide
<220> FEATURE:
<221> NAME/KEY: MISC_FEATURE
<222> LOCATION: (1)..(27)
<223> OTHER INFORMATION: This sequence may encompass 2-9 "Gly Glu Lys" repeating units

<400> SEQUENCE: 142

Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly
1 5 10 15

Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys
20 25

<210> SEQ ID NO 143
<211> LENGTH: 27
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic peptide
<220> FEATURE:
<221> NAME/KEY: MISC_FEATURE
<222> LOCATION: (1)..(27)
<223> OTHER INFORMATION: This sequence may encompass 2-9 "Gly Asp Lys" repeating units

<400> SEQUENCE: 143

Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly
1 5 10 15

Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys
20 25

<210> SEQ ID NO 144
<211> LENGTH: 24
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic peptide
<220> FEATURE:
<221> NAME/KEY: MISC_FEATURE
<222> LOCATION: (1)..(24)
<223> OTHER INFORMATION: This sequence may encompass 2-8 "Gly Glu Lys" repeating units

<400> SEQUENCE: 144

Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly
1 5 10 15

Glu Lys Gly Glu Lys Gly Glu Lys
20

<210> SEQ ID NO 145
<211> LENGTH: 24

-continued

<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic peptide
<220> FEATURE:
<221> NAME/KEY: MISC_FEATURE
<222> LOCATION: (1)..(24)
<223> OTHER INFORMATION: This sequence may encompass 2-8 "Gly Asp Lys" repeating units

<400> SEQUENCE: 145

Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly
1 5 10 15

Asp Lys Gly Asp Lys Gly Asp Lys
20

<210> SEQ ID NO 146
<211> LENGTH: 21
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic peptide
<220> FEATURE:
<221> NAME/KEY: MISC_FEATURE
<222> LOCATION: (1)..(21)
<223> OTHER INFORMATION: This sequence may encompass 2-7 "Gly Glu Lys" repeating units

<400> SEQUENCE: 146

Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly
1 5 10 15

Glu Lys Gly Glu Lys
20

<210> SEQ ID NO 147
<211> LENGTH: 21
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic peptide
<220> FEATURE:
<221> NAME/KEY: MISC_FEATURE
<222> LOCATION: (1)..(21)
<223> OTHER INFORMATION: This sequence may encompass 2-7 "Gly Asp Lys" repeating units

<400> SEQUENCE: 147

Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly
1 5 10 15

Asp Lys Gly Asp Lys
20

<210> SEQ ID NO 148
<211> LENGTH: 18
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic peptide
<220> FEATURE:
<221> NAME/KEY: MISC_FEATURE
<222> LOCATION: (1)..(18)
<223> OTHER INFORMATION: This sequence may encompass 2-6 "Gly Glu Lys" repeating units

-continued

<400> SEQUENCE: 148

Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly
1 5 10 15

Glu Lys

<210> SEQ ID NO 149

<211> LENGTH: 18

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
peptide

<220> FEATURE:

<221> NAME/KEY: MISC_FEATURE

<222> LOCATION: (1)..(18)

<223> OTHER INFORMATION: This sequence may encompass 2-6 "Gly Asp Lys"
repeating units

<400> SEQUENCE: 149

Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly
1 5 10 15

Asp Lys

<210> SEQ ID NO 150

<211> LENGTH: 15

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
peptide

<220> FEATURE:

<221> NAME/KEY: MISC_FEATURE

<222> LOCATION: (1)..(15)

<223> OTHER INFORMATION: This sequence may encompass 2-5 "Gly Glu Lys"
repeating units

<400> SEQUENCE: 150

Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys
1 5 10 15

<210> SEQ ID NO 151

<211> LENGTH: 15

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
peptide

<220> FEATURE:

<221> NAME/KEY: MISC_FEATURE

<222> LOCATION: (1)..(15)

<223> OTHER INFORMATION: This sequence may encompass 2-5 "Gly Asp Lys"
repeating units

<400> SEQUENCE: 151

Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys
1 5 10 15

<210> SEQ ID NO 152

<211> LENGTH: 12

<212> TYPE: PRT

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
peptide

<220> FEATURE:

-continued

```

<221> NAME/KEY: MISC_FEATURE
<222> LOCATION: (1)..(12)
<223> OTHER INFORMATION: This sequence may encompass 2-4 "Gly Glu Lys"
    repeating units

<400> SEQUENCE: 152

Gly Glu Lys Gly Glu Lys Gly Glu Lys Gly Glu Lys
1             5             10

<210> SEQ ID NO 153
<211> LENGTH: 12
<212> TYPE: PRT
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: Description of Artificial Sequence: Synthetic
    peptide
<220> FEATURE:
<221> NAME/KEY: MISC_FEATURE
<222> LOCATION: (1)..(12)
<223> OTHER INFORMATION: This sequence may encompass 2-4 "Gly Asp Lys"
    repeating units

<400> SEQUENCE: 153

Gly Asp Lys Gly Asp Lys Gly Asp Lys Gly Asp Lys
1             5             10

```

What is claimed is:

1. A recombinant cell comprising at least one copy of a heterologous nucleic acid sequence encoding a recombinant polypeptide comprising an amino acid sequence according to a human type 21 alpha 1 collagen, or a fragment thereof.

2. The recombinant cell of claim 1, wherein the recombinant cell is a microbial cell.

3. The recombinant cell of claim 2, wherein the microbial cell is a bacterial cell.

4. The recombinant cell of claim 3, wherein the bacterial cell is of the species *Escherichia coli*.

5. The recombinant cell of claim 1, wherein the heterologous nucleic acid sequence comprises SEQ ID NO: 75.

6. The recombinant cell of claim 1, wherein the heterologous nucleic acid sequence is codon-optimized for expression in a cell.

7. The recombinant cell of claim 1, wherein the recombinant polypeptide is full-length human type 21 alpha 1 collagen.

8. The recombinant cell of claim 1, wherein the recombinant polypeptide is non-naturally occurring.

9. The recombinant cell of claim 1, wherein the recombinant polypeptide comprises an amino acid sequence comprising a truncation relative to a full-length human type 21 alpha 1 collagen.

10. The recombinant cell of claim 9, wherein the truncation comprises a truncation at an N-terminal end relative to a full-length human type 21 alpha 1 collagen, a truncation at a C-terminal end relative to a full-length human type 21 alpha 1 collagen, an internal truncation relative to a full-length human type 21 alpha 1 collagen, or any combination thereof.

11. The recombinant cell of claim 9, wherein the truncation comprises a truncation of between 50 amino acids and 800 amino acids relative to a full-length human type 21 alpha 1 collagen.

12. The recombinant cell of claim 9, wherein the truncation comprises a truncation at an N-terminal end relative to a full-length human type 21 alpha 1 collagen, and a truncation at a C-terminal end relative to a full-length human type 21 alpha 1 collagen.

13. The recombinant cell of claim 12, wherein the truncation at an N-terminal end relative to a full-length human type 21 alpha 1 collagen is a truncation of between 50 amino acids and 600 amino acids.

14. The recombinant cell of claim 12, wherein the truncation at a C-terminal end relative to a full-length human type 21 alpha 1 collagen is a truncation of between 50 amino acids and 250 amino acids.

15. The recombinant cell of claim 9, wherein the truncation is a truncation at an N-terminal end of between 50 amino acids and 600 amino acids relative to a full-length human type 21 alpha 1 collagen, and a truncation at a C-terminal end of between 50 amino acids and 250 amino acids relative to a full-length human type 21 alpha 1 collagen.

16. The recombinant cell of claim 1, wherein the recombinant polypeptide comprises an amino acid sequence according to SEQ ID NO: 76 or a fragment thereof.

17. The recombinant cell of claim 1, wherein the recombinant polypeptide consists of an amino acid sequence according to SEQ ID NO: 76.

18. The recombinant cell of claim 1, wherein the recombinant polypeptide further comprises a secretion tag.

19. The recombinant cell of claim 18, wherein the secretion tag is a DsbA secretion tag.

20. The recombinant cell of claim 18, wherein the recombinant cell is capable of secreting the recombinant polypeptide extracellularly.

21. A composition comprising a recombinant cell of claim 1, and a culture media comprising a recombinant polypeptide comprising an amino acid sequence according to a human type 21 alpha 1 collagen, or a fragment thereof.

22. The composition of claim **21**, wherein the recombinant cell is a bacterial cell of the species *Escherichia coli*.

23. The composition of claim **22**, wherein the heterologous nucleic acid sequence is a nucleic acid sequence according to SEQ ID NO: 75, and the recombinant polypeptide has an amino acid sequence according to SEQ ID NO: 76.

24. A method for producing a recombinant polypeptide, the method comprising:

- a. incubating a recombinant cell of claim **1** in a culture media wherein the recombinant cell secretes the recombinant polypeptide into the culture media;
- b. collecting the culture media comprising the recombinant polypeptide secreted thereto; and
- c. purifying the recombinant polypeptide from the culture media.

25. The method of claim **24**, wherein the recombinant cell is a bacterial cell of the species *Escherichia coli*.

26. The method of claim **25**, wherein the heterologous nucleic acid sequence is a nucleic acid sequence according to SEQ ID NO: 75, and the recombinant polypeptide has an amino acid sequence according to SEQ ID NO: 76.

* * * * *