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- (54) **SECRETED AND TRANSMEMBRANE POLYPEPTIDES AND NUCLEIC ACIDS ENCODING THE SAME**
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(57) **ABSTRACT**

The present invention is directed to novel polypeptides and to nucleic acid molecules encoding those polypeptides. Also provided herein are vectors and host cells comprising those nucleic acid sequences, chimeric polypeptide molecules comprising the polypeptides of the present invention fused to heterologous polypeptide sequences, antibodies which bind to the polypeptides of the present invention and to methods for producing the polypeptides of the present invention.

GGGGCTTCGGCGCCAGCGGCCAGCGCTAGTCGGTCTGGTAAGGATTTACAAAAGGTGCAGGTATG
AGCAGGTCTGAGACTAACATTTTGTGAAGTTGTAAACAGAAAACCTGTTAGAAATGGTGGT
TTCAGCAAGGCTCAGTTTCCCTTCCTTCAGCCCTTGTAATTTGGACATCTGCTGCTTCATATTT
TCATACATTACTGCAGTAACACTCCACCATATAGACCCGGCTTTACCTTATATCAGTGACACTGG
TACAGTAGCTCCAGAAAAATGCTTATTTGGGGCAATGCTAATATTCGGCCAGTTTATGCATTG
CTACCATTTATGTTTCGTTATAGCAAGTTTCATGCTCTGAGTCCCTGAAGAGAACGTTATCATCAA
TTAAACAGGCTGGCCTTGACTTGAATACTGAGTTGTTAGGACTTTCCTATTGTGGCAAACTT
CCAGAAAACACCCCTTTTGTGTCACATGTAAAGTGAGCTGTGCTTACCTTTGGTATGGGCTCAT
TATATATGTTTGTTCAGACCATCCTTCTCACCAAATGCAGCCCAAAATCCATGGCAACAGTC
TTCCTGGATCAGACTGTTGTTGTTATCTGGGTGGAGTAAGTGCACTAGCATGCTGACTTGCTC
ATCAGTTTTCAGACTGGCAATTTTGGGACTGATTTAGAACAGAACTCCATTGGAAACCCCGAGG
ACAAAGGTTATGTGCTTCACATGATCACTACTGCAGCAGATGGTCTATGTCATTTTCTCTTCTT
GGTTTTTCTCTGACTTACATTCGTGATTTTCAGAAAATTTCTTTACGGGTGGAAGCCAAATTACA
TGGATTACCCCTCTATGACACTGCACCTTGGCCCTATTAAAGTAAGCAACGAGCTACTTTCCA
GAGATATTGATGAAGGATAAAATATTCTGTAATGATTATGATCTCAGGGATTGGGGAAGG
TTCACAGAGTTGCTTATTTCTCTGAAAATTTCAACCACTTAATCAAGGCTGACAGTAACACT
GATGATGCTGATAATCAGGAACATGAAGAAGCCATTGATAGATTATCTAAAGGATATCAT
CAAGAGACTATTAAAAACACTATGCCTATCTTTTATCTCAGAAAATAAGTCAAAAGACT
ATG

Related U.S. Application Data

142, filed on Aug. 25, 1999, now abandoned, which is a continuation of application No. 09/380,138, filed on Aug. 25, 1999, now abandoned, which is a continuation of application No. 09/380,137, which is a continuation of application No. 09/311,832, filed on May 14, 1999, which is a continuation of application No. 09/380,139, filed on Aug. 25, 1999, now abandoned.

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May 22, 2000	(WO)	PCT/US00/14042

FIGURE 1

GGGGCTTCGGCGCCAGCGGCCAGCGCTAGTCGGTCTGGTAAGGATTTACAAAAGGTGCAGGTATG
AGCAGGTCTGAAGACTAACATTTTGTGAAGTTGTAAAACAGAAAACCTGTTAGAAATGTGGTGGT
TTCAGCAAGGCCTCAGTTTCCTTCCTTCAGCCCTTGTAATTTGGACATCTGCTGCTTTCATATTT
TCATACATTACTGCAGTAACACTCCACCATATAGACCCGGCTTTACCTTATATCAGTGACACTGG
TACAGTAGCTCCAGAAAAATGCTTATTTGGGGCAATGCTAAATATTGCGGCAGTTTTATGCATTG
CTACCATTTATGTTTCGTTATAAGCAAGTTCATGCTCTGAGTCCTGAAGAGAACGTTATCATCAAA
TTAAACAAGGCTGGCCTTGTACTTGAATACTGAGTTGTTTAGGACTTTCTATTGTGGCAAACCTT
CCAGAAAACAACCCCTTTTTGCTGCACATGTAAGTGGAGCTGTGCTTACCTTTGGTATGGGCTCAT
TATATATGTTTGTTCAGACCATCCTTTCCTACCAAATGCAGCCCCAAAATCCATGGCAAACAAGTC
TTCTGGATCAGACTGTTGTTGGTTATCTGGTGTGGAGTAAGTGCACTTAGCATGCTGACTTGCTC
ATCAGTTTTGCACAGTGGCAATTTTGGGACTGATTTAGAACAGAACTCCATTGGAACCCCGAGG
ACAAAGGTTATGTGCTTCACATGATCACTACTGCAGCAGAATGGTCTATGTCATTTTCCTTCTTT
GGTTTTTTCCTGACTTACATTTCGTGATTTTCAGAAAATTTCTTTACGGGTGGAAGCCAATTTACA
TGGATTAACCTCTATGACACTGCACCTTGCCCTATTAACAATGAACGAACACGGCTACTTTCCA
GAGATATTTTGATGAAAGGATAAAAATATTTCTGTAATGATTATGATTCTCAGGGATTGGGGAAAGG
TTCACAGAAGTTGCTTATTCTTCTCTGAAATTTTCAACCACTTAATCAAGGCTGACAGTAACACT
GATGAATGCTGATAATCAGGAAACATGAAAGAAGCCATTTGATAGATTATTCTAAAGGATATCAT
CAAGAAGACTATTAAAAACACCTATGCCTATACTTTTTTATCTCAGAAAATAAAGTCAAAAAGACT
ATG

FIGURE 2

<subunit 1 of 1, 266 aa, 1 stop

<MW: 29766, pI: 8.39, NX(S/T): 0

MWWFQQGLSFLPSALVIWTSAAFIIFS YITAVTLHHIDPALPYISDTGTVAPEKCLFGAMLNIAAV
LCIATIIYVRYKQVHALSPEENVIIKLNKAGLVLGILSCLGLSIVANFQKTTLFAAHVSGAVLTFG
MGSLYMFVQTILSYQMOPKIHGKQVFWIRLLLVICGVSALSMLTCSSVLHSGNFGTDLEQKLHW
NPEDKGYVLHMITTAAEWSMSFSFFGFFLT YIRDFQKISLRVEANLHGLTLYDTAPCPINNERTR
LLSRDI

Important features:

Type II transmembrane domain:

amino acids 13-33

Other Transmembrane domains:

amino acids 54-73, 94-113, 160-180, 122-141

N-myristoylation sites.

amino acids 57-63, 95-101, 99-105, 124-130, 183-189

FIGURE 3

CGGACGCGTGGGCGGACGCGTGGGGGAGAGCCGCGAGTCCCGGCTGCAGCACCTGGGAGAAGGCAGACC
 GTGTGAGGGGGCCTGTGGCCCCAGCGTGCTGTGGCCTCGGGGAGTGGGAAGTGGAGGCAGGAGCCTTC
 CTTACACTTCGCCATGAGTTTCCTCATCGACTCCAGCATCATGATTACCTCCCAGATACTATTTTTTG
 GATTTGGGTGGCTTTTCTTCATGCGCCAATTGTTTAAAGACTATGAGATACGTCAGTATGTTGTACAG
 GTGATCTTCTCCGTGACGTTTGCAATTTCTTGCAACCATGTTTGAGCTCATCATCTTTGAAATCTTAGG
 AGTATTGAATAGCAGCTCCCGTTATTTTCACTGGAAAATGAACCTGTGTGTAATTCTGCTGATCCTGG
 TTTTCATGGTGCCTTTTACATTGGCTATTTTATTGTGAGCAATATCCGACTACTGCATAAACAACGA
 CTGCTTTTTTCTGTCTCTTATGGCTGACCTTTATGTATTTCTTCGAAAAGTGGAGATCCCTTTCC
 CATTCTCAGCCCCAAAACATGGGATCTTATCCATAGAACAGCTCATCAGCCGGGTGGTGTGATTGGAG
 TGACTCTCATGGCTCTTCTTCTGGATTGGTGCTGTCAACTGCCCATACACTTACATGTCTTACTTC
 CTCAGGAATGTGACTGACACGGATATTCTAGCCCTGGAACGGCGACTGCTGCAACCATGGATATGAT
 CATAAGCAAAAAGAAAAGGATGGCAATGGCACGGAGAACAATGTTCCAGAAGGGGGAAGTGCATAACA
 AACCATCAGGTTTCTGGGGAATGATAAAAAGTGTACCCTTCAGCATCAGGAAGTGAATCTTACT
 CTTATTCAACAGGAAGTGGATGCTTTGGAAGAATTAAGCAGGCAGCTTTTTCTGGAACAGCTGATCT
 ATATGCTACCAAGGAGAGAATAGAATACTCCAAAACCTTCAAGGGGAAATATTTTAATTTTCTTGGTT
 ACTTTTTCTCTATTTACTGTGTTTGGAAAATTTTCATGGCTACCATCAATATTGTTTTTGATCGAGTT
 GGGAAAACGGATCCTGTCAAGAGGCATTGAGATCACTGTGAATTATCTGGGAATCCAATTTGATGT
 GAAGTTTTGGTCCCAACACATTTCTTCTTCTGTTGGAATAATCATCGTCACATCCATCAGAGGAT
 TGCTGATCACTCTTACCAAGTTCTTTTATGCCATCTCTAGCAGTAAGTCTCCAATGTCATTGTCCTG
 CTATTAGCACAGATAAATGGGCATGTACTTTGTCTCCTCTGTGCTGCTGATCCGAATGAGTATGCCTTT
 AGAATACCGCACCATAATCACTGAAGTCCTTGGGAGAACTGCAGTTCAACTTCTATCACCGTTGGTTTG
 ATGTGATCTTCTGGTCAGCGCTCTCTCTAGCATACTCTTCTCTATTTGGCTCACAACAGGCACCA
 GAGAAGCAAATGGCACCTTGAACTTAAGCCTACTACAGACTGTTAGAGGCCAGTGGTTTTCAAAATTTA
 GATATAAGAGGGGGGAAAAATGGAACCAGGGCCTGACATTTTATAAACAACAAAATGCTATGGTAGC
 ATTTTTCACCTTCATAGCATACTCCTTCCCCGTGAGGTGATACTATGACCATGAGTAGCATCAGCCAG
 AACATGAGAGGGGAGAACTCAAGACAATACTCAGCAGAGAGCATCCCGTGTGGATATGAGGCTGG
 GTAGAGGCGGAGAGGAGCCAAAGAACTAAAGGTGAAAAATACACTGGAACCTCTGGGGCAAGACATGT
 CTATGGTAGCTGAGCCAAACACGTAGGATTTCCGTTTTTAAGGTTACATGGAAAAGGTTATAGCTTTG
 CCTTGAGATTGACTCATTAATAATCAGAGACTGTAACAAAAAAAAAAAAAAAAAAAAAGGGCGGCCGCG
 ACTCTAGAGTCGACCTGCAGAAGCTTGCCGCCATGGCCCAACTTGTTTATTGCAGCTTATAATG

FIGURE 4

MSFLIDSSIMITSQILFFGFGWLFFMRQLFKDYEIRQYVQVIFSVTFAFSCTMFELIIFEILGV
LNSSSRYPFWKMNLCVILLILVFMVPFYIGYFIVSNIRLLHKQRLLFSCLLWLTFMYFFWKLGD
FPILSPKHGILSIEQLISRVGIVGVTLMALLSGFGAVNCPYTYMSYFLRNVTDTDILALERRLLQ
TMDMIISKKKRMAMARRTMFQKGEVHNKPSGFWGMIKSVTTSASGSENLTLIQQEVDAL EELSRQ
LFLETADLYATKERIEYSKTFKGKYFNFLGYFFSIYCVWKIFMATINIVFDRVGKTDPVTRGIEI
TVNYLGIQFDVKFWSQHISFILVGIIIVTSIRGLLITLTKFFYAISSSKSSNVIVLLLAQIMGM
YFVSSVLLIRMSMPLEYRTIITEVLGELQFNFYHRWFDVIFLVSALSSILFLYLAHKQAPEKQMAP

Important features:

Signal peptide:

amino acids 1-23

Potential transmembrane domains:

amino acids 37-55, 81-102, 150-168, 288-311, 338-356, 375-398,
425-444

N-glycosylation sites.

amino acids 67-70, 180-183 and 243-246

Eukaryotic cobalamin-binding proteins

amino acids 151-160

FIGURE 5

AGCAGGGAAATCCGGATGTCTCGGTTATGAAGTGGAGCAGTGAGTGTGAGCCTCAACATAGTTCC
 AGAACTCTCCATCCGGACTAGTTATTGAGCATCTGCCTCTCATATCACCAGTGGCCATCTGAGGT
 GTTTCCTGGCTCTGAAGGGGTAGGCACGATGGCCAGGTGCTTCAGCCTGGTGTTGCTTCTCACT
 TCCATCTGGACCACGAGGCTCCTGGTCCAAGGCTCTTTGCGTGCAGAAGAGCTTTCCATCCAGGT
 GTCATGCAGAATTATGGGGATCACCTTGTGAGCAAAAAGGCGAACCAGCAGCTGAATTTACAG
 AAGCTAAGGAGGCTGTAGGCTGCTGGGACTAAGTTTGGCCGGCAAGGACCAAGTTGAAACAGCC
 TTGAAAGCTAGCTTTGAAACTTGCAGCTATGGCTGGGTGGAGATGGATTTCGTGGTCATCTCTAG
 GATTAGCCCCAAACCCCAAGTGTGGGAAAAATGGGGTGGGTGTCCTGATTTGGAAGGTTCCAGTGA
 GCCGACAGTTTGCAGCCTATTGTTACAACCTCATCTGATACTTGGACTAACTCGTGCATTCCAGAA
 ATTATCACCACCAAGATCCCATATTCAACACTCAAACGCAACACAACAGAATTTATTGT
 CAGTGACAGTACCTACTCGGTGGCATCCCCCTTACTCTACAATACCTGCCCCCTACTACTCTCTC
 CTGCTCCAGCTTCCACTTCTATTCCACGGAGAAAAAATTGATTTGTGTACAGAAGTTTTTATG
 GAAACTAGCACCATGTCTACAGAACTGAACCATTTGTTGAAAATAAAGCAGCATTCAAGAATGA
 AGCTGCTGGGTTTGGAGGTGTCCCCACGGCTCTGCTAGTGCTTGCTCTCCTCTTCTTTGGTGCTG
 CAGCTGGTCTTGGATTTTGCTATGTCAAAAGGTATGTGAAGGCCTTCCCTTTTACAAACAAGAAT
 CAGCAGAAGGAAATGATCGAAACCAAGTAGTAAAGGAGGAGAAGGCCAATGATAGCAACCCCTAA
 TGAGGAATCAAAGAAAACCTGATAAAACCCAGAAGAGTCCAAGAGTCCAAGCAAAACTACCGTGC
 GATGCCTGGAAGCTGAAGTTTTAGATGAGACAGAAATGAGGAGACACACCTGAGGCTGGTTTTCTTT
 CATGCTCCTTACCCTGCCCCAGCTGGGGAAATCAAAGGGCCAAAGAACCAAGAAGAAAGTCCA
 CCCTTGGTTCCTAACTGGAATCAGCTCAGGACTGCCATTGGACTATGGAGTGCACCAAGAGAAT
 GCCCTTCTCCTTATTGTAACCCCTGTCTGGATCCTATCCTCCTACCTCCAAAGCTTCCCACGGCCT
 TTCTAGCCTGGCTATGTCTTAATAATATCCCCTGGGAGAAAGGAGTTTTGCAAAGTGCAAGGAC
 CTAAACATCTCATCAGTATCCAGTGGTAAAAAGGCCTCCTGGCTGTCTGAGGCTAGGTGGGTTG
 AAAGCCAAGGAGTCACTGAGACCAAGGCTTCTCTACTGATTCCGCAGCTCAGACCCTTTCTTCA
 GCTCTGAAAGAGAAACACGTATCCCACCTGACATGTCTTCTGAGCCCGGTAAGAGCAAAAGAAT
 GGCAGAAAAGTTTAGCCCCCTGAAAGCCATGGAGATTCTCATAACTTGAGACCTAATCTCTGTAAA
 GCTAAAAATAAGAAATAGAACAAGGCTGAGGATACGACAGTACACTGTCAGCAGGGACTGTAAAC
 ACAGACAGGGTCAAAGTGTTTTCTCTGAACACATTGAGTTGGAATCACTGTTTAGAACACACACA
 CTTACTTTTTCTGGTCTCTACCACTGCTGATATTTCTCTAGGAAATATACTTTTACAAGTAACA
 AAAATAAAAACTCTTATAAATTTCTATTTTATCTGAGTTACAGAAATGATTACTAAGGAAGATT
 ACTCAGTAATTTGTTTAAAAAGTAATAAAATTCACAAACATTTGCTGAATAGCTACTATATGTC
 AAGTGCTGTGCAAGGTATTACACTCTGTAATTGAATATTATTCCTCAAAAAATTGCACATAGTAG
 AACGCTATCTGGGAAGCTATTTTTTTCAGTTTGATATTCTAGCTTATCTACTTCCAAACTAAT
 TTTTATTTTTGCTGAGACTAATCTTATTCATTTTCTCTAATATGGCAACCATTATAACCTTAATT
 TATTATTAACATACCTAAGAAGTACATTGTTACCTCTATATAACAAAGCACATTTTAAAAAGTGCC
 ATTAACAAATGTATCACTAGCCCTCCTTTTTTCCAACAAGAAGGGACTGAGAGATGCAGAAATATT
 TGTGACAAAAAATTAAAGCATTTAGAAAACTT

FIGURE 6

MARCFSLVLLLTISIWTTTRLLVQGSRLAEELSIQVSCRIMGITLVSKKANQQLNFTEAKEACRLLG
LSLAGKDQVETALKASFETCSYGWVGDFVVISRISPNPKCGKNGVGVLIWKVPVSRQFAAYCYN
SSDTWTNSCIPEIITTKDPIFNTQTATQTTEFIVSDSTYSVASPYSTIPAPTTTTPPAPASTSIPR
RKKLICVTEVFMETSTMSTETEPFVENKAAFNEAAGFGGVPTALLVLALLFFGAAAGLGFCYVK
RYVKAFFFTNKNQOKEMIETKVVKEEKANDSNPNEESKKTDKNPEESKSPSKTTVRCLEAEV

Signal sequence:

amino acids 1-16

Transmembrane domain:

amino acids 235-254

N-glycosylation site.

amino acids 53-57, 130-134, 289-293

Casein kinase II phosphorylation site.

amino acids 145-149, 214-218

Tyrosine kinase phosphorylation site.

amino acids 79-88

N-myristoylation site.

amino acids 23-29, 65-71, 234-240, 235-239, 249-255, 253-259

FIGURE 7

CGCCGCGCTCCCGCACCCGCGGCCCCGCCACCGCGCCGCTCCCGCATCTGCACCCGCGAGCCCGGC
GGCCTCCCGGCGGGAGCGAGCAGATCCAGTCCGGCCCCGAGCGCAACTCGGTCCAGTCGGGGCGG
CGGCTGCGGGCGCAGAGCGGAGATGCAGCGGCTTGGGGCCACCCTGCTGTGCCTGCTGCTGGCGG
CGGCGGTCCCCACGGCCCCCGCGCCGCTCCGACGGCGACCTCGGCTCCAGTCAAGCCCGGCCG
GCTCTCAGCTACCCGCGAGGAGGAGCCACCCTCAATGAGATGTTCCGCGAGGTTGAGGAACTGAT
GGAGGACACGCAGCACAAATTGCGCAGCGCGGTGGAAGAGATGGAGGCAGAAGAAGCTGCTGCTA
AAGCATCATCAGAAGTGAACCTGGCAAACCTTACCTCCCAGCTATCACAATGAGACCAACACAGAC
ACGAAGGTTTGAAATAATACCATCCATGTGCACCGAGAAATTACAAGATAACCAACAACACAGAC
TGGACAAATGGTCTTTTTCAGAGACAGTTATCACATCTGTGGGAGACGAAGAAGGCAGAAGGAGCC
ACGAGTGCATCATCGACGAGGACTGTGGGCCCAGCATGTACTGCCAGTTTGCCAGCTTCCAGTAC
ACCTGCCAGCCATGCCGGGGCCAGAGGATGCTCTGCACCCGGGACAGTGAGTGCTGTGGAGACCA
GCTGTGTGTCTGGGGTCACTGCACCAAAATGGCCACCAGGGGAGCAATGGGACCATCTGTGACA
ACCAGAGGGGACTGCCAGCCGGGGCTGTGCTGTGCCTTCCAGAGAGGCCTGCTGTTCCCTGTGTGC
ACACCCCTGCCCCGTGGAGGGCGAGCTTTGCCATGACCCCGCCAGCCGGCTTCTGGACCTCATCAC
CTGGGAGCTAGAGCCTGATGGAGCCTTGGACCGATGCCCTTGTGCCAGTGCCCTCCTCTGCCAGC
CCCACAGCCACAGCCTGGTGTATGTGTGCAAGCCGACCTTCGTGGGGAGCCGTGACCAAGATGGG
GAGATCCTGCTGCCCAGAGAGGTCCCCGATGAGTATGAAGTTGGCAGCTTCATGGAGGAGGTGCG
CCAGGAGCTGGAGGACCTGGAGAGGAGCCTGACTGAAGAGATGGCGCTGGGGGAGCCTGCGGCTG
CCGCCCTGCACTGCTGGGAGGGGAAGAGATTAGATCTGGACCAGGCTGTGGGTAGATGTGCAA
TAGAAATAGCTAATTTATTTCCCCAGGTGTGTGCTTTAGGCGTGGGCTGACCAGGCTTCTTCCTA
CATCTTCTTCCCAGTAAGTTTCCCCCTCTGGCTTGACAGCATGAGGTGTGTGCATTTGTTTCAGCT
CCCCAGGCTGTCTTCCAGGCTTCACAGTCTGGTGCTTGGGAGAGTCAGGCAGGGTTAAACTGCA
GGAGCAGTTTGCCACCCCTGTCCAGATTATTGGCTGCTTTGCCTCTACCAGTTGGCAGACAGCCG
TTTGTCTACATGGCTTTGATAATTGTTTGAGGGGAGGAGATGGAAACAATGTGGAGTCTCCCTC
TGATTGGTTTTGGGGAAATGTGGAGAAGAGTGCCCTGCTTTGCAAACATCAACCTGGCAAAAATG
CAACAAATGAATTTTCCACGCAGTTCTTTCCATGGGCATAGGTAAGCTGTGCCTTCAGCTGTTGC
AGATGAAATGTTCTGTTTACCCTGCATTACATGTGTTTATTTCATCCAGCAGTGTTGCTCAGCTCC
TACCTCTGTGCCAGGGCAGCATTTTTCATATCCAAGATCAATTCCCTCTCTCAGCACAGCCTGGGG
AGGGGGTCATTGTTCTCCTCGTCCATCAGGGATCTCAGAGGCTCAGAGACTGCAAGCTGCTTGCC
CAAGTCACACAGCTAGTGAAGACCAGAGCAGTTTCATCTGGTTGTGACTCTAAGCTCAGTGCTCT
CTCCACTACCCACACCCAGCCTTGGTGCCACCAAAAGTGCTCCCCAAAAGGAAGGAGAATGGGAT
TTTTCTTGAGGCATGCACATCTGGAATTAAGGTCAAATAATTCTCACATCCCTCTAAAAGTAAA
CTACTGTTAGGAACAGCAGTGTTCTCACAGTGTGGGGCAGCCGTCCTTCTAATGAAGACAATGAT
ATTGACACTGTCCCTCTTTGGCAGTTGCATTAGTAACTTTGAAAGGTATATGACTGAGCGTAGCA
TACAGGTTAACCTGCAGAAACAGTACTTAGGTAATTGTAGGGCGAGGATTATAAATGAAATTTGC
AAAATCACTTAGCAGCAACTGAAGACAATTATCAACCACGTGGAGAAAATCAAACCGAGCAGGGC
TGTGTGAAACATGGTTGTAATATGCGACTGCGAACACTGAACTCTACGCCACTCCACAAATGATG
TTTTCAGGTGTCATGGACTGTTGCCACCATGTATTTCATCCAGAGTTCTTAAAGTTTAAAGTTGCA
CATGATTGTATAAGCATGCTTTCTTTGAGTTTAAATTATGTATAAACATAAGTTGCATTTAGAA
ATCAAGCATAAATCACTTCAACTGCAAAAAAAAAAAAAAAAAAAAAAAAAAAAAA

FIGURE 8

MQRLGATLLCLLLAAAVPTAPAPAPTATSAPVKPGPALSYPQEEATLNEMFREVEELMEDTQHKL
RSAVEEMEAEAAAAKASSEVNLANLPPSYHNETNTDTKVGNNTIHVHREIHKITNNQTGQMFSE
TVITSVGDEEGRRSHECIIDEDCGPSMYCQFASFQYTCQPCRGQRMCLTRDSECCGDQLCVWGHC
TKMATRGSNGTICDNQRDCQPLCCAFQRGLLFPVCTPLPVEGELCHDPASRLDLITWELEPDG
ALDRPCASGLLCQPHSHSLVYVCKPTFVGSRDQDGEILLPREVPDEYEVGSGFMEEVRQELEDLE
RSLTEEMALGEPAAAAAALLGEEI

Signal sequence:

amino acids 1-19

N-glycosylation site.

amino acids 96-100, 106-110, 121-125, 204-208

Casein kinase II phosphorylation site.

amino acids 46-50, 67-71, 98-102, 135-139, 206-210, 312-316,
327-331

N-myristoylation site.

amino acids 202-208, 217-223

Amidation site.

amino acids 140-144

FIGURE 9

CGGACGCGTGGGCGGACGCGTGGGGGCTGTGAGAAAGTGCCAATAAATACATCATGCAACCCAC
GGCCACCTTGTGAACTCCTCGTGCCAGGGCTGATGTGCGTCTTCCAGGGCTACTCATCCAAAG
GCCTAATCCAACGTTCTGTCTTCAATCTGCAAATCTATGGGGTCCTGGGGCTCTTCTGGACCCTT
AACTGGGTACTGGCCCTGGGCCAATGCGTCCTCGCTGGAGCCTTTGCCTCCTTCTACTGGGCCTT
CCACAAGCCCCAGGACATCCCTACCTTCCCCTTAATCTCTGCCTTCATCCGCACACTCCGTTACC
ACACTGGGTCATTGGCATTGGAGCCCTCATCCTGACCCTTGTGCAGATAGCCCGGGTCATCTTG
GAGTATATTGACCACAAGCTCAGAGGAGTGCAGAACCCTGTAGCCCGCTGCATCATGTGCTGTTT
CAAGTGCTGCCTCTGGTGTCTGGAAAAATTTATCAAGTTCCTAAACCGCAATGCATACATCATGA
TCGCCATCTACGGGAAGAATTTCTGTGTCTCAGCCAAAAATGCGTTCATGCTACTCATGCGAAAC
ATTGTCAGGGTGGTCGTCTCGGACAAAGTCACAGACCTGCTGCTGTTCTTTGGGAAGCTGCTGGT
GGTCGGAGGCGTGGGGGTCTGTCTCTTTTTTTTTCTCCGGTCGCATCCCGGGGCTGGGTAAAG
ACTTTAAGAGCCCCCACCTCAACTATTACTGGCTGCCCATCATGACCTCCATCCTGGGGGCCTAT
GTCATCGCCAGCGGCTTCTTCAGCGTTTTTCGGCATGTGTGTGGACACGCTCTTCTCTGCTTCCT
GGAAGACCTGGAGCGGAACAACGGCTCCCTGGACCGGCCCTACTACATGTCCAAGAGCCTTCTAA
AGATTCTGGGCAAGAAGAACGAGGCGCCCCCGGACAACAAGAAGAGGAAGAAGTGACAGCTCCGG
CCCTGATCCAGGACTGCACCCACCCCAACCGTCCAGCCATCCAACCTCACTTCGCCTTACAGGT
CTCCATTTTGTGGTAAAAAAAGGTTTTAGGCCAGGCGCCGTGGCTCACGCCTGTAATCCAACACT
TTGAGAGGCTGAGGCGGGCGGATCACCTGAGTCAGGAGTTCGAGACCAGCCTGGCCAACATGGTG
AAACCTCCGTCTCTATTAAAAATACAAAAATTAGCCGAGAGTGGTGGCATGCACCTGTCATCCCA
GCTACTCGGGAGGCTGAGGCAGGAGAATCGCTTGAACCCGGGAGGCAGAGGTTGCAGTGAGCCGA
GATCGCGCCACTGCACTCCAACCTGGGTGACAGACTCTGTCTCCAAAACAAAACAAACAAA
AAGATTTTATTAAAGATATTTTGTTAAGTC

FIGURE 10

RTRGRTRGGCEKVPINTSCNPTAHLVNSSCPGLMCVFQGYSSKGLIQRSVFNLQIYGVLGGLFWTL
NWVLALGQCVLGAFASFYWAFHKPDIPTFPLISAFIRTLRYHTGSLAFGALILTLVQIARVIL
EYIDHKLRGVQNPVARCIMCCFKCCLWCLEKFIKFLNRNAYIMIAIYGKNFCVSAKNAFMLLMRN
IVRVVVLDDKVTDLLLFFGKLLVVGGVGVLSFFFFSGRIPGLGKDFKSPHLNYYWLPIMTSILGAY
VIASGFFSVFGMCVDTLFLCFLEDLERNNGSLDRPYYSKSLKILGKKNEAPPDNKKRKK

Important features:

Transmembrane domains:

amino acids 57-80 (type II), 110-126, 215-231, 254-274

N-glycosylation sites.

amino acids 16-20, 27-31, 289-293

Hypothetical YBR002c family proteins.

amino acids 276-288

Ammonium transporters proteins.

amino acids 204-231

N-myristoylation sites.

amino acids 60-66, 78-84

Amidation site.

amino acids 306-310

FIGURE 11

GCCCCGCGCCCGCGCGCGGGCGCCCGAAGCCGGGAGCCACCGCCATGGGGGCCTGCCTGGGAGCCTGC
TCCCTGCTCAGCTGCGCGTCTGCCTCTGCGGCTCTGCCCCCTGCATCCTGTGCAGCTGCTGCCCCGC
CAGCCGCAACTCCACCGTGAGCCGCCTCATCTTCACGTTCTTCTCTTCTGCGGGTGTGGTGTCCA
TCATTATGCTGAGCCCGGGCGTGAGAGTCAGCTCTACAAGCTGCCCTGGGTGTGTGAGGAGGGGGCC
GGGATCCCCACCGTCTGCAGGGCCACATCGACTGTGGCTCCCTGCTTGGCTACCGCGCTGTCTACCG
CATGTGCTTCGCCACGGCGGCCTTCTTCTTCTTCTTTTACCCTGCTCATGCTCTGCGTGAGCAGCA
GCCGGGACCCCCGGGCTGCCATCCAGAATGGGTTTTGGTTCTTTAAGTTCCTGATCCTGGTGGGCCTC
ACCGTGGGTGCCTTCTACATCCCTGACGGCTCCTTCACCAACATCTGGTTCTACTTCGGCGTGTGGG
CTCCTTCTCTTCATCCTCATCCAGCTGGTGTGCTCATCGACTTTGCGCACTCCTGGAACCAGCGGT
GGCTGGGCAAGCCGAGGAGTGCATTCCCGTGCCTGGTACGCAGGCCTCTTCTTCTTCACTCTCCTC
TTCTACTTGCTGTGCATCGCGCCGTGGCGCTGATGTTTCATGTACTACACTGAGCCAGCGGCTGCCA
CGAGGGCAAGGTCTTCATCAGCCTCAACCTCACCTTCTGTGTCTGCGTGTCCATCGCTGTCTCTGC
CCAAGGTCCAGGACGCCAGCCCAACTCGGGTCTGCTGCAGGCCTCGGTTCATCACCTCTACACCATG
TTGTGCACCTGGTCAGCCCTATCCAGTATCCCTGAACAGAAATGCAACCCCCATTTGCCAACCCAGCT
GGGCAACGAGACAGTTGTGGCAGGCCCCGAGGGCTATGAGACCCAGTGGTGGGATGCCCCGAGCATTG
TGGGCCTCATCATCTTCTCTGTGCACCTCTTTCATCAGTCTGCGCTCCTCAGACCACCGGCAGGTG
AACAGCCTGATGCAGACCGAGGAGTGCCACCTATGCTAGACGCCACACAGCAGCAGCAGCAGCAGGT
GGCAGCCTGTGAGGGCCGGGCCTTTGACAACGAGCAGGACGGCGTCACCTACAGCTACTCCTTCTTCC
ACTTCTGCCTGGTGTGCTGGCCTCACTGCACGTATGATGACGCTCACCAACTGGTACAAGCCCGGTGAG
ACCCGGAAGATGATCAGCACGTGACCGCGTGTGGGTGAAGATCTGTGCCAGCTGGGCAGGGCTGCT
CCTCTACCTGTGGACCCTGGTAGCCCCACTCCTCCTGCGCAACCGCGACTTCAGCTTGAGGCAGCCTCA
CAGCCTGCCATCTGGTGCCTCCTGCCACCTGGTGCCTCTCGGCTCGGTGACAGCCAACCTGCCCCCTC
CCCACACCAATCAGCCAGGCTGAGCCCCACCCTGCCCCAGCTCCAGGACCTGCCCTGAGCCGGGC
CTTCTAGTCGTAGTGCCTTCAGGGTCCGAGGAGCATCAGGCTCCTGCAGAGCCCCATCCCCCGCCAC
ACCCACACGGTGGAGCTGCCTCTTCTTCCCCCTCCTCCCTGTTGCCATACTCAGCATCTCGGATGAA
AGGGCTCCCTTGTCTCAGGCTCCACGGGAGCGGGGCTGCTGGAGAGAGCGGGGAACCTCCACACAG
TGGGGCATCCGGCACTGAAGCCTGGTGTCTTGGTCACGTCCCCAGGGGACCCTGCCCCCTTCTG
GACTTCGTGCCTTACTGAGTCTTAAGACTTTTTCTAATAAACAAGCCAGTGCCTGTAAAAAAA

FIGURE 12

MGACLGACSLLSASCCLCGSAPCILCSCCPASRNSTVSRLIFTFFLFLGVLVSIIMLSPGVESQL
YKLPWVCEEGAGIPTVLQGHIDCGSLLGYRAVYRMCFATAAFFFFFFFFTLLMLCVSSSRDPRAAIQ
NGFWFFKFLILVGLTVGAFYIPDGSFTNIWFYFGVVGSFLFILIQLVLLIDFAHSWNQRWLKAE
ECDSRAWYAGLFFFTLLFYLLSIAAVALMFMYYTEPSGCHEGKVFISLNLTFVCVCSIAAVLPKV
QDAQPNSGLLQASVITLYTMFVTWSALSSIPEQKCNPHLPTQLGNETVVAGPEGYETQWWDA PSI
VGLIIFLLCTLFISLRSSDHRQVNSLMQTEECPPMLDATQQQQQQVAACEGRAFDNEQDGVITYSY
SFFHFCLVLASLHVMMTLTNWYKPGETRKMISTWTAVWVKICASWAGLLLYLWTLVAPLLLRNRD
FS

Signal sequence:

amino acids 1-20

Transmembrane domains:

amino acids 40-58, 101-116, 134-150, 162-178, 206-223, 240-257,
272-283, 324-340, 391-406, 428-444

FIGURE 13

CGGGCCAGCCTGGGGCGGCCGGCCAGGAACCACCCGTTAAGGTGTCTTCTCTTTAGGGATGGTGA
GGTTGGAAAAAGACTCCTGTAAACCCTCCTCCAGGATGAACCACCTGCCAGAAGACATGGAGAACG
CTCTCACCGGGAGCCAGAGCTCCCATGCTTCTCTGCGCAATATCCATTCCATCAACCCACACAA
CTCATGGCCAGGATTGAGTCCTATGAAGGAAGGGAAAAGAAAGGCATATCTGATGTCAGGAGGAC
TTTCTGTTTGTGTTGTACCTTTGACCTCTTATTCTGTAACATTACTGTGGATAATAGAGTTAAATG
TGAATGGAGGCATTGAGAACACATTAGAGAAGGAGGTGATGCAGTATGACTACTATTCTTCATAT
TTTGATATATTTCTTCTGGCAGTTTTTCGATTTAAAGTGTTAATACTTGCATATGCTGTGTGCAG
ACTGCGCCATTGGTGGGCAATAGCGTTGACAACGGCAGTGACCAGTGCCTTTTTACTAGCAAAAG
TGATCCTTTTGAAGCTTTTCTCTCAAGGGGCTTTTGGCTATGTGCTGCCCATCATTTTCATTCATC
CTTGCTGGATTGAGACGTGGTTTCTGGATTTCAAAGTGTTACCTCAAGAAGCAGAAGAAGAAAA
CAGACTCCTGATAGTTTCAAGATGCTTCAGAGAGGGCAGCACTTATACCTGGTGGTCTTTCTGATG
GTCAGTTTTATTCCCTCCTGAATCCGAAGCAGGATCTGAAGAAGCTGAAGAAAAACAGGACAGT
GAGAAACCACTTTTAGAACTATGAGTACTACTTTTGTAAATGTGAAAAACCCTCACAGAAAGTC
ATCGAGGCAAAAAGAGGCAGGCAGTGGAGTCTCCCTGTGACAGTAAAGTTGAAATGGTGACGTC
CACTGCTGGCTTTATTGAACAGCTAATAAAGATTTATTTATTGTAATACCTCACAAACGTTGTAC
CATATCCATGCACATTTAGTTGCCTGCCTGTGGCTGGTAAGGTAATGTCATGATTCATCCTCTCT
TCAGTGAGACTGAGCCTGATGTGTTAACAAATAGGTGAAGAAAGTCTTGTGCTGTATTCCTAATC
AAAAGACTTAATATATTGAAGTAACACTTTTTTAGTAAGCAAGATACCTTTTTATTTCATTCAC
AGAATGGAATTTTTTTGTTTCATGTCTCAGATTTATTTTGTATTTCTTTTTTAACACTCTACATT
TCCCTTGTTTTTTAACTCATGCACATGTGCTCTTTGTACAGTTTTAAAAAGTGTAATAAAATCTG
ACATGTCAATGTGGCTAGTTTTATTTTTCTGTTTTGCATTATGTGTATGGCCTGAAGTGTGGGA
CTTGCAAAAGGGGAAGAAAGGAATTGCGAATACATGTAAAATGTCACCAGACATTTGTATTATTT
TTATCATGAAATCATGTTTTTCTCTGATTGTTCTGAAATGTTCTAAATACTCTTATTTGAATGC
ACAAAATGACTTAAACCATTATATCATGTTTCCTTTGCGTTCAGCCAATTTCAATTAAAAATGAA
CTAAATTAAAAA

FIGURE 14

MNHLPEDMENALTGSQSSHASLRNIHSINPTQLMARIESYEGREKKGISDVRRTFCLFVTFDLLF
VTLLWIIELNVNGGIENTLEKEVMQYDYYSSYFDIFLLAVFRFKVLILAYAVCRLRHWWAIALTT
AVTSAFLLAKVILSKLFSQGAFGYVLPPIISFILAWIETWFLDFKVLQPQAEENRLLIVQDASER
AALIPGGLSDGQFYSPPESEAGSEEAEKQDSEKPLLEL

Important features of the protein:

Signal peptide:

amino acids 1-20

Transmembrane domains:

amino acids 54-72, 100-118, 130-144, 146-166

N-myristoylation sites.

amino acids 14-20, 78-84, 79-85, 202-208, 217-223

FIGURE 15

ACTCGAACCGCAGTTGCTTCGGGACCCAGGACCCCCCTCGGGCCCGACCCGCCAGGAAAGACTGAGG
CCGCGGCCTGCCCCGCCCCGGCTCCCTGCGCCGCCGCCGCTCCCGGGACAGAAGATGTGCTCCAG
GGTCCCTCTGCTGCTGCCGCTGCTCCTGCTACTGCCCCGTTGGGGCTGGGGTGAGGGCTGCCCAT
CCGGCTGCCAGTGACGCCAGCCACAGACAGTCTTCTGCACTGCCCGCCAGGGGACCACGGTGCCC
CGAGACGTGCCACCCGACACGGTGGGGCTGTACGTCTTTGAGAACGGCATCACCATGCTCGACGC
AGGCAGCTTTGCCGGCCTGCCGGGCTGCAGTCCCTGGACCTGTACAGAACCAGATCGCCAGCC
TGCCCAGCGGGGTCTTCCAGCCACTCGCCAACCTCAGCAACCTGGACCTGACGGCCAAACAGGCTG
CATGAAATCACCAATGAGACCTTCCGTGGCCTGCGGCGCCTCGAGCGCCTCTACCTGGGCAAGAA
CCGCATCCGCCACATCCAGCCTGGTGCCCTTCGACACGCTCGACCCGCTCCTGGAGCTCAAGCTGC
AGGACAACGAGCTGCGGGCACTGCCCCCGCTGCGCCTGCCCCGCTGCTGCTGCTGGACCTCAGC
CACAACAGCCTCCTGGCCCTGGAGCCCGGCATCCTGGACACTGCCAACGTGGAGGGCTGCGGCT
GGCTGGTCTGGGGCTGCAGCAGCTGGACGAGGGGCTCTTCAGCCGCTTGCGCAACCTCCACGACC
TGGATGTGTCCGACAACCAGCTGGAGCGAGTGCCACCTGTGATCCGAGGCCTCCGGGGCTGACG
CGCCTGCGGCTGGCCGGCAACACCCGCATTGCCCAGCTGCGGCCGAGGACCTGGCCGGCCTGGC
TGCCCTGCAGGAGCTGGATGTGAGCAACCTAAGCCTGCAGGCCCTGCCCTGGCGACCTCTCGGGCC
TCTTCCCCCGCCTGCGGCTGCTGGCAGCTGCCCGCAACCCCTTCAACTGCGTGTGCCCCCTGAGC
TGGTTTGGCCCCCTGGGTGCGCGAGAGCCACGTACACTGGCCAGCCCTGAGGAGACGCGCTGCCA
CTTCCCGCCCAAGAACGCTGGCCGGCTGCTCCTGGAGCTTGACTACGCCGACTTTGGGTGCCCAG
CCACCACCCACACAGCCACAGTGCCCAACCACGAGGCCCGTGGTGCGGGAGCCACAGCCTTGTCT
TCTAGCTTGGCTCCTACCTGGCTTAGCCCCACAGCGCCGGCCACTGAGGCCCCAGCCCGCCCTC
CACTGCCCCACCGACTGTAGGGCCTGTCCCCAGCCCCAGGACTGCCACCGTCCACCTGCCTCA
ATGGGGGCACATGCCACCTGGGGACACGGCACCACCTGGCGTGCTTGTGCCCGGAAGGCTTCAGC
GGCCTGTACTGTGAGAGCCAGATGGGGCAGGGGACACGGCCAGCCCTACACCAGTCACGCCGAG
GCCACCACGGTCCCTGACCCCTGGGCATCGAGCCGGTGAGCCCCACCTCCCTGCGCGTGGGGCTGC
AGCGCTACCTCCAGGGGAGCTCCGTGCAGCTCAGGAGCCTCCGTCTCACCTATCGCAACCTATCG
GGCCCTGATAAGCGGCTGGTGACGCTGCGACTGCCTGCCTCGCTCGCTGAGTACACGGTCACCCA
GCTGCGGGCCCAACGCCACTTACTCCGTCTGTGTATGCCCTTTGGGGCCCCGGGCGGGTGCCCGAGG
GCGAGGAGGCCTGCGGGGAGGCCCATACACCCCCAGCCGTCCACTCCAACCACGCCCCAGTCACC
CAGGCCCGCGAGGGCAACCTGCCGCTCCTCATTTGCGCCCGCCCTGGCCGCGGTGCTCCTGGCCGC
GCTGGCTGCGGTGGGGGCGAGCCTACTGTGTGCGGCGGGGGCGGGCCATGGCAGCAGCGGCTCAGG
ACAAAGGGCAGGTGGGGCCAGGGGCTGGGGCCCTGGAACTGGAGGGAGTGAAGGTCCCTTGGAG
CCAGGCCCGAAGGCAACAGAGGGCGGTGGAGAGGCCCTGCCCAGCGGGTCTGAGTGTGAGGTGCC
ACTCATGGGCTTCCCAGGGCCTGGCCTCCAGTCACCCCTCCACGCCAAGCCCTACATCTAAGCCA
GAGAGAGACAGGGCAGCTGGGGCCGGGCTCTCAGCCAGTGAGATGGCCAGCCCCCTCCTGCTGCC
ACACCACGTAAGTTCTCAGTCCCAACCTCGGGGATGTGTGCAGACAGGGCTGTGTGACCACAGCT
GGGCCCTGTTCCCTCTGGACCTCGGTCTCCTCATCTGTGAGATGCTGTGGCCAGCTGACGAGCC
CTAACGTCCCCAGAACCGAGTGCCATAGGACAGTGATCCGCCCTGCCCTCCGCAACGTGCAGTC
CCTGGGCACGGCGGGCCCTGCCATGTGCTGGTAACGATGCTCGGTCTGCTGGGCTCTCCAC
TCCAGGCGGACCTTGGGGCCAGTGAAGGAAGCTCCCGGAAGAGCAGAGGGAGAGCGGGTAGGC
GGCTGTGTGACTCTAGTCTTGGCCCCAGGAAGCGAAGGAACAAAAGAAACTGGAAAGGAAGATGC
TTTAGGAACATGTTTTTGCTTTTTTAAAAATATATATATTTATAAGAGATCCTTTCCCATTTATTCT
GGGAAGATGTTTTTCAAACCTCAGAGACAAGGACTTTGGTTTTTGTAAAGACAAACGATGATATGAA
GGCCTTTTGTAAAGAAAAATAAAAGATGAAGTGTGAAA

FIGURE 16

MCSRVP LLLPL LLL LLLALGPGVQGCPSGCQCSQPQTVECTARQGTTVPRDVPPDTVGLYVFENGIT
MLDAGSFAGLPGLQLLDLSQNQIASLPSGVFQPLANLSNLDLTANRLHEITNETFRGLRRLERLY
LGKNRIRHIQPGAFDTLDRLLLEKLQDNELRALPPLRLPRLLLLDLSHNSLLALEPGILDTANVE
ALRLAGLGLQQLDEGLFSRLRNLDLDVSDNQLERVPPVIRGLRGLTRLRLAGNTRIAQLRPEDL
AGLAALQELDVSNLSLQALPGDLSGLFPRLRLLLAAARNPFNCVCPLSWFGPWVRESHVTLASPEE
TRCHFPPKNAGRLLLELDYADFGCPATTTTATVPTTRPVVREPTALSSSLAPTWLSPTAPATEAP
SPPSTAPPTVGPVPQPDCCPSTCLNGGTCHLGRHHLACLCPEGFTGLYCESQMGQGTTPSPTP
VTPRPPRSLTLGIEPVSPSTSLRVGLQRYLQGS SVQLRSRLTYRNLSGPDKRLVTLRLPASLAEY
TVTQLRPNATYSVCMPLGPGRVPEGEEACGEAHTPPAVHSNHAPVTQAREGNLPLLIAPALAAV
LLAALAAVGAAYCVRGRGRAMAAAAQDKGQVGPAGPLELEGVKVPLEPGPKATEGGGEALPSGSE
CEVPLMGFPGLQSPHAKPYI

Important features:

Signal peptide:

amino acids 1-23

Transmembrane domain:

amino acids 579-599

EGF-like domain cysteine pattern signature.

amino acids 430-442

Leucine zipper pattern.

amino acids 197-219, 269-291

N-glycosylation sites.

amino acids 101-105, 117-121, 273-277, 500-504, 528-532

Tyrosine kinase phosphorylation sites.

amino acids 124-131, 337-345

N-myristoylation sites.

amino acids 23-29, 27-33, 70-76, 142-148, 187-193, 348-354,
594-600, 640-646

FIGURE 17

[illegible]

FIGURE 18

MRVRIGLTLLLCVLLSLASASSDEEGSQDESLSKTTLTSDSVKDHTTAGRVVAGQIFLDSEESSEL
ESSIQEEEDSLKSQEGESVTEDISFLESPNPENKDYEEPKKVRKPALTAIEGTAHGEPCHFPPFLFDK
EYDECTSDGREDGRLWCATTYDYKADEKWGFCETEEEAARRQMQEAEMMYQTGMKILNGSNKKSQKR
EAYRYLQKAASMNHTKALERVSYALLFGDYLPQNIQAAREMFEEKLTEEGSPKGTALGFLYASGLGVN
SSQAKALVYYTFGALGGNLIAMVLVSRL

Important features:

Signal peptide:

amino acids 1-21

N-glycosylation sites.

amino acids 195-199, 217-221, 272-276

Tyrosine kinase phosphorylation site.

amino acids 220-228

N-myristoylation sites.

amino acids 120-126, 253-259, 268-274, 270-274, 285-291, 289-295

Glycosaminoglycan attachment site.

amino acids 267-271

Microbodies C-terminal targeting signal.

amino acids 299-303

Type II fibronectin collagen-binding domain protein.

amino acids 127-169

Fructose-bisphosphate aldolase class-II protein.

amino acids 101-119

FIGURE 19

AATTCAGATTTTAAGCCCATTCCTGCAGTGGAATTTTCATGAACTAGCAAGAGGACACCATCTTCTT
GTATTATACAAGAAAGGAGTGTACCTATCACACACAGGGGGAAAAATGCTCTTTTGGGTGCTAGG
CCTCCTAATCCTCTGTGGTTTTCTGTGGACTCGTAAAGGAAAACCTAAAGATTGAAGACATCACTG
ATAAGTACATTTTTATCACTGGATGTGACTCGGGCTTTGGAACTTGGCAGCCAGAACTTTTGAT
AAAAAGGGATTTTCATGTAATCGCTGCCTGTCTGACTGAATCAGGATCAACAGCTTTAAAGGCAGA
AACCTCAGAGAGACTTCGTACTGTGCTTCTGGATGTGACCGACCCAGAGAATGTCAAGAGGACTG
CCCAGTGGGTGAAGAACCAAGTTGGGGAGAAAGTCTCTGGGGTCTGATCAATAATGCTGGTGTT
CCCGGCGTGCTGGCTCCCACTGACTGGCTGACACTAGAGGACTACAGAGAACCTATTGAAGTGAA
CCTGTTTGGACTCATCAGTGTGACACTAAATATGCTTCCTTTGGTCAAGAAAGCTCAAGGGAGAG
TTATTAATGTCTCCAGTGTGGAGGTCGCCTTGCAATCGTTGGAGGGGGCTATACTCCATCCAAA
TATGCAGTGGAAGGTTTCAATGACAGCTTAAGACGGGACATGAAAGCTTTTGGTGTGCACGTCTC
ATGCATTGAACCAGGATTGTTCAAAACAACTTGGCAGATCCAGTAAAGGTAATTGAAAAAAAC
TCGCCATTTGGGAGCAGCTGTCTCCAGACATCAAACAACAATATGGAGAAGGTTACATTGAAAAA
AGTCTAGACAAACTGAAAGGCAATAAATCCTATGTGAACATGGACCTCTCTCCGGTGGTAGAGTG
CATGGACCACGCTCTAACAAGTCTCTTCCCTAAGACTCATTATGCCGCTGGAAAAGATGCCAAAA
TTTTCTGGATACCTCTGTCTCACATGCCAGCAGCTTTGCAAGACTTTTTATTGTTGAAACAGAAA
GCAGAGCTGGCTAATCCCAAGGCAGTGTGACTCAGCTAACCACAAATGTCTCCTCCAGGCTATGA
AATTGGCCGATTTCAAGAACACATCTCCTTTTCAACCCCATTCCTTATCTGCTCCAACCTGGACT
CATTTAGATCGTGCTTATTTGGATTGCAAAGGGAGTCCCACCATCGCTGGTGGTATCCCAGGGT
CCCTGCTCAAGTTTTCTTTGAAAAGGAGGGCTGGAATGGTACATCACATAGGCAAGTCTGCCCT
GTATTTAGGCTTTGCCTGCTTGGTGTGATGTAAGGGAAATTGAAAGACTTGCCCATTCAAAATGA
TCTTTACCGTGGCCTGCCCCATGCTTATGGTCCCCAGCATTACAGTAACCTTGTGAATGTTAAGT
ATCATCTCTTATCTAAATATTAAAAGATAAGTCAACCCAAAAAAAAAAAAAAAAAAAAAAAAAAAA
AAAAAAAAAAAAA

FIGURE 20

MLFWVLGLLILCGFLWTRKGKLIKIEDITDKYIFITGCDSGFGNLAARTFDKKGFHVIAACLTESG
STALKAETSERLRTVLLDVTDPENVKRTAQWVKNOVGEKGLWGLINNAGVPGVLAPTDWLTLEDY
REPIEVNLFGLISVTLNMLPIVKKAQGRVINVSSVGGRLAIVGGGYTPSKYAVEGFNDLRRDMK
AFGVHVSCIEPGLFKTNLADPVKVIKKLAIWEQLSPDIKQQYGEGYIEKSLDKLKGNKSYVNMD
LSPVVECMDHALTSLEPKTHYAAGKDAKIFWIPLSHMPAALQDFLLLKQKAE LANPKAV

Important features of the protein:

Signal peptide:

amino acids 1-17

Transmembrane domain:

amino acids 136-152

N-glycosylation sites.

amino acids 161-163, 187-190 and 253-256

Glycosaminoglycan attachment site.

amino acids 39-42

N-myristoylation sites.

amino acids 36-41, 42-47, 108-113, 166-171, 198-203 and 207-212

FIGURE 21

CTGAGGCGGCGGTAGCATGGAGGGGGAGAGTACGTCGGCGGTGCTCTCGGGCTTTGTGCTCGGCG
 CACTCGCTTTCCAGCACCTCAACACGGACTCGGACACGGAAGGTTTTCTTCTTGGGGAAGTAAAA
 GGTGAAGCCAAGAACAGCATTACTGATTCCCAAATGGATGATGTTGAAGTTGTTTATACAATTGA
 CATTGAGAAATATATTCCATGCTATCAGCTTTTTAGCTTTTATAATTCTTCAGGCGAAGTAAATG
 AGCAAGCACTGAAGAAAATATTATCAAATGTCAAAAAGAATGTGGTAGGTTGGTACAAAATCCGT
 CGTCATTGAGATCAGATCATGACGTTTAGAGAGAGGCTGCTTCACAAAAAAGTGCAGGAGCATT
 TTCAAACCAAGACCTTGTTTTCTGCTATTAACACCAAGTATAATAACAGAAAGCTGCTCTACTC
 ATCGACTGGAACATTTCCTTATATAAACCTCAAAAAGGACTTTTTTACAGGGTACCTTTAGTGGTT
 GCCAATCTGGGCATGTCTGAACAACTGGGTTATAAACTGTATCAGGTTCCCTGTATGTCCACTGG
 TTTTAGCCGAGCAGTACAAACACACAGCTCTAAATTTTTTGAAGAAGATGGATCCTTAAAGGAGG
 TACATAAGATAAATGAAATGTATGCTTCATTACAAGAGGAATTAAAGAGTATATGCAAAAAAGTG
 GAAGACAGTGAACAAGCAGTAGATAAACTAGTAAAGGATGTAAACAGATTAAACGAGAAATTGA
 GAAAAGGAGAGGAGCAGATTTCAGGCAGCAAGAGAGAAGAACATCCAAAAAGACCTCAGGAGA
 ACATTTTTCTTTGTGAGGCATTACGGACCTTTTTTCCAAATCTGAATTTCTTCATTATGTGTT
 ATGTCTTTAAAAAATAGACATGTTTCTAAAGTAGCTGTAACACCAACCACCTCTCGATGTAGT
 AGACAATCTGACCTTAATGGTAGAACACACTGACATTCTGAAGCTAGTCCAGCTAGTACACCAC
 AAATCATTAAGCATAAAGCCTTAGACTTAGATGACAGATGGCAATTCAAGAGATCTCGGTGTTA
 GATACACAAGACAAACGATCTAAAGCAAATACTGGTAGTAGTAACCAAGATAAAGCATCCAAAT
 GAGCAGCCCAGAAACAGATGAAGAAATTGAAAAGATGAAGGGTTTTGGTGAATATTCACGGTCTC
 CTACATTTTGATCCTTTTAACCTTACAAGGAGATTTTTTTATTTGGCTGATGGGTAAAGCCAAAC
 ATTTCTATGTTTTTACTATGTTGAGCTACTTGACAGTAAGTTCATTTGTTTTTACTATGTTCCAC
 TGTTTGAGTAATACACAGATAACTCTTAGTGCATTTACTTCACAAAGTACTTTTTCAAACATCA
 GATGCTTTTATTTCCAAACCTTTTTTTCACCTTTCACCTAAGTTGTTGAGGGGAAGGCTTACACAG
 ACACATTCTTTAGAATTGGAAAAGTGAGACCAGGCACAGTGGCTCACACCTGTAATCCCAGCACT
 TAGGGAAGACAAGTCAGGAGGATTGATTGAAGCTAGGAGTTAGAGACCAGCTGGGCAACGTATT
 GAGACCATGTCTATTAAAAAATAAAATGGAAAAGCAAGAATAGCCTTATTTTCAAATATGGAAA
 GAAATTTATATGAAAATTTATCTGAGTCATTAAATTTCTCCTTAAGTGATACTTTTTTAGAAGTA
 CATTATGGCTAGAGTTGCCAGATAAAATGCTGGATATCATGCAATAAATTTGCAAAACATCATCT
 AAAATTTAAAAA

FIGURE 22

MEGESTSAVLSGFLGALAFQHLNTSDTEGFLGGEVKGEAKNSITDSQMDDVEVVYTIDIQKYI
PCYQLFSFYNSSGEVNEQALKKILSNVKKNVVGWYKERRHSDQIMTFRERLLHKNLQEHFSNQDL
VFLLLTPSIITESCSTHRLEHSLYKPQKGLFHRVPLVVANLGMSEQLGYKTVSGSCMSTGFSSRAV
QTHSSKFFEEEDGSLKEVHKINEMYASLQEEELKSICKKVEDSEQAVDKLVKDVNRLKREIEKRRGA
QIQAAREKNIQKDPQENIFLCQALRTFFPNSEFLHSCVMSLKNRHVSKSSCNYNHHLDVVDNLTL
MVEHTDIPEASPASTPQIIKHKALDLDLRWQFKRSRLDQDKRSKANTGSSNQDKASKMSSPET
DEEIEKMKGFGEYSRSPTF

Important features:

Signal peptide:

amino acids 1-19

N-glycosylation sites.

amino acids 75-79, 322-326

N-myristoylation site.

amino acids 184-154

Growth factor and cytokines receptors family.

amino acids 134-150

FIGURE 23

GGCACAGCCGCGCGGCGGAGGGCAGAGTCAGCCGAGCCGAGTCCAGCCGGACGAGCGGACCAGCGCAGGGCAGCCCAA
 GCAGCGCGCAGCGAACGCCCGCGCCGCCACACCTCTGCGGTCCCCGCGCGCCTGCCACCCTTCCCTCCTTCCCC
 GCGTCCCCGCTCGCCGCGCAGTCAGCTTGCCGGGTTGCTGCCCCGCGAAACCCGAGGTACACAGCCCGCGCCTCT
 GCTTCCCTGGGCGCGCGCCGCTCCACGCCCTCCTTCTCCCCCTGGCCCGCGCCTGGCACCGGGGACCGTTGCTTGA
 CGCGAGGCCCAGCTCTACTTTTTGCCCCGCGTCTCCTCCGCTGCTCGCCTCTTCCACCAACTCCAACCTCCTTCTCCC
 TCCAGTCCACTCGCTAGTCCCCGACTCCGCCAGCCCTCGGCCGCTGCCGTAGCGCCGCTTCCCGTCCGGTCCCAAA
 GGTGGGAACGCGTCCGCCCCGCGCCGACCAATGGCACGGTTCGGCTTGCCCGCGCTTCTCTGCACCCTGGCAGTGCTC
 AGCGCCGCGCTGCTGGCTGCCGAGCTCAAGTCGAAAAGTTGCTCGGAAGTGCACGCTCTTTACGTGTCCAAAGGCTTC
 AACAAGAACGATGCCCCCTCCACGAGATCAACGGTGATCATTTGAAGATCTGTCCCAGGGTTCTACCTGCTGCTCT
 CAAGAGATGGAGGAGAAGTACAGCCTGCAAAAGTAAAGATGATTTCAAAGTGTGGTCAGCGAACAGTGCAATCATTTG
 CAAGCTGCTTTTGTCTTACGTTACAAGAAGTTTGATGAATCTTCAAAGAACTACTTGAAATGCAGAGAAATCCCTG
 AATGATATGTTTGTGAAGACATATGGCCATTTATACATGCAAAATCTGAGCTATTTAAAGATCTCTTCGTAGAGTTG
 AAACGTTACTACGTGGTGGGAAATGTGAACCTGGAAGAAATGCTAAATGACTTCTGGGCTCGCCTCCTGGAGCGGATG
 TTCCGCTGGTGAACCTCCAGTACCCTTTACAGATGAGTATCTGGAATGTGTGAGCAAGTATACGGAGCAGCTGAAG
 CCCTTCGGAGATGTCCTCGCAAATTGAAGCTCCAGGTTACTCGTGCTTTTGTAGCAGCCCGTACTTTCGCTCAAGGC
 TTAGCGGTTGCGGGAGATGTCGTGAGCAAGGTCTCGTGTTAAACCCACAGCCAGTGTAACCATGCCCTGTTGAAG
 ATGATCTACTGCTCCACTGCCGGGTCTCGTGACTGTGAAGCCATGTTACAACCTACTGCTCAAACATCATGAGAGGC
 TGTTTGGCCAACCAAGGGGATCTCGATTTGAATGGAACAATTTCATAGATGCTATGCTGATGGTGGCAGAGAGGCTA
 GAGGGTCTTTTCAACATTGAATCGGTCAATGGATCCCATCGATGTGAAGATTCTGATGCTATTATGAACATGCAGGAT
 AATAGTGTCAAGTGTCTCAGAAGGTTTTCCAGGGATGTGGACCCCCAAGCCCTCCAGCTGGACGAATTTCTCGT
 TCCATCTCTGAAAGTGCCCTCAGTGCTCGCTTCAGACCACATCACCCGAGGAACGCCCAACCACAGCAGCTGGCACT
 AGTTTGGACCGACTGGTTACTGATGTCAAGGAGAAACTGAAACAGGCCAAGAAATTCTGGTCTCCCTTCCGAGCAAC
 GTTTGCAACGATGAGAGGATGGCTGCAGGAAACGGCAATGAGGATGACTGTTGGAATGGGAAAGGCAAAAGCAGGTAC
 CTGTTTGCAGTGACAGGAAATGGATTAGCCAACCAGGGCAACAACCAGAGGTCCAGGTTGACACCAGCAAACCAGAC
 ATACTGATCCTTCGTCAAATCATGGCTCTTCGAGTGATGACCAGCAAGATGAAGAATGCATACAATGGGAACGACGTG
 GACTTCTTTGATATCAGTGATGAAAGTAGTGGAGAAGGAAGTGAAGTGGCTGTGAGTATCAGCAGTGCCCTTCAGAG
 TTTGACTACAATGCCACTGACCATGCTGGGAAGAGTGCCAATGAGAAAGCCGACAGTGCTGGTGTCCGTCTGGGGCA
 CAGGCCTACCTCCTCACTGTCTTCTGCATCTTGTTCTGTTATGCAGAGAGAGTGGAGATTATAATTCTCAAACCTGAG
 AAAAAGTGTTCATCAAAAAGTTAAAAGGCACCAAGTTATCACTTTTCTACCATCCTAGTGACTTTGCTTTTAAATGAA
 TGGACAACAATGTACAGTTTTTACTATGTGGCCACTGGTTTAAAGAAGTGTGACTTTGTTTTCTCATTGAGTTTTGGG
 AGGAAAAGGGACTGTGCATTGAGTTGGTTCTGCTCCCCAAACCATGTAAACGTGGCTAACAGTGTAGGTACAGAA
 CTATAGTTAGTTGTGATTTGTGATTTTATCACTCTATTATTGTTTGTATGTTTTTTCTCATTTCGTTTGTGGGTT
 TTTTTTCCAACGTGATCTCGCCTGTTTCTTACAAGCAAACAGGGTCCCTTCTTGGCACGTAACATGTACGTATT
 TCTGAAATATTAAATAGCTGTACAGAAGCAGGTTTTATTTATCATGTTATCTATTAAAGAAAAAGCCCAAAAAGC

FIGURE 24

MARFGLPALLCTLAVLSAALLAAELKSKSCSEVRRRLYVSKGFNKNDAPLHEINGDHLKICPQGST
CCSQEMEEEKYSLQSKDDFKSVVSEQCNHLQAVFASRYKKFDEFFKELLENAEKSLNDMFVKTYGH
LYMQNSELFKDLFVELKRYVVGNVNLEMLNDFWARLLERMFLVNSQYHFTDEYLECVSKYTE
QLKPFQDVPRKLKLQVTRAFVAARTFAQGLAVAGDVVSKVSVVNPTAQCTHALLKMIYCSHCRGL
VTVKPCYNYCSNIMRGCLANQGDLD FEWNNFIDAMLMAERLEGPFNIESVMDPIDVKISDAIMN
MQDNSVQVSQKVFQCGPPKPLPAGRISRISSESASFARFRPHHPEERPTTAAGTSLDRLVTDVK
EKLKQAKKFWSSSLPSNVCNDERMAAGNGNEDDCWNGKGSRYLFAVTGNGLANQGNNPEVQVDT
KPDILILRQIMALRVMTSKMKNAYNNGNDVDFDISDESSGEGSGSGCEYQQCPSEFDYNATDHAG
KSANEKADSAGVRPGAQAYLLTVFCILFLVMQREWR

Important features:

Signal peptide:

amino acids 1-22

ATP/GTP-binding site motif A (P-loop).

amino acids 515-524

N-glycosylation site.

amino acids 514-518

Glycosaminoglycan attachment sites.

amino acids 494-498, 498-502

N-myristoylation sites.

amino acids 63-69, 224-230, 276-282, 438-444, 497-503, 531-537

Glypicans proteins.

amino acids 54-75, 105-157, 238-280, 309-346, 423-460, 468-506

FIGURE 25

CTCGCCCTCAAATGGGAACGCTGGCCTGGGACTAAAGCATAGACCACCAGGCTGAGTATCCTGAC
CTGAGTCATCCCCAGGGATCAGGAGCCTCCAGCAGGGAACCTTCCATTATATTCTTCAAGCAACT
TACAGCTGCACCGACAGTTGCGATGAAAGTTCTAATCTCTTCCCTCCTCCTGTTGCTGCCACTAA
TGCTGATGTCCATGGTCTCTAGCAGCCTGAATCCAGGGGTCGCCAGAGGCCACAGGGACCGAGGC
CAGGCTTCTAGGAGATGGCTCCAGGAAGGCGGCCAAGAATGTGAGTGCAAAGATTGGTTCCTGAG
AGCCCCGAGAAGAAAATTTCATGACAGTGTCTGGGCTGCCAAGAAGCAGTGCCCCCTGTGATCATT
TCAAGGGCAATGTGAAGAAAACAAGACACCAAAGGCACCACAGAAAGCCAAACAAGCATTCCAGA
GCCTGCCAGCAATTTCTCAAACAATGTCAGCTAAGAAGCTTTGCTCTGCCTTTGTAGGAGCTCTG
AGCGCCCACTCTTCCAATTAAACATTCTCAGCCAAGAAGACAGTGAGCACACCTACCAGACACTC
TTCTTCTCCCACTCACTCTCCCACTGTACCCACCCCTAAATCATTCCAGTGCTCTCAAAAAGCA
TGTTTTTCAAGATCATTTTGTTTGTTGCTCTCTCTAGTGTCTTCTTCTCTCGTCAGTCTTAGCCT
GTGCCCTCCCCCTTACCCAGGCTTAGGCTTAATTACCTGAAAGATTCCAGGAAACTGTAGCTTCCT
AGCTAGTGTCAATTTAACCTTAAATGCAATCAGGAAAGTAGCAAACAGAAGTCAATAAATATTTTT
AAATGTCAAAAAAAAAAAAAAAAAA

FIGURE 26

MKVLISSLLLLLPLMLMSMVSSSLNPGVARGHRDRGQASRRWLQEGGQECECKDWFLRAPRRKFM
TVSGLPKKQCPCDHFKNVKKTRHQRHHRKPNKHSRACQQFLKQCQLRSFALPL

Important features:

Signal peptide:

amino acids 1-22

N-myristoylation sites.

amino acids 27-33, 46-52

FIGURE 27

GGACGCCAGCGCCTGCAGAGGCTGAGCAGGGAAAAAGCCAGTGCCCCAGCGGAAGCACAGCTCAG
AGCTGGTCTGCCATGGACATCCTGGTCCCACCTCCTGCAGCTGCTGGTGCTGCTTCTTACCCCTGCC
CCTGCACCTCATGGCTCTGCTGGGCTGCTGGCAGCCCCTGTGCAAAAGCTACTTCCCCTACCTGA
TGGCCGTGCTGACTCCCAAGAGCAACCGCAAGATGGAGAGCAAGAAACGGGAGCTCTTCAGCCAG
ATAAAGGGGCTTACAGGAGCCTCCGGGAAAGTGGCCCTACTGGAGCTGGGCTGCGGAACCGGAGC
CAACTTTCAGTTCTACCCACCGGGCTGCAGGGTCACCTGCCTAGACCCAAATCCCACCTTTGAGA
AGTTCCTGACAAAGAGCATGGCTGAGAACAGGCACCTCCAATATGAGCGGTTTGTGGTGGCTCCT
GGAGAGGACATGAGACAGCTGGCTGATGGCTCCATGGATGTGGTGGTCTGCACTCTGGTGTGTG
CTCTGTGCAGAGCCCAAGGAAGGTCTGCAGGAGGTCCGGAGAGTACTGAGACCGGGAGGTGTGC
TCTTTTTCTGGGAGCATGTGGCAGAACCATATGGAAGCTGGGCCTTCATGTGGCAGCAAGTTTTC
GAGCCACCTGGAAACACATTGGGGATGGCTGCTGCCTCACCAGAGAGACCTGGAAGGATCTTGA
GAACGCCCAGTTCTCCGAAATCCAAATGGAACGACAGCCCCCTCCCTTGAAGTGGCTACCTGTG
GGCCCCACATCATGGGAAAGGCTGTCAAACAATCTTTCCCAAGCTCCAAGGCACTCATTTGCTCC
TTCCCCAGCCTCCAATTAGAACAAGCCACCCACCAGCCTATCTATCTTCCACTGAGAGGGACCTA
GCAGAATGAGAGAAGACATTCATGTACCACCTACTAGTCCCTCTCTCCCCAACCTCTGCCAGGGC
AATCTCTAACTTCAATCCCGCCTTCGACAGTGAAAAAGCTCTACTTCTACGCTGACCCAGGGAGG
AAACACTAGGACCCTGTGTATCCTCAACTGCAAGTTTCTGGACTAGTCTCCCAACGTTTGCCTC
CCAATGTTGTCCCTTTCCCTTCGTTCCCATGGTAAAGCTCCTCTCGCTTTCCCTCCTGAGGCTACAC
CCATGCGTCTCTAGGAACTGGTCACAAAAGTCATGGTGCCTGCATCCCTGCCAAGCCCCCTGAC
CCTCTCTCCCCACTACCACCTTCTTCCTGAGCTGGGGGCACCAGGGAGAATCAGAGATGCTGGGG
ATGCCAGAGCAAGACTCAAAGAGGCAGAGGTTTGTTCCTCAAATATTTTTTAATAAATAGACGAA
ACCACG

FIGURE 28

MDILVPLLQLLVLLLTLP LHL MALLGCWQPLCKSYFPYLMAVLTPKSNRKMESKKRELF S QIKGL
TGASGKVALLELGCGTGANFQFYPPGCRVTCLDPNPHFEKFLTKSMAENRHLQYERFVVAPGEDM
RQLADGSMDVVVCTLVLC SVQSPRKVLQEVRRVLRPGGV LFFWEHVAEPYGSWAFMWQQVFEPTW
KHIGDGCCLTRETWKDLENAQFSEIQMERQPPPLKWLPVGPHIMGKAVKQSF PSSKALICSFPSL
QLEQATHQPIY LPLRGT

Important features:

Signal peptide:

amino acids 1-23

Leucine zipper pattern.

amino acids 10-32

N-myristoylation sites.

amino acids 64-70, 78-84, 80-86, 91-97, 201-207

FIGURE 29

CAATGTTTGCCTATCCACCTCCCCCAAGCCCCCTTTACCTATGCTGCTGCTAACGCTGCTGCTGCT
GCTGCTGCTGCTTAAAGGCTCATGCTTGGAGTGGGGACTGGTCGGTGCCAGAAAGTCTCTTCTG
CCACTGACGCCCCCATCAGGGATTGGGCCTTCTTTCCCCCTTCCTTTCTGTGTCTCCTGCCTCAT
CGGCCTGCCATGACCTGCAGCCAAGCCCAGCCCCGTGGGGAAGGGGAGAAAGTGGGGGATGGCTTA
AGAAAGCTGGGAGATAGGGAACAGAAGAGGGTAGTGGGTGGGCTAGGGGGGCTGCCTTATTTAAA
GTGGTTGTTTATGATTCTTATACTAATTTATACAAAGATATTAAGGCCCTGTTCAATTAAGAAATT
GTTCCCTTCCCCTGTGTTCAATGTTTGTAAAGATTGTTCTGTGTAAATATGTCTTTATAATAAAC
AGTTAAAAGCTGAAAAAAAAAAAAAAAAAAAAAAAAAAAAA

FIGURE 30

MLLTLLLLLLLLLKGSCLEWGLVGAQKVSSATDAPIRDWAFFPPSFLCLLPHRPAMTCSQAQPRG
EGEKVGDG

Important features:

Signal peptide:

amino acids 1-15

Growth factor and cytokines receptors family:

amino acids 3-18

FIGURE 31

GTTTGAATTCCTTCAACTATACCCACAGTCCAAAAGCAGACTCACTGTGTCCCAGGCTACCAAGTT
 CCTCCAAGCAAGTCATTTCCCTTATTTAACCGATGTGTCCCTCAAACACCTGAGTGCTACTCCCT
 ATTTGCATCTGTTTTGATAAATGATGTTGACACCCTCCACCGAATTCTAAGTGGAAATCATGTCGG
 GAAGAGATACAATCCTTGGCCTGTGTATCCTCGCATTAGCCTTGTCTTTGGCCATGATGTTTACC
 TTCAGATTTCATCACCACCCTTCTGGTTCACATTTTCATTTTCATTGGTTATTTTGGGATTGTTGTT
 TGTCTGCGGTGTTTTATGGTGGCTGTATTATGACTATACCAACGACCTCAGCATAGAATTGGACA
 CAGAAAGGGAAAATATGAAGTGCGTGCTGGGGTTTGCTATCGTATCCACAGGCATCACGGCAGTG
 CTGCTCGTCTTGATTTTTGTTCTCAGAAAGAGAATAAAATTGACAGTTGAGCTTTTCCAAATCAC
 AAATAAAGCCATCAGCAGTGCTCCCTTCCTGCTGTTCCAGCCACTGTGGACATTTGCCATCCTCA
 TTTTCTTCTGGGTCTCTGGGTGGCTGTGCTGCTGAGCCTGGGAAGTGCAGGAGCTGCCCAGGTT
 ATGGAAGGCGGCCAAGTGAATATAAGCCCCCTTCGGGCATTTCGGTACATGTGGTCGTACCATTT
 AATTGGCCTCATCTGGACTAGTGAATTCATCCTTGCGTGCCAGCAAATGACTATAGCTGGGGCAG
 TGGTTACTTGTTATTTCAACAGAAGTAAAAATGATCCTCCTGATCATCCCATCCTTTCGTCTCTC
 TCCATTCTCTTCTTCTACCATCAAGGAACCGTTGTGAAAGGGTCATTTTTAATCTCTGTGGTGAG
 GATTCCGAGAATCATTGTCTATGATGCAAAACGCACTGAAAGAACAGCAGCATGGTGCATTGT
 CCAGGTACCTGTTCCGATGCTGCTACTGCTGTTTCTGGTGTCTTGACAAATACCTGCTCCATCTC
 AACCAGAATGCATATACTACAACCTGCTATTAATGGGACAGATTTCTGTACATCAGCAAAAGATGC
 ATTCAAAATCTTGTCCAAGAAGTCAAGTCACCTTTACATCTATTAAGTCTTTGGAGACTTCATAA
 TTTTCTAGGAAAGGTGTTAGTGGTGTGTTTCACTGTTTTTGGAGGACTCATGGCTTTTAACTAC
 AATCGGGCATTCCAGGTGTGGGCAGTCCCTCTGTTATTGGTAGCTTTTTTGCCTACTTAGTAGC
 CCATAGTTTTTTATCTGTGTTTGAAACTGTGCTGGATGCACTTTTCTGTGTTTTGCTGTTGATC
 TGAAACAAATGATGGATCGTCAGAAAAGCCCTACTTTATGGATCAAGAATTTCTGAGTTTCGTA
 AAAAGGAGCAACAAATTAACAATGCAAGGGCACAGCAGGACAAGCACTCATTAAGGAATGAGGA
 GGGAACAGAACTCCAGGCCATTGTGAGATAGATACCCATTTAGGTATCTGTACCTGGAAAACATT
 TCCTTCTAAGAGCCATTTACAGAATAGAAGATGAGACCACTAGAGAAAAGTTAGTGAATTTTTT
 TTAAGACCTAATAAACCTATTCTTCCTCAAAA

FIGURE 32

MSGRDTILGLCILALALSLAMMFTFRFITLLVHIFISLVILGLLFVCGVLWWLYDYTDLSIE
LDTERENMKCVLGFAIVSTGITAVLLVLI FVLRKRIKLTVELFQITNKAISSAPFLLFQPLWTFE
ILIFFWVLWVAVLLSLGTAGAAQVMEGGQVEYKPLSGIRYMWSYHLIGLIWTSEFILACQOMTIA
GAVVTCYFNRSKNDPPDHPILSSLSILFFYHQGTVVKGSFLISVVRIPRIIVMYMQNALKEQQHG
ALSRYLFRCCYCCFWCLDKYLLHLNQAYTTTAINGTDFCTSAKDAFKILSKNSSHFTSINCFGD
FIIFLGKVLVVCFTVFGGLMAFNYNRAFQVWAVPLLLVAFFAYLVAHSFLSVFETVLDALFLCFA
VDLETNDGSSEKPYFMDQEFLSFVKRSNKLNNARAQQDKHSLRNEEGTELQAIVR

Important features:

Signal peptide:

amino acids 1-20

Putative transmembrane domains:

amino acids 35-54, 75-97, 126-146, 185-204, 333-350, 352-371

N-glycosylation sites.

amino acids 204-208, 295-299, 313-317

N-myristoylation sites.

amino acids 147-153, 178-184, 196-202, 296-275, 342-348

FIGURE 33

[illegible]

FIGURE 34

MRTVVLTMKASVIEMFLVLLVTGVHSNKETAKKIKRPKFTVPQINCDVKAGKIIDPEFIVKCPAG
CQDPKYHVGTDVYASYSSVCGAAVHSGVLDNSGGKILVRKVAGQSGYKGSYSNGVQSLSLPRWR
ESFIVLESKPKKGVITYPSALTYSSSKSPAAQAGETTKAYQRPPIPGTTAQPVTLMQLLAVTVAVA
TPTTLPRPSPSAASTTSIPRPQSVGHRSEQEMDLWSTATYTTSSQNRPRADPGIQRQDPSGAAFQKP
VGADVSLGLVPKEELSTQSLPEVSLGDPNCKIDLSFLIDGSTSIGKRRFRIQKQLLADVAQALDI
GPAGPLMGVVQYGDNPATHFNLKTHNTSRDLKTAIEKITQRGGLSNVGRAISFVTKNFFSKANGN
RSGAPNVVVVMVDGWPTDKVEEASRLARESGINIFFITIEGAAENEKQYVVEPNFANKAVCRTNG
FYSLHVQSWFGLHKTLLQPLVKRVCDTDRILACSKTCLNSADIGFVIDGSSSVGTGNFRTVLQFVTN
LTKEFEISDTRIGAVQYTYEQRLFFGFDKYSSKPDILNAIKRVGYWSGGTSTGAAINFALEQL
FKKSKPNKRKLMILITDGRSYDDVRIPAMAAHLKGVITYAIGVAWAAQEELEVIATHPARHDSFF
VDEFDNLHQYVPRIIQNICTEFN SQPRN

Important features:

Signal peptide:

amino acids 1-26

Transmembrane domain:

amino acids 181-200

N-glycosylation sites.

amino acids 390-394, 520-524

N-myristoylation sites.

amino acids 23-29, 93-99, 115-121, 262-268, 367-373, 389-395,
431-437, 466-472, 509-515, 570-576, 571-577, 575-581, 627-633

Amidation site.

amino acids 304-308

FIGURE 35

CCGAGCACAGGAGATTGCCTGCGTTTAGGAGGTGGCTGCGTTGTGGGAAAAGCTATCAAGGAAGAAATTGC
CAAACCATGTCTTTTTTCTGTTTTTCAGAGTAGTTCACAACAGATCTGAGTGTTTTAATTAAGCATGGAAT
ACAGAAAACAACAAAAAAGCTTAAGCTTTAATTTTCATCTGGAATTCACAGTTTTCTTAGCTCCCTGGACCC
GGTTGACCTGTTGGCTCTTCCCGCTGGCTGCTCTATCACGTGGTGCTCTCCGACTACTCACCCCGAGTGTA
AAGAACCTTCGGCTCGCGTGCTTCTGAGCTGCTGTGGATGGCCTCGGCTCTCTGGACTGTCCTTCCGAGTA
GGATGTCACTGAGATCCCTCAAATGGAGCCTCCTGCTGCTGTCACTCCTGAGTTTCTTTGTGATGTGGTAC
CTCAGCCTTCCCCACTACAATGTGATAGAACGCGTGAAGTGGATGTACTTCTATGAGTATGAGCCGATTTA
CAGACAAGACTTTCACTTCACACTTCGAGAGCATTCAAAGTCTCTCATCAAAATCCATTTCTGGTCATTC
TGGTGACCTCCCAACCTTCAGATGTGAAAGCCAGGCAGGCCATTAGAGTTACTTGGGGTGAAAAAAGTCT
TGGTGGGGATATGAGGTTCTTACATTTTTCTTATTAGGCCAAGAGGCTGAAAAGGAAGACAAAATGTTGGC
ATTGTCTTTAGAGGATGAACACCTTCTTTATGGTGACATAATCCGACAAGATTTTTTAGACACATATAATA
ACCTGACCTTGAAAACCATTTATGGCATTGAGGTGGGTAACTGAGTTTTGCCCAATGCCAAGTACGTAATG
AAGACAGACACTGATGTTTTTCATCAATACTGGCAATTTAGTGAAGTATCTTTTAAACCTAAACCACTCAGA
GAAGTTTTTTCAGAGTTATCCTCTAATTGATAATTATTCCTATAGAGGATTTTACCAAAAACCCATATTT
CTTACCAGGAGTATCCTTTCAAGGTGTTCCCTCCATACTGCAGTGGGTGGGTTATATAATGTCCAGAGAT
TTGGTGCCAAGGATCTATGAAATGATGGGTACGTAAGCAACCCATCAAGTTTGAAGATGTTTATGTGGGAT
CTGTTTGAATTTATTAAAGTGAACATTCATATTCAGAAGACACAAATCTTTTCTTTCTATATAGAATCC
ATTTGGATGTCGTCAACTGAGACGTGTGATTGCAGCCCATGGCTTTTCTTCCAAGGAGATCATCACTTTT
TGGCAGGTCATGCTAAGGAACACCACATGCCATTATTAACTTCACATTCTACAAAAGCCTAGAAGGACAG
GATACCTTGTGGAAAGTGTTAAATAAAGTAGGTACTGTGGAAAATTCATGGGGAGGTCAGTGTGCTGGCTT
ACACTGAACTGAAACTCATGAAAAACCCAGACTGGAGACTGGAGGGTTACACTTGTGATTTATTAGTCAGG
CCCTTCAAAGATGATATGTGGAGGAATTAAATATAAAGGAATTGGAGGTTTTTGTCTAAAGAAATTAATAGG
ACCAACAATTTGGACATGTCATTCTGTAGACTAGAATTTCTTAAAAGGGTGTTACTGAGTTATAAGCTCA
CTAGGCTGTAAAAACAAACAATGTAGAGTTTTATTTATTGAACAATGTAGTCACTTGAAGGTTTTGTGTA
TATCTTATGTGGATTACCAATTTAAAAATATATGTAGTTCTGTGTCAAAAACCTTCTTCACTGAAGTTATA
CTGAACAAAATTTTACCTGTTTTTGGTCATTTATAAAGTACTTCAAGATGTTGCAGTATTTACAGTTATT
ATTATTTAAAATTTACTTCAACTTTGTGTTTTTAAATGTTTTGACGATTTCAATACAAGATAAAAAGGATAG
TGAATCATTCTTTACATGCAACATTTTCCAGTTACTTAACTGATCAGTTTATTATTGATACATCACTCCA
TTAATGTAAAGTCATAGGTCATTATTGCATATCAGTAATCTCTTGGACTTTGTTAAATATTTTACTGTGGT
AATATAGAGAAGAATTAAAGCAAGAAAATCTGAAAA

FIGURE 36

MASALWTVLP SRMSLRSLKWSLLLLLSLLSFFVMWYLSLPHYNVIERVNW MYFYEYEP IYRQDFHF
TLREHSNC SHQNPFLVILVTSHPSDV KARQAIRVTWGEKKS WWGYEVLTF FLLGQEA EKEDKMLA
LSLEDEHLLYGD IIRQDFLD TYNNLT LKTIMAFRWVTEFCPNAKYVMKTD TDVFINTGNLVKYL
NLNHSEKFFTGYPLIDNYSYRGFYQKTHISYQEYFPKVFP PYCSGLGYIMSRDLVPRIYEMMGHV
KPIKFEDVYVGICL NLLKVNIHIPEDTNLFFLYRIHLDVCQLRRVIAAHGFSSKEIITFWQVMLR
NTTCHY

Important features:

Type II transmembrane domain:

amino acids 20-39

N-glycosylation sites.

amino acids 72-76, 154-158, 198-202, 212-216, 326-330

Glycosaminoglycan attachment site.

amino acids 239-243

Ly-6 / u-PAR domain proteins.

amino acids 23-37

N-myristoylation site.

amino acids 271-277

FIGURE 37

CGCTCGGGCACCAGCCGCGGCAAGGATGGAGCTGGGTTGCTGGACGCAGTTGGGGCTCACTTTTCTTCAGCTCCTTCTCATC
TCGTCCTTGCCAAGAGAGTACACAGTCATTAATGAAGCCTGCCCTGGAGCAGAGTGAATATCATGTGTGCGGGAGTGCTGTG
AATATGATCAGATTGAGTGCCTCTGCCCCGGAAGAGGGAGTCTGGGTTATACCATCCCTTGCTGCAGGAATGAGGAGAA
TGAGTGTGACTCCTGCCTGATCCACCCAGGTTGTACCATCTTTGAAAAGTCAAGAGCTGCCGAAATGGCTCATGGGGGGGT
ACCTTGATGACTTCTATGTGAAGGGTTCTACTGTGCAGAGTCCGAGCAGGCTGGTACGGAGGAGACTGCATGCGATGTG
GCCAGGTTCTGCGAGCCCCAAAGGGTCAGATTTTGTGGAAAGCTATCCCTAAATGCTCACTGTGAATGGACCATTTCATGC
TAAACCTGGGTTTGTATCCAACTAAGATTTGTATGTTGAGTCTGGAGTTTGACTACATGTGCCAGTATGACTATGTTGAG
GTTCTGTATGGAGACAACCGCATGGCCAGATCATCAAGCGTGTCTGTGGCAACGAGCGGCCAGCTCCTATCCAGAGCATAG
GATCCTCACTCCAGCTCCTCTTCCACTCCGATGGGCTCCAGAATTTTGACGGTTTCCATGCCATTTATGAGGAGATCACAGC
ATGCTCCTCATCCCCTTGTTTCCATGACGGCACGTGCGTCTTGACAAGGCTGGATCTTACAAGTGTGCCTGCTTGGCAGGC
TATACTGGGCAGCGCTGTGAAAATCTCCTTGAGAAAGAACTGCTCAGACCCTGGGGGCCAGTCAATGGGTACCAGAAAA
TAACAGGGGGCCTGGGCTTATCAACGGACGCCATGCTAAAATTGGCACCGTGGTGTCTTTCTTTTGAACACTCCTATGT
TCTTAGTGGCAATGAGAAAAGAACTTGCCAGCAGAAATGGAGAGTGGTCAGGGAACAGCCCATCTGCATAAAAGCCTGCCGA
GAACCAAGATTTAGACCTGGTGAGAAGGAGAGTTCTTCCGATGCAGGTTCACTCAAGGAGACACCATTACACCAGCTAT
ACTCAGCGGCCCTTCAGCAAGCAGAACTGCAGAGTGCCCTACCAAGAAGCCAGCCCTTCCCTTTGGAGATCTGCCCATGGG
ATACCAACATCTGCATACCCAGCTCCAGTATGAGTGCATCTACCCCTTCTACCGCCGCTGGGCAGCAGCAGGAGGACATGT
CTGAGGACTGGGAAGTGGAGTGGCGGGCACCATCTGCATCCCTATCTGCGGGAAAAATGAGAACATCACTGCTCCAAAGA
CCCAAGGGTTGCGCTGGCCGTGGCAGGCAGCATCTACAGGAGGACCAGCGGGGTGCATGACGGCAGCCTACACAAGGGAGC
GTGGTTCTTAGTCTGCAGCGGTGCCCTGGTGAATGAGCGCACTGTGGTGGTGGCTGCCCACTGTGTTACTGACCTGGGGAG
GTCACCATGATCAAGACAGCAGACCTGAAAGTTGTTTGGGGAAATTCACCGGATGATGACCGGGATGAGAAGACCATCC
AGAGCCTACAGATTTCTGCTATCATTCTGCATCCCAACTATGACCCCATCTGCTTGATGCTGACATGCCATCCTGAAGCT
CCTAGACAAGGCCCGTATCAGCACCCGAGTCCAGCCATCTGCCTCGCTGCCAGTCGGGATCTCAGCACTTCTTCCAGGAG
TCCCACATCACTGTGGCTGGCTGGAAATGTCTGGCAGACGTGAGGAGCCCTGGCTTCAAGAACGACACACTGCGCTCTGGGG
TGGTCAGTGTGGTGGACTCGCTGCTGTGTGAGGAGCAGCATGAGGACCATGGCATCCAGTGAAGTGTCACTGATAACATGTT
CTGTGCCAGCTGGGAACCCACTGCCCTTCTGATATCTGCACTGCAGAGACAGGAGGCATCGCGGCTGTGTCTTCCCGGGA
CGAGCATCTCCTGAGCCACGCTGGCATCTGATGGGACTGGTCAGCTGGAGCTATGATAAAACATGCAGCCACAGGCTCTCCA
CTGCCCTTACCAAGGTGCTGCCTTTTAAAGACTGGATTGAAAGAAATATGAAATGAACCATGCTCATGCACTCCTTGAGAAG
TGTTTCTGTATATCCGTCTGTACGTGTTCATTGCGTGAAGCAGTGTGGGCTGAAGTGTGATTTGGCCTGTGAAGTTGGCT
GTGCCAGGGCTTCTGACTTCAGGGACAAAACCTCAGTGAAGGGTGAGTAGACCTCCATTGCTGGTAGGCTGATGCCGCTCCA
CTACTAGGACAGCCAATTGGAAGATGCCAGGGCTTGCAAGAAGTAAGTTTCTTCAAAGAAGACCATATACAAAACCTCTCCA
CTCCACTGACCTGGTGGTCTTCCCAACTTTTCAGTTATACGAATGCCATCAGCTTGACCAGGGAAGATCTGGGCTTCATGAG
GCCCCTTTGGAGCTCTCAAGTTCTAGAGAGCTGCCTGTGGGACAGCCAGGGCAGCAGAGCTGGGATGTGGTGCATGCCTT
TGTTGTACATGGCCACAGTACAGTCTGGTCCCTTTTCCCTCCCCATCTCTTGTACACATTTTAATAAAATAAGGGTTGGCTTCT
GAACACAAAAA
AAAAA

FIGURE 38

MELGCWTQLGLTFLQLLLISSLPREYTVINEACPGAENIMCRECCEYDQIECVCPGKREVVGYT
IPCCRNEENECDSCLIHPGCTIFENCKSCRNGSWGGLDDFYVKGIFYCAECRAGWYGGDCMRCGQ
VLRAPKGQILLESYPLNAHCEWTIHAKPGFVIQLRFVMLSLEFDYMCQYDYVEVRDGDNRDGOII
KRVCGNERPAPIQSIGSSSLHVLFHSDGSKNFDGFHAIYEEITACSSSPCFHDGTCVLDKAGSYKC
ACLAGYTGQRCENLLEERNCSDPGGPVNGYQKITGGPGLINGRHAKIGTVVSFFCNSYVLSGNE
KRTCQONGEWSGKQPICIKACREPKISDLVRRRVLPQMVSRETPLHQLYSAAFSKQKLQSAPTK
KPALPFGDLPMGYQHLHTQLQYECISPFYRRLGSSRRRTCLRTGKWSGRAPSCIPICGKIENITAP
KTQGLRWPWQAAIYRRTSGVHDGSLHKGAWFLVCSGALVNERTVVVAHCVTDLGKVTMIKTADL
KVVLGKFYRDDDRDEKTIQSLQISAILHPNYDPILLDADIAILKLLDKARISTRVQPICLAASR
DLSTSFQESHITVAGWNVVLADVRSPGFKNDTLRSGVSVSVVDSLLCEEQHEDHGIPVSVTDNMFCA
SWEPTAPSDICTAETGGIAAVSFPGRASPEPRWHLMGLVSWSYDKTCSHRLSTAFTKVLPFKDWI
ERNMK

Important features of the protein:

Signal peptide:

amino acids 1-23

EGF-like domain cysteine pattern signature.

amino acids 260-272

N-glycosylation sites.

amino acids 96-100, 279-283, 316-320, 451-455, 614-618

N-myristoylation sites.

amino acids 35-41, 97-103, 256-262, 284-290, 298-304, 308-314,
474-480, 491-497, 638-644, 666-672

Amidation site.

amino acids 56-60

Serine proteases, trypsin family.

amino acids 489-506

CUB domain proteins profile.

amino acids 150-167

FIGURE 39

GGTTCCTACATCCTCTCATCTGAGAATCAGAGAGCATAATCTTCTTACGGGCCCGTGATTTATTAACGTGGCTTAATC
TGAAGGTTCTCAGTCAAATCTTTGTGATCTACTGATTGTGGGGGCATGGCAAGGTTTGCTTAAAGGAGCTTGGCTGG
TTTGGGCCCTTAGCTGACAGAAGGTGGCCAGGGAGAATGCAGCAGCTGCTCGGAGAATGAAGGCGCTTCTGTTGC
TGGTCTTGCCCTGGCTCAGTCTGCTAACTACATTGACAATGTGGGCAACCTGCACTTCCTGTATTGAGAACTCTGTA
AAGGTGCCCTCCCACTACGGCCTGACCAAAGATAGGAAGAGGCGCTCACAAGATGGCTGTCCAGACGGCTGTGCGAGCC
TCACAGCCACGGCTCCCTCCCCAGAGGTTTCTGCAGCTGCCACCATCTCCTTAATGACAGACGAGCCTGGCCTAGACA
ACCCTGCCTACGTGTCTCGGCAGAGGACGGGAGCCAGCAATCAGCCCAGTGGACTCTGGCCGGAGCAACCGAACTA
GGGCACGGCCCTTTGAGAGATCCACTATTAGAAGCAGATCATTTAAAAAATAAATCGAGCTTTGAGTGTCTTCGAA
GGACAAAGAGCGGGAGTGCAATTGCCAACCATGCCAGCAGGGCAGGGAAAATTCTGAAAACACCACTGCCCTGAAG
TCTTTCCAAGGTTGTACCACTGATTCCAGATGGTGAAATTACCAGCATCAAGATCAATCGAGTAGATCCAGTGAAA
GCCTCTCTATTAGGCTGGTGGGAGGTAGCGAAACCCCACTGGTCCATATCATTATCCAACACATTTATCGTGATGGGG
TGATCGCCAGAGACGGCCGGCTACTGCCAGGAGACATCATTTCTAAAGGTCAACGGGATGGACATCAGCAATGTCCCTC
ACAACTACGCTGTGCGTCTCCTGCGGCAGCCCTGCCAGGTGCTGTGGCTGACTGTGATGCGTGAACAGAAGTTCCGCA
GCAGGAACAATGGACAGGCCCGGATGCCACAGACCCGAGATGACAGCTTTCATGTGATTCTCAACAAAAGTAGCC
CCGAGGAGCAGCTTGAATAAACTGGTGCGCAAGGTGGATGAGCCTGGGGTTTTCATCTTCAATGTGCTGGATGGCG
GTGTGGCATATCGACATGGTCAGCTTGAGGAGAATGACCGTGTGTTAGCCATCAATGGACATGATCTTCGATATGGCA
GCCCAGAAAGTGCGGCTCATCTGATTAGGCCAGTGAAAGACGTGTTACCTCGTGTGCTCCGCCAGGTTCCGGCAGC
GGAGCCCTGACATCTTTAGGAAGCCGGCTGGAACAGCAATGGCAGCTGGTCCCAGGGCCAGGGGAGAGGAGCAACA
CTCCCAAGCCCTCCATCCTACAATTACTTGTGATGAGAAGGTGGTAAATATCCAAAAGACCCCGGTGAATCTCTCG
GCATGACCGTTCGAGGGGAGCATCACATAGAGAATGGGATTGCGCTATCTATGTCATCAGTGTTGAGCCCGGAGGAG
TCATAAGCAGAGATGGAAGAATAAAACAGGTGACATTTTGTGAATGTGGATGGGTCGAACTGACAGAGGTGAGCC
GGAGTGAGGCAGTGGCATTATTGAAAAGAACATCATCCTCGATAGTACTCAAAGCTTTGGAAGTCAAAGAGTATGAGC
CCCAGGAAGACTGCAGCAGCCAGCAGCCCTGGACTCCAACCACAACATGGCCCCACCCAGTGACTGGTCCCCTATCCT
GGGTGATGTGGCTGGAATTACCACGGTGCTTGTATAACTGTAAAGATATTGTATTACGAAGAAACACAGCTGGAAGTC
TGGGCTTCTGCATTGTAGGAGGTTATGAAGAATACATGGAACAAACCTTTTTTTCATCAAATCCATTGTTGAAGGAA
CACCAGCATACAATGATGGAAGAATTAGATGTGGTGATATTCTTCTTGTGTCATGGTAGAAGTACATCAGGAATGA
TACATGCTTGGCTGGCAAGACTGCTGAAGAAGCTTAAAGGAAGAATTACTCTAACTATTGTTTCTTGGCCTGGCACTT
TTTTATAGAATCAATGATGGGTGAGAGGAAAACAGAAAAATCACAATAGGCTAAGAAGTTGAAACACTATATTTATC
TTGTGAGTTTTTATATTTAAAGAAAGAAATACATTGTAAAAATGTCAGGAAAAGTATGATCATCTAATGAAAGCCAGTT
ACACCTCAGAAAATATGATTTCAAAAAAATTAACCTACTAGTTTTTTTTTCAGTGTGGAGGATTTCTCATTACTCTAC
AACATTGTTTATATTTTTTCTATTCAATAAAAAGCCCTAAAACAACCTAAAATGATTGATTGTATACCCCACTGAATT
CAAGCTGATTTAAATTTAAATTTGGTATATGCTGAAGTCTGCCAAGGGTACATTATGGCCATTTTAAATTTACAGCT
AAAATATTTTTTAAATGCATTGCTGAGAAACGTTGCTTTCATCAAACAAGAATAAATATTTTTTCAGAAGTTAAA

FIGURE 40

MKALLLLVLPWLS PANYIDNVGNLHFLYSELCKGASHYGLTKDRKRRSQDGCPCDGCASLTATAPS
PEVSAAATISLMTDEPGLDNPAYVSSAEDGQPAISPVDSGRSNRTRARPFERSTIRSRSFKKINR
ALSVLRRTKSGSAVANHADQGRESENTTAPEVFPRLYHLIPDGEITSIKINRVDPSSESLSIRLV
GGSETPLVHIIIQHIYRDGVIARDGRLLPGDIILKVNGMDISNVPHNYAVRLLRQPCQVLWLTVM
REQKFRSRNNGQAPDAYRPRDDSFHVILNKSSPEEQLGIKLVRKVDEPGVFIFNVLDGGVAYRHG
QLEENDRVLAINGHDLRYGSPESA AHLIQASERRVHLVVSQRQVRQSPDIFQEAGWNSNGSWSPG
PGRSNTPKPLHPTITCHEKVVNIQKDPGESLGMTVAGGASHREWDLPIYVISVEPGGVISRDGR
IKTGDILLNV DGVELTEVSRSEAVALLKRTSSSIVLKALEVKEYEPQEDCSSPAALDSNHNMAPD
SDWSPSWVMWLELPRCLYNCKDIVLRRNTAGSLGFCIVGGYEEYNGNKPFFIKSIVEGTPAYNDG
RIRCGDILLAVNGRSTSGMIHACLARLLKELKGRITLTIVSWPGTFL

Important features:

Signal peptide:

amino acids 1-15

N-glycosylation sites.

amino acids 108-112, 157-161, 289-293, 384-388

Tyrosine kinase phosphorylation sites.

amino acids 433-441, 492-500

N-myristoylation sites.

amino acids 51-57, 141-147, 233-239, 344-350, 423-429, 447-453,
467-473, 603-609

FIGURE 41

ACCAGGCATTGTATCTTCAGTTGTCAAGTTCGCAATCAGATTGGAAAAGCTCAACTTGAAGCTTT
 CTTGCCTGCAGTGAAGCAGAGAGATAGATATTATTACGTAATAAAAAACATGGGCTTCAACCTGACT
 TTCCACCTTTCTACAAATCCGATTACTGTTGCTGTTGACTTTGTGCCTGACAGTGGTTGGGTGGGC
 CACCAGTAATACTTTCGTGGGTGCCATTCAAGAGATTCTTAAAGCAAAGGAGTTCATGGCTAATTTCC
 ATAAGACCCTCATTTTGGGGAAGGGAAAACTCTGACTAATGAAGCATCCACGAAGAAGGTAGAAGTT
 GACAACTGTCCTTCTGTGTCTCCTTACCTCAGAGGCCAGAGCAAGCTCATTTCAAACCAGATCTCAC
 TTTGGAAGAGGTACAGGCAGAAAATCCCAAAGTGTCAGAGGCCGGTATCGCCCTCAGGAATGTAAAG
 CTTTACAGAGGGTCGCCATCCTCGTTCCCCACCGGAACAGAGAGAAACCTGATGTACCTGCTGGAA
 CATCTGCATCCCTTCTGCAGAGGCAGCAGCTGGATTATGGCATCTACGTCATCCACCAGGCTGAAGG
 TAAAAAGTTTAATCGAGCCAACTCTTGAATGTGGGCTATCTAGAAGCCCTCAAGGAAGAAAATTGGG
 ACTGCTTTTATATCCACGATGTGGACCTGGTACCCGAGAATGACTTTAACCTTTACAAGTGTGAGGAG
 CATCCAAGCATCTGGTGGTTGGCAGGAACAGCACTGGGTACAGGTTACGTTACAGTGGATATTTTGG
 GGGTGTTACTGCCCTAAGCAGAGAGCAGTTTTTCAAGGTGAATGGATTCTCTAACAATACTGGGGAT
 GGGGAGGCGAAGACGATGACCTCAGACTCAGGGTTAGCTCCAAAGAATGAAAATTTCCCGGCCCTG
 CCTGAAGTGGGTAAATATACAATGGTCTTCCACACTAGAGACAAAGGCAATGAGGTGAACGCAGAACG
 GATGAAGCTCTTACACCAAGTGTACGAGTCTGGAGAACAGATGGGTTGAGTAGTTGTTCTTATAAAT
 TAGTATCTGTGGAACACAATCCTTTATATATCAACATCACAGTGGATTTCTGGTTTGGTGCAATGACCC
 TGGATCTTTTGGTGATGTTTGGGAAGAACTGATTCTTTGTTTGCAATAATTTGGCCTAGAGACTTCAA
 ATAGTAGCACACATTAAGAACCTGTTACAGCTCATTGTTGAGCTGAATTTTTCCTTTTGTATTTTCT
 TAGCAGAGCTCCTGGTGATGTAGAGTATAAACAGTTGTAACAAGACAGCTTTCTTAGTCATTTTGAT
 CATGAGGGTTAAATATTGTAATATGGATACTTGAAGGACTTTATATAAAAGGATGACTCAAAGGATAA
 AATGAACGCTATTTGAGGACTCTGGTTGAAGGAGATTTATTTAAATTTGAAGTAATATATTATGGGAT
 AAAAGGCCACAGGAAATAAGACTGCTGAATGTCTGAGAGAACCAGAGTTGTTCTCGTCCAAGGTAGAA
 AGGTACGAAGATACAATACTGTTATTCATTTATCCTGTACAATCATCTGTGAAGTGGTGGTGCAGGT
 GAGAAGGCGTCCACAAAAGAGGGGAGAAAAGGCGACGAATCAGGACACAGTGAAGTTGGGAATGAAGA
 GGTAGCAGGAGGGTGGAGTGTGGCTGCAAAGGCAGCAGTAGCTGAGCTGGTTGCAGGTGCTGATAGC
 CTTCAGGGGAGGACCTGCCAGGTATGCCCTTCAGTGATGCCACCAGAGAATACATTCTCTATTAGT
 TTTTAAAGAGTTTTTGTAAAATGATTTGTACAAGTAGGATATGAATTAGCAGTTTACAAGTTTACAT
 ATTAATAATAATAATATGTCTATCAAATACCTCTGTAGTAAAATGTGAAAAAGCAAAA

FIGURE 42

MGFNLT FHL SYKFRLLLLLTLC LTVVGWATSNYFVGAIQEIPKAKEFMANFHKTLILGKGKTLTN
EASTKKVELDNCPSVSPYLRGQSKLIFKPDLTLEEVQAENPKVSRGRYRPQECKALQRVAILVPH
RNREKHLMYLLEHLHPFLQRQQLDYGIYVIHQAEKGKFNRAKLLNVGYLEALKEENWD CFIFHDV
DLVPENDFNLYKCEEHPKHLVVGRNSTGYRLRYSGYFGGVTALSREQFFKVNGFSNNYWG WGGED
DDLRLRVELQRMKISRPLPEVGKYTMVFHTRDKGNEVNAERMKLLHQVSRVWRTDGLSSCSYKLV
SVEHNPLYINITVDFWFGA

Important features:

Signal peptide:

amino acids 1-27

N-glycosylation sites.

amino acids 4-8, 220-224, 335-339

Xylose isomerase proteins.

amino acids 191-202

FIGURE 43

[illegible]

FIGURE 44

MALSSQIWAACLLLLLLLLASLTSGSVFPQQTGQLAELQPQDRAGARASWMPMFQRRRRRDTHFPI
CIFCCGCCHRSKCGMCCKT

Important features:

Signal peptide:

amino acids 1-24

cAMP- and cGMP-dependent protein kinase phosphorylation site.

amino acids 58-59

N-myristoylation site.

amino acids 44-50

Prokaryotic membrane lipoprotein lipid attachment site.

amino acids 1-12

FIGURE 45

GTGGCTTCATTTTCAGTGGCTGACTTCCAGAGAGCAATATGGCTGGTTCCCCAACATGCCTCACCC
TCATCTATATCCTTTGGCAGCTCACAGGGTCAGCAGCCTCTGGACCCGTGAAAGAGCTGGTCGGT
TCCGTTGGTGGGGCCGTGACTTTCCCCCTGAAGTCCAAAGTAAAGCAAGTTGACTCTATTGTCTG
GACCTTCAACACAACCCCTCTTGTCACCATACAGCCAGAAGGGGGCACTATCATAGTGACCCAAA
ATCGTAATAGGGAGAGAGTAGACTTCCCAGATGGAGGCTACTCCCTGAAGCTCAGCAAACCTGAAG
AAGAATGACTCAGGGATCTACTATGTGGGGATATACAGCTCATCACTCCAGCAGCCCTCCACCCA
GGAGTACGTGCTGCATGTCTACGAGCACCTGTCAAAGCCTAAAGTCACCATGGGTCTGCAGAGCA
ATAAGAATGGCACCTGTGTGACCAATCTGACATGCTGCATGGAACATGGGGAAGAGGATGTGATT
TATACCTGGAAGGCCCTGGGGCAAGCAGCCAATGAGTCCCATAATGGGTCCATCCTCCCCATCTC
CTGGAGATGGGGAGAAAGTGATATGACCTTCATCTGCGTTGCCAGGAACCCTGTCAGCAGAACT
TCTCAAGCCCCATCCTTGCCAGGAAGCTCTGTGAAGGTGCTGCTGATGACCCAGATTCTCCATG
GTCCTCCTGTGTCTCCTGTTGGTGCCCTCCTGCTCAGTCTCTTTGTACTGGGGCTATTTCTTTG
GTTTCTGAAGAGAGAGAGACAAGAAGAGTACATTGAAGAGAAGAAGAGAGTGGACATTTGTCGGG
AAACTCCTAACATATGCCCCATCTGGAGAGAAACACAGAGTACGACACAATCCCTCACACTAAT
AGAACAATCCTAAAGGAAGATCCAGCAAATACGGTTTACTCCACTGTGGAAATACCGAAAAAGAT
GGAAAATCCCCACTCACTGCTCACGATGCCAGACACACCAAGGCTATTTGCCTATGAGAATGTTA
TCTAGACAGCAGTGCCTCCCTAAGTCTCTGCTCA

FIGURE 46

MAGSPTCLTLIYILWQLTGSAASGPVKELVGSVGGAVTFPLKSKVKQVDSIVWTFNTTPLVTIQP
EGGTIIIVTQNRNRERVDFPDGGYSLKLSKLLKNDSGIYYVGIYSSSLQQPSTQEYVLHVYEHLSK
PKVTMGLQSNKNGTCVTNLTCCMEHGEEDEVITWKALGQAANESHNGSILPISWRWGESDMTFIC
VARNPVSRNFSSPILARKLCEGAADDPDSSMVLLCLLLVPLLLSLFVLGLFLWFLKRERQEEYIE
EKKRVDICRETPNICPHSGENTHEYDTIPHTNRTILKEDPANTVYSTVEIPKKMENPHSLLTMPDT
PRLEAYENVI

Important features:

Signal peptide:

amino acids 1-22

Transmembrane domain:

amino acids 224-250

Leucine zipper pattern.

amino acids 229-251

N-glycosylation sites.

amino acids 98-102, 142-146, 148-152, 172-176, 176-180, 204-208,
291-295

FIGURE 47

GGCTCGAGCGTTTCTGAGCCAGGGGTGACCATGACCTGCTGCGAAGGATGGACATCCTGCAATGG
ATTGAGCCTGCTGGTTCTACTGCTGTTAGGAGTAGTTCTCAATGCGATACCTCTAATTGTCAGCT
TAGTTGAGGAAGACCAATTTTCTCAAAACCCCATCTCTTGCTTTGAGTGGTGGTTCAGGAATT
ATAGGAGCAGGTCTGATGGCCATTCCAGCAACAACAATGTCCTTGACAGCAAGAAAAAGAGCGTG
CTGCAACAACAGAACTGGAATGTTTCTTTCATCATTTTTTCAGTGTGATCACAGTCATTGGTGCTC
TGTATTGCATGCTGATATCCATCCAGGCTCTCTTAAAGGTCCTCTCATGTGTAATTCTCCAAGC
AACAGTAATGCCAATTGTGAATTTTCATTGAAAAACATCAGTGACATTCATCCAGAATCCTTCAA
CTTGCAAGTGGTTTTTCAATGACTCTTGTGCACCTCCTACTGGTTTTCAATAAAACCCACCAGTAACG
ACACCATGGCGAGTGGCTGGAGAGCATCTAGTTTCCACTTCGATTCTGAAGAAAAACAAACATAGG
CTTATCCACTTCTCAGTATTTTTAGGTCTATTGCTTGTTGGAATTCTGGAGGTCCTGTTGGGCT
CAGTCAGATAGTCATCGGTTTCCTTGGCTGTCTGTGTGGAGTCTCTAAGCGAAGAAGTCAAATTG
TGTAGTTTAATGGGAATAAAATGTAAGTATCAGTAGTTTGAAAAAAAAA

FIGURE 48

MTCCEGWTS CNGFSL LVLLLLGVVLNAIPLIVSLVEEDQFSQNPISCFEWWFPGIIGAGLMAIPA
TTMSLTARKRACCNRTGMFLSSFFSVITVIGALYCM LISIQALLKGPLMCNPSNSNANCEFSL
KNISDIHPESFNLQWFFNDSCAPPTGFNKPTSNDTMASGWRASSFHFDSEENKHRLIHFSVFLGL
LLVGILEVLFGLSQIVIGFLGCLCGVSKRRSQIV

Important features:

Transmembrane domains:

amino acids 10-31 (type II), 50-72, 87-110, 191-213

N-glycosylation sites.

amino acids 80-84, 132-136, 148-152, 163-167

cAMP- and cGMP-dependent protein kinase phosphorylation site.

amino acids 223-227

N-myristoylation sites.

amino acids 22-28, 54-60, 83-89, 97-103, 216-222

Prokaryotic membrane lipoprotein lipid attachment site.

amino acids 207-218

TNFR/NGFR family cysteine-rich region protein.

amino acids 4-12

FIGURE 49

ATCCGTTCTCTGCGCTGCCAGCTCAGGTGAGCCCTCGCCAAGGTGACCTCGCAGGACACTGGTGA
AGGAGCAGTGAGGAACCTGCAGAGTCACACAGTTGCTGACCAATTGAGCTGTGAGCCTGGAGCAG
ATCCGTGGGCTGCAGACCCCGCCCCAGTGCCTCTCCCCCTGCAGCCCTGCCCCCTCGAACTGTGA
CATGGAGAGAGTGACCCTGGCCCTTCTCCTACTGGCAGGCCTGACTGCCTTGGAAGCCAATGACC
CATTTGCCAATAAAGACGATCCCTTCTACTATGACTGGAAAAACCTGCAGCTGAGCGGACTGATC
TGCGGAGGGCTCCTGGCCATTGCTGGGATCGCGGCAGTTCTGAGTGGCAAATGCAAATACAAGAG
CAGCCAGAAGCAGCACAGTCCTGTACCTGAGAAGGCCATCCCACTCATCACTCCAGGCTCTGCCA
CTACTTGCTTGAGCACAGGACTGGCCTCCAGGGATGGCCTGAAGCCTAACACTGGCCCCCAGCACC
TCCTCCCCTGGGAGGCCTTATCCTCAAGGAAGGACTTCTCTCCAAGGGCAGGCTGTTAGGCCCCCT
TTCTGATCAGGAGGCTTCTTTATGAATTAACTCGCCCCACCACCCCTCA

FIGURE 50

MERVTLALLLLAGLTALEANDPFANKDDPFYYDWKNLQLSGLICGGLLAIAGIAAVLSGKCKYKS
SQKQHSPVPEKAIPPLITPGSATTC

Important features:

Signal peptide:

amino acids 1-16

Transmembrane domain:

amino acids 36-59

N-myristoylation sites.

amino acids 41-47, 45-51, 84-90

Extracellular proteins SCP/Tpx-1/Ag5/PR-1/Sc7.

amino acids 54-67

FIGURE 51

[illegible]

FIGURE 52

MKFQGPPLACLLLLALCLGSGEAGPLQSGEESTGTNIGEALGHGLGDALSEGVGKAIGKEAGGAAGSKVS
EALGQGTREAVGTGVRQVPGFGAADALGNRVGEAAHALGNTGHEIGRQAEDVIRHGADAVRGSWQGV
GHSGAWETSGGHGIFGSQGGGLGGQGNPGGGLGTPWVHGYPGNSAGSFGMNPQGAPWGQGGNGGPPNF
GTNTQGAVAQPGYGSVRASNQNEGCTNPPPSGSGGGSSNSGGGSGSQSGSSGSGSNGDNNNGSSSSGGS
SSGSSSSGSSSGGSSGGSSGSSGSSGSSGSGSRGDSGSESSWGSSTGSSSGNHGSSGGGNGHKPGCEKPGNE
ARGSGESGIQGFRGQGVSSNMREISKEGNRLGGSGLDNYRGQGSWGSGGGDAVGGVNTVNSETSPGM
FNFDTFWKNFKSKLGFINWDAINKDQRSSRIP

Signal peptide:

amino acids 1-21

N-glycosylation site.

amino acids 265-269

Glycosaminoglycan attachment site.

amino acids 235-239, 237-241, 244-248, 255-259, 324-328, 388-392

Casein kinase II phosphorylation site.

amino acids 26-30, 109-113, 259-263, 300-304, 304-308

N-myristoylation site.

amino acids 17-23, 32-38, 42-48, 50-56, 60-66, 61-67, 64-70, 74-80,
90-96, 96-102, 130-136, 140-146, 149-155, 152-158, 155-161,
159-165, 163-169, 178-184, 190-196, 194-200, 199-205, 218-224,
236-242, 238-244, 239-245, 240-246, 245-251, 246-252, 249-252,
253-259, 256-262, 266-272, 270-276, 271-277, 275-281, 279-285,
283-289, 284-290, 287-293, 288-294, 291-297, 292-298, 295-301,
298-304, 305-311, 311-317, 315-321, 319-325, 322-328, 323-329,
325-331, 343-349, 354-360, 356-362, 374-380, 381-387, 383-389,
387-393, 389-395, 395-401

Cell attachment sequence.

amino acids 301-304

FIGURE 53

GGAGAAGAGGTTGTGTGGGACAAGCTGCTCCCGACAGAAGGATGTCGCTGCTGAGCCTGCCCTGG
CTGGGCCTCAGACCGGTGGCAATGTCCCATGGCTACTCCTGCTGCTGGTTGTGGGCTCCTGGCT
ACTCGCCCGCATCCTGGCTTGGACCTATGCCTTCTATAACAAC TGCCGCCGGCTCCAGTGTTTCC
CACAGCCCCCAAACGGAAC TGGTTTTGGGGTCACCTGGGCCTGATCACTCCTACAGAGGAGGGC
TTGAAGGACTCGACCCAGATGTGCGCCACCTATTCCCAGGGCTTTACGGTATGGCTGGGTCCCAT
CATCCCCCTTCATCGTTTTATGCCACCCTGACACCATCCGGTCTATCACCAATGCCTCAGCTGCCA
TTGCACCCAAGGATAATCTCTTCATCAGTTTCCTGAAGCCCTGGCTGGGAGAAGGGATACTGCTG
AGTGGCGGTGACAAGTGGAGCCGCCACCGTCGGATGCTGACGCCCCCTTCCATTTCAACATCCT
GAAGTCCTATATAACGATCTTCAACAAGAGTGCAAACATCATGCTTGACAAGTGGCAGCACCTGG
CCTCAGAGGGCAGCAGTCGTCTGGACATGTTTGAGCACATCAGCCTCATGACCTTGGACAGTCTA
CAGAAATGCATCTTCAGCTTTGACAGCCATTGTCAGGAGAGGCCAGTGAATATATTGCCACCAT
CTTGGAGCTCAGTGCCCTTGTAGAGAAAAGAAGCCAGCATATCCTCCAGCACATGGACTTTCTGT
ATTACCTCTCCCATGACGGGCGGCGCTTCCACAGGGCCTGCCGCCTGGTGATGACTTCACAGAC
GCTGTTCATCCGGGAGCGGCGTCGCACCCTCCCCACTCAGGGTATTGATGATTTTTTCAAAGACAA
AGCCAAGTCCAAGACTTTGGATTTTATTGATGTGCTTCTGCTGAGCAAGGATGAAGATGGGAAGG
CATTGTCAGATGAGGATATAAGAGCAGAGGCTGACACCTTCATGTTTGGAGGCCATGACACCACG
GCCAGTGGCCTCTCCTGGGTCTGTACAACCTTGCGAGGCACCCAGAATACCAGGAGCGCTGCCG
ACAGGAGGTGCAAGAGCTTCTGAAGGACCGCGATCCTAAAGAGATTGAATGGGACGACCTGGCCC
AGCTGCCCTTCCTGACCATGTGCGTGAAGGAGAGCCTGAGGTTACATCCCCCAGCTCCCTTCATC
TCCCGATGCTGCACCCAGGACATTGTTCTCCCAGATGGCCGAGTCATCCCCAAAGGCATTACCTG
CCTCATCGATATTATAGGGGTCCATCACAAACCAACTGTGTGGCCGATCCTGAGGTCTACGACC
CCTTCCGCTTTGACCCAGAGAACAGCAAGGGGAGGTCACCTCTGGCTTTTATTCCCTTTCTCCGCA
GGGCCCAGGAACTGCATCGGGCAGGCGTTTCCCATGGCGGAGATGAAAGTGGTCCTGGCGTTGAT
GCTGCTGCACTTCCGGTTCCTGCCAGACCACACTGAGCCCCGCAGGAAGCTGGAATTGATCATGC
GCGCCGAGGGCGGGCTTTGGCTGCGGGTGGAGCCCCTGAATGTAGGCTTGCACTGACTTTCTGAC
CCATCCACCTGTTTTTTTGCAGATTGTCATGAATAAAACGGTGCTGTCAA

FIGURE 54

MSLLSLPWLGLRPVAMSPWLLLLLVGWSLLARILAWTYAFYNNCRRLQCFPPKRNWFWGHLG
LITPTEEGLKDSTQMSATYSQGFTVWLGPPIPFIVLCHPDTIRSITNASAAIAPKDNLFIRFLKP
WLGEIGILLSGGDKWSRHRMLTPAFHFNILKSYITIFNKSANIMLDKWQHLASEGSSRLDMFEHI
SLMTLDSLQKCIFSFDSHCQERPSEYIATILELSALVEKRSQHILQHMDFLYYLSHDGRRFHRAC
RLVHDFTDVIRERRRTLPTQGIDDFKDKAKSKTLDLDFIDVLLLSKDEDGKALSDEDIRAEADTF
MFGGHDTTASGLSWVLYNLARHPEYQERCQEVQELLKDRDPKEIEWDDLAQLPFLTMCVKESLR
LHPPAPFISRCCTQDIVLPDGRVIPKGITCLIDIIGVHHNPTVWPDPEVYDPFRFDPENSKGRSP
LAFIPFSAGPRNCIGQAFAMAEMKVVLALMLLHFRFLPDHTEPRRKLELIMRAEGGLWLRVEPLN
VGLQ

Important features:

Transmembrane domains:

amino acids 13-32 (type II), 77-102

Cytochrome P450 cysteine heme-iron ligand signature.

amino acids 461-471

N-glycosylation sites.

amino acids 112-116, 168-172

FIGURE 55

[illegible]

FIGURE 56

MGPVKQLKRMFEPTRLIATIMVLLCFALTLCSAFWWHNKGLALIFCILQSLALTWYSLSFIPFAR
DAVKKCFVCLA

Important features:

Signal peptide:

amino acids 1-33

Type II fibronectin collagen-binding domain protein.

amino acids 30-72

FIGURE 57

[illegible]

FIGURE 58

MLCLCLYVPVIGEAEQTEFQYFESKGLPAELKSIFKLSVFIPSQEFSTYRQWKQKIVQAGDKDLDG
QLDFEEFVHYLQDHEKKRLRVFKILDKKNDGRIDAQEIMQSLRDLGVKISEQQAEKILKSMDKNG
TMTIDWNEWRDYHLLHPVENIPEIILYWKHSTIFDVGENLTVPDEFTVEERQTGMWWRHLVAGGG
AGAVSRTCTAPLDRLKVLMOVHASRSNNMGIVGGFTQMIREGGARSLWRNGINVLKIAPESAIK
FMAYEQIKRLVGSDQETLRIHERLVAGSLAGAIAQSSIYPMEVLKTRMALRKTGQYSGMLDCARR
ILAREGVAAFYKGYVPNMLGIIPYAGIDLAVYETLKNAWLQHYAVNSADPGVFVLLACGTMSSTC
GQLASYPLALVRTRMQAQASIEGAPEVTMSSLFKHILRTEGAFGLYRGLAPNFMKVIPAVSISYV
VYENLKITLGVQSR

Important features:

Signal peptide:

amino acids 1-16

Putative transmembrane domains:

amino acids 284-304, 339-360, 376-394

Mitochondrial energy transfer proteins signature.

amino acids 206-215, 300-309

N-glycosylation sites.

amino acids 129-133, 169-173

Elongation Factor-hand calcium-binding protein.

amino acids 54-73, 85-104, 121-140

FIGURE 59

GGAAGGCAGCGGCAGCTCCACTCAGCCAGTACCCAGATACGCTGGGAACCTTCCCCAGCCATGGC
TTCCCTGGGGCAGATCCTCTTCTGGAGCATAATTAGCATCATCATTATTCTGGCTGGAGCAATTG
CACTCATCATTGGCTTTGGTATTTTCAAGGAGACACTCCATCACAGTCACTACTGTGCGCTCAGCT
GGGAACATTGGGGAGGATGGAATCCTGAGCTGCACCTTTTGAACCTGACATCAAACCTTTCTGATAT
CGTGATACAATGGCTGAAGGAAGGTGTTTTAGGCTTGGTCCATGAGTTCAAAGAAGGCAAAGATG
AGCTGTGCGAGCAGGATGAAATGTTTCAAGGCGCGACAGCAGTGTGTGCTGATCAAGTGATAGTT
GGCAATGCCTCTTTGCGGCTGAAAAACGTGCAACTCACAGATGCTGGCACCTACAAATGTTATAT
CATCACTTCTAAAGGCAAGGGGAATGCTAACCTTGAGTATAAACTGGAGCCTTCAGCATGCCGG
AAGTGAATGTGGACTATAATGCCAGCTCAGAGACCTTGGCGGTGTGAGGCTCCCCGATGGTTCCCC
CAGCCCACAGTGGTCTGGGCATCCCAAGTTGACCAGGGAGCCAACTTCTCGGAAGTCTCCAATAC
CAGCTTTGAGCTGAACTCTGAGAATGTGACCATGAAGGTTGTGTCTGTGCTCTACAATGTTACGA
TCAACAACACATACTCCTGTATGATTGAAAATGACATTGCCAAAGCAACAGGGGATATCAAAGTG
ACAGAATCGGAGATCAAAAGGCGGAGTCACCTACAGCTGCTAAACTCAAAGGCTTCTCTGTGTGT
CTCTTCTTTCTTTGCCATCAGCTGGGCACCTTCTGCCTCTCAGCCCTTACCTGATGCTAAAATAAT
GTGCCTTGGCCACAAAAAAGCATGCAAAGTCATTGTTACAACAGGGATCTACAGAACTATTTTAC
CACCAGATATGACCTAGTTTTATATTTCTGGGAGGAAATGAATTCATATCTAGAAGTCTGGAGTG
AGCAAACAAGAGCAAGAAACAAAAAGAAGCCAAAAGCAGAAGGCTCCAATATGAACAAGATAAAT
CTATCTTCAAAGACATATTAGAAGTTGGGAAAATAATTCATGTGAAC TAGACAAGTGTGTTAAGA
GTGATAAGTAAATGCACGTGGAGACAAGTGCATCCCCAGATCTCAGGGACCTCCCCCTGCCTGT
CACCTGGGGAGTGAGAGGACAGGATAGTGCATGTTCTTTGTCTCTGAATTTTTAGTTATATGTGC
TGTAATGTTGCTCTGAGGAAGCCCCCTGGAAAGTCTATCCCAACATATCCACATCTTATATTCCAC
AAATTAAGCTGTAGTATGTACCCTAAGACGCTGCTAATTGACTGCCACTTCGCAACTCAGGGGCG
GCTGCATTTTAGTAATGGGTCAAATGATTCACTTTTTTATGATGCTTCCAAAGGTGCCTTGGCTTC
TCTTCCCAACTGACAAATGCCAAAGTTGAGAAAAATGATCATAATTTTAGCATAAACAGAGCAGT
CGGGGACACCGATTTTATAAATAAACTGAGCACCTTCTTTTTAAACAAAAAAAAAAAAAAAAAAAA
AAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAA

FIGURE 6o

MASLGQILFWSIISIISIIIIILAGAIALIIGFGISGRHSITVTTVASAGNIGEDGILSCTFEPDIKLS
DIVIQWLKEGVLGLVHEFKEGKDELSEQDEMFRGRTAVFADQVIVGNASRLKKNVQLTDAGTYKC
YIITSKGKGKNANLEYKTGAFSMPEVNVDYNASSETLRCEAPRWFPQPTVVWASQVDQGANFSEVS
NTSFELNSENVTMKVVSVLNVNTINNTYSCMIENDIAKATGDIKVTESI KRRSHLQLLNSKASL
CVSSFFAISWALLPLSPYMLK

Important features:

Signal peptide:

amino acids 1-28

Transmembrane domain:

amino acids 258-281

N-glycosylation sites.

amino acids 112-116, 160-164, 190-194, 196-200, 205-209, 216-220,
220-224

N-myristoylation sites.

amino acids 52-58, 126-132, 188-194

FIGURE 61

TGACGTCAGAATCACCATGGCCAGCTATCCTTACCGGCAGGGCTGCCCAGGAGCTGCAGGACAAG
CACCAGGAGCCCCCTCCGGGTAGCTACTACCCTGGACCCCCCAATAGTGGAGGGCAGTATGGTAGT
GGGCTACCCCTGGTGGTGGTTATGGGGGTCTGCCCTGGAGGGCCTTATGGACCACCAGCTGG
TGGAGGGCCCTATGGACACCCCAATCCTGGGATGTTCCCCTCTGGAACCTCAGGAGGACCATATG
GCGGTGCAGCTCCCGGGGGCCCCCTATGGTGCAGCCACCTCCAAGTTCTACGGTGCCCAGCAGCCT
GGGCTTTATGGACAGGGTGGCGCCCCCTCCCAATGTGGATCCTGAGGCCTACTCCTGGTTCCAGTC
GGTGGACTCAGATCACAGTGGCTATATCTCCATGAAGGAGCTAAAGCAGGCCCTGGTCAACTGCA
ATTGGTCTTCATTCAATGATGAGACCTGCCTCATGATGATAAACATGTTTGACAAGACCAAGTCA
GGCCGCATCGATGTCTACGGCTTCTCAGCCCTGTGGAAATTCATCCAGCAGTGGAAGAACCCTCTT
CCAGCAGTATGACCGGGACCGCTCGGGCTCCATTAGCTACACAGAGCTGCAGCAAGCTCTGTCCC
AAATGGGCTACAACCTGAGCCCCCAGTTCAACCAGCTTCTGGTCTCCCGCTACTGCCACGCTCT
GCCAATCCTGCCATGCAGCTTGACCGCTTCATCCAGGTGTGCACCCAGCTGCAGGTGCTGACAGA
GGCCTTCCGGGAGAAGGACACAGCTGTACAAGGCAACATCCGGCTCAGCTTCGAGGACTTCGTCA
CCATGACAGCTTCTCGGATGCTATTGACCCAACCATCTGTGGAGAGTGGAGTGCACCAGGGACCTT
TCCTGGCTTCTTAGAGTGAGAGAAGTATGTGGACATCTCTTCTTTCTGTCCCTCTAGAAGAAC
ATTCTCCCTTGCTTGATGCAACACTGTTCCAAAAGAGGGTGGAGAGTCCTGCATCATAGCCACCA
AATAGTGAGGACCGGGGCTGAGGCCACACAGATAGGGGCCCTGATGGAGGAGAGGATAGAAGTTGA
ATGTCCTGATGGCCATGAGCAGTTGAGTGGCACAGCCTGGCACCAGGAGCAGGTCCTTGTAATGG
AGTTAGTGTCCAGTCAGCTGAGCTCCACCCTGATGCCAGTGGTGAGTGTTTCATCGGCCTGTTACC
GTTAGTACCTGTGTTCCCTCACCAGGCCATCCTGTCAAACGAGCCCATTTTCTCCAAAGTGGAAT
CTGACCAAGCATGAGAGAGATCTGTCTATGGGACCAGTGGCTTGGATTCTGCCACACCCATAAAT
CCTTGTTGTTAACTTCTAGCTGCCTGGGGCTGGCCCTGCTCAGACAAATCTGCTCCCTGGGCAT
CTTTGGCCAGGCTTCTGCCCCCTGCAGCTGGGACCCCTCACTTGCTGCCATGCTCTGCTCGGCT
TCAGTCTCCAGGAGACAGTGGTCACCTCTCCCTGCCAATACTTTTTTAAATTTGCATTTTTTTTC
ATTTGGGGCCAAAAGTCCAGTGAAATTGTAAGCTTCAATAAAAGGATGAAACTCTGA

FIGURE 62

MASYPYRQGCPGAAGQAPGAPPGSYYPGPPNSGGQYGSGLPPGGGYGGPAPGGPYGPPAGGGPYG
HPNPGMFPSGTPGGPYGGAAPGGPYGQPPSSYGAQQPGLYGQGGAPPNVDP EAYSWFQSVDS DH
SGYISMKELKQALVNCNWSSFNDETC LMMINMFDKTKSGRIDVYGFSALWKFIQQWKNLFQQYDR
DRSGSISYTELQQALSQMGYNLSPQFTQLLVSRYCPRSANPAMQLDRFIQVCTQLQVLTEAFREK
DTAVQGNIRLSFEDFVTMTASRML

Important features of the protein:

Signal peptide:

amino acids 1-19

N-glycosylation site.

amino acids 147-150

Casein kinase II phosphorylation sites.

amino acids 135-138, 150-153, 202-205, 271-274

N-myristoylation sites.

amino acids 9-14, 15-20, 19-24, 33-38, 34-39, 39-44, 43-48, 61-
66, 70-75, 78-83, 83-88, 87-92, 110-115

FIGURE 63

[illegible]

FIGURE 64

MQGRVAGSCAPLGLLLVCLHLPGLFARSIGVVEEKVSQNFGTNLPQLGQPSSTGPSNSEHPQPAL
DPRSNDLARVPLKLSVPPSDGFPPAGGSQVQRWPPSWGLPAMDSWPPEDPWQMAAAAEDRLGEA
LPEELSYLSSAAALAPGSGPLPGESSPDATGLSPEASLLHQDSESRRRLPRSNSLGAGGKILSQRP
PWSLIHRVLPDHPWGTNLNPSVSWGGGGPGTGWGTRMPHPHEGIWGINNQPPGTSGWNINRYPGGS
WGNINRYPGGSWGNINRYPGGSWGNIHLYPGINNPFPFPGVLRPPGSSWNIPAGFPNPPSPRLQWG

Important features of the protein:

Signal peptide:

amino acids 1-26

Casein kinase II phosphorylation sites.

amino acids 56-59, 155-158

N-myristoylation sites.

amino acids 48-53, 220-225, 221-226, 224-229, 247-252, 258-263,
259-264, 269-274, 270-275, 280-285, 281-286, 305-310

FIGURE 65

AAGGAGAGGCCACCGGGACTTCAGTGTCTCCTCCATCCCAGGAGCGCAGTGGCCACTATGGGGTC
TGGGCTGCCCCCTTGTCTCCTCTTGACCCTCCTTGGCAGCTCACATGGAACAGGGCCGGGTATGA
CTTTGCAACTGAAGCTGAAGGAGTCTTTTCTGACAAATTCCTCCTATGAGTCCAGCTTCCTGGAA
TTGCTTGAAAAGCTCTGCCTCCTCCTCCATCTCCCTTCAGGGACCAGCGTCACCCTCCACCATGC
AAGATCTCAACACCATGTTGTCTGCAACACATTGACAGCCATTGAAGCCTGTGTCCTTCTTGGCCC
GGGCTTTTGGGCGGGGATGCAGGAGGCAGGCCCCGACCCTGTCTTTCAGCAGGCCCCCACCCTC
CTGAGTGGCAATAAATAAAATTCGGTATGCTG

FIGURE 66

MGSGPLPLVLLLTLLGSSSHGTGPGMTLQLKLKESFLTNSSESSFLELLEKLCLLLHLPSTSVTL
HHARSQHHVVCNT

Important features:

Signal peptide:

amino acids 1-19

N-glycosylation site.

amino acids 37-41

N-myristoylation sites.

amino acids 15-21, 19-25, 60-66

FIGURE 67

ACGGACCGAGGGTTTCGAGGGAGGGACACGGACCAGGAACCTGAGCTAGGTCAAAGACGCCCCGGGC
CAGGTGCCCCGTCGCAGGTGCCCCCTGGCCGGAGATGCGGTAGGAGGGGCGAGCGCGAGAAGCCCC
TTCCTCGGCGCTGCCAACCCGCCACCCAGCCCATGGCGAACCCCGGGCTGGGGCTGCTTCTGGCG
CTGGGCCTGCCGTTCTGCTGGCCCCGCTGGGGCCGAGCCTGGGGGCAAATACAGACCACTTCTGC
AAATGAGAATAGCACTGTTTTGCCTTCATCCACCAGCTCCAGCTCCGATGGCAACCTGCGTCCGG
AAGCCATCACTGCTATCATCGTGGTCTTCTCCCTCTTGCTGCCTTGCTCCTGGCTGTGGGGCTG
GCACTGTTGGTGCGGAAGCTTCGGGAGAAAGCGGCAGACGGAGGGCACCTACCGGCCCAGTAGCGA
GGAGCAGTTCTCCCATGCAGCCGAGGCCCCGGGCCCCCTCAGGACTCCAAGGAGACGGTGCAGGGCT
GCCTGCCCATCTAGGTCCCCCTCTCCTGCATCTGTCTCCCTTCATTGCTGTGTGACCTTGGGGAAA
GGCAGTGCCCTCTCTGGGCAGTCAGATCCACCCAGTGCTTAATAGCAGGGAAGAAGGTACTTCAA
AGACTCTGCCCCCTGAGGTCAAGAGAGGATGGGGCTATTCACTTTTATATATTTATATAAAATTAG
TAGTGAGATGTAAAAAAAAAAAAAAAAAAAAA

FIGURE 68

MANPGLGLLLALGLPFLRLRWGRAWGQIQTTSANENSTVLPSTSSSSDGNLRPEAITAIIVVFS
LLAALLLAVGLALLVRKLREKRQTEGTYRPSSEEQFSHAAEARAPQDSKETVQGCLPI

Important features:

Signal peptide:

amino acids 1-19

Transmembrane domain:

amino acids 56-80

N-glycosylation site.

amino acids 36-40

cAMP- and cGMP-dependent protein kinase phosphorylation site.

amino acids 86-90

Tyrosine kinase phosphorylation site.

amino acids 86-94

N-myristoylation sites.

amino acids 7-13, 26-32

FIGURE 69

GCCAGGAATAACTAGAGAGGAAACA**ATG**GGGTTATTAGAGGTTTGTTCCTCTTAGTTCTGTGCCTGCTGCCACGATCAAATACTTCTTCATTAGCTGAATAATAATGGCTTTGAAGATATTGTATTGTTATAGATCCTAGTGTGCCAGAAATGATGAAAAATAATTAACAAATAGAGGATATGGTGACTACAGCTTCTACGTACTTGTGAAGCGCAAAAAAGATTTTTTTTCAAAAATGTATCTATATTCTGAGAAATTGGAAGGAAATCTCTAGTACAAAAGGCCAAACATGAAACCATAAATCATGCTGATGTTTATAGTTGCACCACCTACACTCCCAGGTAGAGATGAACCATACCCAAGCAGTTCCAGAAATGTGGAGAGAAAGGCGAATACATTCACCTTCACCCCTGACCTTCTACTTGGAAAAAACAAAATGAATATTGGACCAACAGGCAAACCTGTTTGTCCATGAGTGGGCTCACCTCCGGTGGGGAGTGTGTGATGAGTACAATGAAGTACAGCCTTCTACCGCTGTCAAGTCAAAAATCGAAGCAACAAGTGTCCCGCAGGTATCTCTGTGAGAAATAGAGTTTATAAGTGTCAAGGAGGCAGCTGCTTAGTAGAGCATGCAAGATTGATCTACAACAAAATGTATGGAAAAGATTGTCAATTCCTTCTGTATAAAGTACAAACAGAAAAAGCATCCATAATGTTTATGCAAAGTATTGATTCGTGTGTGAATTTGTGAACGAAAAAACCCATAATCAAGAAGCTCCAAGCCTACAAAACATAAAGTGCAATTTAGAAGTACATGGGAGGTGATTAGCAATCTGTAGGAGTTTAAAAAACCCATACCCATGGTGACACCCACTCCTCCACCTGTCTTCATTGGTGAAGTACGTCAAAGAAATGTGTGCTTAGTTCTTGATAAGTCTGGAAGCATGGGGGTAAGGACCGCTAAATCGAATGAATCAAGCAGCAAAACATTTCTGTCTGCAGACTGTTGAAAAATGGATCCTGGGTGGGGATGGTTCACCTTTGATAGTACTGCCACTTATTGTAATAAGCTAAATCAAAATAAAAAAGCAGTGATGAAGAAACACACTCATGGCAGGATTACCTACATATCCTCTGGGAGGAACCTTCATCTGCTGTGAATTAATATGCATTTCCAGGTGATGGAGAGCTACATCCCAACTCGATGGATCGGAAGTACTGCTGCTGACTGATGGGGAGGATAACACTGCAAGTTCTGTATTGATGAAGTGAAACAAAGTGGGGCATTGTTTCATTTATTGCTTTGGGAAGAGCTGCTGATGAAGCAGTAATAGAGATGAGCAAGATAACAGGAGGAAGTCATTTTATGTTTCAGATGAAGCTCAGAACAAATGGCCTCATTTGATGCTTTGGGGCTCTTACATCAGGAAATACGTATCTCTCCAGAGTCCCTTCAGCTCGAAAGTAAGGGATTAAACACTGAATAGTAATGCCCTGGATGAACGACACTGTCTATAATTGATAGTACA GTGGGAAAGGACAGCTTCTTTCTCATCATGGAACAGTCTGCCTCCAGTATTTCTCTCTGGGATCCGAGTGGGAACAATAATGAAAAATTTACAGATGGATGCAACTTCCAAAAATGGCCTATCTCAGTATTCCAGGAACGCAAGGTGGGCACTTGGGCATACAATCTTCAAGCCAAAGCGAACCCAGAAACATTAACTATTACGTAACCTTCTCGAGCGCAAAATCTTCTGTGCCTCCAATCAGATGAATGCTAAAATGAATAAGGCATTAACAGTTTCCCCAGCCCCAATGATTGTTACGCAGAAATTTCTACAGGATATGTACCTGTTCTGGAGCCAATTGACTGCTTCTTATGTAACACAGAATGGACATTACAGAAGTTTGGAACTTTTGGATAATGGTGACAGGCGCTGATTCTTTCAAGAATGATGGAGTCTACTCCAGGTATTTTACAGCATATACAGAAAATGGCAGATATAGCTTAAAAGTTCCGGCTCATGGAGGAGCAACACTGCCGGCTAAAATACGGCCTCCA CTGAATAGAGCCGCTACATACAGGCTTGGGTAGTGAAAGCGGGAAATGAAGCAAAACCCGCAAGACTGAAATCTGATGAGGATACTCAGACCACCTTGGAGGATTCGAGCGCAACAGCATCTCGGAGGTGCAATTTGTGATACAGATCCCAAGCTTCCCTTGCCTGACCAATACCCCAAGTCAAAATCAGACAGCTTGTGATGCCAGGTTTCTGAGGATAAGATTATTCTTACATGGACAGCACCAGGAGATAATTTGATGTTGGAAAAGTTCAACGTTATATCATAAGAAATAGTGCAAGTATTCTTGATCTAAGAGACAGTTTGTGATGCTCTTCAAGTAATACTACTGATCTGCACCAAGGAGGCCAATCTCAAGGAAAGCTTTGCATTTAAACCAAGAAATATCTCAGAAGAAATGCAACCCACATATTTATTGCCATTTAAAGTATAGATAAAGCAATTTGACATCAAAAGTATCAACATTTGCAAGAACTTGTGTTATCCCTCAAGCAAATCCTGATGACATTGATCCTACACCTACTCCTACTCCTACTCCTACTCCTGATAAAAGTCATAATCTGGAGTTTAATATTTCTACGCTGGTATGCTGTGATTTGGGTCTGTGTAATTTGTTAACTTTATTTTAAGTACCACCATTTGAACCTTAACGAAGAAAAAATCTTCAAGTAGACCTAGAAGAGAGTTTAAAAAACAACAAATGTAGTAAGTAAGGATATCTGTAATCTTAAATTCATCCCATGTGTGATCATAAATCATAAAAATAATTTTAAGATGTGGGAAAGGATACCTTTGATTAATAAAAAACCTCATGGATATGTAAGAACTGTCAAGATTAAAAATTAATAGTTTCATTTATTTGTTATTTTATTGTAAGAAATAGTGATGAACAAAGATCCTCTTTTTCATCATGATACCTGGTGTGATATTATTTGATGCAACAGTTTCTGAAATGATATTTCAAATTTGCATCAAGAAATTAATCATCTATCTGAGTAGTCAAAATACAAGTAAGAGGAGCAAAATACAACAACTTTGGAAAAAATAAAAAA

FIGURE 70

MGLFRGFVFLLVLCLLHQSNSTFIKLNNGFEDIVIVIDPSVPEDEKIIIEQIEDMVTASTYLFE
ATEKRFFFKNVSILIPENWKENPQYKRPKHENHKKHADVIVAPPTLPGRDEPYTKQFTECGEKGEY
IHFTPDLLLGGKQNEYGPPGKLFVHEWAHLRWGVFDEYNEDQPFYRAKSKKIEATRCISAGISGRN
RVYKCGGSGCLSRACRIDSTTKLYGKDCQFFPDKVQTEKASIMFMQSIDSVVEFCNEKTHNQEAP
SLQNIKCNFRSTWEVISNSEDFKNTIPMVTPPPPPVFSLKISQRIVCLVLDKSGSMGGKDRLNR
MNQAAKHFLQLQTVENGSWVGMVHFDSTATIVNKLIQIKSSDERNTLMAGLPTYPLGGTSICSGIK
YAFQVIGELHSQLDGSEVLLLTGDNNTASSCIDEVKQSGAIVHFIALGRAADEAVIEMSKITGG
SHFYVSDEAQNNGLIDAFGALTSGNTDLSQKSLQLESKGLTLNSNAWMNDTVIIDSTVGKDTFFL
ITWNSLPSPISLWDPSGTIMENFTVDATSKMAYLSIPGTAKVGTWAYNLQAKANPETLTITVTSR
AANSSVPPITVNAKMNDVNSFPSPMIVYAEILQGYVPVLGANVTAFIESQNGHTEVLELLDNGA
GADSEKNDGVYSRYFTAYTENGRYSLKVRHGGANTARLKLRPPLNRAAYIPGWVNGEIEANPP
RPEIDEDTQTTLEDFSRASGGAFVVSQVPSLPLPDQYPPSQITDLDATVHEDKIIILTWAPGDN
FDVGKVQRYIIIRISASILDLRDSFDDALQVNTDLSPEANSKESFAFKPENISEENATHIFIAI
KSIDKSNLTSKVSANIAQVTLFIPQANPDDIDPTPTPTPTPTPKSHNSGVNISTLVLSVIGSVVI
VNFILSTTI

Signal peptide:

amino acids 1-21

Putative transmembrane domains:

amino acids 284-300, 617-633

Leucine zipper pattern.

amino acids 469-491, 476-498

N-glycosylation site.

amino acids 20-24, 75-79, 340-344, 504-508, 542-546, 588-592,
628-632, 811-815, 832-836, 837-841, 852-856, 896-900

FIGURE 71

CTCTTAGGTGAAACCTTGGGAGTAGAGTACTGACAGCAAAGACCGGGAAAGACCATACGTCCCCGGGCAGGGGTGA
 CAACAGGTGTCATCTTTTTGATCTCGTGTGTGGCTGCCTTCTATTTCAGGAAAGACGCCAAGTAATTTTGACCCA
 GAGGAGCAATGATGTAGCCACCTCCTAACCTTCCCTTCTTGAACCCCGATTATGCCAGGATTTACTAGAGAGTGTCA
 ACTCAACCAGCAAGCGGCTCCTTCGGCTTAACCTTGTGGTTGGAGGAGAGAACCTTTGTGGGGCTGCGTTCTCTTAGCA
 GTGCTCAGAAGTGACTTGCCTGAGGGTGGACCAGAAGAAAGGAAAGGTCCCCTCTTGCTGTTGGCTGCACATCAGGAA
 GGCTGTGATGGGAATGAAGGTGAAGAACTTGGAGATTTCACTTCAGTCATTGCTTCTGCCGTGCAAGATCATCCTTTAAA
 AGTAGAGAAGCTGCTCTGTGTGGTGGTTAACTCCAAGAGGCGAGAAGCTCGTCTAGAAGGAAATGGATGCAAGCAGCTC
 CGGGGGCCCCAAACGATGCTTCCTGTGGTCTAGCCCAGGGAAGCCCTTCCGTGGGGGGCCCCGGCTTTGAGGGATGCC
 ACCGGTTCTGGACGCATGGCTGATTCCTGAATGATGATGGTTCCGCCGGGGGCTGCTTGGCTGGATTTCGGGGTGGTG
 GTTTTGTGGTGCTCCTCTGCTGTGCTATCTCTGCTGTACATGTTGGCCTGCACCCCAAAGGTGACGAGGAGCAG
 CTGGCACTGCCCAGGGCCAACAGCCCCACGGGGAAGGAGGGGTACCAGGCCGTCTTCAGGAGTGGGAGGAGCAGCAC
 CGCAACTACGTGAGCAGCCTGAAGCGGCAGATCGCACAGCTCAAGGAGGAGCTGCAGGAGAGGAGTGCAGCAGCTCAGG
 AATGGGCAGTACCAAGCCAGCGATGCTGCTGGCCTGGGTCTGGACAGGAGCCCCCAGAGAAAACCCAGGCCGACCTC
 CTGGCCTTCTGCACTCGCAGGTGGACAAGGCAGAGGTGAATGCTGGCGTCAAGCTGGCCACAGAGTATGCAGCAGTG
 CCTTTCGATAGCTTTACTCTACAGAAGGTGTACCAGCTGGAGACTGGCCTTACC GCCCACCCGAGGAGAAGCCTGTG
 AGGAAGGACACAGCGGGATGAGTTGGTGGAGGCCATTGAATCAGCCTTGGAGACCCTGAACAATCCTGCAGAGAACAGC
 CCCAATCACCGTCTTACACGGCCTCTGATTTTATAGAAGGGATCTACCGAACAGAAAGGGACAAAGGGACATTGTAT
 GAGCTCACCTTCAAAGGGGACCACAAACACGAATTCAAACGGCTCATCTTATTTGACCATTCAGCCCCATCATGAAA
 GTGAAAATGAAAAGCTCAACATGGCCAAACACGCTTATCAATGTTATCGTGCCTCTAGCAAAAAGGGTGGACAAGTTC
 CGGCAGTTCATGCAGAATTTAGGGAGATGTGCAATGAGCAGGATGGGAGAGTCCATCTCATGTTGTTTACTTTGGG
 AAAGAAGAAATAAATGAAGTCAAAGGAATACTTGAAAACACTTCCAAGCTGCCAATTCAGGAATTTACCTTCATC
 CAGCTGAATGGAGAATTTTCTCGGGGAAAGGGACTTGATGTTGGAGCCGCTTCTGGAAGGGAAGCAACGTCCTTCTC
 TTTTCTGTGTGTGGACATCTACTTCACATCTGAATTCCTCAATACGTGTAGGCTGAATACACAGCCAGGGAAGAAG
 GTATTTTATCCAGTTCCTTTTTCAGTCAGTACAATCCTGGCATAATATACGGCCACCATGATGCAGTCCCTCCCTTGGAA
 CAGCAGCTGGTCATAAAGAAGGAACTGGATTTTGGAGAGACTTTGGATTTGGGATGACGTGTGATGATCGGTGCAGAC
 TTCATCAATATAGGTGGGTTTGTCTGGACATCAAAGGCTGGGGCGGAGAGGATGTGCACCTTTATCGCAAGTATCTC
 CACAGCAACCTCATAGTGGTACGGACGCTGTGCGAGGACTTTCCACCTCTGGCATGAGAAGCGTGCATGGACGAG
 CTGACCCCGAGCAGTACAAGATGTGCATGCAGTCCAAGGCCATGAACGAGGCATCCCACGGCCAGCTGGGCATGCTG
 GTGTTGAGGCACGAGATAGAGGCTCACCTTCGCAACAGAAACAGAGACAAGTAGCAAAAAACATGAAGTCCAGAA
 GAAGGATTGTGGGAGACACTTTTCTTCTTTTGGCAATTACTGAAAGTGGCTGCAACAGAGAAAAGACTTCCATATA
 GGACGACAAAAGAAATGGACTGATGGGTGAGAGATGAGAAAGCCTCCGATTTCTCTCTGTTGGGCTTTTACAAACAGA
 AATCAAAATCTCCGCTTTGCCTGCAAAAGTAACCCAGTTGCACCTGTGAAGTGTCTGACAAAGGCAGAAATGCTGTG
 AGATTATAAGCCTAATGGTGTGGAGGTTTTGATGGTGTTTACAACTACCTGAGACCTGTTGTTTTGTGTGCTCATGGA
 AATATTCATGATTTAAGAGCAGTTTTGTAAAAATTCATTAGCATGAAAGGCAAGCATATTTCTCCTCATATGAATGA
 GCCTATCAGCAGGGCTCTAGTTTCTAGGAATGCTAAAAATATCAGAAGGCAGGAGAGGAGATAGGCTTATTATGATACT
 AGTGACTACATTAAGTAAATAAAATGACCCAGAAAACAGAAACCAATAATATCGTGTATATTTTCCCCAAGAT
 TAACCAAAAATAATCTGCTTATCTTTTGGTGTCTTTTAACTGTCTCCGTTTTTTCTTTTATTTAAAAATGCACT
 TTTTTTCCCTTGTGAGTTATAGTCTGCTTATTTAATTACCCTTTGCAAGCCTTACAAGAGAGCACAAGTTGGCCTAC
 ATTTTTATATTTTTAAGAAGATACTTTGAGATGCATTATGAGAATTTTCAGTTCAAAGCATCAAATTGATGCCATAT
 CCAAGGACATGCCAAATGCTGATTCTGTGAGGCATGAATGTGAGGCATTGAGACATAGGGAAGGAATGGTTTGTACT
 AATACAGACGTACAGATACTTTCTCTGAAGAGTATTTTTCGAAGAGGAGCAACTGAACACTGGAGGAAAAGAAAATGAC
 ACTTTCTGCTTTTACAGAAAAGGAACTCATTGAGACTGCTGATATCGTGATGTACCTAAAAGTCAGAAACACATTTT
 CTCCTCAGAAGTAGGGACCGCTTTCTTACCTGTTTAAATAAACCAAGTATACCGTGTGAACCAACAAATCTCTTTTC
 AAAACAGGGTGCCTCCTGGCTTCTGGCTTCCATAAGAAGAAATGGAGAAAAATATATATATATATATATATATGTT
 GAAAGATCAATCCATCTGCCAGAACTAGTGGGATGGAAGTTTTTGTCTACATGTTATCCACCCAGGCCAGGTGGAAG
 TAACTGAATTATTTTTAAATAAGCAGTTCTACTCAATCACCAGATGCTTCTGAAAATTGCATTTTATTACCATT
 CAACTATTTTTTAAAAATAAATACAGTTAATAGAGTGGTTTCTTCATTCATGTGAAAATTATTAGCCAGCACCAG
 ATGCATGAGCTAATATCTCTTTGAGTCTTGTCTGTTGCTCACAGTAAACTCATTGTTTAAAGCTTCAAGAAC
 ATTCAAGCTGTTGGTGTGTTAAAAATGCATTGTATTGATTTGTACTGGTAGTTTATGAAATTTAATTAAACACAGG
 CCATGAATGGAAGGTGGTATTGCACAGCTAATAAAATATGATTTGTGGATATGAA

FIGURE 72

MMMVRRGLLAWISRVVLLVLLCCAISVLYMLACTPKGDEEQLALPRANSPTGKEGYQAVLQEW
EQHRNYVSSSLKRQIAQLKEELQERSEQLRNGQYQASDAAGLGLDRSPPEKTQADLLAFLHSQVDK
AEVNAGVKLATEYAAVPFDSFTLQKVYQLETGLTRHPEEKPVKDKRDELVEAIESALETNNPA
ENSPNHRPYTASDFIEGIYRTERDKGTLTYELTFKGDHKHEFKRLILFRPFSPIMKVKNEKLNMAN
TLINIVIVPLAKRVDKFRQFMQNFREMCIEQDGRVHLTVVYFGKEEINEVKGILENTSKAANFRNF
TFIQLNGEFSRGKGLDVGARFWKGSNVLLFFCDVDIYFTSEFLNTCRLNTQPGKKVFYPVLFSQY
NPGIIYGHHDVPPLEQQLVIKKETGFWRDFGFGMTCQYRSDFINIGGFDLDIKGWGGEDVHLYR
KYLHSNLIVVRTPVRLFWHEKRCMDELTPEQYKCMQSKAMNEASHGQLGMLVFRHEIEAHL
RKQKQKTSSKKT

Important features:

Signal peptide:

amino acids 1-27

N-glycosylation sites.

amino acids 315-319, 324-328

N-myristoylation sites.

amino acids 96-102, 136-142, 212-218, 311-317, 339-345, 393-399

Amidation site.

amino acids 377-381

FIGURE 73

GAGACTGCAGAGGGAGATAAAGAGAGAGGGCAAAGAGGCAGCAAGAGATTTGTCCTGGGGATCCA
 GAAACCCATGATACCCTACTGAACACCGAATCCCCTGGAAGCCCACAGAGACAGAGACAGCAAGA
 GAAGCAGAGATAAATACACTCAGGCCAGGAGCTCGCTCGCTCTCTCTCTCTCTCTCACTCCTC
 CCTCCCTCTCTCTCTGCCTGTCCTAGTCCTCTAGTCCTCAAATTTCCAGTCCCCTGCACCCCTTC
 CTGGGACACTTATGTTGTTCTCCGCCCTCCTGCTGGAGGTGATTTGGATCCTGGCTGCAGATGGGG
 GTCAACACTGGACGTATGAGGGCCACATGGTCAGGACCATTGGCCAGCCTCTTACCCTGAGTGT
 GGAAACAATGCCCAGTCGCCCATCGATATTCAGACAGACAGTGTGACATTTGACCCTGATTTGCC
 TGCTCTGCAGCCCCACGGATATGACCAGCCTGGCACCGAGCCTTTGGACCTGCACAACAATGGCC
 ACACAGTGCAACTCTCTCTGCCCTCTACCCTGTATCTGGGTGGACTTCCCCGAAAATATGTAGCT
 GCCCAGCTCCACCTGCACTGGGGTCAGAAAGGATCCCCAGGGGGGTGAGAACACCAGATCAACAG
 TGAAGCCACATTTGCAGAGCTCCACATTGTACATTATGACTCTGATTCCTATGACAGCTTGAGTG
 AGGCTGCTGAGAGGCCTCAGGGCCTGGCTGTCTGGGCATCCTAATTGAGGTGGGTGAGACTAAG
 AATATAGCTTATGAACACATTCTGAGTCACTTGCATGAAGTCAGGCATAAAGATCAGAAGACCTC
 AGTGCCTCCCTTCAACCTAAGAGAGCTGCTCCCCAACAGCTGGGGCAGTACTTCCGCTACAATG
 GCTCGCTCACAACTCCCCCTTGCTACCAGAGTGTGCTCTGGACAGTTTTTTATAGAAGGTCCCAG
 ATTTCAATGGAACAGCTGGAAAAGCTTCAGGGGACATTGTTCTCCACAGAAGAGGAGCCCTCTAA
 GCTTCTGGTACAGAACTACCGAGCCCTTCAGCCTCTCAATCAGCGCATGGTCTTTGCTTCTTTCA
 TCCAAGCAGGATCCTCGTATACCACAGGTGAAATGCTGAGTCTAGGTGTAGGAATCTTGTTGGC
 TGTCTCTGCCTTCTCCTGGCTGTTTATTTTCATTGCTAGAAAGATTCCGGAAGAAGAGGCTGGAAAA
 CCGAAAGAGTGTGGTCTTCACCTCAGCACAAGCCACGACTGAGGCATTAAATTCCTTCTCAGATAC
 CATGGATGTGGATGACTTCCCTTCATGCCTATCAGGAAGCCTCTAAAATGGGGTGTAGGATCTGG
 CCAGAAACACTGTAGGAGTAGTAAGCAGATGTCTCCTTCCCCTGGACATCTCTTAGAGAGGAAT
 GGACCCAGGCTGTCAATCCAGGAAGAACTGCAGAGCCTTCAGCCTCTCCAAACATGTAGGAGGAA
 ATGAGGAAATCGCTGTGTTGTTAATGCAGAGANCAAACTCTGTTTAGTTGCAGGGGAAGTTTGGG
 ATATACCCCAAAGTCTCTACCCCCCTCACTTTTATGGCCCTTCCCTAGATATACTGCGGGATCT
 CTCCTTAGGATAAAGAGTTGCTGTTGAAGTTGTATATTTTGGATCAATATATTTGGAATTAAAG
 TTTCTGACTTT

FIGURE 74

MLFSALLLEVIWILAADGGQHWTYEGPHGQDHWPA SYPECGNNAQSPIDIQTDSVTFDPDLPALQ
PHGYDQPGTEPLDLHNNGHTVQLSLPSTLYLGGLPRKYVAAQLHLHWGQKGSPGGSEHQINSEAT
FAELHIVHYDSYDSLSEAAERPQGLAVLGILIEVGETKNIAYEHILSHLHEVRHKDQKTSVPP
FNLRELLPKQLGQYFRYNGSLTTPPCYQSVLWTVFYRRSQISMEQLEKLQGTLFSTEEEPSKLLV
QNYRALQPLNQRMVFASFIQAGSSYTTGEMLSLGVGILVGCLCLLLAVYFIARKIRKKRLENRKS
VVFTSAQATTEA

Important features of the protein:

Signal peptide:

amino acids 1-15

Transmembrane domain:

amino acids 291-310

N-glycosylation site.

amino acids 213-216

Eukaryotic-type carbonic anhydrases proteins

amino acids 197-245, 104-140, 22-69

FIGURE 75

TGCCGCTGCCGCCGCTGCTGCTGTTGCTCCTGGCGGCGCCTTGGGGACGGGCAGTTCCTGTGTC
TCTGGTGGTTTGCTAAACCTGCAAACATCACCTTCTTATCCATCAACATGAAGAATGTCCTACA
ATGGACTCCACCAGAGGGTCTTCAAGGAGTTAAAGTTACTTACACTGTGCAGTATTTTCATCACAA
ATTGGCCCCACCAGAGGTGGCACTGACTACAGATGAGAAGTCCATTTCTGTTGTCTGACAGCTCC
AGAGAAGTGAAGAGAAATCCAGAAGACCTTCCTGTTTCCATGCAACAAATATACTCCAATCTGA
AGTATAACGTGTCTGTGTTGAATACTAAATCAAACAGAACGTGGTCCCAGTGTGTGACCAACCAC
ACGCTGGTGCTCACCTGGCTGGAGCCGAACACTCTTTACTGCGTACACGTGGAGTCTTTCGTCCC
AGGGCCCCCTCGCCGTGCTCAGCCTTCTGAGAAGCAGTGTGCCAGGACITTTGAAAGATCAATCAT
CAGAGTTCAAGGCTAAATCATCTTCTGGTATGTTTTGCCCATATCTATTACCGTGTTCITTTTT
TCTGTGATGGGCTATTCCATCTACCGATATATCCACGTTGGCAAAGAGAAACACCCAGCAAATTT
GATTTTGATTTATGGAAATGAATTTGACAAAAGATTCTTTGTGCCTGCTGAAAAATCGTGATTA
ACTTTATCACCTCAATATCTCGGATGATTCTAAATTTCTCATCAGGATATGAGTTTACTGGGA
AAAAGCAGTGATGTATCCAGCCTTAATGATCCTCAGCCCAGCGGGAACCTGAGGCCCCCTCAGGA
GGAAGAGGAGGTGAAACATTTAGGGTATGCTTCGCATTTGATGGAATTTTTTGTGACTCTGAAG
AAAACACGGAAGGTACTTCTCTCACCCAGCAAGAGTCCCTCAGCAGAACAATACCCCCGGATAAA
ACAGTCATTGAATATGAATATGATGTCAGAACCACTGACATTTGTGCGGGGCTGAAGAGCAGGA
GCTCAGTTTGCAGGAGGAGGTGTCCACACAAGGAACATTATTGGAGTGCAGGCAGCGTTGGCAG
TCTTGGGCCCCGAAACGTTACAGTACTCATACACCCCTCAGCTCCAAGACTTAGACCCCTGGCG
CAGGAGCACACAGACTCGGAGGAGGGGCCGAGGAAGAGCCATCGACGACCTGGTTCGACTGGGA
TCCCCAACTGGCAGGCTGTGTATTCCTTCGCTGTCCAGCTTCGACCAGGATTCAGAGGGCTGCG
AGCCTTCTGAGGGGATGGGCTCGGAGAGGAGGGTCTTCTATCTAGACTCTATGAGGAGCCGGCT
CCAGACAGGCCACCAGGAGAAAATGAAACCTATCTCATGCAATTCATGGAGGAATGGGGTTATA
TGTGCAGATGGAAACTGATGCCAACACTTCCTTTTGCTTTTGTTCCTGTGCAACAAGTGAG
TCACCCCTTTGATCCCAGCCATAAAGTACCTGGGATGAAAGAAGTTTTTCCAGTTTGTGAGTGT
CTGTGAGAATTACTTATTTCTTTTCTCTATTCTCATAGCACGTGTGTGATTGGTTCATGCATGTA
GGTCTCTTAACAATGATGGTGGGCCTCTGGAGTCCAGGGGCTGGCCGTTGTTCTATGCAGAGAA
AGCAGTCAATAAATGTTTGCCAGACTGGGTGCAGAATTATTCAGGTGGGTGT

FIGURE 76

MSYNGLHQRVFKELKLLTLCSSISQIGPPEVALTTDEKSI SVVLTAPEKWKRNPEDLPVSMQQIY
SNLKYNVSVLNTKSNRTWSQCVTNHTLVLTWLEPNTLYCVHVESFVPGPPRAQPSEKQCARTLK
DQSSEFKAKIIFWYVLPISITVFLFSVMGYSIYRYIHVGKEKHPANLILYIGNEFDKRFFVPAEK
IVINFITLNISSDKISHQDMSLLGKSSDVSSLNDPQPSGNLRPPQEEEEVKHLGYASHLMEIFC
DSEENTEGTSLTQQESLRTIPDPKTVIEYEYDVRTDICAGPEEQELSLQEEVSTQGTLLSQA
ALAVLGPQTLQYSYTPQLQDLPLAQEHTDSEEGPEEEPSTTLVDWDPQTGRLCIPSLSSFDQDS
EGCEPSEGDGLGEEGLLSRLYEAPAPDRPPGENETYLMQFMEEWGLYVQMEN

Important features:

Signal peptide:

amino acids 1-28

Transmembrane domain:

amino acids 140-163

N-glycosylation sites.

amino acids 71-74, 80-83, 89-92, 204-207, 423-426

FIGURE 77

GAGGAGCGGGCCGAGGACTCCAGCGTGCCAGGTCTGGCATCCTGCACTTGCTGCCCTCTGACAC
 CTGGGAAGATGGCCGGCCCGTGACCTTCACCCCTTCTCTGTGGTTTGCTGGCAGCCACCTTGATC
 CAAGCCACCCTCAGTCCCAGTTCAGTTCTCATCCTCGGCCCAAAAGTCATCAAAGAAAAGCTGAC
 ACAGGAGCTGAAGGACCACAACGCCACCAGCATCCTGCAGCAGCTGCCGCTGCTCAGTGCCATGC
 GGGAAAAGCCAGCCGGAGGCATCCCTGTGCTGGGCAGCCTGGTGAACACCGTCCTGAAGCACATC
 ATCTGGCTGAAGGTCATCACAGCTAACATCCTCCAGCTGCAGGTGAAGCCCTCGGCCAATGACCA
 GGAGCTGCTAGTCAAGATCCCCCTGGACATGGTGGCTGGATTCAACACGCCCCCTGGTCAAGACCA
 TCGTGGAGTTCCACATGACGACTGAGGCCAAAGCCACCATCCGCATGGACACCAGTGCAAGTGGC
 CCCACCCGCTGGTCCTCAGTGACTGTGCCACCAGCCATGGGAGCCTGCGCATCCAAGTCTGTGTA
 TAAGCTCTCCTTCCTGGTGAACGCCTTAGCTAAGCAGGTTCATGAACCTCCTAGTGCCATCCCTGC
 CCAATCTAGTGAAAAACCAGCTGTGTCCCGTGATCGAGGCTTCCTTCAATGGCATGTATGCAGAC
 CTCCTGCAGCTGGTGAAGGTGCCATTTCCCTCAGCATTGACCGTCTGGAGTTTGACCTTCTGTA
 TCCTGCCATCAAGGGTGACACCATTAGCTCTACCTGGGGGCCAAGTTGTTGGACTCACAGGGAA
 AGGTGACCAAGTGGTTCAATAACTCTGCAGCTTCCTGACAATGCCCCACCTGGACAACATCCCG
 TTCAGCCTCATCGTGAGTCAGGACGTGGTGAAAGCTGCAGTGGCTGCTGTGCTCTCTCCAGAAGA
 ATTCATGGTCCTGTTGGACTCTGTGCTTCCTGAGAGTGCCCATCGGCTGAAGTCAAGCATCGGGC
 TGATCAATGAAAAGGCTGCAGATAAGCTGGGATCTACCCAGATCGTGAAGATCCTAACTCAGGAC
 ACTCCCGAGTTTTTTATAGACCAAGGCCATGCCAAGGTGGCCCAACTGATCGTGCTGGAAGTGTT
 TCCCTCCAGTGAAGCCCTCCGCCCTTTGTTACCCCTGGGCATCGAAGCCAGCTCGGAAGCTCAGT
 TTTACACCAAAGGTGACCAACTTATACTCAACTTGAATAACATCAGCTCTGATCGGATCCAGCTG
 ATGAACTCTGGGATTGGCTGGTTCCAACCTGATGTTCTGAAAAACATCATCACTGAGATCATCCA
 CTCCATCCTGCTGCCGAACCAGAATGGCAAATTAAGATCTGGGGTCCCAGTGTCATTGGTGAAGG
 CCTTGGGATTCGAGGCAGCTGAGTCCTCACTGACCAAGGATGCCCTTGTGCTTACTCCAGCCTCC
 TTGTGGAAACCCAGCTCTCCTGTCTCCAGTGAAGACTTGATGGCAGCCATCAGGGAAAGGCTGG
 GTCCAGCTGGGAGTATGGGTGTGAGCTCTATAGACCATCCCTCTCTGCAATCAATAAACACTTG
 CCTGTGAAAAA

FIGURE 78

MAGPWTFLLCGLLAATLIQATLSPTAVLILGPKVIKEKLTQELKDHNATSILQQLPLLSAMREK
PAGGIPVLGSLVNTVLKHIIWLKVITANILQLQVKPSANDQELLVKIPLDMVAGFNTPLVKTIVE
FHMTTEAQATIRMDTSASGPTRLVLSDCATSHGSLRIQLLYKLSFLVNALAKQVMNLLVPSLPNL
VKNQLCPVIEASFNGMYADLLQLVKVPISLSIDRLEFDLLYPAIKGDITIQLYLGAKLKLLDSQGKVT
KWFNNSAASLTMPITLDNIPFSLIVSQDVVKAAVAVALSPPEEFMVLLDSVLPESAHLKSSIGLIN
EKAADKLGSTQIVKILTQDTPEFFIDQGHAKVAQLIVLEVFPSSSEALRPLFTLGIEASSEAQFYT
KGDQLILNLNNISSDRIQLMNSGIGWFQPDVLKNIITEIIHSILLPNQNGKLRSQVPSVSLVKALG
FEAAESSLTkdALVLTpASLWKPSSPVSQ

Important features of the protein:

Signal peptide:

amino acids 1-21

N-glycosylation sites.

amino acids 48-51, 264-267, 401-404

Glycosaminoglycan attachment site.

amino acids 412-415

LBP / BPI / CETP family proteins.

amino acids 407-457

FIGURE 79

GAGAGAAGTCAGCCTGGCAGAGAGACTCTGAAATGAGGGATTAGAGGTGTTCAAGGAGCAAGAGC
TTCAGCCTGAAGACAAGGGAGCAGTCCCTGAAGACGCTTCTACTGAGAGGTCTGCCATGGCCTCT
CTTGGCCTCCAAC TTGTGGGCTACATCCTAGGCCTTCTGGGGCTTTTGGGCACACTGGTTGCCAT
GCTGCTCCCCAGCTGGAAAACAAGTTCTTATGTGGTGCCAGCATTGTGACAGCAGTTGGCTTCT
CCAAGGGCCTCTGGATGGAATGTGCCACACACAGCACAGGCATCACCCAGTGTGACATCTATAGC
ACCCTTCTGGGCCTGCCCCGCTGACATCCAGGCTGCCCAGGCCATGATGGTGACATCCAGTGCAAT
CTCCTCCCTGGCCTGCATTATCTCTGTGGTGGGCATGAGATGCACAGTCTTCTGCCAGGAATCCC
GAGCCAAAGACAGAGTGCGGTAGCAGGTGGAGTCTTTTTCATCCTTGGAGGCCTCCTGGGATTC
ATTCTGTTGCCTGGAATCTTCATGGGATCCTACGGGACTTCTACTCACCAGTGGTGCTGACAG
CATGAAATTTGAGATTGGAGAGGCTCTTTACTTGGGCATTATTTCTTCCCTGTTCTCCCTGATAG
CTGGAATCATCCTCTGCTTTTCTGCTCATCCAGAGAAATCGCTCCAAC TACTACGATGCCTAC
CAAGCCCCAACCCTTTGCCACAAGGAGCTCTCCAAGGCCTGGTCAACCTCCCAAAGTCAAGAGTGA
GTTCAATTCCTACAGCCTGACAGGGTATGTGTGAAGAACCAGGGGCCAGAGCTGGGGGGTGGCTG
GGTCTGTGAAAAACAGTGGACAGACCCCCGAGGGCCACAGGTGAGGGACACTACCAGTGGATCGT
GTCAGAAAGGTGCTGCTGAGGATAGACTGACTTTGGCCATTGGATTGAGCAAAGGCAGAAATGGGG
GCTAGTGTAACAGCATGCAGGTTGAATTGCCAAGGATGCTCGCCATGCCAGCCTTTCTGTTTTCC
TCACCTTGCTGCTCCCCTGCCCTAAGTCCCCAACCCTCAACTTGAAACCCCATTCCTTAAGCCA
GGACTCAGAGGATCCCTTTGCCCTCTGGTTTACCTGGGACTCCATCCCCAAACCCACTAATCACA
TCCCACTGACTGACCCTCTGTGATCAAAGACCCTCTCTCTGGCTGAGGTTGGCTCTTAGCTCATT
GCTGGGGATGGGAAGGAGAAGCAGTGGCTTTTGTGGGCATTGCTCTAACCTACTTCTCAAGCTTC
CCTCCAAAGAACTGATTGGCCCTGGAACCTCCATCCCACTCTTGTTATGACTCCACAGTGTCCA
GACTAATTTGTGCATGAACTGAAATAAAACCATCCTACGGTATCCAGGGAACAGAAAGCAGGATG
CAGGATGGGAGGACAGGAAGGCAGCCTGGGACATTTAAAAAATA

FIGURE 8o

MASLGLQLVGYILGLLGLLGTLVAMLLPSWKTSSYVGASIVTAVGFSKGLWMECATHSTGITQCD
IYSTLLGLPADIQAAQAMMTSSAIISSLACIISVVGMRCTVFCQESRAKDRVAVAGGVFFILGGL
LGFIPVAWNLHGILRDFYSPLVPDSMKFEIGEALYLGIISSLFSLIAGIILCFSCSSQRNRSNYY
DAYQAQPLATRSSPRPGQPPKVKSEFNSYSLTGYV

Important features of the protein:

Signal peptide:

amino acids 1-24

Transmembrane domains:

amino acids 82-102, 117-140, 163-182

N-glycosylation site.

amino acids 190-193

PMP-22 / EMP / MP20 family proteins.

amino acids 46-59

FIGURE 81

CCCACGCGTCCGCGCCTCTCCCTTCTGCTGGACCTTCCTTCGTCTCTCCATCTCTCCCTCCTTTC
 CCCGCGTTCTCTTTCCACCTTTCTCTTCTTCCACCTTAGACCTCCCTTCCTGCCCTCCTTTCCCT
 GCCCACCGCTGCTTCCTGGCCCTTCTCCGACCCCGCTCTAGCAGCAGACCTCCTGGGGTCTGTGG
 GTTGATCTGTGGCCCCCTGTGCCTCCGTGTCCTTTTCGTCTCCCTTCCTCCCGACTCCGCTCCCGG
 ACCAGCGGCCTGACCCTGGGGAAGGATGGTTCCCGAGGTGAGGGTCCTCTCCTCCTTGCTGGGA
 CTCGCGCTGCTCTGGTTCCCCCTGGACTCCCACGCTCGAGCCCGCCAGACATGTTCTGCCTTTT
 CCATGGGAAGAGATACTCCCCCGGCGAGAGCTGGCACCCCTACTTGAGCCACAAGGCCTGATGT
 ACTGCCTGCGCTGTACCTGCTCAGAGGGCGCCCATGTGAGTTGTTACCGCCTCCACTGTCCGCCT
 GTCCACTGCCCCAGCCTGTGACGGAGCCACAGCAATGCTGTCCCAAGTGTGTGGAACCTCACAC
 TCCCTCTGGACTCCGGGCCCCACCAAAGTCTGCCAGCACAAACGGGACCATGTACCAACACGGAG
 AGATCTTCAGTGCCCATGAGCTGTTCCCTCCCGCCTGCCCAACCAGTGTGTCTCTGCAGCTGC
 ACAGAGGGCCAGATCTACTGCGGCCTCACAACTGCCCCGAACCAGGCTGCCAGCACCCCTCCC
 ACTGCCAGACTCCTGCTGCCAAGCCTGCAAAGATGAGGCAAGTGAGCAATCGGATGAAGAGGACA
 GTGTGCAGTCGCTCCATGGGGTGAGACATCCTCAGGATCCATGTTCCAGTGATGCTGGGAGAAAG
 AGAGGCCCGGGCACCCAGCCCCACTGGCCTCAGCGCCCTCTGAGCTTCATCCCTCGCCACTT
 CAGACCCAAGGGAGCAGGCAGCACAACTGTCAAGATCGTCCTGAAGGAGAAACATAAGAAAGCCT
 GTGTGCATGGCGGGAAGACGTACTCCACGGGGAGGTGTGGCACCCGGCCTTCCGTGCCTTCGGC
 CCCTTGCCCTGCATCCTATGCACCTGTGAGGATGGCCGCCAGGACTGCCAGCGTGTGACCTGTCC
 CACCGAGTACCCCTGCCGTACCCCGAGAAAGTGGCTGGGAAGTGCTGCAAGATTTGCCAGAGG
 ACAAAGCAGACCCTGGCCACAGTGAGATCAGTTCTACCAGGTGTCCAAGGCACCGGGCCGGGTCT
 CTCGTCCACACATCGGTATCCCCAAGCCCAGACAACCTGCGTCGCTTTGCCCTGGAACACGAGGC
 CTCGGACTTGGTGGAGATCTACCTCTGGAAGCTGGTAAAAGATGAGGAACTGAGGCTCAGAGAG
 GTGAAGTACCTGGCCCAAGGCCACACAGCCAGAATCTTCCACTTGACTCAGATCAAGAAAGTCAG
 GAAGCAAGACTTCCAGAAAGAGGCACAGCACTTCCGACTGCTCGCTGGCCCCCACGAAGGTCCT
 GGAACGTCTTCTAGCCCAGACCCTGGAGCTGAAGGTCACGGCCAGTCCAGACAAAGTGACCAAG
 ACATAACAAAGACCTAACAGTTGCAGATATGAGCTGTATAATTGTTGTTATTATATATTAATAAA
 TAAGAAGTTGCATTACCCTCAAAAAAAAAAAAAAAAAAAAAA

FIGURE 82

MVPEVRVLSSLLGLALLWFPLDSHARARPDMFCLFHGKRYSPGESWHPYLEPQGLMYCLRCTCSE
GAHVSCYRLHCPPVHCPQPVTEPQQCCPKCVEPHTPSGLRAPPKSCQHNGTMYQHGEIFSAHELF
PSRLPNQCVLCSCTEGQIYCGLTTCPEPGCPAPLPLPDSCCQACKDEASEQSDEEDSVQSLHGVR
HPQDPCSSDAGRKRGPPTAPTGLSAPLSFIPRHFRPKGAGSTTVKIVLKEKHKKACVHGGKTYS
HGEVWHPAFRAFGPLPCILCTCEDGRQDCQRVTCPTTEYPCRHPKQVAGKCKICPEDKADFGHSE
ISSTRCPKAPGRVLVHTSVSPSPDNLRRFALEHEASDLVEIYLWKLVKDEETEAQRGEVPGPRPH
SQNLPLDSDQESQEARLPERGTALPTARWPPRRSLERLPSDPGAEGHGQSRQSDQDITKT

Signal peptide:

amino acids 1-25

FIGURE 83

GACAGCTGTGTCTCGATGGAGTAGACTCTCAGAACAGCGCAGTTTGCCCTCCGCTCACGCAGAGCCTCTCC
 GTGGCTTCCGCACCTTGAGCATTAGGCCAGTTCTCCTCTTCTCTCTAATCCATCCGTCACCTCTCCTGTCA
 TCCGTTTCCATGCCGTGAGGTCCATTACAGAACACATCCATGGCTCTCATGCTCAGTTTGGTTCTGAGTC
 TCCTCAAGCTGGGATCAGGGCAGTGGCAGGTGTTTGGGCCAGACAAGCCTGTCCAGGCCTTGGTGGGGAG
 GACGCAGCATTCTCCTGTTTCTGTCTCCTAAGACCAATGCAGAGGCCATGGAAGTGC GGTTCTTCAGGGG
 CCAGTTCTCTAGCGTGGTCCACCTCTACAGGGACGGGAAGGACCAGCCATTTATGCAGATGCCACAGTATC
 AAGGCAGGACAAAACCTGGTGAAGGATTCTATTGCGGAGGGGCGCATCTCTCTGAGGCTGGAAAACATTACT
 GTGTTGGATGCTGGCCTCTATGGGTGCAGGATTAGTTCCCAGTCTTACTACCAGAAGGCCATCTGGGAGCT
 ACAGGTGTGAGCACTGGGCTCAGTTCTCTCATTTCATCACGGGATATGTTGATAGAGACATCCAGCTAC
 TCTGTCACTCCTCGGGCTGGTTCCCCCGGGCCACAGCGAAGTGGAAAGGTCCACAAGGACAGGATTTGTCC
 ACAGACTCCAGGACAAACAGAGACATGCATGGCCTGTTTGATGTGGAGATCTCTCTGACCGTCCAAGAGAA
 CGCCGGGAGCATATCCTGTTCCATGCGGCATGCTCATCTGAGCCGAGAGGTGGAATCCAGGGTACAGATAG
 GAGATACCTTTTTTCGAGCCTATATCGTGGCACCTGGCTACCAAAGTACTGGGAATACTCTGCTGTGGCCTA
 TTTTTTGGCATTGTTGGACTGAAGATTTTCTTCTCCAAATTCCAGTGGAAAATCCAGGCGGAACCTGGACTG
 GAGAAGAAAGCACGGACAGGCAGAATTGAGAGACGCCCGGAAACACGCAGTGGAGGTGACTCTGGATCCAG
 AGACGGCTCACCCGAAGCTCTGCGTTTCTGATCTGAAAACCTGTAACCCATAGAAAAGCTCCCCAGGAGGTG
 CCTCACTCTGAGAAGAGATTTACAAGGAAGAGTGTGGTGGCTTCTCAGAGTTTCCAAGCAGGGAAACATTA
 CTGGGAGGTGGACGGAGGACACAATAAAAGGTGGCGCTGGGAGTGTGCCGGGATGATGTGGACAGGAGGA
 AGGAGTACGTGACTTTGTCTCCCGATCATGGGTACTGGGTCTCAGACTGAATGGAGAACATTTGTATTTT
 ACATTAAATCCCCGTTTTATCAGCGTCTTCCCCAGGACCCACCTACAAAATAGGGGTCTTCTTGGACTA
 TGAGTGTGGGACCATCTCCTTCTTCAACATAAATGACCAGTCCCTTATTTATACCTGACATGTGGTTTG
 AAGGCTTATTGAGGCCCTACATTGAGTATCCGTCCTATAATGAGCAAAATGGAATCCCATAGTCATCTGC
 CCAGTCACCCAGGAATCAGAGAAAGAGGCCTCTTGGCAAAGGGCCTCTGCAATCCCAGAGACAAGCAACAG
 TGAGTCCTCCTCACAGGCAACCACGCCCTTCTCCCCAGGGGTGAAATGTAGGATGAATCACATCCCACAT
 TCTTCTTTAGGGATATTAAGGTCTCTCTCCAGATCCAAAGTCCCGCAGCAGCCGGCCAAGGTGGCTTCCA
 GATGAAGGGGGACTGGCCTGTCCACATGGGAGTCAGGTGTCATGGCTGCCCTGAGCTGGGAGGGAAGAAGG
 CTGACATTACATTTAGTTTGTCTCACTCCATCTGGCTAAGTGATCTTGAAATACCACCTCTCAGGTGAAG
 AACCGTCAGGAATCCCATCTCACAGGCTGTGGTGTAGATTAAGTAGACAAGGAATGTGAATAATGCTTAG
 ATCTTATTGATGACAGAGTGATCCTAATGGTTTGTTCATTATATTACACTTTCAGTAAAAAAA

FIGURE 84

MALMLSIVLSLLKLGSGQWQVFGPDKPVQALVGEDAAAFSCFLSPKTNAEAMEVRFFRGQFSSVVH
LYRDGKDQPFMQMPQYQGRTKLVKDSIAEGRISLRLENITVLDAGLYGCRISQSYQKAIWELQ
VSALGSVPLISITGYVDRDIQLLCQSSGWFFRPTAKWKGPQGQDLSTDSRTNRDMHGLFDVEISL
TVQENAGSISCSMRHAHLSREVESRVQIGDTFFEPISWHLATKVLGILCCGLFFGIVGLKIFFSK
FQWKIQAEILDWRRKHGQAEIRDARKHAVEVTLPETAHPKLCVSDLKTVTHRKAQEVPHSEKRF
TRKSVVASQS FQAGKHYWEVDGGHNKRWRVGVCRDDVDRRKEYVTLSPDHGYWVLRNLNGEHLFT
LNPRFISVFPRTPTKIGVFLDYECGTISFFNINDQSLIYTLTCRFEGLLRPYIEYPSYNEQNGT
PIVICPVTQESEKEASWQRASAIPESTNSSESSQATTPLPRGEM

Signal peptide:

amino acids 1-17

Transmembrane domain:

amino acids 239-255

FIGURE 85

AACAGACGTTCCCTCGCGGCCCTGGCACCTCTAACCCAGACATGCTGCTGCTGCTGCTGCTGCCCCCT
GCTCTGGGGGAGGGAGAGGGCGGAAGGACAGACAAGTAACTGCTGACGATGCAGAGTTCCGTGA
CGGTGCAGGAAGGCCCTGTGTGTCCATGTGCCCTGCTCCTTCTCCTACCCCTCGCATGGCTGGATT
TACCCTGGCCCAGTAGTTCATGGCTACTGGTTCCGGGAAGGGGCCAATACAGACCAGGATGCTCC
AGTGGCCACAAACAACCCAGCTCGGGCAGTGTGGGAGGAGACTCGGGACCGATTCCACCTCCTTG
GGGACCCACATACCAAGAATTGCACCCTGAGCATCAGAGATGCCAGAAGAAGTGATGCGGGGAGA
TACTTCTTTTCGTATGGAGAAAGGAAGTATAAAATGGAATTATAAACATCACCGGCTCTCTGTGAA
TGTGACAGCCTTGACCCACAGGCCAACATCCTCATCCCAGGCACCCCTGGAGTCCGGCTGCCCCC
AGAATCTGACCTGCTCTGTGCCCTGGGCCTGTGAGCAGGGGACACCCCTATGATCTCCTGGATA
GGGACCTCCGTGTCCCCCTGGACCCCTCCACCACCCGCTCCTCGGTGCTCACCTCATCCCACA
GCCCCAGGACCATGGCACCAGCCTCACCTGTCAGGTGACCTTCCCTGGGGCCAGCGTGACCACGA
ACAAGACCGTCCATCTCAACGTGTCCTACCCGCCTCAGAACTTGACCATGACTGTCTTCCAAGGA
GACGGCACAGTATCCACAGTCTTGGGAAATGGCTCATCTCTGTCACTCCCAGAGGGCCAGTCTCT
GCGCCTGGTCTGTGCAGTTGATGCAGTTGACAGCAATCCCCCTGCCAGGCTGAGCCTGAGCTGGA
GAGGCCTGACCCTGTGCCCCTCACAGCCCTCAAACCCGGGGGTGCTGGAGCTGCCTTGGGTGCAC
CTGAGGGATGCAGCTGAATTCACCTGCAGAGCTCAGAACCTCTCGGCTCTCAGCAGGTCTACCT
GAACGTCTCCCTGCAGAGCAAAGCCACATCAGGAGTGACTCAGGGGGTGGTCGGGGGAGCTGGAG
CCACAGCCCTGGTCTTCCTGTCTTCTGCGTCATCTTCGTTGTAGTGAGGTCCTGCAGGAAGAAA
TCGGCAAGGCCAGCAGCGGGCGTGGGAGATACGGGCATAGAGGATGCAAACGCTGTGAGGGGTC
AGCCTCTCAGGGGCCCCCTGACTGAACCTTGGGCAGAAGACAGTCCCCCAGACCAGCCTCCCCCAG
CTTCTGCCCCGCTCCTCAGTGGGGGAAGGAGAGCTCCAGTATGCATCCCTCAGCTTCCAGATGGTG
AAGCCTTGGGACTCGCGGGGACAGGAGGCCACTGACACCGAGTACTCGGAGATCAAGATCCACAG
ATGAGAACTGCAGAGACTCACCTGATTGAGGGATCACAGCCCCTCCAGGCAAGGGAGAAGTCA
GAGGCTGATTCTTGTAGAATTAACAGCCCTCAACGTGATGAGCTATGATAACACTATGAATTATG
TGCAGAGTGAAAAGCACACAGGCTTTAGAGTCAAAGTATCTCAAACCTGAATCCACACTGTGCCC
TCCCTTTTATTTTTTTAACTAAAAGACAGACAAATTCCTA

FIGURE 86

MLLLLLPLLWGRERAEGQTSKLLTMQSSVTVQEGLCVHVPCSFSPSHGWIYPGPVVHGYWFREG
ANTDQDAPVATNNPARAVWEETRDRFHLLGDPHTKNCTLSIRDARRSDAGRYFFRMEKGSIKWNY
KHHRLSVNVTALTHRPNILIPGTLESGCPQNLTCVWPWACEQGTPPMISWIGTSVSPLDPSTTRS
SVLTLIPQPQDHGTSLTCQVTFPGASVTTNKTVHLNVSYPQNLMTVFAQDGTVSTVLGNGSSL
SLPEGQSLRLVCAVDAVDSNPPARLSLSWRGLTLCPSQPSNPGVLELPWVHLRDAAEFTCRAQNP
LGSQQVYLNVSLSKATSGVTQGVVGGAGATALVFLSFCVIFVVVRSCRKKSARPAAGVGDGTGIE
DANAVRGASASQGPLEPWAEDSPPDQPPASARSSVGEGELQYASLSFQMKPWDSRGQEATDTE
YSEIKIHR

Signal peptide:

amino acids 1-15

Transmembrane domain:

amino acids 351-370

FIGURE 87

AGAAAGCTGCACTCTGTTGAGCTCCAGGGCGCAGTGGAGGGAGGGAGTGAAGGAGCTCTCTGTAC
CCAAGGAAAGTGCAGCTGAGACTCAGACAAGATTACAATGAACCAACTCAGCTTCCTGCTGTTTC
TCATAGCGACCACCAGAGGATGGAGTACAGATGAGGCTAATACTTACTTCAAGGAATGGACCTGT
TCTTCGTCTCCATCTCTGCCCAGAAGCTGCAAGGAAATCAAAGACGAATGTCCTAGTGCATTTGA
TGGCCTGTATTTTCTCCGCACTGAGAATGGTGTATCTACCAGACCTTCTGTGACATGACCTCTG
GGGTGGCGGCTGGACCCTGGTGGCCAGCGTGCATGAGAATGACATGCGTGGGAAGTGCACGGTG
GGCGATCGCTGGTCCAGTCAGCAGGGCAGCAAAGCAGACTACCCAGAGGGGGACGGCAACTGGGG
CAACTACAACACCTTTGGATCTGCAGAGGCGGCCACGAGCGATGACTACAAGAACCCTGGCTACT
ACGACATCCAGGCCAAGGACCTGGGCATCTGGCACGTGCCCCAATAAGTCCCCCATGCAGCACTGG
AGAAACAGCTCCCTGCTGAGGTACCGCACGGACACTGGCTTCCTCCAGACACTGGGACATAATCT
GTTTGGCATCTACCAGAAATATCCAGTGAAATATGGAGAAGGAAAGTGTTGGACTGACAACGGCC
CGGTGATCCCTGTGGTCTATGATTTTGGCGACGCCCAGAAAACAGCATCTTATTACTCACCTAT
GGCCAGCGGGAATTCAGTGCGGGATTTGTTCAAGTTCAGGGTATTTAATAACGAGAGAGCAGCCAA
CGCCTTGTGTGCTGGAATGAGGGTCACCGGATGTAACACTGAGCATCACTGCATTGGTGGAGGAG
GATACTTTCCAGAGGCCAGTCCCCAGCAGTGTGGAGATTTTCTGGTTTTGATTGGAGTGGATAT
GGAACATCATGTTGGTTACAGCAGCAGCCGTGAGATAACTGAGGCAGCTGTGCTTCTATTCTATCG
TTGAGAGTTTTGTGGGAGGGAACCCAGACCTCTCCTCCCAACCATGAGATCCCAAGGATGGAGAA
CAACTTACCCAGTAGCTAGAATGTTAATGGCAGAAGAGAAAACAATAAATCATATTGACTCAAGA
AAAAAA

FIGURE 88

MNQLSFLFLIATTRGWSTDEANTYFKEWTCSSSPSLPRSCKEIKDECPSAFDGLYFLRTENGVI
YQTFCDMTSGGGGWTLVASVHENDMRGKCTVGDRWSSQQGSKADYPEGDGNWANYNTFGSAEAAT
SDDYKNPGYYDIQAKDLGIWHVFNKSPMQHWRNSSLLRYRTDTGFLQTLGHNLEFGIYQKYPVKYG
EGKCWTDNGPVIPIVVYDFGDAQKTASYISPYGQREFTAGFVQFRVENNERAANALCAGMRVTGCN
TEHHCIGGGGYFPEASPPQCGDFSGFDWSGYGTHVGYSSSREITEAAVLLFYR

Important features:

Signal peptide:

amino acids 1-16

N-glycosylation site.

amino acids 163-167

Glycosaminoglycan attachment sites.

amino acids 74-78, 289-293

N-myristoylation sites.

amino acids 76-82, 115-121, 124-130, 253-259, 292-298

FIGURE 89

CTAGATTTGTCGGCTTGCGGGGAGACTTCAGGAGTCGCTGTCTCTGAACTTCCAGCCTCAGAGAC
CGCCGCCCTTGTCCTCCGAGGGCCATGGGCCGGGTCTCAGGGCTTGTGCCCTCTCGCTTCCTGACG
CTCCTGGCGCATCTGGTGGTCGTCATCACCTTATTCTGGTCCCGGGACAGCAACATACAGGCCTG
CCTGCCTCTCACGTTACCCCCGAGGAGTATGACAAGCAGGACATTAGCTGGTGGCCGCGCTCT
CTGTCACCCTGGGCCTCTTTGCAGTGGAGCTGGCCGGTTTCCTCTCAGGAGTCTCCATGTTCAAC
AGCACCCAGAGCCTCATCTCCATTGGGGCTCACTGTAGTGCATCCGTGGCCCTGTCTTCTTCAT
ATTGAGCGTTGGGAGTGCACTACGTATTGGTACATTTTTGTCTTCTGCAGTGCCCTTCCAGCTG
TCACTGAAATGGCTTTATTTCGTCACCGTCTTTGGGCTGAAAAGAAACCCTTCTTGATTACCTTCA
TGACGGGAACCTAAGGACGAAGCCTACAGGGGCAAGGGCCGCTTCGTATTCCTGGAAGAAGGAAG
GCATAGGCTTCGGTTTTCCCTCGGAAACTGCTTCTGCTGGAGGATATGTGTTGGAATAATTACG
TCTTGAGTCTGGGATTATCCGCATTGTATTTAGTGCTTTGTAATAAAATATGTTTGTAGTAACA
TTAAGACTTATATACAGTTTTAGGGGACAATTAAAAAAAAAAAA

FIGURE 90

MGRVSGLVPSRFLTLLAHLVVVITLFWSRDSNIQACLPLTFTPEEYDKQDIQLVAALSVTLGLFA
VELAGFLSGVSMFNSTQSLISIGAHCSASVALSFFIFERWECTTYWYIFVFCSALPAVTEMALFV
TVFGLKKKPF

Transmembrane domain:

amino acids 12-28 (type II), 51-66, 107-124

FIGURE 91

CTGGGACCCCGAAAAGAGAAGGGGAGAGCGAGGGGACGAGAGCGGAGGAGGAAGATGCAACTGAC
 TCGCTGCTGCTTCGTGTTCCCTGGTGCAGGGTAGCCTCTATCTGGTCATCTGTGGCCAGGATGATG
 GTCCTCCCGGCTCAGAGGACCCTGAGCGTGATGACCACGAGGGCCAGCCCCGGCCCCGGGTGCCT
 CGGAAGCGGGGCCACATCTCACCTAAGTCCCGCCCCATGGCCAATTCCACTCTCCTAGGGCTGCT
 GGCCCCGCTGGGGAGGCTTGGGGCATTCTTGGGCAGCCCCCAACCGCCGAACCACAGCCCC
 CACCCTCAGCCAAGGTGAAGAAAATCTTTGGCTGGGGCGACTTCTACTCCAACATCAAGACGGTG
 GCCCTGAACCTGCTCGTCACAGGGAAGATTGTGGACCATGGCAATGGGACCTTCAGCGTCCACTT
 CCAACACAATGCCACAGGCCAGGGAAACATCTCCATCAGCCTCGTGCCCCCAGTAAAGCTGTAG
 AGTTCCACCAGGAACAGCAGATCTTCATCGAAGCCAAGGCCTCCAAAATCTTCAACTGCCGGATG
 GAGTGGGAGAAGGTAGAACGGGGCGCCGGACCTCGCTTTGCACCCACGACCCAGCCAAGATCTG
 CTCCCGAGACCACGCTCAGAGCTCAGCCACCTGGAGCTGCTCCCAGCCCTTCAAAGTCGTCTGTG
 TCTACATCGCCTTCTACAGCACGGACTATCGGCTGGTCCAGAAGGTGTGCCCAGATTACAACCTAC
 CATAGTGATACCCCCTACTACCCATCTGGGTGACCCGGGGCAGGCCACAGAGGCCAGGCCAGGGC
 TGAAGGACAGGCCTGCCCATGCAGGAGACCATCTGGACACCGGGCAGGGAAGGGGTGGGCCTC
 AGGCAGGGAGGGGGGTGGAGACGAGGAGATGCCAAGTGGGGCCAGGGCCAAGTCTCAAGTGGCAG
 AGAAAGGGTCCCAAGTGCTGGTCCCAACCTGAAGCTGTGGAGTGACTAGATCACAGGAGCACTGG
 AGGAGGAGTGGGCTCTCTGTGCAGCCTCACAGGGCTTTGCCACGGAGCCACAGAGAGATGCTGGG
 TCCCCGAGGCCTGTGGGCAGGCCGATCAGTGTGGCCCCAGATCAAGTCATGGGAGGAAGCTAAGC
 CCTTGGTTCTTGCCATCCTGAGGAAAGATAGCAACAGGGAGGGGGAGATTTTCATCAGTGTGGACA
 GCCTGTCAACTTAGGATGGATGGCTGAGAGGGCTTCCTAGGAGCCAGTCAGCAGGGTGGGGTGGG
 GCCAGAGGAGCTCTCCAGCCCTGCCTAGTGGGCGCCCTGAGCCCTTGTCTGTGTGCTGAGCATGG
 CATGAGGCTGAAGTGGCAACCCTGGGGTCTTTGATGTCTTGACAGATTGACCATCTGTCTCCAGC
 CAGGCCACCCCTTTCCAAAATTCCCTCTTCTGCCAGTACTCCCCCTGTACCACCCATTGCTGATG
 GCACACCCATCCTTAAGCTAAGACAGGACGATTGTGGTCTCCACACTAAGGCCACAGCCCATC
 CGCGTGCTGTGTGTCCCTCTTCCACCCCAACCCCTGCTGGCTCCTCTGGGAGCATCCATGTCCCG
 GAGAGGGGTCCCTCAACAGTCAGCCTCACCTGTGACACCGGGGTTCTCCCGGATCTGGATGGCGC
 CGCCCTCTCAGCAGCGGGCACGGGTGGGGCGGGGCCGGGCCGAGAGCATGTGCTGGATCTGTTT
 TGTGTGTCTGTCTGTGGGTGGGGGAGGGGAGGGAAGTCTTGTGAAACCGCTGATTGCTGACTTT
 TGTGTGAAGAATCGTGTCTTGGAGCAGGAAATAAAGCTTGCCCCGGGGCA

FIGURE 92

MQLTRCCFVFLVQGSLYLVICGQDDGPPGSEDPERDDHEGQPRPRVPRKRGHISPKSRPMANSTL
LGLLAPPGEAWGILGQPPNRPNHSPPPSAKVKKIFGWGDFYSNIKTVALNLLVTGKIVDHGNGTF
SVHFQHNATGQGNISISLVPPSKAVEFHQEQQIFIEAKASKIFNCRMWEKVERGRRTSLCTHDP
AKICSRDHAQSSATWSCSQPFKVVCVYIAFYSTDYRLVQKVC PDYNYHSDTPYYPSG

Important features of the protein:

Signal peptide:

amino acids 1-14

N-glycosylation sites.

amino acids 62-65, 127-130, 137-140, 143-146

2-oxo acid dehydrogenases acyltransferase

amino acids 61-71

FIGURE 93

CGGTGGCCATGACTGCGGCCGTGTTCTTCGGCTGCGCCTTCATTGCCTTCGGGCCTGCGCTCGCC
CTTTATGTCTTCACCATCGCCATCGAGCCGTTGCGTATCATCTTCCTCATCGCCGGAGCTTTCTT
CTGGTTGGTGTCTCTACTGATTTTCGTCCCTTGTTTGGTTCATGGCAAGAGTCATTATTGACAACA
AAGATGGACCAACACAGAAATATCTGCTGATCTTTGGAGCGTTTGTCTCTGTCTATATCCAAGAA
ATGTTCCGATTTGCATATTATAAACTCTTAAAAAAGCCAGTGAAGGTTTGAAGAGTATAAACCC
AGGTGAGACAGCACCCCTCTATGCGACTGCTGGCCTATGTTTCTGGCTTGGGCTTTGGAATCATGA
GTGGAGTATTTTCCTTTGTGAATACCCTATCTGACTCCTTGGGGCCAGGCACAGTGGGCATTTCAT
GGAGATTCTCCTCAATTCTTCCTTTATTCAGCTTTCATGACGCTGGTCATTATCTTGCTGCATGT
ATTCTGGGGCATTGTATTTTTTGATGGCTGTGAGAAGAAAAAGTGGGGCATCCTCCTTATCGTTC
TCCTGACCCACCTGCTGGTGTGAGCCAGACCTTCATAAGTTCTTATTATGGAATAAACCTGGCG
TCAGCATTTATAATCCTGGTGCTCATGGGCACCTGGGCATTCTTAGCTGCGGGAGGCAGCTGCCG
AAGCCTGAAACTCTGCCTGCTCTGCCAAGACAAGAACTTTCTTCTTTACAACCAGCGCTCCAGAT
AACCTCAGGGAACCAGCACTTCCCAAACCGCAGACTACATCTTTAGAGGAAGCACAACTGTGCCT
TTTTCTGAAAATCCCTTTTTCTGGTGGAATTGAGAAAGAAATAAACTATGCAGATA

FIGURE 94

MTAAVFFGCAFIAGFPALALYVFTIAIEPLRIIFLIAGAFFWLVSLLISSLVWFMARVIIDNKDG
PTQKYLLIFGAFVSVYIQEMFRFAYYKLLKKASEGLKSINPGETAPSMRLLAYVSGLGFGIMSGV
FSFVNTLSDSLPGPTVGIHGDSPOFFLYSAFMTLVIIILLHVFWGIVFFDGCEKKKWGILLIVLLT
HLLVSAQTFISSYYGINLASAFIILVLMGTWAFILAAGGSCRSLKLCLLCQDNFLLYNQSR

Important features of the protein:

Signal peptide:

amino acids 1-19

Transmembrane domains:

amino acids 32-51, 119-138, 152-169, 216-235

Glycosaminoglycan attachment site.

amino acids 120-123

Sodium:neurotransmitter symporter family protein

amino acids 31-65

FIGURE 95

AATTTTTCACCAGAGTAAACTTGAGAAACCAACTGGACCTTGAGTATTGTACATTTTGCCTCGTG
GACCCAAAGGTAGCAATCTGAAACATGAGGAGTACGATTCTACTGTTTTGTCTTCTAGGATCAAC
TCGGTCATTACCACAGCTCAAACCTGCTTTGGGACTCCCTCCCACAAAACCTGGCTCCGGATCAGG
GAACACTACCAAACCAACAGCAGTCAAATCAGGTCTTTCCTTCTTTAAGTCTGATACCATTAACA
CAGATGCTCACACTGGGGCCAGATCTGCATCTGTAAATCCTGCTGCAGGAATGACACCTGGTAC
CCAGACCCACCCATTGACCCTGGGAGGGTTGAATGTACAACAGCAACTGCACCCACATGTGTTAC
CAATTTTGTGACACAACCTTGAGCCCAGGGCACTATCCTAAGCTCAGAGGAATTGCCACAAATC
TTCACGAGCCTCATCATCCATTCTTGTTCCTGGGAGGCATCCTGCCACCAGTCAGGCAGGGGC
TAATCCAGATGTCCAGGATGGAAGCCTTCCAGCAGGAGGAGCAGGTGTAAATCCTGCCACCCAGG
GAACCCAGCAGGCCGCCTCCCAACTCCCAGTGGCACAGATGACGACTTTGCAGTGACCACCCCT
GCAGGCATCCAAAGGAGCACACATGCCATCGAGGAAGCCACCACAGAATCAGCAAATGGAATTCA
GTAAGCTGTTTCAAATTTTTTCAACTAAGCTGCCTCGAATTTGGTGATACATGTGAATCTTTATC
ATTGATTATATTATGGAATAGATTGAGACACATTGGATAGTCTTAGAAGAAATTAATTCTTAATT
TACCTGAAAATATTCTTGAAATTTAGAAAATATGTTCTATGTAGAGAATCCCAACTTTTAAAAA
CAATAATTCAATGGATAAATCTGTCTTTGAAATATAACATTATGCTGCCTGGATGATATGCATAT
TAAACATATTTGGAACCTGGAA
AAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAA

FIGURE 96

MRSTILLFCLLGSTRSLPQLKPALGLPPTKLAPDQGTLPNQQQSNQVFPSLSLIPLTQM
LTLGPDLHLLNPAAGMTPGTQTHPLTLGGLNVQQQLHPHVLPIFVTQLGAQGTTLSSEE
LPQIFTSLIIHSLFPGGILPTSQAGANPDVQDGS LPAGGAGVNPATQGTPAGRLPTPSG
TDDDFAVTTPAGIQRSTHAIEEATTESANGIQ

Signal peptide:

amino acids 1-16

FIGURE 97

GCTCAAGTGCCCTGCCTTGCCCCACCCAGCCCAGCCTGGCCAGAGCCCCCTGGAGAAGGAGCTCT
CTTCTTGCTTGGCAGCTGGACCAAGGGAGCCAGTCTTGGGCGCTGGAGGGCCTGTCTGACCATG
GTCCCTGCCTGGCTGTGGCTGCTTTGTGTCTCCGTCCCCCAGGCTCTCCCAAGGCCCAGCCTGC
AGAGCTGTCTGTGGAAGTTCCAGAAAACCTATGGTGGAATTTCCCTTTATACCTGACCAAGTTGC
CGCTGCCCCGTGAGGGGGCTGAAGGCCAGATCGTGCTGTGAGGGGACTCAGGCAAGGCAACTGAG
GGCCCCATTGCTATGGATCCAGATTCTGGCTTCCTGCTGGTGACCAGGGCCCTGGACCGAGAGGA
GCAGGCAGAGTACCAGCTACAGGTCAACCCTGGAGATGCAGGATGGACATGTCTTGTGGGGTCCAC
AGCCTGTGCTTGTGCACGTGAAGGATGAGAATGACCAGGTGCCCCATTCTCTCAAGCCATCTAC
AGAGCTCGGCTGAGCCGGGGTACCAGGCCTGGCATCCCCTTCCTCTTCCTTGAGGCTTCAGACCG
GGATGAGCCAGGCACAGCCAACTCGGATCTTCGATTCCACATCCTGAGCCAGGCTCCAGCCCAGC
CTTCCCCAGACATGTTCCAGCTGGAGCCTCGGCTGGGGGCTCTGGCCCTCAGCCCCAAGGGGAGC
ACCAGCCTTGACCACGCCCTGGAGAGGACCTACCAGCTGTTGGTACAGGTCAAGGACATGGGTGA
CCAGGCCTCAGGCCACCAGGCCACTGCCACCGTGGAAGTCTCCATCATAGAGAGCACCTGGGTGT
CCCTAGAGCCTATCCACCTGGCAGAGAATCTCAAAGTCTTATACCCGCCACCACATGGCCCAGGTA
CACTGGAGTGGGGGTGATGTGCACTATCACCTGGAGAGCCATCCCCGGGACCCTTTGAAGTGAA
TGCAGAGGGAAACCTCTACGTGACCAGAGAGCTGGACAGAGAAGCCCAGGCTGAGTACCTGTCTC
AGGTGCGGGCTCAGAATTCCCATGGCGAGGACTATGCGGGCCCTCTGGAGCTGCACGTGCTGGTG
ATGGATGAGAATGACAACGTGCCATCTGCCCTCCCCGTGACCCACAGTCAGCATCCCTGAGCT
CAGTCCACCAGGTACTGAAGTGACTAGACTGTGAGCAGAGGATGCAGATGCCCCCGGCTCCCCCA
ATTCCACGTTGTGTATCAGCTCCTGAGCCCTGAGCCTGAGGATGGGGTAGAGGGGAGAGCCTTC
CAGGTGGACCCCACTTCAGGCAGTGTGACGCTGGGGGTGCTCCCACTCCGAGCAGGCCAGAACAT
CCTGCTTCTGGTGCTGGCCATGGACCTGGCAGGCGCAGAGGGTGGCTTCAGCAGCACGTGTGAAG
TCGAAGTCGAGTCACAGATATCAATGATCACGCCCTGAGTTTCATCACTTCCAGATTGGGCCT
ATAAGCCTCCCTGAGGATGTGGAGCCCGGACTCTGGTGGCCATGCTAACAGCCATTGATGCTGA
CCTCGAGCCCGCCTTCCGCCTCATGGATTTTGCCATTGAGAGGGGAGACACAGAAGGGACTTTTG
GCCTGGATTGGGAGCCAGACTCTGGGCATGTTAGACTCAGACTCTGCAAGAACCTCAGTTATGAG
GCAGCTCCAAGTCATGAGGTGGTGGTGGTGGTGCAGAGTGTGGCGAAGCTGGTGGGGCCAGGCCC
AGGCCCTGGAGCCACCGCCACGGTGAAGTGTGCTAGTGGAGAGAGTGTGCCACCCCCAAGTTGG
ACCAGGAGAGCTACGAGGCCAGTGTCCCCATCAGTGCCCCAGCCGGCTCTTTCCTGTGACCATC
CAGCCCTCCGACCCCATCAGCCGAACCCCTCAGGTTCTCCCTAGTCAATGACTCAGAGGGCTGGCT
CTGCATTGAGAAATTCTCCGGGGAGGTGCACACCGCCAGTCCCTGCAGGGCGCCAGCCTGGGG
ACACCTACACGGTGCTTGTGGAGGGCCAGGATACAGCCCTGACTCTTGCCCTGTGCCCTCCCAA
TACCTCTGCACACCCCGCCAAGACCATGGCTTGATCGTGAGTGGACCCAGCAAGGACCCGATCT
GGCCAGTGGGCACGGTCCCTACAGCTTCACCCCTTGGTCCCAACCCACGGTGCACGGGATTGGC
GCCTCCAGACTCTCAATGGTTCCCATGCCACCTCACCTTGGCCCTGCATTGGGTGGAGCCACGT
GAACACATAATCCCCGTGGTGGTCAGCCACAATGCCAGATGTGGCAGCTCCTGGTTCGAGTGAT
CGTGTGTCGCTGCAACGTGGAGGGGCAGTGCATGCGCAAGGTGGGCCGATGAAGGGCATGCCCA
CGAAGCTGTCGGCAGTGGGCATCCTTGTAGGCACCCTGGTAGCAATAGGAATCTTCTCATCCTC
ATTTTCACCCACTGGACCATGTCAAGGAAGAAGGACCCGGATCAACCAGCAGACAGCGTGCCCCCT
GAAGGCGACTGTC**TGA**ATGGCCCAGGCAGCTCTAGCTGGGAGCTTGGCCTCTGGCTCCATCTGAG
TCCCCCTGGGAGAGAGCCAGCACCCCAAGATCCAGCAGGGGACAGGACAGAGTAGAAGCCCCCTCCA
TCTGCCCTGGGGTGGAGGCACCATCACCATCACCAGGCATGTCTGCAGAGCCTGGACACCAACTT
TATGGACTGCCCATGGGAGTGCTCCAAATGTCAGGGTGTTCGCCAATAATAAAGCCCCAGAGAA
CTGGGCTGGGCCCTATGGGAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAG

FIGURE 98

MVPAWLWLLCVSVPQALPKAQP AELSVEVPENYGGNFPLYLTKLPLPREGAEGQIVLSGDSGKAT
EGPFAMDPDSGFLLVTRALDREEQAEYQLQVTLEMQDGHVLWGPPVVLVHVKDENDQVP HFSQAI
YRARLSRGTRPGIPFLFLEASDRDEPGTANSDLRFHILSQAPAQSPDMFQLEPRLGALALSPKG
STSLDHALERTYQLLVQVKMDGDAQSGHQATATVEVSI IESTWVSLEPIHLAENLKVLYPHHMAQ
VHWSGGDVHYHLESHPPGPF EVNAEGNLYVTRELDREAQAEYLLQVRAQNSHGEDIAAPLELHVL
VMDENDNVPICPPRDPTVSIPELSPPGTEVTRL SAEDADAPGSPNSHV VYQLLSPEPEDGVEGRA
FQVDPTSGSVTLGVLPLRAGQNILLVLAMDLAGAEGGFSSTCEVEVAVTDINDHAPEFITSQIG
PISLPEDVEPGTLVAMLT AIDADLEPAFRLMDFAIERGDTEGTFGLDWEPDSGHVRLRLCKNLSY
EAAPSHEVVVVVQSVAKLVGPGPGGATATVTVLVERVMPPPKLDQESYEASVPISAPAGSFLLT
IQPSDPISRTLRFSLVNDSEGWLCIEKFSGEVHTAQSLQGAQPGDTYTVLVEAQDTALT LAPVPS
QYLCTPRQDHGLIVSGPSKDPDLASGHGPYSFTLGPNPTVQRDWRLQTLNGSHAYLT LALHWVEP
REHIIPVVVSHNAQMWQLLVRVIVCRCNVEGQCMRKVGRMKGMPTKLSAVGILVGT LVAIGIFLI
LIFTHWTMSRKKDPDQPADSVPLKATV

Signal peptide:

amino acids 1-18

Transmembrane domain:

amino acids 762-784

FIGURE 99

GGCTGACCGTGCTACATTGCCTGGAGGAAGCCTAAGGAACCCAGGCATCCAGCTGCCCCACGCCTG
 AGTCCAAGATTCTTCCCAGGAACACAAACGTAGGAGACCCACGCTCCTGGAAGCACCAGCCTTTA
 TCTCTTCACCTTCAAGTCCCCTTTCTCAAGAATCCTCTGTTCTTTGCCCTCTAAAGTCTTGGTAC
 ATCTAGGACCCAGGCATCTTGCTTTCCAGCCACAAAGAGACAGATGAAGATGCAGAAAAGGAAATG
 TTCTCCTTATGTTTGGTCTACTATTGCATTTAGAAGCTGCAACAAATTCCAATGAGACTAGCACC
 TCTGCCAACACTGGATCCAGTGTGATCTCCAGTGGAGCCAGCACAGCCACCAACTCTGGGTCCAG
 TGTGACCTCCAGTGGGGTCAGCACAGCCACCATCTCAGGGTCCAGCGTGACCTCCAATGGGGTCA
 GCATAGTCACCAACTCTGAGTTCCATACAACCTCCAGTGGGATCAGCACAGCCACCAACTCTGAG
 TTCAGCACAGCGTCCAGTGGGATCAGCATAGCCACCAACTCTGAGTCCAGCACAACTCCAGTGG
 GGCCAGCACAGCCACCAACTCTGAGTCCAGCACACCCTCCAGTGGGGCCAGCACAGTCCACCAACT
 CTGGGTCCAGTGTGACCTCCAGTGGAGCCAGCACTGCCACCAACTCTGAGTCCAGCACAGTGTCC
 AGTAGGGGCCAGCATGCCACCAACTCTGAGTCTAGCACACTCTCCAGTGGGGCCAGCACAGCCAC
 CAACTCTGACTCCAGCACAACTCCAGTGGGGCTAGCACAGCCACCAACTCTGAGTCCAGCACAA
 CCTCCAGTGGGGCCAGCACAGCCACCAACTCTGAGTCCAGCACAGTGTCCAGTAGGGCCAGCACT
 GCCACCAACTCTGAGTCCAGCACAACTCCAGTGGGGCCAGCACAGCCACCAACTCTGAGTCCAG
 AACGACCTCCAATGGGGCTGGCACAGCCACCAACTCTGAGTCCAGCACGACCTCCAGTGGGGCCA
 GCACAGCCACCAACTCTGACTCCAGCACAGTGTCCAGTGGGGCCAGCACTGCCACCAACTCTGAG
 TCCAGCACGACCTCCAGTGGGGCCAGCACAGCCACCAACTCTGAGTCCAGCACGACCTCCAGTGG
 GGCTAGCACAGCCACCAACTCTGACTCCAGCACAACTCCAGTGGGGCCGGCACAGCCACCAACT
 CTGAGTCCAGCACAGTGTCCAGTGGGATCAGCACAGTCCCAATTCTGAGTCCAGCACACCCTCC
 AGTGGGGCCAACACAGCCACCAACTCTGAGTCCAGTACGACCTCCAGTGGGGCCAACACAGCCAC
 CAACTCTGAGTCCAGCACAGTGTCCAGTGGGGCCAGCACTGCCACCAACTCTGAGTCCAGCACAA
 CCTCCAGTGGGGTCAGCACAGCCACCAACTCTGAGTCCAGCACAACTCCAGTGGGGCTAGCACA
 GCCACCAACTCTGACTCCAGCACAACTCCAGTGGGGCCAGCACAGCCACCAACTCTGAGTCTAG
 CACAGTGTCCAGTGGGATCAGCACAGTCCCAATTCTGAGTCCAGCACAACTCCAGTGGGGCCA
 ACACAGCCACCAACTCTGGGTCCAGTGTGACCTCTGCAGGCTCTGGAACAGCAGCTCTGACTGGA
 ATGCACACAACTTCCCATAGTGCATCTACTGCAGTGAAGGCAAAGCCTGGTGGGTCCCTGGT
 GCCGTGGGAAATCTTCTCATCACCTGGTCTCGGTTGTGGCGGCCGTGGGGCTCTTTGCTGGGC
 TCTTCTTCTGTGTGAGAAACAGCCTGTCCCTGAGAAACACCTTTAACACAGTGTCTACCACCCT
 CATGGCCTCAACCATGGCCTTGGTCCAGGCCCTGGAGGGAATCATGGAGCCCCCACAGGCCAG
 GTGGAGTCCTAACTGGTTCTGGAGGAGACCAGTATCATCGATAGCCATGGAGATGAGCGGGAGGA
 ACAGCGGGCCCTGAGCAGCCCCGGAAGCAAGTGCCGCATTCTTCAGGAAGGAAGAGACCTGGGCA
 CCCAAGACCTGGTTTCTTTTCATTTCATCCAGGAGACCCCTCCCAGCTTTGTTTGGATCCTGAA
 AATCTTGAAGAAGGTATTCTCACCTTTCTTGCCTTTACCAGACACTGGAAAGAGAATACTATAT
 TGCTCATTTAGCTAAGAAATAAATACATCTCATCTAACACACACGACAAAGAGAAGCTGTGCTTG
 CCCCAGGGTGGGTATCTAGCTCTGAGATGAACTCAGTTATAGGAGAAAACCTCCATGCTGGACTC
 CATCTGGCATTCAAAATCTCCACAGTAAAATCCAAAGACCTCAAAAAAAAAAAAAAAAAAAAAA
 AAAAAAAAAAAAAAAAAAAAAAAAAAAAAA

FIGURE 100

MKMQKGNVLLMFGLLLHLEAATNSNETSTTSANTGSSVSSGASTATNSGSSVTSSGVSTATISGS
SVTSNGVSIVTNSEFHTTSSGISTATNSEFSTASSGISIATNSESSTTSSGASTATNSESSTPSS
GASTVTNSGSSVTSSGASTATNSESSTVSSRASTATNSESSTLSSGASTATNSDSSTTSSGASTA
TNSESSTTSSGASTATNSESSTVSSRASTATNSESSTTSSGASTATNSESRTTSNGAGTATNSES
STTSSGASTATNSDSSTVSSGASTATNSESSTTSSGASTATNSESSTTSSGASTATNSDSSTTSS
GAGTATNSESSTVSSGISTVTNSESSTPSSGANTATNSESSTTSSGANTATNSESSTVSSGASTA
TNSESSTTSSGVSTATNSESSTTSSGASTATNSDSSTTSSEASTATNSESSTVSSGISTVTNSES
STTSSGANTATNSGSSVTSAGSGTAALTGMHTTSHSASTAVSEAKPGGSLVPWEIFLITLVSVVA
AVGLFAGLFFCVRNSLSLRNTFNTAVYHPHGLNHGLGPGPGGNHGAPHRPRWSPNWFWRPVSII
AMEMSGRNSGP

Signal peptide:

amino acids 1-20

Transmembrane domain:

amino acids 510-532

FIGURE 101

GGCCGGACGCCTCCGCGTTACGGGATGAATTAACGGCGGGTTCCGCACGGAGGTTGTGACCCCTA
CGGAGCCCCAGCTTGCCACGCACCCCACTCGGCGTCGCGCGGCGTGCCCTGCTTGTCACAGGTG
GGAGGCTGGAACATATCAGGCTGAAAAACAGAGTGGGTACTCTCTTCTGGGAAGCTGGCAACAAAT
GGATGATGTGATATATGCAATTCCAGGGGAAGGGAATTTGTGGTGCTTCTGAACCCATGGTCAATT
AACGAGGCAGTTTCTAGCTACTGCACGTACTTCATAAAGCAGGACTCTAAAAGCTTTGGAATCAT
GGTGTGTCATGGAAAGGGATTTACTTTATACTGACTCTGTTTTGGGGAAGCTTTTTTGGGAAGCATTT
TCATGCTGAGTCCCTTTTTTACCTTTGATGTTTGTAAACCCATCTTGGTATCGCTGGATCAACAAC
CGCCTTGTGGCAACATGGCTCACCTACCTGTGGCATTATTGGAGACCATGTTTGGTGTAAGT
GATTATAACTGGGGATGCATTTGTTCTGGAGAAAGAGTGTCAATTATCATGAACCATCGGACAA
GAATGGACTGGATGTTCTGTGGAATTGCCTGATGCGATATAGCTACCTCAGATTGGAGAAAATT
TGCCTCAAAGCGAGTCTCAAAGGTGTTCTGGATTTGGTTGGGCCATGCAGGCTGCTGCCTATAT
CTTCATTCATAGGAAATGGAAGGATGACAAGAGCCATTTCGAAGACATGATTGATTACTTTTGTG
ATATTCACGAACCACTTCAACTCCTCATATTTCCAGAAGGGACTGATCTCACAGAAAACAGCAAG
TCTCGAAGTAATGCATTTGCTGAAAAAATGGACTTCAGAAATATGAATATGTTTTACATCCAAG
AACTACAGGCTTTACTTTTGTGGTAGACCGTCTAAGAGAAGGTAAGAACCTTGATGCTGTCCATG
ATATCACTGTGGCGTATCCTCACAACATTCCTCAATCAGAGAAGCACCTCCTCCAAGGAGACTTT
CCCAGGGAAATCCACTTTCACGTCCACCGGTATCCAATAGACACCTCCCCACATCCAAGGAGGA
CCTTCAACTCTGGTGCCACAAACGGTGGGAAGAGAAAGAAGAGAGGCTGCGTTCCTTCTATCAAG
GGGAGAAGAATTTTTATTTTACCGGACAGAGTGTCAATTCACCTTGCAAGTCTGAACTCAGGGTC
CTTGTGGTCAAATTGCTCTCTATACTGTATTGGACCCTGTTAGCCCTGCAATGTGCCTACTCAT
ATATTTGTACAGTCTTGTTAAGTGGTATTTTATAATCACCATTGTAATCTTTGTGCTGCAAGAGA
GAATATTTGGTGGACTGGAGATCATAGAACTTGCATGTTACCGACTTTTACACAAACAGCCACAT
TTAAATTCAAAGAAAAATGAGTAAGATTATAAGGTTTGCCATGTGAAAACCTAGAGCATATTTTG
GAAATGTTCTAAACCTTTCTAAGCTCAGATGCATTTTTGCATGACTATGTGCAATATTTCTTACT
GCCATCATTATTTGTTAAAGATATTTTGCACCTTAATTTTGTGGGAAAAATATTGCTACAATTTTT
TTTAATCTCTGAATGTAATTTGATACTGTGTACATAGCAGGGAGTGATCGGGGTGAAATAACTT
GGGCCAGAATATTATTAAACAATCATCAGGCTTTTAAA

FIGURE 102

MHSRGREIVVLLNPWSINEAVSSYCTYFIKQDSKSFSGIMVSWKGIYFILTLFWGSFFGSIFMLSP
FLPLMFVNPSWYRWINNRLVATWLTLPVALLETMFGVKVIITGDAFVPGERSVIIMNHRTRMDWM
FLWNCLMRYSYLRLEKICLKASLKGVPFGFGWAMQAAAYIFIHRKWKDDKSHFEDMIDYFCDIHEP
LQLLIFPEGTDLTENSKSRSNFAEKNGLQKYEYVLHPRTTGFTFVVDRLREGKNLDAVHDITVA
YPHNIPQSEKHLLQGDFPREIHFHVHRYPIDTLPTSKEDLQLWCHKRWEEKEERLRSFYQGEKNF
YFTGQSVIPPCKSELRLVLVVKLLSILYWTLESPAMCLLIYLYSLVKWYFIITIVIFVLQERIFGG
LEIIELACYRLLHKQPHLNSKKNE

Important features of the protein:

Signal peptide:

amino acids 1-22

Transmembrane domains:

amino acids 44-63, 90-108, 354-377

FIGURE 103

CGGCTCGAGCGGCTCGAGTGAAGAGCCTCTCCACGGCTCCTGCGCCTGAGACAGCTGGCCTGACC
TCCAAATCATCCATCCACCCCTGCTGTCTCATCTGTTTTTCATAGTGTGAGATCAACCCACAGGAATA
TCCATGGCTTTTGTGCTCATTGTTGTTCTCAGTTTCTACGAGCTGGTGTGAGGACAGTGGCAAGT
CACTGGACCGGGCAAGTTTGTCCAGGCCTTGGTGGGGGAGGACGCCGTGTTCTCCTGCTCCCTCT
TTCCTGAGACCAGTGCAGAGGCTATGGAAGTGGGTTCTTCAGGAATCAGTTCCATGCTGTGGTCT
CACCTCTACAGAGATGGGGAAGACTGGGAATCTAAGCAGATGCCACAGTATCGAGGGGAGAAGTGA
GTTTGTGAAGGACTCCATTGCAGGGGGGCGTGTCTCTCTAAGGCTAAAAACATCACTCCCTCGG
ACATCGGCCTGTATGGGTGCTGGTTCAGTTCCCAGATTTACGATGAGGAGGCCACCTGGGAGCTG
CGGGTGGCAGCACTGGGCTCACTTCTCTCATTCCATCGTGGGATATGTTGACGGAGGTATCCA
GTTACTCTGCCTGTCTCAGGCTGGTTCCTCCAGCCACAGCCAAGTGGAAAGGTCCACAAGGAC
AGGATTTGTCTTCAGACTCCAGAGCAAATGCAGATGGGTACAGCCTGTATGATGTGGAGATCTCC
ATTATAGTCCAGGAAAATGCTGGGAGCATATTGTGTTCCATCCACCTTGCTGAGCAGAGTCAATGA
GGTGGAAATCCAAGGTATTGATAGGAGAGACGTTTTTCCAGCCCTCACCTTGGCGCCTGGCTTCTA
TTTTACTCGGGTTACTCTGTGGTGCCTGTGTGGTGTGTGTCATGGGGATGATAATTGTTTTCTTC
AAATCCAAAGGGAAAATCCAGGCGGAAGTGGACTGGAGAAGAAAGCACGGACAGGCAGAATTGAG
AGACGCCCCGAAACACGCAGTGGAGGTGACTCTGGATCCAGAGACGGCTCACCCGAAGCTCTGCG
TTTCTGATCTGAAAAGTGAACCCATAGAAAAGCTCCCCAGGAGGTGCCTCACTCTGAGAAGAGA
TTTACAAGGAAGAGTGTGGTGGCTTCTCAGGGTTTCCAAGCAGGGAGACATTACTGGGAGGTGGA
CGTGGGACAAAATGTAGGGTGGTATGTGGGAGTGTGTGGGATGACGTAGACAGGGGGAAGAACA
ATGTGACTTTGTCTCCCAACAATGGGTATTGGGTCTCAGACTGACAACAGAACATTTGTATTTT
ACATTCAATCCCCATTTTATCAGCCTCCCCCCCCAGCACCCCTCCTACACGAGTAGGGGTCTTCT
GGACTATGAGGGTGGGACCATCTCCTTCTCAATACAAATGACCAGTCCCTTATTTATACCCTGC
TGACATGTGAGTTTGAAGGCTTGTTGAGACCCTATATCCAGCATGCGATGTATGACGAGGAAAAG
GGGACTCCCATATTATATGTCCAGTGTCTGGGGATGAGACAGAGAAGACCCTGCTTAAAGGGC
CCCACACCACAGACCCAGACACAGCCAAGGGAGAGTGTCTCCGACAGGTGGCCCCAGCTTCTCT
CCGGAGCCTGCGCACAGAGAGTCACGCCCCCACTCTCCTTTAGGGAGCTGAGGTCTTCTGCCC
TGAGCCCTGCAGCAGCGGCAGTCACAGCTTCCAGATGAGGGGGGATTGGCCTGACCCTGTGGGAG
TCAGAAGCCATGGCTGCCCTGAAGTGGGGACGGAATAGACTCACATTAGGTTTAGTTTGTGAAAA
CTCCATCCAGCTAAGCGATCTTGAACAAGTCACAACCTCCCAGGCTCCTCATTGTGCTAGTCACGG
ACAGTGATTCTGCTCACAGGTGAAGATTAAAGAGACAACGAATGTGAATCATGCTTGCAGGTT
TGAGGGCACAGTGTGCTAATGATGTGTTTTTATATTATACATTTTCCCACCATAAACTCTGTT
TGCTTATTCACATTAATTTACTTTTCTCTATACCAAATCACCCATGGAATAGTTATTGAACACC
TGCTTTGTGAGGCTCAAAGAATAAAGAGGAGGTAGGATTTTTCACTGATTCTATAAGCCCAGCAT
TACCTGATACCAAAACCAGGCAAAGAAAACAGAAGAAGAGGAAGGAAAACCTACAGGTCCATATCC
CTCATTAAACACAGACACAAAAATTTCTAAATAAAATTTTAAACAAATTAAGTAAACAATATATTTA
AAGATGATATATACTACTCAGTGTGGTTTGTCCACAAATGCAGAGTTGGTTTTAATATTTAAAT
ATCAACCAGTGTAATTCAGCACATTAATAAAGTAAAAAAGAAAACCATAAAAAAAAAAAAAA

FIGURE 104

MAFVLILVLSFYELVSGQWQVTGPGKFVQALVGEDAVFSCSLFPETSAEAMEVRFFRNQFHAVVH
LYRDGEDWESKQMPQYRGRTEFVKDSIAGGRVSLRLKNITPSDIGLYGCWFSSQIYDEEATWELR
VAALGSLPLISIVGYVDGGIQLLCLSSGWFPQPTAKWKGPQGQDLSSDSRANADGYSLYDVEISI
IVQENAGSILCSIHLEQSHEVESKVLIGETFFQPSPWRLASILLGLLCGALCGVVMGMIIVFFK
SKGKIQAELDWRRKHGQAEIRDARKHAVEVTLDPETAHPKLCVSDLKTVTHRKAPEVPHSEKRF
TRKSVVASQGFQAGRHYWEVDVGQNVGWYVGVCRDDVDVRGKNNVTLSPNNGYWVLRRLTTEHLYFT
FNPHFISLPPSTPPTRVGVFLDYEGGTISFFNTNDQSLIYTLLTCQFEGLLRPYIQHAMYDEEKG
TPIFICPVSWG

Signal peptide:

amino acids 1-17

Transmembrane domains:

amino acids 131-150, 235-259

FIGURE 105

CCTTCACAGGACTCTTCATTGCTGGTTGGCAATGATGTATCGGCCAGATGTGGTGAGGGCTAGGAAAAGAG
 TTTGTTGGGAACCCCTGGGTTATCGGCCTCGTCATCTTCATATCCCTGATTGTCCTGGCAGTGTGCATTGGA
 CTCCTGTTCATTATGTGAGATATAATCAAAGAAGACCTACAATTACTATAGCACATTGTCATTTACAAC
 TGACAAACTATATGCTGAGTTTGGCAGAGAGGCTTCTAACAATTTTACAGAAATGAGCCAGAGACTTGAAT
 CAATGGTGAAAAATGCATTTTATAAATCTCCATTAAGGGAAGAATTTGTCAAGTCTCAGGTTATCAAGTTC
 AGTCAACAGAAGCATGGAGTGTGGCTCATATGCTGTTGATTTGTAGATTTCACTCTACTGAGGATCCTGA
 AACTGTAGATAAAATTGTTCAACTTGTTTTACATGAAAAGCTGCAAGATGCTGTAGGACCCCTAAAGTAG
 ATCCTCACTCAGTTAAATTAATAAATCAACAAGACAGAAACAGACAGCTATCTAAACCATTGCTGCGGA
 ACACGAAGAAGTAAACTCTAGGTGAGTCTCAGGATCGTTGGTGGGACAGAAGTAGAAGAGGGTGAATG
 GCCCTGGCAGGCTAGCCTGCAGTGGGATGGGAGTCATCGCTGTGGAGCAACCTTAATTAATGCCACATGGC
 TTGTGAGTGCTGCTCACTGTTTTACAACATATAAGAACCCTGCCAGATGGACTGCTTCCTTTGGAGTAACA
 ATAAACCTTCGAAAATGAAACGGGGTCTCCGGAGAATAATTGTCCATGAAAAATACAAACACCCATCACA
 TGACTATGATATTTCTCTTGCGAGAGCTTTCTAGCCCTGTTCCCTACACAAATGCAGTACATAGAGTTTGTCT
 TCCCTGATGCATCCTATGAGTTTCAACCAGGTGATGTGATGTTTGTGACAGGATTTGGAGCACTGAAAAAT
 GATGTTTACAGTCAAAATCATCTTCGACAAGCACAGGTGACTCTCATAGACGCTACAACCTGCAATGAACC
 TCAAGCTTACAATGACGCCATAACTCCTAGAATGTTATGTGCTGGCTCCTTAGAAGGAAAAACAGATGCAT
 GCCAGGGTGACTCTGGAGGACCACTGGTTAGTTCAGATGCTAGAGATATCTGGTACCTTGCTGGAATAGTG
 AGCTGGGGAGATGAATGTGCGAAACCCAACAAGCCTGGTGTTTATACTAGAGTTACGGCCTTGCGGGACTG
 GATTACTTCAAAAACCTGGTATCTTAAGAGACAAAAGCCTCATGGAACAGATAACATTTTTTTTTGTTTTTTG
 GGTGTGGAGGCCATTTTTAGAGATACAGAATTGGAGAAGACTTGCAAAACAGCTAGATTTGACTGATCTCA
 ATAAACTGTTTGCTTGATGCATGTATTTCTTCCAGCTCTGTTCCGCACGTAAGCATCCTGCTTCTGCCA
 GATCAACTCTGTCATCTGTGAGCAATAGTTGAAACTTTATGTACATAGAGAAATAGATAATACAATATTAC
 ATTACAGCCTGTATTCAATTTGTTCTCTAGAAGTTTTGTGAGAATTTTGACTTGTTGACATAAATTTGTAAT
 GCATATATACAATTTGAAGCACTCCTTTCTTCAGTTCCTCAGCTCCTCTCATTTTCAAGCAATATCCATTT
 TCAAGGTGCAGAACAAGGAGTGAAAGAAAAATATAAGAAGAAAAAAATCCCCTACATTTTATTGGCACAGAA
 AAGTATTAGGTGTTTTTCTTAGTGGAATATTAGAAATGATCATATTCAATTATGAAAGGTCAAGCAAAGACA
 GCAGAATACCAATCACTTCATCATTTAGGAAGTATGGGAAGTAAAGTAAAGGAAGTCCAGAAAGAAGCCAAG
 ATATATCCTTATTTTCATTTCCAAACAACACTACTATGATAAATGTGAAGAAGATTCTGTTTTTTTGTGACCT
 ATAATAATTATACAAACTTCATGCAATGTACTTGTCTAAGCAAATTAAGCAAATATTTATTTAACATTG
 TTACTGAGGATGTCAACATATAACAATAAAATATAAATCACCCA

FIGURE 106

MMYRPDVVRARKRVCWEPWVIGLVIFISLIVLAVCIGLTVHYVRYNQKKTYNYSTLSFTTDKLY
AEFGREASNNFTEMSQRLESMVKNAFYKSPLREEFVKSQVIKFSQQKHGVLAHMLLICRFHSTED
PETVDKIVQLVLHEKLQDAVGPPKVDPHSVKIKKINKTETDSYLNHCCGTRRSKTLGQSLRIVGG
TEVEEGEPWQASLQWDGSHRCGATLINATWLVSAAHCFTTYKNPARWTASFGVTIKPSKMKRGL
RRIIVHEKYKHPSHDYDISLAELSSPVPTYNAVHRVCLPDASYEFQPGDVMFVTGFGALKNDGYS
QNHLRQAQVTLIDATTCNEPQAYNDAITPRMLCAGSLEGKTDACQGDSGGPLVSSDARDIWYLAG
IVSWGDECAKPNKPGVYTRVTALRDWITSKTGI

Transmembrane domain:

amino acids 21-40 (type II)

FIGURE 107

AGAGAAAGAAGCGTCTCCAGCTGAAGCCAATGCAGCCCTCCGGCTCTCCGCGAAGAAGTTCCCTG
 CCCCAGATGAGCCCCCGCGTGCCTCCCCGACTATCCCCAGGCGGGCGTGGGGCACCAGGGCCCAGC
 GCGGACGATCGCTGCCGTTTTGCCCTTGGGAGTAGGATGTGGTGAAAGGATGGGGCTTCTCCCTT
 ACGGGGCTCACAAATGSCCAGAGAAGATTCCGTGAAGTGTCTGCGCTGCCTGCTCTACGCCCTCAA
 TCTGCTCTTTTTGGTTAATGTCCATCAGTGTGTTGGCAGTTTTCTGCTTGGATGAGGGACTACCTAA
 ATAATGTTCTCACTTTAACTGCAGAAACGAGGGTAGAGGAAGCAGTCATTTTGACTTACTTTTCCT
 GTGGTTCATCCGCTCATGATTGCTGTTTGTGTTTTCCTTATCATTGTGGGGATGTTAGGATATTG
 TGAACGGTGAAAAGAAATCTGTTGCTTCTTGTCATGGTACTTTGGAAGTTTGTCTGTCAATTTTCT
 GTGTAGAACTGGCTTGTGGCGTTTGGACATATGAACAGGAACCTTATGGTTCCAGTACAATGGTCA
 GATATGGTCACTTTGAAAGCCAGGATGACAAATTATGGATTACCTAGATATCGGTGGCTTACTCA
 TGCTTGGAAATTTTTTTCAGAGAGAGTTTAAAGTGTGTGGAGTAGTATATTTCACTGACTGGTTGG
 AAATGACAGAGATGGACTGGCCCCCAGATTCTGTCTGTGTTAGAGAATTCAGGATGTTCCAAA
 CAGGCCACCAGGAAGATCTCAGTGACCTTTATCAAGAGGGTTGTGGGAAGAAAATGTATTCTCTT
 TTTGAGAGGAACCAAAACAACTGCAGGTGCTGAGGTTTCTGGGAATCTCCATTGGGGTGACACAAA
 TCCTGGCCATGATTCTCACCATTACTCTGCTCTGGGCTCTGTATTATGATAGAAGGGAGCCTGGG
 ACAGACCAAAATGATGTCCCTGAAGAATGACAACTCTCAGCACCTGTCATGTCCCTCAGTAGAACT
 GTTGAAACCAAGCCTGTCAAGAATCTTTGAACACACATCCATGGCAAACAGCTTTAATACACACT
 TTGAGATGGAGGAGTTATATAAAGAAATGTCACAGAAGAAAACCACAAACTTGTTTTATTGGACT
 TGTGAATTTTTGAGTACATACTATGTGTTTCAGAAATATGTAGAAATAAAATGTTGCCATAAAA
 TAACACCTAAGCATATACTATTCTATGCTTTAAATGAGGATGGAAAAGTTTCATGTCATAAGTC
 ACCACCTGGACAATAATTGATGCCCTTAAATGCTGAAGACAGATGTCATACCCACTGTGTAGCC
 TGTGTATGACTTTTACTGAACACAGTTATGTTTTGAGGCAGCATGGTTTGATTAGCATTTCCGCA
 TCCATGCAAACGAGTCACATATGGTGGGACTGGAGCCATAGTAAAGGTTGATTTACTTCTACCAA
 CTAGTATATAAAGTACTAATTAAATGCTAACATAGGAAGTTAGAAAATACTAATAACTTTTATTA
 CTCAGCGATCTATTCTTCTGATGCTAAATAAATTATATATCAGAAAACCTTTCAATATTGGTGACT
 ACCTAAATGTGATTTTTGCTGGTTACTAAAATATTCTTACCCTTAAAAGAGCAAGCTAACACAT
 TGTCTTAAGCTGATCAGGGATTTTTTGTATATAAGTCTGTGTTAAATCTGTATAATTCAGTCGAT
 TTCAGTTCTGATAATGTTAAGAATAACCATTATGAAAAGGAAAATTTGTCCTGTATAGCATCATT
 ATTTTTAGCCTTTCTGTGTTAATAAAGCTTTACTATTCTGTCCTGGGCTTATATTACACATATAAC
 TGTTATTTAAATACTTAACCACTAATTTTGAATAATTACCAGTGTGATACATAGGAATCATTATTC
 AGAATGTAGTCTGGTCTTTAGGAAGTATTAATAAGAAAATTTGCACATAACTTAGTTGATTGAGA
 AAGGACTTGTATGCTGTTTTCTCCCAAATGAAGACTCTTTTTGACACTAAACACTTTTTAAAAA
 GCTTATCTTTGCCCTTCTCCAAACAAGAAGCAATAGTCTCCAAGTCAATATAAATTCTACAGAAAA
 TAGTGTCTTTTTCTCCAGAAAAATGCTTGTGAGAATCATTAAAACATGTGACAAATTTAGAGATT
 CTTTGTTTTATTTCACTGATTAAATATACTGTGGCAAATTACACAGATTATTAAATTTTTTACAA
 GAGTATAGTATATTTATTTGAAATGGGAAAAGTGCATTTTACTGTATTTTGTGTATTTTGTGTTAT
 TTCTCAGAATATGGAAAGAAAATTAAATGTGTCAATAAATATTTTCTAGAGAGTAA

FIGURE 108

MAREDSVKCLRCLLYALNLLFWLMSISVLAVSAWMRDYLNNVLTTLTAETRVEEA VILTYFPVVHP
VMIAVCCFLIIVGMLGYCGTVKRNLLLLAWYFGSLLVIFCVELACGVWTYEQELMVPVQWSDMVT
LKARMTNYGLPRYRWLTHAWNFFQREFKCCGVVYFTDWLEMTMDWPPDSCCVREFPGCSKQAHQ
EDLS DLYQEGCGKKMYSFLRGTKQLQVLRFLGISIGVTQILAMILTITLLWALYYDRREP GTDQM
MSLKNDNSQHLSCPSVELLKPSLSRIFEHTSMANSFNTHFEMEEL

Signal peptide:

amino acids 1-33

Transmembrane domains:

amino acids 12-35, 57-86, 94-114, 226-248

FIGURE 109

CCAAGGCCAGAGCTGTGGACACCTTATCCCACTCATCCTCATCCTCTTCTCTGATAAAGCCCCCTACCAGTGCT
GATAAAGTCTTTCTCGTGAGAGCCTAGAGGCCTTAAAAAAGTGCTTGAAAGAGAAGGGGACAAAGGAACA
CCAGTATTAAGAGGATTTTCCAGTGTCTTGGCAGTTGGTCCAGAAGGATGCTCCTCCATTCTGCTTCTCACCTG
CCTCTTCATCACAGGCACCTCCGTGTCAACCCGTGGCCCTAGATCCTTGTCTGCTTACATCAGCCTGAATGAGC
CCTGGAGGAACACTGACCACCAGTTGGATGAGTCTCAAGGTCCTCCTCTATGTGACAACCATGTGAATGGGGAG
TGGTACCACCTTACGGGCATGGCGGGAGATGCCATGCCTACCTTCTGCATACCAGAAAACCACTGTGGAACCCA
CGCACCTGTCTGGCTCAATGGCAGCCACCCCTAGAAGGCGACGGCATTGTGCAACGCCAGGCTTGTGCCAGCT
TCAATGGGAAGTGTCTGTCTGTGAACACCACGGTGGAAGTCAAGGCTTGCCCTGGAGGCTACTATGTGTATCGT
CTGACCAAGCCAGCGTCTGTCTTCCACGTCTACTGTGGTCATTTTTATGACATCTGCGACGAGGACTGCCATGG
CAGCTGCTCAGATACCAGCGAGTGCACATGCGCTCCAGGAAGTGTGCTAGGCCCTGACAGGCAGACATGCTTTG
ATGAAAATGAATGTGAGCAAAACAACGGTGGCTGCAGTGAGATCTGTGTGAACCTCAAAAACCTCTACCGCTGT
GAGTGTGGGGTTGGCCGTGTGCTAAGAAGTGATGGCAAGACTTGTGAAGACGTTGAAGGATGCCACAATAACAA
TGGTGGCTGCAGCCACTCTTGCCCTGGATCTGAGAAAGGCTACCAGTGTGAATGTCCCCGGGGCCTGGTGCTGT
CTGAGGATAACCACACTTGCCAAGTCCCTGTGTTGTGCAAAATCAAATGCCATTGAAAGTGAACATCCCCAGGGAG
CTGGTTGGTGGCTGGAGCTCTTCTGACCAACACCTCCTGCCGAGGAGTGTCCAACGGACCCATGTCAACAT
CCTCTTCTCTCTCAAGACATGTGGTACAGTGGTGCATGTGGTGAATGACAAGATTGTGGCCAGCAACCTCGTGA
CAGGTCTACCCAAGCAGACCCCGGGGAGCAGCGGGGACTTCATCATCCGAACCAGCAAGCTGCTGATCCCGGTG
ACCTGCGAGTTTCCACGCCTGTACACCATTCTGAAGGATACGTTCCCAACCTTCGAAACTCCCCACTGGAAT
CATGAGCCGAAATCATGGGATCTTCCCATCTCACTCTGGAGATCTTCAAGGACAATGAGTTTGAAGAGCCTTACC
GGGAAGCTCTGCCACCCCTCAAGCTTCGTGACTCCCTCTACTTTGGCATTGAGCCCCTGGTGCACGTGAGCGGC
TTGGAAAGCTTGGTGGAGAGCTGCTTTGCCACCCCACTCCAAGATCGACGAGGTCTGAAATACTACCTCAT
CCGGGATGGCTGTGTTTCAAGTGTGTTGGGCAAGACCACAAGGAAGTGTCTGCACTGCCGGGTTCTTGTCTGTGGAGTG
TTGGACGAGCGTTCCCGCTGTGCCAGGGTTGCCACCGGCAATGCGTCGTGGGGCAGGAGGAGAGGACTCAGC
CGGTCTACAGGGCCAGACGCTAACAGGCGGCCGATCCGCATCGACTGGGAGGACTAGTTTCGTAGCCATACCTC
GAGTCCCTGCATTGGACGGCTCTGCTCTTTGGAGCTTCTCCCCCACCGCCCTCTAAGAACATCTGCCAACAGC
TGGGTTTACAGACTTACACTGTGAGTTCAGACTCCAGCACCAACTCACTCTGATTCTGGTCCATTCACTGGGCA
CAGGTCACAGCACTGCTGAACAATGTGGCCTGGGTGGGGTTTCATCTTCTAGGGTTGAAAATAAACTGTCCA
CCCAGAAAGACTCACCCATTTCCTCATTTCTTCTTCTACTTAAATACCTCGTGTATGGTGCAATCAGAC
CACAAAATCAGAAGCTGGGTATAATATTTCAAGTTACAAACCCTAGAAAAATTAAACAGTTACTGAAATTATGA
CTTAAATACCAATGACTCCTTAAATATGTAAATTATAGTTATACCTTGAAATTTCAATTCAAATGCAGACTAA
TTATAGGGAATTTGGAAGTGATCAATAAACAGTATATAATTTT

FIGURE 110

MPPFLLLTCLFITGTSVSPVALDPCSAYISLNEPWRNTDHLDESQGPPLCDNHVNGEWYHFTGMAGDAMP
TFCIPENHCGTHAPVWLNNGSHPLEGDGIVQRQACASFNGNCCLWNTTVEVKACPGGYVYRLTKPSVCFHV
YCGHFDICDEDCHGSCSDTSECTCAPGTVLGPDRQTCFDENECEQNNGGCSEICVNLKNSYRCECGVGRV
LRSDGKTCEDVEGCHNNNGGCSHSLGSEKGYQCECPRGLVLSNHTCQVPVLCKSNAIEVNIIPRELVGG
LELFLTNTSCRGVSNNGTHVNILFSLKTCGTVVDVNDKIVASNLVTGLPKQTPGSSGDFIIRTSKLLIPVT
CEFPRLYTISEGYVPNLRNSPLEIMSRNHGIFPFTLEIFKDNEFEOPYREALPTLKLKRLSLYFGIEPVVHV
SGLESVLVESCFATPTSKIDEVLKYYLIRDGCVSDDSVKQYTSRDHLAKHFQVPVFKFVGKDHKEVFLHCRV
LVCGVLDERSRCAQGCCHRRMRGAGGEDSAGLQGQTLTGGPIRIDWED

Important features of the protein:

Signal peptide:

amino acids 1-16

N-glycosylation sites.

amino acids 89-93, 116-120, 259-263, 291-295, 299-303

Tyrosine kinase phosphorylation sites.

amino acids 411-418, 443-451

N-myristoylation sites.

amino acids 226-232, 233-239, 240-246, 252-258, 296-302, 300-306,
522-528, 531-537

Aspartic acid and asparagine hydroxylation site.

amino acids 197-209

ZP domain proteins.

amino acids 431-457

Calcium-binding EGF-like proteins.

amino acids 191-212, 232-253

FIGURE 111

GAGAGAGGCAGCAGCTTGCTCAGCGGACAAGGATGCTGGGCGTGAGGGACCAAGGCCTGCCCTGCACTCGG
 GCCTCCTCCAGCCAGTGCTGACCAGGGACTTCTGACCTGCTGGCCAGCCAGGACCTGTGTGGGGAGGCCCT
 CCTGCTGCCTTGGGGTGACAATCTCAGCTCCAGGCTACAGGGAGACCGGGAGGATCACAGAGCCAGCATGT
 TACAGGATCCTGACAGTGATCAACCTCTGAACAGCCTCGATGTCAAACCCCTGCGCAAACCCCGTATCCCC
 ATGGAGACCTTCAGAAAGGTGGGGATCCCCATCATCATAGCACTACTGAGCCTGGCGAGTATCATCATTGT
 GGTGTCTCTCATCAAGGTGATTCTGGATAAATACTACTTCTCTGCGGGCAGCCTCTCCACTTCATCCCGA
 GGAAGCAGCTGTGTGACGGAGAGCTGGACTGTCCCTTGGGGGAGGACGAGGAGCACTGTGTCAAGAGCTTC
 CCCGAAGGGCCTGCAGTGGCAGTCCGCCCTCTCCAAGGACCGATCCACACTGCAGGTGCTGGACTCGGCCAC
 AGGGAAGTGGTTCTCTGCCTGTTTCGACAACCTTCACAGAAGCTCTCGCTGAGACAGCCTGTAGGCAGATGG
 GCTACAGCAGAGCTGTGGAGATTGGCCCAGACCAGGATCTGGATGTTGTTGAAATCACAGAAAACAGCCAG
 GAGCTTCGCATGCGGAACTCAAGTGGGCCCTGTCTCTCAGGCTCCCTGGTCTCCCTGCACTGTCTTGCCTG
 TGGGAAGAGCCTGAAGACCCCCCGTGTGGTGGGTGGGAGGAGGCCTCTGTGGATTCTTGGCCTTGGCAGG
 TCAGCATCCAGTACGACAAACAGCACGTCTGTGGAGGGAGCATCCTGGACCCCCACTGGGTCTCTACGGCA
 GCCACTGCTTCAGGAAACATACCGATGTGTTCAACTGGAAGGTGCGGGCAGGCTCAGACAACTGGGCAG
 CTTCCCATCCCTGGCTGTGGCCAAGATCATCATCATTGAATTCAACCCCATGTACCCCAAAGACAATGACA
 TCGCCCTCATGAAGCTGCAGTTCCCACTCACTTTCTCAGGCACAGTCAGGCCCATCTGTCTGCCCTTCTTT
 GATGAGGAGCTCACTCCAGCCACCCCACTCTGGATCATTGGATGGGGCTTTACGAAGCAGAATGGAGGGAA
 GATGTCTGACATACTGCTGCAGGCGTCAGTCCAGGTCATTGACAGCACACGGTGCAATGCAGACGATGCGT
 ACCAGGGGGAAGTCACCGAGAAGATGATGTGTGCAGGCATCCCGGAAGGGGGTGTGGACACCTGCCAGGGT
 GACAGTGGTGGGCCCCTGATGTACCAATCTGACCAGTGGCATGTGGTGGGCATCGTTAGCTGGGGCTATGG
 CTGCGGGGGCCCGAGCACCCCAAGGAGTATACACCAAGGTCTCAGCCTATCTCAACTGGATCTACAATGTCT
 GGAAGGCTGAGCTGTAATGCTGCTGCCCCCTTTCAGTGCTGGGAGCCGCTTCCTTCCTGCCCTGCCACCT
 GGGGATCCCCCAAAGTCAGACACAGAGCAAGAGTCCCCTTGGGTACACCCCTCTGCCCACAGCCTCAGCAT
 TTCTTGGAGCAGCAAAGGGCCTCAATTCCTGTAAGAGACCCTCGCAGCCCAGAGGCGCCAGAGGAAGTCA
 GCAGCCCTAGCTCGGCCCACTTGGTGTCTCCAGCATCCCAGGGAGAGACACAGCCCACTGAACAAGGTCT
 CAGGGGTATTGCTAAGCCAAGAAGGAACTTTCCACACTACTGAATGGAAGCAGGCTGTCTTGTAAGGCC
 CAGATCACTGTGGGCTGGAGAGGAGAAGGAAAGGGTCTGCCCCAGCCCTGTCCGTCTTACCCATCCCCAA
 GCCTACTAGAGCAAGAAACAGTTGTAATATAAAATGCACTGCCCTACTGTTGGTATGACTACCGTTACCT
 ACTGTTGTCATTGTTATTACAGCTATGGCCACTATTATTAAAGAGCTGTGTAACATCTCTGGCAAAAAAA
 AAAA

FIGURE 112

MLQDPDSQPLNSLDVKPLRKPRIPMETFRKVGIPIIIIALLSLASIIIVVVLIKVILDKYYFLCG
QPLHFIPRKQLCDGELDCPLGEDEEHCVKSFPEGPAVAVRLSKDRSTLQVLDSATGNWFSACFDN
FTEALAEACRQMGYSRAVEIGPDQDLDVVEITENSQELMRNSSGPCLSGSLVSLHCLACGKSL
KTPRVVGEEASVDSWPWQVSIQYDKQHVCSSILDPHWVLTAAHCFRKHTDVFNWKVRAGSDKL
GSFPSLAVAKIIIIIEFNPMYPKDNDIALMKLQFPLTFSGTVRPICLPFFDEELTPATPLWIIGWG
FTKQNGGKMSDILLQASVQVIDSTRCNADDAAYQGEVTEKMMCAGIPEGGVDTCCQGDSSGGLMYQS
DQWHVVGIVSWGYGCGGPSTPGVYTKVSAYLNWIYNVWKAEL

Transmembrane domain:

amino acids 32-53 (typeII)

FIGURE 113

GGCTGGACTGGAACCTCCTGGTCCCAAGTGATCCACCCGCCTCAGCCTCCCAAGGTGCTGTGATTA
TAGGTGTAAGCCACCGTGTCTGGCCTCTGAACAACCTTTTTCAGCAACTAAAAAGCCACAGGAGT
TGAAGTGTAGGATTCTGACTATGCTGTGGTGGCTAGTGCTCCTACTCCTACCTACATTAAATC
TGTTTTTTGTTCTCTTGTAAGTACCTTTACCTTCCTAACACAGAGGATCTGTCACTGTGGCTCT
GGCCCCAACCTGACCTTCACTCTGGAACGAGAACAGAGGTTTCTACCCACACCGTCCCCTCGAAG
CCGGGGACAGCCTCACCTTGCTGGCCTCTCGCTGGAGCAGTGCCCTCACCAACTGTCTCACGTCT
GGAGGCACTGACTCGGGCAGTGCAGGTAGCTGAGCCTCTTGGTAGCTGCGGCTTTCAAGGTGGGC
CTTGCCCTGGCCGTAGAAGGGATTGACAAGCCCGAAGATTTCATAGGCGATGGCTCCCCTGCCC
AGGCATCAGCCTTGCTGTAGTCAATCACTGCCCTGGGGCCAGGACGGGCGTGGACACCTGCTCA
GAAGCAGTGGGTGAGACATCACGCTGCCCCGCCATCTAACCTTTTCATGTCCTGCACATCACCTG
ATCCATGGGCTAATCTGAACTCTGTCCCAAGGAACCCAGAGCTTGAGTGAGCTGTGGCTCAGACC
CAGAAGGGGTCTGCTTAGACCACCTGGTTTATGTGACAGGACTTGCAATTCTCCTGGAACATGAGG
GAACGCCCGAGGAAAGCAAAGTGGCAGGGAAGGAACCTTGCGCAAATTATGGGTGAGAAAAGATG
GAGGTGTTGGGTTATCACAAGGCATCGAGTCTCCTGCATTCACTGGACATGTGGGGGAAGGGCTG
CCGATGGCGCATGACACACTCGGGACTCACCTCTGGGGCCATCAGACAGCCGTTTCCGCCCCGAT
CCACGTACCAGCTGCTGAAGGGCAACTGCAGGCCGATGCTCTCATCAGCCAGGCAGCAGCCAAAA
TCTGCGATCACCAGCCAGGGGCAGCCGTCTGGGAAGGAGCAAGCAAAGTGACCATTTCTCCTCCC
CTCCTTCCCTCTGAGAGGCCCTCCTATGTCCCTACTAAAGCCACCAGCAAGACATAGCTGACAGG
GGCTAATGGCTCAGTGTTGGCCCAGGAGGTGAGCAAGGCCTGAGAGCTGATCAGAAGGGCCTGCT
GTGCGAACACGGAAATGCCTCCAGTAAGCACAGGCTGCAAAATCCCCAGGCAAAGGACTGTGTGG
CTCAATTTAAATCATGTTCTAGTAATTGGAGCTGTCCCCAAGACCAAAGGAGCTAGAGCTTGGTT
CAAATGATCTCCAAGGGCCCTTATACCCAGGAGACTTTGATTTGAATTTGAAACCCCAAATCCA
AACCTAAGAACCAGGTGCATTAAGAATCAGTTATTGCCGGGTGTGGTGGCCTGTAATGCCAACAT
TTTGGGAGGCCGAGGCGGGTAGATCACCTGAGGTCAGGAGTTCAAGACCAGCCTGGCCAACATGG
TGAAACCCCTGTCTCTACTAAAAATACAAAAAACTAGCCAGGCATGGTGGTGTGTGCCTGTATC
CCAGCTACTCGGGAGGCTGAGACAGGAGAATTACTTGAACCTGGGAGGTGAAGGAGGCTGAGACA
GGAGAATCACTTCAGCCTGAGCAACACAGCGAGACTCTGTCTCAGAAAAATAAAAAAGAATTA
TGGTTATTTGTAA

FIGURE 114

MLWWLVLLLLPTLKSVFCSLVTSLYLPNTEDLSLWLWPKPDLHSGTRTEVSTHTVPSKPGTASPC
WPLAGAVPSPTVSRLEALTRAVQVAEPLGSCGFQGGPCPGRRRD

Signal peptide:

amino acids 1-15

FIGURE 115

CAGCAGTGGTCTCTCAGTCCTCTCAAAGCAAGGAAAGAGTACTGTGTGCTGAGAGACCATGGCAA
AGAATCCTCCAGAGAATTGTGAAGACTGTCACATTCTAAATGCAGAAGCTTTTAAATCCAAGAAA
ATATGTAAATCACTTAAGATTTGTGGACTGGTGTTTGGTATCCTGGCCCTAACTCTAATTGTCCT
GTTTGGGGGAGCAAGCACTTCTGGCCGGAGGTACCCAAAAAGCCTATGACATGGAGCACACTT
TCTACAGCAATGGAGAGAAGAAGAAGATTTACATGGAAATTGATCCTGTGACCAGAAGTCAAATA
TTCAGAAGCGGAAATGGCACTGATGAAACATTGGAAGTGCACGACTTTAAAAACGGATACACTGG
CATCTACTTCGTGGGTCTTCAAAAATGTTTTATCAAAACTCAGATTAAAGTGATTCTGAATTTT
CTGAACCAGAAGAGGAAATAGATGAGAATGAAGAAATTACCACAACCTTTCTTTGAACAGTCAGTG
ATTTGGGTCCCAGCAGAAAAGCCTATTGAAAACCGAGATTTTCTTAAAAATTCCAAAATCTGGA
GATTTGTGATAACGTGACCATGTATTGGATCAATCCCACTCTAATATCAGTTTCTGAGTTACAAG
ACTTTGAGGAGGAGGGAGAAGATCTTCACTTTCTGCCAACGAAAAAAGGGATTGAACAAAAT
GAACAGTGGGTGGTCCCTCAAGTGAAAGTAGAGAAGACCCGTACGCCAGACAAGCAAGTGAGGA
AGAACTTCCAATAAATGACTATACTGAAAATGGAATAGAATTTGATCCCATGCTGGATGAGAGAG
GTTATTGTTGTATTTACTGCCGTGAGGCAACCGCTATTGCCGCCGCGTCTGTGAACCTTTACTA
GGCTACTACCCATATCCATACTGCTACCAAGGAGGACGAGTCATCTGTCGTGTCATCATGCCTTG
TAACTGGTGGGTGGCCCGCATGCTGGGGAGGGTCTAATAGGAGGTTTGAGCTCAAATGCTTAAAC
TGCTGGCAACATATAATAAATGCATGCTATTCAATGAATTTCTGCCTATGAGGCATCTGGCCCT
GGTAGCCAGCTCTCCAGAATTACTTGTAGGTAATTCCTCTCTTCATGTTCTAATAAACTTCTACA
TTATCACCAAAAAAAAAAAAAAAAAAAAAA

FIGURE 116

MAKNPPENCEDCHILNAEAFKSKKICKSLKICGLVFGILALTILIVLFWGSKHFWPEVPKKAYDME
HTFYSNGEKKKIYMEIDPVTRTEIFRSGNGTDETLEVHDFKNGYTGIIYFVGLQKCFIKTQIKVIP
EFSEPEEEIDENEEITTTFFEQSVIWWPAEKPIENRDFLKNKILEICDNVTMYWINPTLISVSE
LQDFEEEGEDLHFPAANEKKGIEQNEQWVVPQVKVEKTRHARQASEEELPINDYTENGIEFDPMLD
ERGYCCIIYCRGNRYCRRVCEPLLGYYPYPYCYQGGRVICRVIMPCNWWVARMLGRV

Important features of the protein:

Signal peptide:

amino acids 1-40

Transmembrane domain:

amino acids 25-47 (type II)

N-glycosylation sites.

amino acids 94-97, 180-183

Glycosaminoglycan attachment sites.

amino acids 92-95, 70-73, 85-88, 133-136, 148-151, 192-195, 239-242

N-myristoylation sites.

amino acids 33-38, 95-100, 116-121, 215-220, 272-277

Microbodies C-terminal targeting signal.

amino acids 315-317

Cytochrome c family heme-binding site signature.

amino acids 9-14

FIGURE 117

GAGCTCCCCCTCAGGAGCGCGTTAGCTTCACACCTTCGGCAGCAGGAGGGCGGCAGCTTCTCGCAGGCGGCA
GGGCGGGCGGCCAGGATCATGTCCACCACCACATGCCAAGTGGTGGCGTTCCCTCCTGTCCATCCTGGGGCT
GGCCGGCTGCATCGCGGCCACCGGGATGGACATGTGGAGCACCCAGGACCTGTACGACAACCCCGTCACCT
CCGTGTTCCAGTACGAAGGGCTCTGGAGGAGCTGCGTGAGGCAGAGTTCAGGCTTCACCGAATGCAGGCCC
TATTTACCATCCTGGGACTTCCAGCCATGCTGCAGGCAGTGCAGGCCCTGATGATCGTAGGCATCGTCCT
GGGTGCCATTGGCCTCCTGGTATCCATCTTTGCCCTGAAATGCATCCGCATTGGCAGCATGGAGGACTCTG
CCAAAGCCAACATGACACTGACCTCCGGGATCATGTTTATTGTCTCAGGTCTTTGTGCAATTGCTGGAGTG
TCTGTGTTTGCCAAACATGCTGGTGAATACTTCTGGATGTCCACAGCTAACATGTACACCGGCATGGGTGG
GATGGTGCAGACTGTTTTCAGACCAGGTACACATTTGGTGCGGCTCTGTTCTGTTGGGCTGGGTCTGTTGGAGGCC
TCACACTAATTGGGGGTGTGATGATGTGCATCGCCTGCCGGGGCTGGCACCAGAAGAAACCAACTACAAA
GCCGTTTCTTATCATGCCTCAGGCCACAGTGTTCCTTACAAGCCTGGAGGCTTCAAGGCCAGCACTGGCTT
TGGGTCCAACACCAAAAAACAAGAAGATATACGATGGAGGTGCCCGCACAGAGGACGAGGTACAATCTTATC
CTTCCAAGCACGACTATGTGTAATGTCTTAAGACCTCTCAGCACGGGCGGAAGAACTCCCGGAGAGCTCA
CCCCAAAAACAAGGAGATCCCATCTAGATTTCTTCTTGCTTTTGACTCACAGCTGGAAGTTAGAAAAGCCT
CGATTTTCATCTTTGGAGAGGCCAAATGGTCTTAGCCTCAGTCTCTGTCTCTAAATATTCCACCATAAAAA
GCTGAGTTATTTATGAATTAGAGGCTATAGCTCACATTTTCAATCCTCTATTCTTTTTTTAAATATAACT
TTCTACTCTGATGAGAGAATGTGGTTTTAATCTCTCTCTCACATTTTGATGATTTAGACAGACTCCCCCTC
TTCCTCCTAGTCAATAAACCCATTGATGATCTATTCCCAGCTTATCCCCAAGAAAACCTTTTGAAAGGAAA
GAGTAGACCCAAAGATGTATTTTCTGCTGTTGAATTTTGTCTCCCCACCCCCAACTTGGCTAGTAATAA
ACACTTACTGAAGAAGAAGCAATAAGAGAAAGATATTTGTAATCTCTCCAGCCCATGATCTCGGTTTTCTT
ACACTGTGATCTTAAAAGTTACCAAACCAAAGTCATTTTCAGTTTGAGGCAACCAAACCTTTCTACTGCTG
TTGACATCTTCTTATTACAGCAACACCATTCTAGGAGTTTCTGAGCTCTCCACTGGAGTCTCTTTCTGT
CGCGGGTCAGAAATGTCCCTAGATGAATGAGAAAATTATTTTTTTTAAATTTAAGTCCTAAATATAGTTAA
AATAAATAATGTTTTAGTAAATGATACACTATCTCTGTGAAATAGCCTCACCCCTACATGTGGATAGAAG
GAAATGAAAAAATAATTGCTTTGACATTGTCTATATGGTACTTTGTAAAGTCATGCTTAAGTACAAATCC
ATGAAAAGCTCACACCTGTAATCCTAGCAC'TTTGGGAGGCTGAGGAGGAAGGATCACTTGAGCCCAGAAGT
TCGAGACTAGCCTGGGCAACATGGAGAAGCCCTGTCTCTACAAAATACAGAGAGAAAAAATCAGCCAGTCA
TGGTGGCATAACCTGTAGTCCCAGCATTCGGGAGGCTGAGGTGGGAGGATCACTTGAGCCCAGGGAGGT
TGGGGCTGCAGTGAGCCATGATCACACCCTGCACTCCAGCCAGGTGACATAGCGAGATCCTGTCTAAAAA
AATAAAAAATAAATAATGGAACACAGCAAGTCCTAGGAAGTAGGTTAAACTAATTCTTTAA

FIGURE 118

MSTTTCQVVAFLLSILGLAGCIAATGMDMWSTQDLYDNPVTSVFQYEGLRSCVRQSSGFTECRP
YFTILGLPAMLQAVRALMIVGIVLGAIGLLVSIFALKCIRIGSMEDSAKANMTLTSGIMFIVSGL
CAIAGVSVFANMLVTNFWMSTANMYTGMGGMVQTVQTRYTFGAALFVGWVAGGLTLIGGVMMCIA
CRGLAPEETNYKAVSYHASGHSVAYKPGGFKASTGFGSNTKNKKIYDGGARTEDEVQSYPSKHDY
V

Signal peptide:

amino acids 1-23

Transmembrane domains:

amino acids 81-100, 121-141, 173-194

FIGURE 119

GGAAAACTGTTCTCTTCTGTGGCACAGAGAACCCTGCTTCAAAGCAGAAGTAGCAGTTCGGGAGTCC
 AGCTGGCTAAAACTCATCCCAGAGGATAATGGCAACCCATGCCTTAGAAATCGCTGGGCTGTTTCTTG
 GTGGTGTGGAATGGTGGGCACAGTGGCTGTCACTGTCATGCCTCAGTGGAGAGTGTGGCCTTCATT
 GAAAACAACATCGTGGTTTTTGAACCTTCTGGGAAGGACTGTGGATGAATTGCGTGAGGCAGGCTAA
 CATCAGGATGCAGTGCAAAATCTATGATTCCCTGCTGGCTCTTTCTCCGGACCTACAGGCAGCCAGAG
 GACTGATGTGTGCTGCTTCCGTGATGTCCTTCTTGGCTTTCATGATGGCCATCCTTGGCATGAAATGC
 ACCAGGTGCACGGGGGACAATGAGAAGGTGAAGGCTCACATTCTGCTGACGGCTGGAATCATCTTCAT
 CATCACGGGCATGGTGGTGTCTATCCCTGTGAGCTGGGTGCCAATGCCATCATCAGAGATTTCTATA
 ACTCAATAGTGAATGTTGCCCAAAACGTGAGCTTGGAGAAGCTCTCTACTTAGGATGGACCACGGCA
 CTGGTGTGATTGTTGGAGGAGCTCTGTTCTGCTGCGTTTTTTGTTGCAACGAAAAGAGCAGTAGCTA
 CAGATACTCGATACCTTCCCATCGCACAAACCCAAAAAGTTATCACACCGGAAAGAAGTCACCGAGCG
 TCTACTCCAGAAGTCAGTATGTGTAGTTGTGTATGTTTTTTAACTTTACTATAAAGCCATGCAAATG
 ACAAAAATCTATATTACTTTCTCAAATGGACCCCAAAGAACTTTGATTTACTGTTCTTAACTGCCT
 AATCTTAATTACAGGAAGTGTGCATCAGCTATTTATGATTCTATAAGCTATTTACGAGAATGAGATA
 TTAAACCAATGCTTTGATTGTTCTAGAAAGTATAGTAATTTGTTTTCTAAGGTGGTTCAAGCATCTA
 CTCTTTTTATCATTTACTTCAAATGACATTGCTAAAGACTGCATTATTTTACTACTGTAATTTCTCC
 ACGACATAGCATTATGTACATAGATGAGTGTAAACATTTATATCTCACATAGAGACATGCTTATATGGT
 TTTATTTAAAATGAAATGCCAGTCCATTACACTGAATAAATAGAACTCAACTATTGCTTTTCAGGGAA
 ATCATGGATAGGGTTGAAGAAGGTTACTATTAATTGTTTAAAAACAGCTTAGGGATTAATGTCCTCCA
 TTTATAATGAAGATTAAAATGAAGGCTTTAATCAGCATTGTAAAGGAAATTGAATGGCTTTCTGATAT
 GCTGTTTTTTAGCCTAGGAGTTAGAAATCCTAACTTCTTTATCCTCTTCTCCCAGAGGCTTTTTTTTT
 CTTGTGTATTAAATTAACATTTTTTAAACGCAGATATTTGTCAAGGGGCTTTCATTCAAACCTGCTT
 TTCCAGGGCTATACTCAGAAGAAAGATAAAAGTGTGATCTAAGAAAAAGTGATGGTTTTAGGAAAGTG
 AAAATATTTTTGTTTTTGTATTTGAAGAAGAATGATGCATTTTGACAAGAAATCATATATGTATGGAT
 ATATTTTAATAAGTATTTGAGTACAGACTTTGAGGTTTCATCAATATAAATAAAAGAGCAGAAAAATA
 TGTCTTGGTTTTTCATTTGCTTACCAAAAAACAACAACAAAAAAGTTGTCCTTTGAGAACTTCACCT
 GCTCCTATGTGGGTACCTGAGTCAAAATTGTCATTTTTGTTCTGTGAAAAATAAATTTCTTCTTGTA
 CCATTTCTGTTTAGTTTTACTAAAATCTGTAAATACTGTATTTTTCTGTTTATCCAAATTTGATGAA
 ACTGACAATCCAATTTGAAAGTTTGTGTGACGTCTGTCTAGCTTAAATGAATGTGTTCTATTTGCTT
 TATACATTTATATTAATAAATTTGTACATTTTTTCTAATT

FIGURE 120

MATHALEIAGLFLGGVGMVGTVAVTVMPPQWRVSAFIENNIVVFENFW EGLWMNCVRQANIRMQCK
IYDSLLALSPDLQAARGLMCAASVMSFLAFMMAILGMKCTRCTGDNEKVKAHILLTAGIIFIITG
MVVLIPVSWVANAIIRDFYNSIVNVAQKRELGEALYLGWTTALVLIVGGALFCCVFCCNEKSSSY
RYSIPSHRTTQKSYHTGKKSPSVYSRSQYV

Signal peptide:

amino acids 1-17

Transmembrane domains:

amino acids 82-101, 118-145, 164-188

FIGURE 121

GGAGAGAGGCGCGCGGGTGAAAGGCGCATTGATGCAGCCTGCGGCGGCCTCGGAGCGCGGCGGAG
CCAGACGCTGACCACGTTCCCTCCTCGGTCTCCTCCGCCTCCAGCTCCGCGCTGCCCCGGCAGCC
GGGAGCCATGCGACCCAGGGCCCCGCGCCTCCCCGCAGCGGCTCCGCGGCCTCCTGCTGCTCC
TGCTGCTGCAGCTGCCCCGCGCCGTCGAGCGCCTCTGAGATCCCCAAGGGGAAGCAAAAGGCGCAG
CTCCGGCAGAGGGAGGTGGTGGACCTGTATAATGGAATGTGCTTACAAGGGCCAGCAGGAGTGCC
TGGTCGAGACGGGAGCCCTGGGGCCAATGTTATTCCGGGTACACCTGGGATCCCAGGTCGGGATG
GATTCAAAGGAGAAAAGGGGGAATGTCTGAGGGAAAGCTTTGAGGAGTCCTGGACACCCAACTAC
AAGCAGTGTTTCATGGAGTTCATTGAATTATGGCATAGATCTTGGGAAAATTGCGGAGTGTACATT
TACAAAGATGCGTTCAAATAGTGCTCTAAGAGTTTTGTTTCAGTGGCTCACTTCGGCTAAAATGCA
GAAATGCATGCTGTGAGCGTTGGTATTTACATTCAATGGAGCTGAATGTTTCAGGACCTCTTCCC
ATTGAAGCTATAATTTATTTGGACCAAGGAAGCCCTGAAATGAATTCAACAATTAATATTCATCG
CACTTCTTCTGTGGAAGGACTTTGTGAAGGAATTGGTGCTGGATTAGTGGATGTTGCTATCTGGG
TTGGCACTTGTTTCAGATTACCCAAAAGGAGATGCTTCTACTGGATGGAATTCAGTTTCTCGCATC
ATTATTGAAGAACTACCAAAATAAATGCTTTAATTTTCATTTGCTACCTCTTTTTTTATTATGCC
TTGGAATGGTTCACTTAAATGACATTTTAAATAAGTTTATGTATACATCTGAATGAAAAGCAAAG
CTAAATATGTTTACAGACCAAAGTGTGATTTACACTGTTTTTAAATCTAGCATTATTCATTTTG
CTTCAATCAAAAGTGGTTTCAATATTTTTTTTTAGTTGGTTAGAATACTTTCTTCATAGTCACATT
CTCTCAACCTATAATTTGGAATATTGTTGTGGTCTTTTGTTTTTTCTCTTAGTATAGCATTTTTTA
AAAAATATAAAAGCTACCAATCTTTGTACAATTTGTAAATGTTAAGAATTTTTTTTATATCTGT
TAAATAAAAATTATTTCCAACA

FIGURE 122

MRPQGPAASPQRLRGLLLLLLLQLPAPSSASEIPKGKQKAQLRQREVVDLYNGMCLQGPAGVPGR
DGSPGANVIPGTPGIPGRDGFKEKGECLRESFEESWTPNYKQCSWSSLNYGIDLGKIAECTFTK
MRSNSALRVLFSGSLRLKCRNACCQRWYFTFNGAECSGPLPIEAIYLDQGSPEMNSTINIHRIS
SVEGLCEGIGAGLVDVAIWVGTCSDYPKGDASTGWNSVSRIIEELPK

Signal peptide:

amino acids 1-30

Transmembrane domain:

amino acids 195-217

FIGURE 123

GCTGAGCGTGTGCGCGGTACGGGGCTCTCCTGCCTTCTGGGCTCCAACGCAGCTCTGTGGCTGAA
CTGGGTGCTCATCACGGGAACCTGCTGGGCTATGGAATACAGATGTGGCAGCTCAGGTAGCCCCAA
ATTGCCTGGAAGAATACATCATGTTTTTCGATAAGAAGAAATTGTAGGATCCAGTTTTTTTTTTA
ACCGCCCCCTCCCCACCCCCCAAAAAAAGCTGTAAAGATGCAAAAACGTAATATCCATGAAGATCC
TATTACCTAGGAAGATTTTGATGTTTTGCTGCCAATGCGGTGTTGGGATTTATTTGTTCTTGGAG
TGTTCTGCGTGGCTGGCAAAGAATAATGTTCCAAAATCGGTCCATCTCCCAAGGGGTCCAATTTT
TCTTCTGCGTGTGAGCGAGCCCTGACTCACTACAGTGCAGCTGACAGGGGCTGTCATGCAACTG
GCCCCTAAGCCAAAGCAAAAGACCTAAGGACGACCTTTGAACAATACAAAGGATGGGTTTCAATG
TAATTAGGCTACTGAGCGGATCAGCTGTAGCACTGGTTATAGCCCCACTGTCTTACTGACAATG
CTTTCTTCTGCCGAACGAGGATGCCCTAAGGGCTGTAGGTGTGAAGGCAAAATGGTATATTGTGA
ATCTCAGAAAATTACAGGAGATACCCTCAAGTATATCTGCTGGTTGCTTAGGTTTGTCCCTTCGCT
ATAACAGCCTTCAAAAACTTAAGTATAATCAATTTAAAGGGCTCAACCAGCTCACCTGGCTATAC
CTTGACCATAACCATATCAGCAATATTGACGAAAATGCTTTTAATGGAATACGCAGACTCAAAGA
GCTGATTCTTAGTTCCAATAGAATCTCCTATTTTCTTAACAATACCTTCAGACCTGTGACAAATT
TACGGAACCTGGATCTGTCTTATAATCAGCTGCATTCTCTGGGATCTGAACAGTTTCGGGGCTTG
CGGAAGCTGCTGAGTTTACATTTACGGTCTAACTCCCTGAGAACCATCCCTGTGCGAATATTCCA
AGACTGCCGCAACCTGGAACTTTTGGACCTGGGATATAACCGGATCCGAAGTTTAGCCAGGAATG
TCTTTGCTGGCATGATCAGACTCAAAGAACTTCACCTGGAGCACAATCAATTTTCCAAGCTCAAC
CTGGCCCTTTTTCCAAGTTTGGTCAGCCTTCAGAACCCTTACTTGCACTGGAATAAAATCAGTGT
CATAGGACAGACCATGTCTGGACCTGGAGCTCCTTACAAAGGCTTGATTTATCAGGCAATGAGA
TCGAAGCTTTCAGTGGACCCAGTGTTCAGTGTGTCCGAATCTGCAGCGCCTCAACCTGGAT
TCCAACAAGCTCACATTTATTGGTCAAGAGATTTTGGATTCTTGGATATCCCTCAATGACATCAG
TCTTGCTGGGAATATATGGGAATGCAGCAGAAATATTTGCTCCCTTGTAAGCTGGCTGAAAAGTT
TTAAAGGTCTAAGGGAGAATACAATTATCTGTGCCAGTCCCAAAGAGCTGCAAGGAGTAAATGTG
ATCGATGCAGTGAAGAACTACAGCATCTGTGGCAAAAGTACTACAGAGAGGTTTGATCTGGCCAG
GGCTCTCCCAAAGCCGACGTTTAAAGCCCAAGCTCCCCAGGCCGAAGCATGAGAGCAAACCCCTT
TGCCCCCGACGGTGGGAGCCACAGAGCCCGGCCAGAGACCGATGCTGACGCCGAGCACATCTCT
TTCCATAAAATCATCGCGGGCAGCGTGGCGCTTTTCTGTCCGTGCTCGTCATCTCTGCTGGTTAT
CTACGTGTCATGGAAGCGGTACCTGCGAGCATGAAGCAGCTGCAGCAGCGCTCCCTCATGCGAA
GGCACAGGAAAAAGAAAAGACAGTCCCTAAAGCAAATGACTCCAGCACCCAGGAATTTTATGTA
GATTATAAACCACCAACACGGAGACCAGCGAGATGCTGCTGAATGGGACGGGACCCCTGCACCTA
TAACAAATCGGGCTCCAGGGAGTGTGAGGTATGAACCATTGTGATAAAAAGAGCTCTTAAAAGCT
GGGAAATAAGTGGTGCTTTATTGAACTCTGGTGACTATCAAGGGAACGCGATGCCCCCCTCCCC
TTCCCTCTCCCTCTCACTTTGGTGGCAAGATCCTTCTTGTCCGTTTTAGTGCATTACATAACT
GGTCATTTTCTCTCATACATAATCAACCCATTGAAATTTAAATACCACAATCAATGTGAAGCTT
GAACTCCGGTTTAAATATAATACCTATTGTATAAGACCCTTTACTGATTCCATTAATGTGCATTT
GTTTTAAGATAAAACTTCTTTTCATAGGTAAAAA

FIGURE 124

MGFNVIRLLSGSAVALVIAPTVLLTMLSSAERGCPKGCRCGKMVYCESQKLQEIPSSISAGCLG
LSLRYNLSQKLKYNQFKGLNQLTWLYLDHNNHISNIDENAFNGIRRLKELILSSNRISYFLNNTFR
PVTNLRNLDLSYNQLHSLGSEQFRGLRKLSSLHLRSNSLRITPVRIQDCRNLELLDLGYNRIRS
LARNVFAGMIRLKEHLEHNQFSKLNALFPRLVSLQNLYLQWNKISVIGQTMSWTWSSLQRLDL
SGNEIEAFSGPSVFQCVPNLQRLNLDNKLTFIQEILDSWISLNDISLAGNIWECSRNICSLVN
WLKSFKGLRENTIICASPKELGVNVIDAVKNYSICGKSTTERFDLARALPKPTFKPKLPRPKHE
SKPPLPPTVGATEPGPETDADAEHISFHKIIAGSVALFLSVLVILLVIYVSWKRYPASMKQLQQR
SLMRRHRKKKRQSLKQMTPTSTQEFYVDYKPTNTETSEMLLNGTGPCITYNKSGSRECEV

Important features of the protein:

Signal peptide:

amino acids 1-33

Transmembrane domain:

amino acids 420-442

N-glycosylation sites.

amino acids 126-129, 357-360, 496-499, 504-507

cAMP- and cGMP-dependent protein kinase phosphorylation site.

amino acids 465-468

Tyrosine kinase phosphorylation site.

amino acids 136-142

N-myristoylation sites.

amino acids 11-16, 33-38, 245-250, 332-337, 497-502, 507-512

FIGURE 125

CCGTTATCGTCTTGCGCTACTGCTGAATGTCCGTCCCGGAGGAGGAGGAGAGGCTTTTGCCGCTG
ACCCAGAGATGGCCCCGAGCGAGCAAATTCCTACTGTCCGGCTGCGCGGCTACCGTGGCCGAGCT
AGCAACCTTTCCCCTGGATCTCACAAAACTCGACTCCAAATGCAAGGAGAAGCAGCTCTTGCTC
GGTTGGGAGACGGTGCAAGAGAATCTGCCCCCTATAGGGGAATGGTGCGCACAGCCCTAGGGATC
ATTGAAGAGGAAGGCTTTCTAAAGCTTTGGCAAGGAGTGACACCCGCCATTTACAGACACGTAGT
GTATTCTGGAGGTCGAATGGTCACATATGAACATCTCCGAGAGGTTGTGTTTGGCAAAAGTGAAG
ATGAGCATTATCCCCCTTTGGAAATCAGTCATTGGAGGGATGATGGCTGGTGTATTGGCCAGTTT
TTAGCCAATCCAACTGACCTAGTGAAGGTTGAGATGCAAATGGAAGGAAAAAGGAACTGGAAGG
AAAACCATTTGCGATTTTCGTGGTGTACATCATGCATTTGCAAAAATCTTAGCTGAAGGAGGAATAC
GAGGGCTTTGGGCAGGCTGGGTACCCAATATACAAAGAGCAGCACTGGTGAATATGGGAGATTTA
ACCACTTATGATACAGTGAAACACTACTTGGTATTGAATACACCACTTGAGGACAATATCATGAC
TCACGGTTTATCAAGTTTATGTTCTGGACTGGTAGCTTCTATTCTGGGAACACCAGCCGATGTCA
TCAAAGCAGAATAATGAATCAACCACGAGATAAACAAGGAAGGGGACTTTTGTATAAATCATCG
ACTGACTGCTTGATTCAAGGCTGTTCAAGGTGAAGGATTCATGAGTCTATATAAAGGCTTTTACC
ATCTTGGCTGAGAATGACCCCTTGGTCAATGGTGTCTGGCTTACTTATGAAAAATCAGAGAGA
TGAGTGGAGTCAGTCCATTTTAA

FIGURE 126

MSVPEEEERLLPLTQRWPRASKFLLSGCAATVAELATFPLDLTKTRLQMGEAALARLGDGARES
APYRGMVRTALGIIEEEGFLKLWQGVTPAIYRHVVYSGGRMVTYEHLREVVFVKSEDEHYPLWKS
VIGGMMAGVIGQFLANPTDLVKVQMMEGKRRKLEGKPLRFRGVHHAFAKILAEGGIRGLWAGWVP
NIQRAALVNMGDLTTYDTVKHYLVLNTPLEDNIMTHGLSSLCGLVASILGTPADVIKSRIMNQP
RDKQGRGLLYKSSTDCLIQAVQGEFMSLYKGFLPSWLRMTPWSMVFWLTYEKIREMSGVSPF

Transmembrane domains:

amino acids 25-38, 130-147, 233-248

FIGURE 127

CGCGGATCGGACCCAAGCAGGTCGGCGGGCGGCGGCAGGAGAGCGGGCCGGGCGTCAGCTCCTCGAC
CCCCGTGTGGGCTAGTCCAGCGAGGCGGACGGGCGGCGTGGGCCCATGGCCAGGCCCGGCATGG
AGCGGTGGCGCGACCGGCTGGCGCTGGTGACGGGGGCGCTCGGGGGGCATCGGCGCGGCCGTGGCC
CGGGCCCTGGTCCAGCAGGGACTGAAGGTGGTGGGCTGCGCCCGCACTGTGGGCAACATCGAGGA
GCTGGCTGCTGAATGTAAGAGTGCAGGCTACCCCGGGACTTTGATCCCCTACAGATGTGACCTAT
CAAATGAAGAGGACATCCTCTCCATGTTCTCAGCTATCCGTTCTCAGCACAGCGGTGTAGACATC
TGCATCAACAATGCTGGCTTGGCCCGGCCTGACACCCTGCTCTCAGGCAGCACCAAGTGGTTGGAA
GGACATGTTCAATGTGAACGTGCTGGCCCTCAGCATCTGCACACGGGAAGCCTACCAGTCCATGA
AGGAGCGGAATGTGGACGATGGGCACATCATTAACATCAATAGCATGTCTGGCCACCGAGTGTTA
CCCCTGTCTGTGACCCACTTCTATAGTGCCACCAAGTATGCCGTCACTGCGCTGACAGAGGGACT
GAGGCAAGAGCTTCGGGAGGCCCAGACCCACATCCGAGCCACGTGCATCTCTCCAGGTGTGGTGG
AGACACAATTGCGCTTCAAACCTCCACGACAAGGACCCTGAGAAGGCAGCTGCCACCTATGAGCAA
ATGAAGTGTCTCAAACCCGAGGATGTGGCCGAGGCTGTTATCTACGTCTCAGCACCCCCGCACA
CATCCAGATTGGAGACATCCAGATGAGGCCCACGGAGCAGGTGACCTAGTGACTGTGGGAGCTCC
TCCTTCCCTCCCCACCCTTCATGGCTTGCCTCCTGCCTCTGGATTTTAGGTGTTGATTTCTGGAT
CACGGGATACCACTTCCTGTCCACACCCCGACCAGGGGCTAGAAAATTTGTTTGAGATTTTATA
TCATCTTGTCAAATTGCTTCAGTTGTAAATGTGAAAAATGGGCTGGGGAAAGGAGGTGGTGTCCC
TAATTGTTTTACTTGTTAACTTGTTCTTGTTGCCCTGGGCACTTGGCCTTTGTCTGCTCTCAGTG
TCTTCCCTTTGACATGGGAAAGGAGTTGTGGCCAAAATCCCCTCTTCTTGACCTCAACGTCTG
TGGCTCAGGGCTGGGGTGGCAGAGGGAGGCCTTACCTTATATCTGTGTTGTTATCCAGGGCTCC
AGACTTCCTCCTCTGCCTGCCCCACTGCACCCTCTCCCCCTTATCTATCTCCTTCTCGGCTCCCC
AGCCCAGTCTTGGCTTCTTGTCCCCTCCTGGGGTCATCCCTCCACTCTGACTCTGACTATGGCAG
CAGAACACCAGGGCCTGGCCCAAGTGGATTTTCATGGTGATCATTAAAAAAGAAAAATCGCAACCAA
AAAAAAAAA

FIGURE 128

MARPGMERWRDRLALVTGASGGIGA AAVARALVQQGLKVVG CARTVGNIEELAAECKSAGYPGTLI
PYRCDLSNEEDILSMFSAIRSQHSGVDICINNAGLARPD TLLSGSTSGWKDMFNVNVLALSICTR
EAYQSMKERNVDDGHIININSMGHRVLP LSVTHFY SATKYAVTALTEGLRQELREAQTHIRATC
ISPGVVETQFAFKLHDKDPEKAAATYEQMKCLKPEDVAEAVIYVLSTPAHIQIGDIQMRPTEQVT

Important features of the protein:

Signal peptide:

amino acids 1-17

N-myristoylation sites.

amino acids 18-24, 21-27, 22-28, 24-30, 40-46, 90-96, 109-115,
199-205

Short-chain alcohol dehydrogenase.

amino acids 30-42, 104-114

FIGURE 129

AACTTCTACATGGGCCTCCTGCTGCTGGTGCTCTTCCTCAGCCTCCTGCCGGTGGCCTACACCAT
CATGTCCCTCCCACCCTCCTTTGACTGCGGGCCGTTCAGGTGCAGAGTCTCAGTTGCCCGGGAGC
ACCTCCCCTCCCGAGGCAGTCTGCTCAGAGGGCCTCGGCCCAGAATTCCAGTTCTGGTTTCATGC
CAGCCTGTAAAAGGCCATGGAACTTTGGGTGAATCACCGATGCCATTTAAGAGGGTTTTCTGCCA
GGATGGAAATGTTAGGTCGTTCTGTGTCTGCGCTGTTCAATTCAGTAGCCACCAGCCACCTGTGG
CCGTTGAGTGCTTGAAATTGAGGAAGTGAAGAAATTAATTTCTCATGTATTTTTCTCATTTATTTA
TTAATTTTTTAAGTATAGTTGTACATATTTGGGGGTACATGTGATATTTGGATACATGTATACAA
TATATAATGATCAAATCAGGGTAACTGGGATATCCATCACATCAAACATTTATTTTTTATTCTTT
TTAGACAGAGTCTCACTCTGTCACCCAGGCTGGAGTGCAGTGGTGCCATCTCAGCTTACTGCAAC
CTCTGCCTGCCAGGTTCAAGCGATTCTCATGCCTCCACCTCCCAAGTAGCTGGGACTACAGGCAT
GCACCACAATGCCCAACTAATTTTTGTATTTTTAGTAGAGACGGGGTTTTGCCATGTTGCCCAGG
CTGGCCTTGAACCTCGGCCTCAAACAATCCACTTGCCTCGGCCTCCCAAAGTGTTATGATTACA
GGCGTGAGCCACCGTGCCTGGCCTAAACATTTATCTTTTCTTTGTGTTGGGAACTTTGAAATTAT
ACAATGAATTATTGTAACTGTCATCTCCCTGCTGTGCTATGGAACACTGGGACTTCTTCCCTCT
ATCTAACTGTATATTTGTACCAGTTAACCAACCGTACTTCATCCCCACTCCTCTCTATCCTTCCC
AACCTCTGATCACCTCATTCTACTCTCTACCTCCATGAGATCCACTTTTTTAGCTCCCACATGTG
AGTAAGAAAATGCAATATTTGTCTTTCTGTGCCTGGCTTATTTCACTTAACATAATGACTTCCTG
TTCCATCCATGTTGCTGCAAATGACAGGATTTGTTCTTAATTTCAATTAAAATAACCACACATG
GCAAAAA

FIGURE 130

MGLLLLVLFLSLLPVAYTIMSLPPSFDCGPFRVSVAREHLPSRGSLLRGPRPRIPVLVSCQPV
KGGHTLGESPMFPKRVFCQDGNVRSFCVCAVHFSSHQPPVAVECLK

Important features of the protein:

Signal peptide:

amino acids 1-18

N-myristoylation site.

amino acids 86-92

Zinc carboxypeptidases, zinc-binding region 2 signature.

amino acids 68-79

FIGURE 131

TTCTGAAGTAACGGAAGCTACCTTGTATAAAGACCTCAACACTGCTGACCATGATCAGCGCAGCCTGGAGC
 ATCTTCCTCATCGGGACTAAAATTGGGCTGTTCCCTTCAAGTAGCACCTCTATCAGTTATGGCTAAATCCTG
 TCCATCTGTGTGTCGCTGCGATGCGGGTTTCATTTACTGTAATGATCGCTTTCTGACATCCATTCCAACAG
 GAATACCAGAGGATGCTACAACCTCTTACCTTCAGAACAACCAAATAAATAATGCTGGGATTCTTCAGAT
 TTGAAAACTTGCTGAAAGTAGAAGAATATACCTATACCACAACAGTTTAGATGAATTTCTACCAACCT
 CCCAAAGTATGTAAAGAGTTACATTTGCAAGAAAATAACATAAGGACTATCACTTATGATTCACTTTCAA
 AAATTTCCCTATCTGGAAGAATTACATTTAGATGACAACCTCTGTCTCTGCAGTTAGCATAGAAGAGGGAGCA
 TTCCGAGACAGCAACTATCTCCGACTGCTTTTCCCTGTCCCGTAATCACCTTAGCACAATTCCCTGGGGTTT
 GCCCAGGACTATAGAAGAACTACGCTTGGATGATAATCGCATATCCACTATTTTCATCACCATCTCTTCAAG
 GTCTCACTAGTCTAAAACGCCTGGTTCTAGATGGAACCTGTTGAACAATCATGGTTTAGGTGACAAAGTT
 TTCTTCAACCTAGTTAATTTGACAGAGCTGTCCCTGGTGCAGAAATTCCTGACTGCTGCACCAGTAAACCT
 TCCAGGCACAAACCTGAGGAAGCTTTATCTTCAAGATAACCACATCAATCGGGTGCCCCAAATGCTTTTT
 CTTATCTAAGGCAGCTCTATCGACTGGATATGTCCAATAATAACCTAAGTAATTTACCTCAGGGTATCTTT
 GATGATTTGGACAATATAACACAACCTGATTCTTCGCAACAATCCCTGGTATTGCGGGTGCAAGATGAAATG
 GGTACGTGACTGGTTACAATCACTACCTGTGAAGGTCAACGTGCGTGGGCTCATGTGCCAAGCCCCAGAAA
 AGGTTCGTGGGATGGCTATTAAGGATCTCAATGCAGAACTGTTTGATTGTAAGGACAGTGGGATTGTAAGC
 ACCATTAGATAACCACTGCAATACCCAACACAGTGTATCCTGCCCAAGGACAGTGGCCAGCTCCAGTGAC
 CAAACAGCCAGATATTAAGAACCCCAAGCTCACTAAGGATCAACAAACCACAGGGAGTCCCTCAAGAAAAA
 CAATTACAATTACTGTGAAGTCTGTACCTCTGATACCATTATATCTCTTGGAACTTGCTCTACCTATG
 ACTGCTTTGAGACTCAGCTGGCTTAAACTGGGCCATAGCCCGGCATTTGGATCTATAACAGAAACAATTGT
 AACAGGGGAACGCAGTGAGTACTTGGTCACAGCCCTGGAGCCTGATTCACCCTATAAAGTATGCATGGTTC
 CCATGGAAACCAGCAACCTCTACCTATTTGATGAAACTCCTGTTTGTATTGAGACTGAACTGCACCCCTT
 CGAATGTACAACCCTACAACCACCTCAATCGAGAGCAAGAGAAAGAACCTTACAAAAACCCCAATTTACC
 TTTGGCTGCCATCATTGGTGGGGCTGTGGCCCTGTTTACCATTGCCCTTCTTGCTTTAGTGTGTGGTATG
 TTCATAGGAATGGATCGCTCTTCTCAAGGAAGTGTGCATATAGCAAAGGGAGGAGAAGAAAGGATGACTAT
 GCAGAAGCTGGCACTAAGAAGGACAACCTCTATCCTGGAAATCAGGGAACTTCTTTTCAGATGTTACCAAT
 AAGCAATGAACCCATCTCGAAGGAGGAGTTTGTAAATACACACCATATTTCTCCTAATGGAATGAATCTGT
 ACAAAAACAATCACAGTGAAAGCAGTAGTAACCGAAGCTACAGAGACAGTGGTATTCCAGACTCAGATCAC
 TCACACTCATGATGCTGAAGGACTCACAGCAGACTTGTTGTTTTGGGTTTTTTAAACCTAAGGGAGGTGATG
 GT

FIGURE 132

MISAAWSIFLIGTKIGLFLQVAPLSVMAKSCPSVCRC DAGFIYCND RFLTSIPTGIPEDATTLYL
QNNQINNAGIPSDLKNLLKVERIYLYHNSLDEFPTNLPKYVKELHLQENNIRTITYDSLSKIPYL
EELHDDNSVSAVSIEEGAFRDSNYLRLLFLSRNHLSTIPWGLPRTIEELRLDDNRISTISSPSL
QGLTSLKRLVLDGNLLNNHGLGDKVFFNLVNLTELSLVRNSLTAAPVNLPGTNLRKLYLQDNHIN
RVPPNAFSYLRQLYRLDMSNNNLSNLPQGIFDDLDNITQLILRNNPWYCGCKMKWVRDWLQSLPV
KVNVRGLMCQAPEKVRGMAIKDLNAELFDCKDSGIVSTIQITTAIPNTVYPAQGQWPAPVTKQPD
IKNPKLTKDQQTGTGSPSRKTITITVKSVTSDTIHISWKLALPMTALRLSWLKLGHSPAFGSITET
IVTGERSEYLVTALEPDSPYKVCMPMETSNNLYLFDETPVC IETETAPLRMYNP TTTTLNREQEKE
PYKNPNLPLAAIIGGAVALVTIALLALVCWYVHRNGSLFSRNCAYSKGRRRKDDYAEAGTKKDNS
ILEIRETSFQMLPISNEPISKEEFVIHTIFPPNGMNLKNNHSESSSNRSYRDSGIPDSDHSHS

Important features of the protein:

Signal peptide:

amino acids 1-28

Transmembrane domain:

amino acids 531-552

N-glycosylation sites.

amino acids 226-229, 282-285, 296-299, 555-558, 626-629, 633-636

Tyrosine kinase phosphorylation site.

amino acids 515-522

N-myristoylation sites.

amino acids 12-17, 172-177, 208-213, 359-364, 534-539, 556-561,
640-645

Amidation site.

amino acids 567-570

Leucine zipper pattern.

amino acids 159-180

Phospholipase A2 aspartic acid active site.

amino acids 34-44

FIGURE 133

CCGTCATCCCCCTGCAGCCACCCTTCCCAGAGTCCTTTGCCCAGGCCACCCCAGGCTTCTTGGCA
 GCCCTGCCGGGGCCACTTGTCTTCAATGTCTGCCAGGGGGAGGTGGGAAGGAGGTGGGAGGAGGGCG
 TGCAGAGGCAGTCTGGGCTTGGCCAGAGCTCAGGGTGCTGAGCGTGTGACCAGCAGTGAGCAGAG
 GCCGGCCATGGCCAGCCTGGGGCTGCTGCTCCTGCTCTTACTGACAGCACTGCCACCGCTGTGGT
 CCTCCTCACTGCCTGGGCTGGACACTGCTGAAAGTAAAGCCACCATTGCAGACCTGATCCTGTCT
 GCGCTGGAGAGAGCCACCGTCTTCTAGAACAGAGGCTGCCTGAAATCAACCTGGATGGCATGGT
 GGGGGTCCGAGTGCTGGAAGAGCAGCTAAAAAGTGTCCGGGAGAAGTGGGCCAGGAGCCCCCTGC
 TGCAGCCGCTGAGCCTGCGCGTGGGGATGCTGGGGGAGAAGCTGGAGGCTGCCATCCAGAGATCC
 CTCCACTACCTCAAGCTGAGTGATCCCAAGTACCTAAGAGAGTTCCAGCTGACCCTCCAGCCCCG
 GTTTTGGAAAGCTCCACATGCCTGGATCCACACTGATGCCTCCTTGGTGTACCCACGTTCCGGG
 CCCAGGACTCATTCTCAGAGGAGAGAAGTGACGTGTGCCTGGTGCAGCTGCTGGGAACCGGGACG
 GACAGCAGCGAGCCCTGCGGCCTCTCAGACCTCTGCAGGAGCCTCATGACCAAGCCCCGGCTGCTC
 AGGCTACTGCCTGTCCCACCAACTGCTCTTCTTCTCTGGGCCAGAATGAGGGGATGCACACAGG
 GACCACTCCAACAGAGCCAGGACTATATCAACCTCTTCTGCGCCAACATGATGGACTTGAACCGC
 AGAGCTGAGGCCATCGGATACGCCTACCTTACCCGGGACATCTTCATGGAAAACATCATGTTCTG
 TGGAAATGGGCGGCTTCTCCGACTTCTACAAGCTCCGGTGGCTGGAGGCCATTCTCAGCTGGCAGA
 AACAGCAGGAAGGATGCTTCGGGGAGCCTGATGCTGAAGATGAAGAATTATCTAAAGCTATTCAA
 TATCAGCAGCATTTTTTCGAGGAGAGTGAAGAGGCGAGAAAAACAATTTCCAGATTCTCGCTCTGT
 TGCTCAGGCTGGAGTACAGTGGCGCAATCTCGGCTCACTGCAACCTTTGCCTCCTGGGTTCAAGC
 AATTCTCTTGCCTCATCCTCCCGAGTAGCTGGGACTACAGGAGCGTGCCACCATACTGGCTAAT
 TTTTATATTTTTTTAGTAGAGACAGGGTTTCATCATGTTGCTCATGCTGGTCTCGAACTCCTGAT
 CTCAAGAGATCCGCCCACCTCAGGCTCCCAAAGTGTGGGATTATTAGGTGTGAGCCACCGTGTCTG
 GCTGAAAAGCACTTTCAAAGAGACTGTGTTGAATAAAGGGCCAAGGTTCTTGCCACCCAGCACTC
 ATGGGGGCTCTCTCCCTAGATGGCTGCTCCTCCCACAACACAGCCACAGCAGTGGCAGCCCTGG
 GTGGCTTCCTATACATCCTGGCAGAATACCCCCAGCAAACAGAGAGCCACACCCATCCACACCG
 CCACCACCAAGCAGCCGCTGAGACGGACGGTTCCATGCCAGCTGCCTGGAGGAGGAACAGACCCC
 TTTAGTCCTCATCCCTTAGATCCTGGAGGGCACGGATCACATCCTGGGAAGAAGGCATCTGGAGG
 ATAAGCAAAGCCACCCCGACACCCAATCTTGGAAGCCCTGAGTAGGCAGGGCCAGGGTAGGTGGG
 GGCCGGGAGGGACCCAGGTGTGAACGGATGAATAAAGTTCAACTGCAACTGAAAAAAAAA

FIGURE 134

MSARGRWEGGGRRACRGS LGLARAQGAERTSSEQRPAMASLG LLLLLLLLTALPPLWSSSLPGLD
TAESKATIADLILSALERATVFLEQRLPEINLDGMVGVRVLEEQLKSVREKWAQEPLLOPLSLRV
GMLGEKLEAAIQRS LHYLKLSDPKYLREFQLTLP GFWKLP HAWIHTDASLVYPTFGPQDSFSEE
RSDVCLVQLLGTGTDSS EPCGLSDLCRSLMTKPGCSGYCLSHQLLFFLWARMRGCTQG PLQQSQD
YINLFCANMMDLNRRAEAIGYAYPTRDIFMENIMFCGMGGFSDFYKLRWLEAILS WQKQEGCFG
EPDAEDEELSKAIQYQQHFSRRVKRREKQFPDSRSVAQAGVQWRNLGSLQPLPPGFKQFSCLILP
SSWDYRSVPPYLANFYIFLVETGFHHVAHAGLELLISRD PPTSGSQSVGL

Important features of the protein:

Signal peptide:

amino acids 1-26

Transmembrane domain:

amino acids 39-56

Tyrosine kinase phosphorylation sites.

amino acids 149-156, 274-282

N-myristoylation sites.

amino acids 10-16, 20-26, 63-69, 208-214

Amidation site.

amino acids 10-14

Glycoprotein hormones beta chain signature 1.

amino acids 230-237

FIGURE 135

GGTCTGAGTGCAGAGCTGCTGTCAATGGCGGCCGCTCTGTGGGGCTTCTTTCCCGTCCTGCTGCTG
CTGCTGCTATCGGGGGATGTCCAGAGCTCGGAGGTGCCCGGGGCTGCTGCTGAGGGATCGGGAGG
GAGTGGGGTCGGCATAGGAGATCGCTTCAAGATTGAGGGGCGTGCA GTTGTTCAGGGGTGAAGC
CTCAGGACTGGATCTCGGCGGCCCGAGTGCTGGTAGACGGAGAAGAGCACGTGCGTTTCCTTAAG
ACAGATGGGAGTTTTGTGGTTCATGATATACCTTCTGGATCCTTATGTAGTGGAAGTTGTATCTCC
AGCTTACAGATTTGATCCCGTTCGAGTGGATATCACTTCGAAAGGAAAAATGAGAGCAAGATATG
TGAATTACATCAAAACATCAGAGGTTGTCAGACTGCCCTATCCTCTCCAAATGAAATCTTCAGGT
CCACCTTCTTACTTTATTAAAAGGGAATCGTGGGGCTGGACAGACTTTCTAATGAACCCAATGGT
TATGATGATGGTTCTTCCCTTTATTGATATTTGTGCTTCTGCCTAAAGTGGTCAACACAAGTGATC
CTGACATGAGACGGGAAATGGAGCAGTCAATGAATATGCTGAATTCCAACCATGAGTTGCCTGAT
GTTTCTGAGTTCATGACAAGACTCTTCTCTTCAAATCATCTGGCAAATCTAGCAGCGGCAGCAG
TAAAACAGGCAAAAGTGGGGCTGGCAAAGGAGGTAGTCAGGCCGTCCAGAGCTGGCATTTCAC
AAACACGGCAACACTGGGTGGCATCCAAGTCTTGAAAACCGTGTGAAGCAACTACTATAAACTT
GAGTCATCCCGACGTTGATCTCTTACAACCTGTGTATGTT
AACTTTTTAGCACATGTTTTGTACTTGGTACACGAGAAAACCCAGCTTTCATCTTTTGTCTGTAT
GAGGTCAATATTGATGTCAC TGAATTAATTACAGTGTCTATAGAAAATGCCATTAATAAATTAT
ATGAACTACTATACATTATGTATATTAATTAAACATCTTAATCCAGAAATCAAAAAAAAAAAAAA
AAAAAAAAAAAAAAAAA

FIGURE 136

MAAALWGFFPVLLLLLLSGDVQSSEVPGAAAEGSGSGVGIGDRFKIEGRAVVPGVKPDWISAA
RVLVDGEEHVGFLKTDGSFVVHDIPSGSYVVEVVS PAYRFDPVRVDITSGGKMRARYVNYIKTSE
VVRLPYPLQMKSSGPPSYFIKRESWGWTDFLMNPMVMMVLPLLIFVLLPKVVNTSDPDMRREME
QSMNMLNSNHELDPDVSEFMTRLFSSKSSGKSSSGSSKTGKSGAGKRR

Important features of the protein:

Signal sequence:

amino acids 1-23

Transmembrane domain:

amino acids 161-182

N-glycosylation site.

amino acids 184-187

Glycosaminoglycan attachment sites.

amino acids 37-40, 236-239

cAMP- and cGMP-dependent protein kinase phosphorylation site.

amino acids 151-154

N-myristoylation sites.

amino acids 33-38, 36-41, 38-44, 229-234

Amidation site.

amino acids 238-241

ATP/GTP-binding site motif A (P-loop).

amino acids 229-236

FIGURE 137

GATGGCGCAGCCACAGCTTCTGTGAGATTTCGATTTCTCCCCAGTTCCCCTGTGGGTCTGAGGGGA
CCAGAAGGGTGAGCTACGTTGGCTTTCTGGAAGGGGAGGCATATGCGTCAATCCCCAAAACAA
GTTTTGACATTTCCCCTGAAATGTCATTCTCTATCTATTCACTGCAAGTGCCTGCTGTTCCAGGC
CTTACCTGCTGGGCACTAACGGCGGAGCCAGGATGGGGACAGAATAAAGGAGCCACGACCTGTGC
CACCAACTCGCACTCAGACTCTGAACTCAGACCTGAAATCTTCTCTTCACGGGAGGCTTGGCAGT
TTTTCTTACTCCTGTGGTCTCCAGATTTCAGGCCTAAGATGAAAGCCTCTAGTCTTGCCTTCAGC
CTTCTCTCTGCTGCGTTTTATCTCCTATGGACTCCTTCCACTGGACTGAAGACACTCAATTTGGG
AAGCTGTGTGATCGCCACAAACCTTCAGGAAATACGAAATGGATTTTCTGAGATACGGGGCAGTG
TGCAAGCCAAAGATGGAACATTGACATCAGAATCTTAAGGAGGACTGAGTCTTTGCAAGACACA
AAGCCTGCGAATCGATGCTGCCTCCTGCGCCATTTGCTAAGACTCTATCTGGACAGGGTATTTAA
AAACTACCAGACCCCTGACCATTATACTCTCCGGAAGATCAGCAGCCTCGCCAATTCCTTTCTTA
CCATCAAGAAGGACCTCCGGCTCTCTCATGCCCACATGACATGCCATTGTGGGGAGGAAGCAATG
AAGAAATACAGCCAGATTCTGAGTCACTTTGAAAAGCTGGAACCTCAGGCAGCAGTTGTGAAGGC
TTTGGGGGAACTAGACATTCTTCTGCAATGGATGGAGGAGACAGAAATAGGAGGAAAGTGATGCTG
CTGCTAAGAATATTCGAGGTCAAGAGCTCCAGTCTTCAATACCTGCAGAGGAGGCATGACCCCAA
ACCACCATCTCTTTACTGTACTAGTCTTGTGCTGGTCACAGTGTATCTTATTTATGCATTACTTG
CTTCCTTGCATGATTGTCTTTATGCATCCCCAATCTTAATTGAGACCATACTTGTATAAGATTTT
TGTAATATCTTTCTGCTATTGGATATATTTATTAGTTAATATATTTATTTATTTTTTGCTATTTA
ATGTATTTATTTTTTTACTTGGACATGAACTTTAAAAAAATTCACAGATTATATTTATAACCTG
ACTAGAGCAGGTGATGTATTTTTTATACAGTAAAAAAAACCTTGTAATTC TAGAAGAGTGG
CTAGGGGGGTATTTCATTTGTATTCACTAAGGACATATTTACTCATGCTGATGCTCTGTGAGAT
ATTTGAAATTGAACCAATGACTACTTAGGATGGGTGTGGAATAAGTTTTGATGTGGAATTGCAC
ATCTACCTTACAATTACTGACCATCCCCAGTAGACTCCCCAGTCCCATAATTGTGTATCTTCCAG
CCAGGAATCCTACACGGCCAGCATGTATTTCTACAAATAAAGTTTTCTTTGCATACCAAAAAAAA
AAAAA

FIGURE 138

MRQFPKTSFDISPEMSFSIYSLQVPAVPG LTCWALTAEPGWGQNGATT CATNSHSDSELRPEIF
SSREAWQFFLLWSPDFRPKM KASSLA FSLLSAAFYLLWTPSTGLKTLNLGSCVIATNLQEIRNG
FSEIRGSVQAKDGNIDIRILRRTESLQDTKPANRCCLLRHLLRLYLDRVFKNYQTPDHYTLRKIS
SLANSFLT IKKDLRLSHAHMTCHCGEEAMKKYSQILSHFEKLEPQA AVVKALGELDILLQWMEET
E

Important features of the protein:

Signal peptide:

amino acids 1-42

cAMP- and cGMP-dependent protein kinase phosphorylation sites.

amino acids 192-195, 225-228

N-myristoylation sites.

amino acids 42-47, 46-51, 136-141

FIGURE 139

CCTGGAGCCGGAAGCGCGGCTGCAGCAGGGCGAGGCTCCAGGTGGGGTCGGTTCCGCATCCAGCC
TAGCGTGTCCACGATGCGGCTGGGCTCCGGGACTTTTCGCTACCTGTTGCGTAGCGATCGAGGTGC
TAGGGATCGCGGTCTTCTTCGGGGATTCTTCCCGGCTCCCGTTCGTTCTCTGCCAGAGCGGAA
CACGGAGCGGAGCCCCAGCGCCCCGAACCTCGGCTGGAGCCAGTTCTAACTGGACCACGCTGCC
ACCACCTCTCTTCAGTAAAGTTGTTATTGTTCTGATAGATGCCTTGAGAGATGATTTTGTGTTTG
GGTCAAAGGGTGTGAAATTTATGCCCTACACAACCTTACCTTGTGGAAAAAGGAGCATCTCACAGT
TTTGTGGCTGAAGCAAAGCCACCTACAGTTACTATGCCTCGAATCAAGGCATTGATGACGGGGAG
CCTTCTGGCTTTGTGCGACGTATCAGGAACCTCAATTCTCCTGCACTGCTGGAAGACAGTGTGA
TAAGACAAGCAAAAGCAGCTGGAAAAAGAATAGTCTTTTATGGAGATGAAACCTGGGTAAATTA
TTCCCAAAGCATTTTGTGGAATATGATGGAACAACCTCATTTTTCGTGTCAGATTACACAGAGGT
GGATAATAATGTCACGAGGCATTGGGATAAAGTATTAAGAGAGGAGATTGGGACATATTAATCC
TCCACTACCTGGGGCTGGACCACATTGGCCACATTTCAGGGCCCAACAGCCCCCTGATTGGGCAG
AAGCTGAGCGAGATGGACAGCGTGTGATGAAGATCCACACCTCACTGCAGTCGAAGGAGAGAGA
GACGCCTTTACCCAATTTGCTGGTTCTTTGTGGTGACCATGGCATGTCTGAAACAGGAAGTCACG
GGGCCTCCTCCACCGAGGAGGTGAATACACCTCTGATTTTAAATCAGTTCTGCGTTTGAAAGGAAA
CCCGGTGATATCCGACATCCAAAGCACGTCCAATAGACGGATGTGGCTGCGACACTGGCGATAGC
ACTTGGCTTACCGATTCCAAAAGACAGTGTAGGGAGCCTCCTATTCCAGTTGTGGAAGGAAGAC
CAATGAGAGAGCAGTTGAGATTTTACATTTGAATACAGTGCAGCTTAGTAAACTGTTGCAAGAG
AATGTGCCGTCATATGAAAAGATCCTGGGTTTGAGCAGTTTAAATGTCAGAAAGATTGCATGG
GAACTGGATCAGACTGTACTTGGAGGAAAAGCATTGAGAAGTCCTATTCAACCTGGGCTCCAAGG
TTCTCAGGCAGTACCTGGATGCTCTGAAGACGCTGAGCTTGTCCCTGAGTGCACAAGTGGCCCAG
TTCTCACCCCTGCTCCTGCTCAGCGTCCCACAGGCACTGCACAGAAAGGCTGAGCTGGAAGTCCCA
CTGTCACTCTCCTGGGTTTTCTCTGCTCTTTTATTTGGTGATCCTGGTTCTTTCGGCCGTTACAGT
CATTTGTGTGCACCTCAGCTGAAAGTTCTGTGCTACTTCTGTGGCCTCTCGTGGCTGGCGGCAGGCT
GCCTTTTCGTTTACCAGACTCTGGTTGAACACCTGGTGTGTGCCAAGTGTGGCAGTGCCTTGGAC
AGGGGGCCTCAGGGAAGGACGTGGAGCAGCCTTATCCCAGGCCCTGGGTGTCCCGACACAGGTG
TTCACATCTGTGCTGTGAGGTGAGATGCCTCAGTTCTTGGAAGCTAGGTTCTTGGACTGTTAC
CAAGGTGATTGTAAAGAGCTGGCGGTACAGAGGAACAAGCCCCCAGCTGAGGGGGTGTGTGAA
TCGGACAGCCTCCCAGCAGAGGTGTGGGAGCTGCAGCTGAGGGAAGAAGAGACAATCGGCCTGGA
CACTCAGGAGGGTCAAAGGAGACTTGGTCGCACCACTCATCCTGCCACCCCAGAATGCATCCT
GCCTCATCAGGTCCAGATTTCTTTCCAAGGCGGACGTTTTCTGTTGGAATTCTTAGTCTTGGCC
TCGGACACCTTCATTTCGTTAGCTGGGGAGTGGTGGTGAGGCAGTGAAGAAGAGCGGATGGTCAC
ACTCAGATCCACAGAGCCCAGGATCAAGGGACCCACTGCAGTGGCAGCAGGACTGTTGGGCCCCC
ACCCCAACCCTGCACAGCCCTCATCCCTCTTGGCTTGAGCCGTCAGAGGCCCTGTGCTGAGTGT
CTGACCGAGACACTCACAGCTTTGTATCAGGGCACAGGCTTCCTCGGAGCCAGGATGATCTGTG
CCACGCTTGCACCTCGGGCCCATCTGGGCTCATGCTCTCTCCTGCTATTGAATTAGTACCTAG
CTGCACACAGTATGTAGTTACCAAAAGAATAAACGGCAATAATTGAGAAAAAAA

FIGURE 140

MRLGSGTTFATCCVAIEVLGIAVFLRGFFPAPVRSSARAEGAEPPAPEPSAGASSNWTTLPPPLF
SKVVIVLIDALRDDDFVFGSKGVKFMPTTYLVEKGASHSFVAEAKPPTVTMPRIKALMTGSLPGF
VDVIRNLNSPALLEDVIRQAKAAGKRIVFYGDETWVKLFPKHFVEYDGTTSFFVSDYTEVDNNV
TRHLDKVLKRGDWDILILHYLGLDHIGHISGPN SPLIGQKLSEMDSVLMKIHTSLQSKERETPLP
NLLVLCGDHGMSETGSHGASSTEEVNTPLILISSAFERKPGDIRHPKHVQ

Important features of the protein:

Signal peptide:

amino acids 1-34

Transmembrane domain:

amino acids 58-76

N-glycosylation sites.

amino acids 56-60, 194-198

N-myristoylation sites.

amino acids 6-12, 52-58, 100-106, 125-131, 233-239, 270-276,
275-281, 278-284

Amidation site.

amino acids 154-158

Cell attachment sequence.

amino acids 205-208

FIGURE 141

GGCACGAGGCAAGCCTTCCAGGTTATCGTGACGCACCTTGAAAGTCTGAGAGCTACTGCCCTACA
GAAAGTTACTAGTGCCCTAAAGCTGGCGCTGGCACTGATGTTACTGCTGCTGTTGGAGTACAAC
TCCCTATAGAAAACAACCTGCCAGCACCTTAAGACCACTCACACCTTCAGAGTGAAGAACTTAAAC
CCGAAGAAATTCAGCATTCATGACCAGGATCACAAAGTACTGGTCCTGGACTCTGGGAATCTCAT
AGCAGTTCCAGATAAAAACTACATACGCCCAGAGATCTTCTTTGCATTAGCCTCATCCTTGAGCT
CAGCCTCTGCGGAGAAAGGAAGTCCGATTCTCCTGGGGGTCTCTAAAGGGGAGTTTTGTCTCTAC
TGTGACAAGGATAAAGGACAAAGTCATCCATCCCTTCAGCTGAAGAAGGAGAACTGATGAAGCT
GGCTGCCCAAAGGAATCAGCACGCCGGCCCTTCATCTTTTATAGGGCTCAGGTGGGCTCCTGGA
ACATGCTGGAGTCGGCGGCTCACCCCGGATGGTTCATCTGCACCTCCTGCAATTGTAATGAGCCT
GTTGGGGTGACAGATAAATTTGAGAACAGGAAACACATTGAATTTTCATTTCAACCAGTTTGCAA
AGCTGAAATGAGCCCCAGTGAGGTCAGCGATTAGGAAACTGCCCCATTGAACGCCTTCCTCGCTA
ATTTGAACTAATTGTATAAAAAACACCAAACCTGCTCACT

FIGURE 142

MLLLLLLEYNFPIENNCQHLKTTHTFRVKNLNPKKFSIHDQDHKVLVLDSGNLIAVPDKNYIRPEI
FFALASSLSSASAEGKSPILLGVSKGEFCLYCDKDKGQSHPSLQLKKEKLMKLAAQKESARRPFI
FYRAQVGSWNMLESAAHPGWFICTSCNCNEPVGVTDKFENRKHIEFSFQPVCKAEMSPSEVSD

cAMP- and cGMP-dependent protein kinase phosphorylation site.

amino acids 33-36

N-myristoylation site.

amino acids 50-55, 87-92

Interleukin-1

amino acids 37-182

FIGURE 143

CTAGAGAGTATAGGGCAGAAGGATGGCAGATGAGTGACTCCACATCCAGAGCTGCCTCCCTTTAA
TCCAGGATCCTGTCCTTCCTGTCCTGTAGGAGTGCCGTGTTGCCAGTGTGGGGTGAGACAAGTTTG
TCCCACAGGGCTGTCTGAGCAGATAAGATTAAGGGCTGGGTCTGTGCTCAATTAACCTGTGGG
CACGGGGGCTGGGAAGAGCAAAGTCAGCGGTGCCTACAGTCAGCACCATGCTGGGCCTGCCGTGG
AAGGGAGGTCTGTCCTGGGCGCTGCTGCTGCTTCTCTTAGGCTCCCAGATCCTGCTGATCTATGC
CTGGCATTTCACAGAGCAAAGGGACTGTGATGAACACAATGTCATGGCTCGTTACCTCCCTGCCA
CAGTGGAGTTTGCTGTCCACACATTCAACCAACAGAGCAAGGACTACTATGCCTACAGACTGGGG
CACATCTTGAATTCCCTGGAAGGAGCAGGTGGAGTCCAAGACTGTATTCTCAATGGAGCTACTGCT
GGGGAGAAGTAGGTGTGGGAAATTTGAAGACGACATTGACAAGTGCATTTCCAAGAAAGCACAG
AGCTGAACAATACTTTCACCTGCTTCTTCACCATCAGCACCAGGCCCTGGATGACTCAGTTCAGC
CTCCTGAACAAGACCTGCTTGGAGGGATTCCACTGAGTGAAACCCACTCACAGGCTTGTCCATGT
GCTGCTCCCACATTCCGTGGACATCAGCACTACTCTCCTGAGGACTCTTCAGTGGCTGAGCAGCT
TTGGACTTGTTTGTATCCTATTTTGCATGTGTTTGAGATCTCAGATCAGTGTTTTAGAAAATCC
ACACATCTTGAGCCTAATCATGTAGTGTAGATCATTAACATCAGCATTTTAAGAAAAAAAAAAAA
AAA

FIGURE 144

MLGLPWKGGLSWALLLLLLLGSQILLIYAWHFHEQRDCDEHNVMARYLPATVEFAVHTFNQQSKDY
YAYRLGHILNSWKEQVESKTVFSMELLLGRTRCGKFEDDIDNCHFQESTELNNTFTCFETISTRP
WMTQFSLLNKTCLEGFH

Important features of the protein:

Signal peptide:

amino acids 1-25

N-glycosylation sites.

amino acids 117-121, 139-143

N-myristoylation site.

amino acids 9-15

FIGURE 145

CTGTGCAGCTCGAGGCTCCAGAGGCACACTCCAGAGAGAGCCAAGGTTCTGACGCGATGAGGAAG
CACCTGAGCTGGTGGTGGCTGGCCACTGTCTGTCATGCTGCTCTTCAGCCACCTCTCTGCGGTCCA
GACGAGGGGCATCAAGCACAGAATCAAGTGGAAACCGGAAGGCCCTGCCCAGCACTGCCCAGATCA
CTGAGGCCCAGGTGGCTGAGAACCGCCCCGGGAGCCTTCATCAAGCAAGGCCGCAAGCTCGACATT
GACTTCGGAGCCGAGGGCAACAGGTACTACGAGGCCAACTACTGGCAGTTCCCCGATGGCATCCA
CTACAACGGCTGCTCTGAGGCTAATGTGACCAAGGAGGCATTTGTCACCGGCTGCATCAATGCCA
CCCAGGCGGCGAACCAGGGGGAGTTCCAGAAGCCAGACAACAAGCTCCACCAGCAGGTGCTCTGG
CGGCTGGTCCAGGAGCTCTGCTCCCTCAAGCATTGCGAGTTTTGGTTGGAGAGGGGCGCAGGACT
TCGGGTCACCATGCACCAGCCAGTGCTCCTCTGCCTTCTGGCTTTGATCTGGCTCATGGTGAAAT
AAGCTTGCCAGGAGGCTGGCAGTACAGAGCGCAGCAGCGAGCAAATCCTGGCAAGTGACCCAGCT
CTTCTCCCCCAAACCCACGCGTGTCTGAAGGTGCCCAGGAGCGGCGATGCACTCGCACTGCAAA
TGCCGCTCCCACGTATGCGCCCTGGTATGTGCCTGCGTTCTGATAGATGGGGGACTGTGGCTTCT
CCGTCACTCCATTCTCAGCCCCTAGCAGAGCGTCTGGCACACTAGATTAGTAGTAAATGCTTGAT
GAGAAGAACACATCAGGCACTGCGCCACCTGCTTCACAGTACTTCCCAACAACCTCTTAGAGGTAG
GTGTATTCCCGTTTTACAGATAAGGAAACTGAGGCCCAGAGAGCTGAAGTACTGCACCCAGCATC
ACCAGCTAGAAAGTGGCAGAGCCAGGATTCAACCCCTGGCTTGTCTAACCCAGGTTTTCTGCTCT
GTCCAATTCCAGAGCTGTCTGGTGATCACTTTATGTCTCACAGGGACCCACATCCAAACATGTAT
CTCTAATGAAATTGTGAAAGCTCCATGTTTAGAAATAAATGAAAACACCTGA

FIGURE 146

MRKHLSWWLATVCMLLFSHLSAVQTRGIKHRIKWNRKALPSTAQITEAQVAENRPGAFIKQGRK
LDIDFGAEGNRYYEANYWQFPDGIHYNGCSEANVTKEAFVTGCINATQAANQGEFOKPDNKLHQQ
VLWRLVQELCSLKHCFWLERGAGLRVTMHQPVLLCLLALIWLMMVK

Important features of the protein:

Signal peptide:

amino acids 1-26

Transmembrane domain:

amino acids 157-171

N-glycosylation sites.

amino acids 98-102, 110-114

Tyrosine kinase phosphorylation site.

amino acids 76-83

N-myristoylation sites.

amino acids 71-77, 88-94, 93-99, 107-113, 154-160

Amidation site.

amino acids 62-66

FIGURE 147

GCCTTGGCCTCCCAAAGGGCTGGGATTATAGGCGTGACCACCATGTCTGGTCCAGAGTCTCATTT
CCTGATGATTTATAGACTCAAAGAAAACTATGTTCAGAAGCTCTCTTCTCTTCTGGCCTCCTCT
CTGTCTTCTTTCCCTCTTTCTTCTTATTTTAATTAGTAGCATCTACTCAGAGTCATGCAAGCTGG
AAATCTTTCATTTTGCTTGTCAGTGGGGTAGGTCAGTCTTAGTTTTATTTTTTGAAATTT
CAACTTTCAGATTCAGGGGGTACATGTGAAGGTTTGTGTTTATGAGTATATTGCATGATGCTGAGG
TTTGGGGT

FIGURE 148

MFRSSLLFWPPLCLLSLFLLLILISSIYSESKLEIFHFACQWGRSLSLSFYFLKFQLSDSGGTCE
GLFYEYIA

Important features of the protein:

Signal peptide:

amino acids 1-25

N-myristoylation site.

amino acids 62-68

FIGURE 149

GTCTCCGCGTCACAGGAACCTTCAGCACCCACAGGGCGGACAGCGCTCCCCCTCTACCTGGAGACTTGAC
TCCCGCGCGCCCCAACCCCTGCTTATCCCTTGACCGTCGAGTGTGAGAGATCCTGCAGCCGCCAGTCC
CGGCCCCCTCTCCCGCCCCACACCCACCCCTCCTGGCTCTTCCTGTTTTTACTCCTCCTTTTCATTGATA
ACAAAAGCTACAGCTCCAGGAGCCCAGCGCCGGGCTGTGACCCAAGCCGAGCGTGGAAGAATGGGGTT
CCTCGGGACCGGCACTTGGATTCTGGTGTAGTGCTCCCGATTCAAGCTTTCCCCAACCTGGAGGAA
GCCAAGACAAATCTCTACATAATAGAGAATTAAGTGCAGAAAGACCTTTGAATGAACAGATTGCTGAA
GCAGAAGAAGACAAGATTAAAAAACATATCCTCCAGAAAACAAGCCAGGTGAGAGCAACTATTCTTT
TGTTGATAACTTGAACCTGCTAAAGGCAATAACAGAAAAGGAAAAAATTGAGAAAGAAAGACAATCTA
TAAGAAGCTCCCCACTTGATAATAAGTTGAATGTGGAAGATGTTGATTCAACCAAGAATCGAAAACTG
ATCGATGATTATGACTCTACTAAGAGTGGATTGGATCATAAATTTCAAGATGATCCAGATGGTCTTCA
TCAACTAGACGGGACTCCTTTAACCGCTGAAGACATTGTCCATAAAATCGCTGCCAGGATTTATGAAG
AAAATGACAGAGCCGTGTTTGACAAGATTGTTTCTAACTACTTAATCTCGGCCTTATCACAGAAAGC
CAAGCACATACACTGGAAGATGAAGTAGCAGAGGTTTTACAAAAATTAATCTCAAAGGAAGCCAACAA
TTATGAGGAGGATCCCAATAAGCCACAAGCTGGACTGAGAATCAGGCTGGAAAAATACCAGAGAAAG
TGACTCCAATGGCAGCAATTCAAGATGGTCTTGCTAAGGGAGAAAACGATGAAACAGTATCTAACACA
TTAACCTTGACAAATGGCTTGGAAGGAGAACTAAAACCTACAGTGAAGACAACCTTGAGGAACTCCA
ATATTTCCCAAATTTCTATGCGCTACTGAAAAGTATTGATTGAGAAAAAGAACAAAGAGAAAGAAA
CACTGATTACTATCATGAAAACACTGATTGACTTTGTGAAGATGATGGTGAAATATGGAACAATATCT
CCAGAAGAAGGTGTTTCTACCTTGAAAACCTGGATGAAATGATTGCTCTTCAGACCAAAAACAAGCT
AGAAAAAATGCTACTGACAATATAAGCAAGCTTTTCCAGCACCATCAGAGAAGAGTCATGAAGAAA
CAGACAGTACCAAGGAAGAAGCAGCTAAGATGGAAAAGGAATATGGAAGCTTGAAGGATTCCACAAAA
GATGATAACTCCAACCCAGGAGGAAAGACAGATGAACCCAAAGGAAAAACAGAAGCCTATTTGGAAGC
CATCAGAAAAAATATTGAATGGTTGAAGAAACATGACAAAAGGGAAATAAAGAAGATTATGACCTTT
CAAAGATGAGAGACTTCATCAATAAACAAGCTGATGCTTATGTGGAGAAAGGCATCCTTGACAAGGAA
GAAGCCGAGGCCATCAAGCGCATTTATAGCAGCCTGTAAAAATGGCAAAAGATCCAGGAGTCTTTCAA
CTGTTTCAGAAAACATAATATAGCTTAAAACACTTCTAATTCTGTGATTAAAAATTTTTTGACCCAAGG
GTTATTAGAAAGTGCTGAATTTACAGTAGTTAACCTTTTACAAGTGGTTAAAACATAGCTTTCTTCCC
GTAAAAACTATCTGAAAGTAAAGTTGTATGTAAGCTGAAAAAAAAAAAAAAAAAAAAA

FIGURE 150

MGFLGTGTWILVLVLPIQAFPKPGGSQDKSLHNRELSAERPLNEQIAEAEEDKIKKTYPPENKPG
QSNYSFVDNLNLLKAITEKEKIEKERQSIRSSPLDNKLNVEDVDSTKNRKLIDDYDSTKSGLDHK
FQDDPDGLHQLDGTPLTAEDIVHKIAARIYEENDRAVFDKIVSKLLNLGLITESQAHTLEDEVAE
VLQKLISKEANNYEEDPNKPTSWTENQAGKIKEKVTMAAIQDGLAKGENDETVSNTLTLTNGLE
RRTKTYSEDNFEELQYFPNIFYALLKSIDSEKEAKEKETLITIMKTLIDFVKMMVKYGTISPEEGV
SYLENLDEMIALQTKNKLEKNATDNISKLFPAPEKSHEETDSTKEEAAKMEKEYGSLKDSTKDD
NSNPGGKTDEPKGKTEAYLEAIRKNIEWLKKHKDKGNKEDYDLSKMRDFINKQADAYVEKGILDK
EEAEAIKRIYSSL

N-glycosylation sites:

amino acids 68-71, 346-349, 350-353

Casein kinase II phosphorylation site:

amino acids 70-73, 82-85, 97-100, 125-128, 147-150, 188-191, 217-
220, 265-268, 289-292, 305-308, 320-323, 326-329, 362-365, 368-
341, 369-372, 382-385, 386-389, 387-390

N-myristoylation sites:

amino acids 143-148, 239-244

FIGURE 151

CGGCTCGAGGCTCCCGCCAGGAGAAAGGAACATTCTGAGGGGAGTCTACACCCTGTGGAGCTCAA
GATGGTCTCTGAGTGGGGCGCTGTGCTTCCGAATGAAGGACTCGGCATTGAAGGTGCTTTATCTGC
 ATAATAACCAGCTTCTAGCTGGAGGGCTGCATGCAGGGAAGGTCATTAAAGGTGAAGAGATCAGC
 GTGGTCCCCAATCGGTGGCTGGATGCCAGCCTGTCCCCCGTCATCCTGGGTGTCCAGGGTGAAG
 CCAGTGCCTGTCTGTGGGGTGGGGCAGGAGCCGACTCTAACACTAGAGCCAGTGAACATCATGG
 AGCTCTATCTTGGTGCCAAGGAATCCAAGAGCTTACCTTCTACCGGCGGGACATGGGGCTCACC
 TCCAGCTTCGAGTCGGCTGCCACCCGGGCTGGTTCCTGTGCACGGTGCCTGAAGCCGATCAGCC
 TGTGAGACTCAGCCAGCTTCCCGAGAATGGTGGCTGGAATGCCCCATCACAGACTTCTACTTCC
 AGCAGTGTGAC**TAG**GGCAACGTGCCCCCAGAACTCCCTGGGCAGAGCCAGCTCGGGTGAGGGGT
 GAGTGGAGGAGACCCATGGCGGACAATCACCTCTCTGCTCTCAGGACCCCCACGTCTGACTTAG
 TGGGCACCTGACCACTTTGTCTTCTGGTTCCTCAGTTTGGATAAAATTCTGAGATTTGGAGCTCAGT
 CCACGGTCCCTCCCCCACTGGATGGTGCTACTGCTGTGGAACCTTGTA AAAACCATGTGGGGTAAA
 CTGGGAATAACATGAAAAGATTTCTGTGGGGGTGGGGTGGGGGAGTGGTGGGAATCATTCCTGCT
 TAATGGTAACTGACAAGTGTACCCTGAGCCCCGAGGCCAACCCATCCCCAGTTGAGCCTTATA
 GGGTCAGTAGCTCTCCACATGAAGTCTGTCACTCACCACCTGTGCAGGAGAGGGAGGTGGTCATA
 GAGTCAGGGATCTATGGCCCTTGGCCCAGCCCCACCCCTTCCCTTTAATCCTGCCACTGTCATA
 TGCTACCTTTCCCTATCTCTTCCCTCATCATCTTGTGTGGGCATGAGGAGGTGGTGATGTCAGAA
 GAAATGGCTCGAGCTCAGAAGATAAAAGATAAGTAGGGTATGCTGATCCTCTTTTAAAAACCCAA
 GATACAATCAAAATCCCAGATGCTGGTCTCTATTCCCATGAAAAAGTGCTCATGACATATTGAGA
 AGACCTACTTACAAAGTGGCATATATTGCAATTTATTTTAATTA AAAAGATACCTATTTATATATT
 TCTTTATAGAAAAAGTCTGGAAGAGTTTACTTCAATTGTAGCAATGTCAGGGTGGTGGCAGTAT
 AGGTGATTTTTCTTTTAATTCTGTTAATTTATCTGTATTTCCCTAATTTTTCTACAATGAAGATGA
 ATTCCTTGATATAAAAATAAGAAAAGAAATTAATCTTGAGGTAAGCAGAGCAGACATCATCTCTGA
 TTGTCTCAGCCTCCACTTCCCCAGAGTAAATTCAAATGAATCGAGCTCTGCTGCTCTGGTTGG
 TTGTAGTAGTGATCAGGAAACAGATCTCAGCAAAGCCACTGAGGAGGAGGCTGTGCTGAGTTTGT
 GTGGCTGGAATCTCTGGGTAAGGAACCTAAAGAACAAAAATCATCTGGTAATTCTTTCCTAGAAG
 GATCACAGCCCCCTGGGATTCCAAGGCATTGGATCCAGTCTCTAAGAAGGCTGCTGTACTGGTTGA
 ATTGTGTCCCCCTCAAATTCACATCCTTCTTGGAATCTCAGTCTGTGAGTTTATTTGGAGATAAG
 GTCTCTGCAGATGTAGTTAGTTAAGACAAGGTCTGCTGGATGAAGGTAGACCTAAATTCATAT
 GACTGGTTTCCCTTGATGAAAAGGAGAGGACACAGAGACAGAGGAGACGCGGGGAAGACTATGTA
 AAGATGAAGGCAGAGATCGGAGTTTTGCAGCCACAAGCTAAGAAACACCAAGGATTGTGGCAACC
 ATCAGAAGCTTGGAAGAGGCAAAGAAGAATCTTCCCTAGAGGCTTTAGAGGGATAACGGCTCTG
 CTGAAACCTTAATCTCAGACTTCCAGCCTCCTGAACGAAGAAAGAATAAATTTGGCTGTTTTAA
 GCCACCAAGGATAATTGGTTACAGCAGCTCTAGGAACTAATAACAGCTGCTAAAATGATCCCTGT
 CTCCTCGTGTTTACATTCTGTGTGTGTCCCCTCCACAATGTACCAAAGTTGTCTTTGTGACCAA
 TAGAATATGGCAGAAGTGATGGCATGCCACTTCCAAGATTAGGTTATAAAAAGACACTGCAGCTTC
 TACTTGAGCCCTCTCTCTGTGCCACCCACCGCCCCCAATCTATCTTGGCTCACTCGCTCTGGGGG
 AAGCTAGCTGCCATGCTATGAGCAGGCCTATAAAGAGACTTACGTGGTAAAAAATGAAGTCTCCT
 GCCCACAGCCACATTAGTGAACCTAGAAGCAGAGACTCTGTGAGATAATCGATGTTTGTGTTTT
 AAGTTGCTCAGTTTTTGGTCTAACTTGTTATGCAGCAATAGATAAATAATATGCAGAGAAAAG

FIGURE 152

MVLSGALCFRMKDSALKVLYLHNNQLLAGGLHAGKVIKGEELSVVPSNRWLDASLSPVILGVQGGG
QCLSCGVGQEPTLTLEPVNIMELYLGAKESKSFTFYRRDMGLTSSFESAAYPGWFLCTVPEADQP
VRLTQLPENGGWNAPITDFYFQQCD

N-myristoylation sites.

amino acids 29-34, 30-35, 60-65, 63-68, 73-78, 91-96, 106-111

Interleukin-1 signature.

amino acids 111-131

Interleukin-1 proteins.

amino acids 8-29, 83-120, 95-134, 64-103

FIGURE 153

CTTCAGAACAGGTTCTCCTTCCCCAGTCACCAGTTGCTCGAGTTAGAATTGTCTGCAATGGCCGC
CCTGCAGAAATCTGTGAGCTCTTTCCCTTATGGGGACCCTGGCCACCAGCTGCCTCCTTCTCTTGG
CCCTCTTGGTACAGGGAGGAGCAGCTGCGCCCATCAGCTCCCACTGCAGGCTTGACAAGTCCAAC
TTCCAGCAGCCCTATATCACCAACCGCACCTTCATGCTGGCTAAGGAGGCTAGCTTGGCTGATAA
CAACACAGACGTTTCGTCTCATTGGGGAGAACTGTTCCACGGAGTCAGTATGAGTGAGCGCTGCT
ATCTGATGAAGCAGGTGCTGAACTTCACCCCTGAAGAAGTGCTGTTCCCTCAATCTGATAGGTTC
CAGCCTTATATGCAGGAGGTGGTGCCCTTCCTGGCCAGGCTCAGCAACAGGCTAAGCACATGTCA
TATTGAAGGTGATGACCTGCATATCCAGAGGAATGTGCAAAAGCTGAAGGACACAGTGAAAAAGC
TTGGAGAGAGTGGAGAGATCAAAGCAATTGGAGAACTGGATTTGCTGTTTATGTCTCTGAGAAAT
GCCTGCATTTTGACCAGAGCAAAGCTGAAAAATGAATAACTAACCCCCCTTCCCTGCTAGAAATAA
CAATTAGATGCCCCAAAGCGATTTTTTTTTAACCAAAAGGAAGATGGGAAGCCAACTCCATCATG
ATGGGTGGATTCCAAATGAACCCCTGCGTTAGTTACAAAGGAAACCAATGCCACTTTTGTTTATA
AGACCAGAAGGTAGACTTTCTAAGCATAGATATTTATTGATAACATTTTCATTGTAAGTGGTGTTT
TATACACAGAAAACAATTTATTTTTTAAATAATTGTCTTTTTCCATAAAAAAGATTACTTTCCAT
TCCTTTAGGGGAAAAAACCCCTAAATAGCTTCATGTTTCCATAATCAGTACTTTATATTTATAAA
TGTATTTATTATTATTATAAGACTGCATTTTATTTATATCATTTTATTAATATGGATTTATTTAT
AGAAACATCATTCGATATTGCTACTTGAGTGTAAGGCTAATATTGATATTTATGACAATAATTAT
AGAGCTATAACATGTTTATTTGACCTCAATAAACACTTGGATATCCC

FIGURE 154

MAALQKSVSSFLMGTLATSCLLLLLALLVQGGAAAPISSHCRLDKSNFQQPYITNRTFMLAKEASL
ADNNTDVRLIGEKLFGVSMSERCYLMKQVLNFTLEEVLFPQSDRFQPYMQEVVPFLARLSNRLS
TCHIEGDDLHIQRNVQKLKDTVKKLGESGEIKAIGELDLLFMSLRNACI

Important features of the protein:

Signal peptide:

amino acids 1-33

N-glycosylation sites.

amino acids 54-58, 68-72, 97-101

N-myristoylation sites.

amino acids 14-20, 82-88

Prokaryotic membrane lipoprotein lipid attachment site.

amino acids 10-21

FIGURE 155

GGCTTGCTGAAAATAAAATCAGGACTCCTAACCTGCTCCAGTCAGCCTGCTTCCACGAGGCCTGT
CAGTCAGTGCCCGACTTGTGACTGAGTGTGCAGTGCCAGCATGTACCAGGTCAGTGCAGAGGGC
TGCCTGAGGGCTGTGCTGAGAGGGAGAGGAGCAGAGATGCTGCTGAGGGTGGAGGGAGGCCAAGC
TGCCAGGTTTGGGGCTGGGGGCCAAGTGGAGTGAGAACTGGGATCCCAGGGGGAGGGTGCAGAT
GAGGGAGCGACCCAGATTAGGTGAGGACAGTTCTCTCATTAGCCTTTTCCTACAGGTGGTTGCAT
TCTTGGCAATGGTCATGGGAACCCACACCTACAGCCACTGGCCCAGCTGCTGCCCCAGCAAAGGG
CAGGACACCTCTGAGGAGCTGCTGAGGTGGAGCACTGTGCCTGTGCCTCCCCTAGAGCCTGCTAG
GCCCACCGCCACCCAGAGTCCTGTAGGGCCAGTGAAGATGGACCCCTCAACAGCAGGGCCATCT
CCCCCTGGAGATATGAGTTGGACAGAGACTTGAACCGGCTCCCCCAGGACCTGTACCACGCCCGT
TGCCTGTGCCCGCACTGCGTCAGCCTACAGACAGGCTCCCACATGGACCCCCGGGGCAACTCGGA
GCTGCTCTACCACAACCAGACTGTCTTCTACAGGCGGCCATGCCATGGCGAGAAGGGCACCCACA
AGGGCTACTGCCTGGAGCGCAGGCTGTACCGTGTTTCCCTTAGCTTGTGTGTGTGTGCGGGCCCCGT
GTGATGGGCTAGCCGGACCTGCTGGAGGCTGGTCCCTTTTTGGGAAACCTGGAGCCAGGTGTACA
ACCACTTGCCATGAAGGGCCAGGATGCCCAGATGCTTGGCCCCCTGTGAAGTGCTGTCTGGAGCAG
CAGGATCCCGGGACAGGATGGGGGGCTTTGGGGAAAACCTGCACTTCTGCACATTTTGAAAAGAG
CAGCTGCTGCTTAGGGCCGCCGGAAGCTGGTGTCTGTCTATTTCTCTCAGGAAAGGTTTTCAAA
GTTCTGCCCATTTCTGGAGGCCACCACTCCTGTCTCTTCCCTCTTTTCCCATCCCCTGCTACCCTG
GCCCAGCACAGGCACTTCTAGATATTTCCCCCTTGCTGGAGAAGAAAGAGCCCCCTGGTTTTATT
TGTTTGTTTACTCATCACTCAGTGAGCATCTACTTTGGGTGCATTCTAGTGTAGTTACTAGTCTT
TTGACATGGATGATTCTGAGGAGGAAGCTGTTATTGAATGTATAGAGATTTATCCAAATAAATAT
CTTTATTTAAAAATGAAAAA

FIGURE 156

MRERPRLGEDSSLISLFLQVVAFLAMVMGTHTYSHWPSCCPSKGQDTSEELLRWSTVPVPPLEPA
RPNRHPESCRASEDGPLNSRAISPWRYELDRDLNRLPQDLYHARCLCPHCVSLQTGSHMDPRGNS
ELLYHNQTVFYRRPCHGEKGTGKGYCLERRLYRVSLACVCVRPRVMG

Important features of the protein:

Signal peptide:

amino acids 1-32

N-glycosylation site.

amino acids 136-140

Tyrosine kinase phosphorylation site.

amino acids 127-135

N-myristoylation sites.

amino acids 44-50, 150-156

FIGURE 157

CCGGCGATGTCGCTCGTCTGCTAAGCCTGGCCGCGCTGTGCAGGAGCGCCGTACCCCGAGAGCC
GACCGTTCAATGTGGCTCTGAACTGGGCCATCTCCAGAGTGGATGCTACAACATGATCTAATCC
CCGGAGACTTGAGGGACCTCCGAGTAGAACCTGTTACAAC TAGTGTGCAACAGGGGACTATTCA
ATTTTGATGAATGTAAGCTGGGTACTCCGGGCAGATGCCAGCATCCGCTTGTTGAAGGCCACCAA
GATTTGTGTGACGGGCAAAAGCAACTTCCAGTCCTACAGCTGTGTGAGGTGCAATTACACAGAGG
CCTTCCAGACTCAGACCAGACCCTCTGGTGGTAAATGGACATTTTCCTACATCGGCTTCCCTGTA
GAGCTGAACACAGTCTATTTTCATTGGGGCCCATAATATTCCTAATGCAAATATGAATGAAGATGG
CCCTTCCATGTCTGTGAATTTACCTCACCAGGCTGCCTAGACCACATAATGAAATATAAAAAA
AGTGTGTCAAGGCCGGAAGCCTGTGGGATCCGAACATCACTGCTTGTGAAGAAGATGAGGAGACA
GTAGAAGTGAAC TTCACAACCACTCCCCTGGGAAACAGATACATGGCTCTTATCCAACACAGCAC
TATCATCGGGTTTTCTCAGGTGTTTGAGCCACACCAGAAGAAACAAACGCGAGCTTCAGTGGTGA
TTCCAGTGACTGGGGATAGTGAAGGTGCTACGGTGCAGCTGACTCCATATTTTCCTACTTGTGGC
AGCGACTGCATCCGACATAAAGGAACAGTTGTGCTCTGCCACAAACAGGCGTCCCTTTCCCTCT
GGATAACAACAAAAGCAAGCCGGGAGGCTGGCTGCCTCTCCTCCTGCTGTCTCTGCTGGTGGCCA
CATGGGTGCTGGTGGCAGGGATCTATCTAATGTGGAGGCACGAAAGGATCAAGAAGACTTCCTTT
TCTACCACCACACTACTGCCCCCATTAAGGTTCTTGTGGTTTACCCATCTGAAATATGTTTCCA
TCACACAATTTGTTACTTCCTGAATTTCTTCAAACCATTCGAGAAGTGAGGTCACTCTTGAAA
AGTGGCAGAAAAAGAAAATAGCAGAGATGGGTCCAGTGCAGTGGCTTGCCACTCAAAAGAAGGCA
GCAGACAAAGTCGTCTTCTTCTTCCAATGACGTCAACAGTGTGTGCGATGGTACCTGTGGCAA
GAGCGAGGGCAGTCCCAGTGAGAACTCTCAAGACCTCTTCCCCCTTGCCTTTAACCTTTTCTGCA
GTGATCTAAGAAGCCAGATTCATCTGCACAAATACGTGGTGGTCTACTTTAGAGAGATTGATACA
AAAGACGATTACAATGCTCTCAGTGTCTGCCCCAAGTACCACCTCATGAAGGATGCCACTGCTTT
CTGTGCAGAACTTCTCCATGTCAAGCAGCAGGTGTCAGCAGGAAAAAGATCACAAGCCTGCCACG
ATGGCTGCTGCTCCTTGTAG

FIGURE 158

MSLVLLSLAALCRSAVPREPTVQCGSETGPSPEWMLQHDLPGLRDLRVEPVTTTSVATGDYSILMNVS
WVLRADASIRLLKATKICVTKGSNFSYSCVRCNYTEAFQTQTRPSGGKWTFSYIGFPVELNTVYFIGAHNIP
NANMNEDGPSMSVNFTSPGCLDHIMKYKKKCVKAGSLWDPNITACKKNEETVEVNFTTTPLGNRYMALIQH
STIIGFSQVFEPHQKKQTRASVVIPVTGDSEGATVQLTPYFPTCGSDCIRHKGTVVLCPTGVPFPLDNNK
SKPGGWLPLLLLSLLVATWVLVAGIYLMWRHERIKKTSFSTTTLLPPIKVLVVYPSEICFHHTICYFTEFL
QNHCRSEVILEKWQKKKIAEMGPVQWLATQKKAADKVVFLSNDVNSVCDGTCGKSEGSPSENSQDLFPLA
FNLFCSDLRSQIHLHKYVVVYFREIDTKDDYNALSVC PKYHLMKDATAFCAELLHVKKQVSAGKRSQACHD
GCCSL

Important features of the protein:

Signal peptide:

amino acids 1-14

Transmembrane domain:

amino acids 290-309

N-glycosylation sites.

amino acids 67 - 71, 103 - 107, 156 - 160, 183 - 187, 197 - 201 and 283
- 287

cAMP- and cGMP-dependent protein kinase phosphorylation sites.

amino acids 228 - 232 and 319 - 323

Casein kinase II phosphorylation sites.

amino acids 178 - 182, 402 - 406, 414 - 418 and 453 - 457

N-myristoylation site.

amino acids 116-122

Amidation site.

amino acids 488-452

FIGURE 159

AGCCACCAGCGCAACATGACAGTGAAGACCCTGCATGGCCCAGCCATGGTCAAGTACTTGCTGCT
GTCGATATTGGGGCTTGCCTTTCTGAGTGAGGCGGCAGCTCGGAAAATCCCCAAAGTAGGACATA
CTTTTTTCCAAAAGCCTGAGAGTTGCCCGCCTGTGCCAGGAGGTAGTATGAAGCTTGACATTGGC
ATCATCAATGAAAACCAGCGCGTTTCCATGTCACGTAACATCGAGAGCCGCTCCACCTCCCCCTG
GAATTACACTGTCACTTGGGACCCCAACCGGTACCCCTCGGAAGTTGTACAGGCCCAGTGTAGGA
ACTTGGGCTGCATCAATGCTCAAGGAAAGGAAGACATCTCCATGAATTCCGTTCCCATCCAGCAA
GAGACCCTGGTCGTCCGGAGGAAGCACCAAGGCTGCTCTGTTTCTTTCCAGTTGGAGAAGGTGCT
GGTGACTGTTGGCTGCACCTGCGTCACCCCTGTCATCCACCATGTGCAGTAAGAGGTGCATATCC
ACTCAGCTGAAGAAG

FIGURE 160

MTVKTLHGPMVKYLLLSILGLAFLSEAAARKIPKVGHTFFQKPESCPPVPGGSMKLDIGIINEN
QRVMSRNIESRSTSPWNYTWTWDPNRYPSSEVVQAQCRNLGCINAQGKEDISMNSVPIQQETLVV
RRKHQGCSVSFQLEKVLVTVGCTCVTPVIHHVQ

Signal sequence:

amino acids 1-30

N-glycosylation site.

amino acids 83-87

N-myristoylation sites.

amino acids 106-111, 136-141

FIGURE 161

A C A C T G G C C A A A C A A A A A C G A A A G C A C T C C G T G C T G G A A G T A G G A G G A G A G T C A G G A C T C C C A G G
 A C A G A G A G T G C A C A A A C T A C C C A G C A C A G C C C C C T C C G C C C C C T C T G G A G G C T G A A G A G G G A T T C
 C A G C C C C T G C C A C C C A C A G A C A C G G G C T G A C T G G G G T G T C T G C C C C C C T T G G G G G G G G C A G C A C
 A G G G C C T C A G G C C T G G G T G C C A C C T G G C A C C T A G A A G A T G C C T G T G C C C T G G T T C T T G C T G T C C T
 T G G C A C T G G G C C G A A G C C C A G T G G T C C T T T C T C T G G A G A G G C T T G T G G G G C C T C A G G A C G C T A C C
 C A C T G C T C T C C G G G C C T C T C C T G C C G C C T C T G G G A C A G T G A C A T A C T C T G C C T G C C T G G G G A C A T
 C G T G C C T G C T C C G G G C C C C G T G C T G G C G C C T A C G C A C C T G C A G A C A G A G C T G G T G C T G A G G T G C C
 A G A A G G A G A C C G A C T G T G A C C T C T G T C T G C G T G T G G C T G T C C A C T T G G C C G T G C A T G G G C A C T G G
 G A A G A G C C T G A A G A T G A G G A A A G T T T G G A G G A G C A G C T G A C T C A G G G G T G G A G G A G C C T A G G A A
 T G C C T C T C T C C A G G C C C A A G T C G T G C T C T C C T T C C A G G C C T A C C C T A C T G C C C G C T G C G T C C T G C
 T G G A G G T G C A A G T G C C T G C T G C C C T T G T G C A G T T T G G T C A G T C T G T G G G C T C T G T G G T A T A T G A C
 T G C T T C G A G G C T G C C C T A G G G A G T G A G G T A C G A A T C T G G T C C T A T A C T C A G C C C A G G T A C G A G A A
 G G A A C T C A A C C A C A C A C A G C A G C T G C C T G C C C T G C C C T G G C T C A A C G T G T C A G C A G A T G G T G A C A
 A C G T G C A T C T G G T T C T G A A T G T C T C T G A G G A G C A G C A C T T C G G C C T C T C C C T G T A C T G G A A T C A G
 G T C C A G G G C C C C C A A A A C C C G G T G G C A C A A A A C C T G A C T G G A C C G C A G A T C A T T A C C T T G A A
 C C A C A C A G A C C T G G T T C C C T G C C T C T G T A T T C A G G T G T G G C C T C T G G A A C C T G A C T C C G T T A G G A
 C G A A C A T C T G C C C C T T C A G G G A G G A C C C C G C G C A C A C C A G A A C C T C T G G C A A G C C G C C C G A C T G
 C G A C T G C T G A C C C T G C A G A G C T G G C T G C T G G A C G C A C C G T G C T C G C T G C C C G C A G A A G C G G C A C T
 G T G C T G G C G G G C T C C G G G T G G G G A C C C C T G C C A G C C A C T G G T C C C A C C G C T T T C C T G G G A G A A C G
 T C A C T G T G G A C A A G G T T C T C G A G T T C C C A T T G C T G A A A G G C C A C C C T A A C C T C T G T G T T C A G G T G
 A A C A G C T C G G A G A A G C T G C A G C T G C A G G A G T G C T T G T G G G C T G A C T C C C T G G G G C C T C T C A A A G A
 C G A T G T G C T A C T G T T G G A G A C A C G A G G C C C C A G G A C A A C A G A T C C C T C T G T G C C T T G G A A C C C A
 G T G G C T G T A C T T C A C T A C C C A G C A A A G C C T C C A C G A G G G C A G C T C G C C T T G G A G A G T A C T T A C T A
 C A A G A C C T G C A G T C A G G C C A G T G T C T G C A G C T A T G G G A C G A T G A C T T G G G A G C G C T A T G G G C C T G
 C C C C A T G G A C A A A T A C A T C C A C A A G C G C T G G G C C C T C G T G T G G C T G G C C T G C C T A C T C T T T G C C G
 C T G C G C T T T C C C T C A T C C T C C T T C T C A A A A A G G A T C A C G C G A A A G G G T G G C T G A G G C T C T T G A A A
 C A G G A C G T C C G C T C G G G G G C G G C C G C C A G G G G C C G C G C G G C T C T G C T C C T A C T C A G C C G A T G A
 C T C G G G T T T C G A G C G C C T G G T G G G C G C C C T G G C G T C G G C C C T G T G C C A G C T G C C G C T G C G C G T G G
 C C G T A G A C C T G T G G A G C C G T C G T G A A C T G A G C G C G C A G G G G C C C G T G G C T T G G T T T C A C G C G C A G
 C G G C G C C A G A C C C T G C A G G A G G G C G G C G T G G T G G T C T T G C T C T T C T C C C G G T G C G G T G G C G C T
 G T G C A G C G A G T G G C T A C A G G A T G G G G T G T C C G G G C C C G G G G C G C A C G G C C C G C A C G A C G C C T T C C
 G C G C C T C G C T C A G C T G C G T G C T G C C C G A C T T C T T G C A G G G C C G G G C G C C C G G C A G C T A C G T G G G G
 G C C T G C T T C G A C A G G C T G C T C C A C C C G G A C G C C G T A C C C G C C C T T T T C C G C A C C G T G C C C G T C T T
 C A C A C T G C C C T C C C A A C T G C C A G A C T T C C T G G G G G C C C T G C A G C A G C C T C G C G C C C C G C G T C C G
 G G C G G C T C C A A G A G A G A G C G G A G C A A G T G T C C C G G G C C C T T C A G C C A G C C C T G G A T A G C T A C T T C
 C A T C C C C C G G G G A C T C C C G C G C C G G A C G C G G G T G G G A C C A G G G C G G G A C C T G G G G C G G G G A
 C G G G A C T T A A A T A A A G G C A G A C G C T G T T T T T C T A A A A A A

FIGURE 162

MPVPWFLLSLALGRSPVVLSELERLVGPQDATHCSPGLSCRLWDSILCLPGDIVPAPGPVLAPTHLQTELV
LRCQKETDCDLCLRVAVHLAVHGHWEPEDEEKFGGAADSGVEEPRNASLQAQVVLSTQAYPTARCVLLEV
QVPAALVQFGQSVGSVVYDCFEAALGSEVRIWSYTQPRYEKELNHTQQLPALPWLNVNSADGDNVHLVLNVS
EEQHFGLSLYWNQVQGPPKPRWHKNLTGPIITLNHTDLVPCLCIQVWPLEPDSVRTNICPFREDPRAHQN
LWQAARLRLTLQSWLLDAPCSLPAEAALCWRAPGGDPCQPLVPPLSWENVTVDKVLEFPLLKGHPNLCVQ
VNSSEKLQEQECLWADSLGPLKDDVLLLETRGPQDNRSILCALEPSGCTSLPSKASTRAARLGEYLLQDLQS
GQCLQLWDDDLGALWACPMCKYIHKRWALVWLACLLFAAALSLILLKKDHAKGWLRLKQDVRSGAAARG
RAALLYSADDSGFERLVGALASALCQLPLRVAVDLWSRRELSAQGPVAVFHAQRRQTLQEGGVVLLFSP
GAVALCSEWLQDGVSGPGAHGPHDAFRASLSCVLPDFLQGRAPGSYVGACFDRLHPDAVPALFRTVPVFT
LPSQLPDFLGALQQPRAPRSGRLQERAEQVSRAALQPALDSYFHPPGTPAPGRGVGPGAGPGAGDGT

Signal sequence:

amino acids 1-20

Transmembrane domain.

amino acids 453-475

N-glycosylation sites.

amino acids 118-121, 186-189, 198-201, 211-214, 238-241, 248-251,
334-337, 357-360, 391-394

Glycosaminoglycan attachment site.

amino acids 583-586

cAMP- and cGMP-dependent protein kinase phosphorylation site.

amino acids 552-555

N-myristoylation sites.

amino acids 107-112, 152-157, 319-324, 438-443, 516-521, 612-617,
692-697, 696-701, 700-705

FIGURE 163

GGGAGGGCTCTGTGCCAGCCCCGATGAGGACGCTGCTGACCATCTTGACTGTGGGATCCCTGGCT
GCTCAGCCCCCTGAGGACCCCTCGGATCTGCTCCAGCACGTGAAATTCCAGTCCAGCAACTTTGA
AAACATCCTGACGTGGGACAGCGGGCCAGAGGGCACCCAGACACGGTCTACAGCATCGAGTATA
AGACGTACGGAGAGAGGGACTGGGTGGCAAAGAAGGGCTGTCAGCGGATCACCCGGAAGTCCTGC
AACCTGACGGTGGAGACGGGCAACCTCACGGAGCTCTACTATGCCAGGGTCACCGCT
GTCAGTGCGGGAGGCGGTCAGCCACCAAGATGACTGACAGGTTGAGTCTCTGCAGCACACTAC
CCTCAAGCCACCTGATGTGACCTGTATCTCCAAAGTGAGATCGATTGAGATGATTGTTTCATCCTA
CCCCACGCCAATCCGTGCAGGCGATGGCCACCGGCTAACCCCTGGAAGACATCTTCCATGACCTG
TTCTACCACTTAGAGCTCCAGGTCAACCGCACCTACCAAATGCACCTTGGAGGGAAGCAGAGAGA
ATATGAGTTCTTCGGCCTGACCCCTGACACAGAGTTTCTTGGCACCATCATGATTTGCGTTCCCA
CCTGGGCCAAGGAGAGTGCCCCCTACATGTGCCGAGTGAAGACACTGCCAGACCGGACATGGACC
TACTCCTTCTCCGGAGCCTTCTGTCTCCATGGGCTTCTCGTCGCAGTACTCTGCTACCTGAG
CTACAGATATGTACCAAGCCGCTGCACCTCCCAACTCCCTGAACGTCCAGCGAGTCTTGACTT
TCCAGCCGCTGCGCTTCATCCAGGAGCACGTCTGATCCCTGTCTTTGACCTCAGCGGCCCCAGC
AGTCTGGCCAGCCTGTCCAGTACTCCAGATCAGGGTGTCTGGACCCAGGGAGCCCGCAGGAGC
TCCACAGCGGCATAGCCTGTCCGAGATCACCTACTTAGGGCAGCCAGACATCTCCATCCTCCAGC
CCTCCAACGTGCCACCTCCCCAGATCCTCTCCCCACTGTCTATGCCCCAAACGCTGCCCTGAG
GTCGGGCCCCCATCTATGCACCTCAGGTGACCCCCGAAGCTCAATTCCCATTTACGCCCCACA
GGCCATCTCTAAGGTCCAGCCTTCTCCTATGCCCCCTCAAGCCACTCCGGACAGCTGGCCTCCCT
CCTATGGGGTATGCATGGAAGGTTCTGGCAAAGACTCCCCACTGGGACACTTTCTAGTCTTAA
CACCTTAGGCCTAAAGGTGAGCTTCAGAAAGAGCCACCAGCTGGAAGCTGCATGTTAGGTGGCCT
TTCTCTGCAGGAGGTGACCTCCTTGGCTATGGAGGAATCCCAAGAAGCAAAATCATTGCACCAGC
CCCTGGGGATTGTGCACAGACAGAACATCTGACCCAAATGTGCTACACAGTGGGGAGGAAGGGACA
CCACAGTACCTAAAGGGCCAGCTCCCCCTCCTCTCCTCAGTCCAGATCGAGGGCCACCCATGTC
CCTCCCTTTGCAACCTCCTTCCGGTCCATGTTCCCCCTCGGACCAAGGTCCAAGTCCCTGGGGCC
TGCTGGAGTCCCTTGTGTGTCCCAAGGATGAAGCCAAGAGCCCAGCCCCTGAGACCTCAGACCTG
GAGCAGCCCACAGAACTGGATTCTCTTTTCAGAGGCCTGGCCCTGACTGTGCAGTGGGAGTCTTG
AGGGGAATGGGAAAGGCTTGGTGCTTCTCCTGTCCCTACCCAGTGTACATCCTTGGCTGTCA
ATCCCATGCCTGCCCATGCCACACACTCTGCGATCTGGCCTCAGACGGGTGCCCTTGAGAGAAGC
AGAGGGAGTGGCATGCAGGGCCCCCTGCCATGGGTGCGCTCCTCACCGGAACAAAGCAGCATGATA
AGGACTGCAGCGGGGGAGCTCTGGGGAGCAGCTTGTGTAGACAAGCGCGTGTCTCGTGTAGCCCTG
CAAGGCAGAAATGACAGTGCAAGGAGGAAATGCAGGGAAACTCCCGAGGTCCAGAGCCCCACCTC
CTAACACCATGGATTCAAAGTGCTCAGGGAATTTGCTCTCCTTGCCCCATTCTGGCCAGTTTC
ACAATCTAGCTCGACAGAGCATGAGGCCCCCTGCCTCTCTGTGTCATTGTTCAAAGGTGGGAAGAGA
GCCTGGAAAAGAACCAGGCCTGGAAAAGAACCAGAAGGAGGCTGGGCAGAACCAGAACAACCTGC
ACTTCTGCCAAGGCCAGGGCCAGCAGGACGGCAGGACTCTAGGGAGGGGTGTGGCCTGCAGCTCA
TTCCCAGCCAGGGCAACTGCCTGACGTTGCACGATTTGAGCTTCATTCTCTGATAGAACAAAGC
GAAATGCAGGTCCACCAGGGAGGGAGACACACAAGCCTTTTCTGCAGGCAGGAGTTTCAGACCCT
ATCCTGAGAATGGGGTTTGAAAGGAAGGTGAGGGCTGTGGCCCCCTGGACGGGTACAATAACACAC
TGTAATGATGTACAACTTTGCAAGCTCTGCCTTGGGTTGAGCCCATCTGGGCTCAAATTCAGC
CTCACCCTCACAAGCTGTGTGACTTCAAACAAATGAAATCAGTGCCAGAACCTCGGTTTCTCTC
ATCTGTAATGTGGGGATCATAACACCTACCTCATGGAGTTGTGGTGAAGATGAAATGAAGTCATG
TCTTTAAAGTGCTTAATAGTGCTGGTACATGGGCAGTGCCCAATAAACGGTAGCTATTTAAAAA
AAAAAAA

FIGURE 164

MRTLLTILTVGSLAAHAPEDPSDLLQHVKFQSSNFENILTWDSGPEGTPDTPVYSIEYKTYGERDW
VAKKGCQRITRKSCNLTVETGNLTELYARVTAVSAGGRSATKMTDRFSSLQHTTLKPPDVTCLIS
KVRSIQMIVHPTPTPIRAGDGHRLTLEDIFHDLFYHLELQVNRITYQMHLGGKQREYEFFGLTPDT
EFLGTIMICVPTWAKESAPYMCRVKTLPDRTWTYSFSGAFLFSMGFLVAVLCYLSYRYVTKPPAP
PNSLNVQRVLTFQPLRFIQEHVLIPVFDLSGPSSLAQPVQYSQIRVSGPREPAGAPQRHSLSEIT
YLGQPDISILQPSNVPPPQILSPLSYAPNAAPEVGPPSYAPQVTPEAQFPFYAPQAISKVQPSSY
APQATPDSWPPSYGVCMESGKDSPTGTLSSPKHLRPKGQLQKEPPAGSCMLGGLSLQEVTSIAM
EESQEAQSLHQPLGICTDRTSDPNVLHSGEETGPQYLKGQLPLLSSVQIEGHPMSLPLQPPSGPC
SPSDQGPSPWGLLESILVCPKDEAKSPAPETSDLEQPTELDSLFRGLALTQWES

Signal sequence.

amino acids 1-17

Transmembrane domain.

amino acids 233-250

N-glycosylation sites.

amino acids 80-83, 87-90, 172-175

N-myristoylation sites.

amino acids 11-16, 47-52, 102-107, 531-536, 565-570

FIGURE 165

TGGCCTACTGGAAAAAAAAAAAAAAAAAAGTCACCCGGGCCCGCGGTGGCCACAACATGG
CTGCGGCGCCGGGGCTGCTCTTCTGGCTGTTTCGTGCTGGGGGCGCTCTGGTGGGTCCCGGGCCAG
TCGGATCTCAGCCACGGACGGCGTTTCTCGGACCTCAAAGTGTGCGGGGACGAAGAGTGCAGCAT
GTTAATGTACCGTGGGAAAGCTCTTGAAGACTTCACGGGCCCTGATTGTGCTTTTGTGAATTTTA
AAAAAGGTGACGATGTATATGTCTACTACAACTGGCAGGGGGATCCCTTGAACTTTGGGCTGGA
AGTGTGTAACACAGTTTTTGGATATTTTCCAAAAGATTTGATCAAGGTACTTCATAAATACACGGA
AGAAGAGCTACATATTCCAGCAGATGAGACAGACTTTGTCTGCTTTGAAGGAGGAAGAGATGATT
TTAATAGTTATAATGTAGAAGAGCTTTTAGGATCTTTGGAACTGGAGGACTCTGTACCTGAAGAG
TCGAAGAAAGCTGAAGAAGTTTCTCAGCACAGAGAGAAATCTCCTGAGGAGTCTCGGGGGCGTGA
ACTTGACCCTGTGCCTGAGCCCGAGGCATTGAGAGCTGATTGAGAGGATGGAGAAGGTGCTTTCT
CAGAGAGCACCGAGGGGCTGCAGGGACAGCCCTCAGCTCAGGAGAGCCACCCTCACACCAGCGGT
CCTGCGGCTAACGCTCAGGGAGTGCAGTCTTCGTTGGACACTTTTGAAGAAATTCTGCACGATAA
ATTGAAAGTGCCGGGAAGCGAAAGCAGAACTGGCAATAGTTCTCCTGCCTCGGTGGAGCGGGAGA
AGACAGATGCTTACAAAGTCCTGAAAACAGAAATGAGTCAGAGAGGAAGTGGACAGTGC GTTATT
CATTACAGCAAAGGATTTTCGTTGGCATCAAAATCTAAGTTTGTTTTACAAAGATTGTTTTTAGTA
CTAAGCTGCCTTGGCAGTTTGCATTTTGTAGCCAAACAAAAATATATTATTTCCCTTCTAAGTA
AAAAAAAAAAAAAAAAAAAAA

FIGURE 166

MAAAPGLLFWLFWLGALWWVPGQSDLSHGRRFSDLKVCGDEEC SMLMYRGKALEDFTGPDCRFVN
FKKGDDVYVYYKLAGGSLELWAGSVEHSFGYFPKDLIKVLHKYTEELHIPAETDFVCFEGGRD
DFNSYNVEELLGSLELED SVPEESKKAEEVSQHREKSP EESRGRELDPVPEPEAFRA DSEGE GA
FSESTEG LQGQPSAQESH PHTSGPAANAQGVQSSLDTFEEILHDKLKVPGSESRTGNSSPASVER
EKTDAYKVLKTEM SQRGSGQCVIHYSKGFRWHQNL SLFYKDCF

Important features of the protein:

Signal peptide:

amino acids 1-22

N-glycosylation site.

amino acids 294-298

cAMP- and cGMP-dependent protein kinase phosphorylation site.

amino acids 30-34

Tyrosine kinase phosphorylation site.

amino acids 67-76

N-myristoylation sites.

amino acids 205-211, 225-231, 277-283

Amidation site.

amino acids 28-32

FIGURE 167

CCAGGACCAGGGCGCACCGGCTCAGCCTCTCATTGTGTCAGAGGCCGGGGAAGAGAAGCAAAGCGC
AACGGTGTGGTCCAAGCCGGGGCTTCTGCTTCGCCTCTAGGACATACACGGGACCCCTAACTTC
AGTCCCCCAAACGCGCACCCCTCGAAGTCTTGAAGTCCAGCCCCGCACATCCACGCGCGGCACAGG
CGCGGCAGGCGGCAGGTCCCGGCCGAAGGCGATGCGCGCAGGGGGTTCGGGCAGCTGGGCTCGGGC
GGCGGGAGTAGGGCCCGGCAGGGAGGCAGGGAGGCTGCATATTAGAGTTCGCGGGCTGCGCCCTG
GGCAGAGGCCCGCCCTCGCTCCACGCAACACCTGCTGCTGCCACCGCGCCGCGATGAGCCGCGTGG
TCTCGCTGCTGCTGGGCGCCGCGCTGCTCTGCGGCCACGGAGCCTTCTGCCGCGCGTGGTTCAGC
GGCCAAAAGGTGTGTTTTGCTGACTTCAAGCATCCCTGCTACAAAATGGCCTACTTCCATGAACT
GTCCAGCCGAGTGAGCTTTAGGAGGCACGCCTGGCTTGTGAGAGTGAGGGAGGAGTCTCTCTCA
GCCTTGAGAATGAAGCAGAACAGAAGTTAATAGAGAGCATGTTGCAAAACCTGACAAAACCCGGG
ACAGGGATTTCTGATGGTGATTTCTGGATAGGGCTTTGGAGGAATGGAGATGGGCAAACATCTGG
TGCCTGCCCAGATCTCTACCAGTGGTCTGATGGAAGCAATTCCCAGTACCGAAACTGGTACACAG
ATGAACCTTCTGCGGAAGTGAAAAGTGTGTTGTGATGTATCACCACCAACTGCCAATCCTGGC
CTTGGGGGTCCCTACCTTTACCAGTGGAAATGATGACAGGTGTAACATGAAGCACAATTATATTTG
CAAGTATGAACCAGAGATTAATCCAACAGCCCCTGTAGAAAAGCCTTATCTTACAAATCAACCAG
GAGACACCCATCAGAATGTGGTGTACTGAAGCAGGTATAATTCCAATCTAATTTATGTTGTT
ATACCAACAATACCCCTGCTCTTACTGATACTGGTTGCTTTTGGAACTGTTGTTTCCAGATGCT
GCATAAAAGTAAAGGAAGAACAATACTAGTCCAAACCAGTCTACACTGTGGATTTCAAAGAGTA
CCAGAAAAGAAAGTGGCATGGAAGTATAAATACTCATTGACTTGGTTCCAGAATTTTGTAATTCT
GGATCTGTATAAGGAATGGCATCAGAACAATAGCTTGGAAATGGCTTGAAATCACAAAGGATCTGC
AAGATGAAGTGAAGCTCCCCCTTGAGGCAAATATTAAAGTAATTTTTATATGTCTATTATTTCA
TTTAAAGAATATGCTGTGCTAATAATGGAGTGAGACATGCTTATTTTGCTAAAGGATGCACCCAA
ACTTCAAACCTCAAGCAAATGAAATGGACAATGCAGATAAAGTTGTTATCAACACGTCGGGAGTA
TGTGTGTTAGAAGCAATTCCTTTTATTTCTTTACCTTTTATAAGTTGTTATCTAGTCAATGTAA
TGTATATTGTATTGAAATTTACAGTGTGCAAAAGTATTTTACCTTTGCATAAGTGTGTGATAAAA
ATGAACTGTTCTAATATTTATTTTATGGCATCTCATTTTTCAATACATGCTCTTTTGATTAAAG
AACTTATTACTGTTGTCAACTGAATTCACACACACACAAATATAGTACCATAGAAAAAGTTGT
TTTCTCGAAATAATTCATCTTTCAGCTTCTCTGCTTTTGGTCAATGTCTAGGAAATCTCTTCAGA
AATAAGAAGCTATTTTCAATTAAGTGTGATATAAACCTCCTCAAACATTTTACTTAGAGGCAAGGAT
TGTCTAATTTCAATTGTGCAAGACATGTGCCTTATAATTATTTTATAGCTTAAATTAACAGATT
TTGTAATAATGTAACCTTGTAAATAGGTGCATAAACACTAATGCAGTCAATTTGAACAAAAGAAG
TGACATACACAATATAAATCATATGTCTTCACACGTTGCCTATATAATGAGAAGCAGCTCTCTGA
GGGTTCTGAAATCAATGTGGTCCCTCTCTTGCCCACTAAACAAAGATGGTTGTTTCGGGGTTTGGG
ATTGACACTGGAGGCAGATAGTTGCAAAGTTAGTCTAAGGTTTCCCTAGCTGTATTTAGCCTCTG
ACTATATTAGTATACAAAGAGGTCATGTGGTTGAGACCAGGTGAATAGTCACTATCAGTGTGGAG
ACAAGCACAGCACACAGACATTTTAGGAAGGAAAGGAACACGAAATCGTGTGAAAATGGGTG
AACCATCAGTGATCGCATATTCATTGATGAGGGTTTGCCTTGAGATAGAAAATGGTGGCTCCTTT
CTGTCTTATCTCCTAGTTTCTTCAATGCTTACGCCTTGTCTTCTCAAGAGAAAGTTGTAACCT
CTGGTCTTCATATGTCCCTGTGCTCCTTTTAACCAAATAAAGAGTTCTTGTCTTCTGGGGGAAAAA
AAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAA

FIGURE 168

MSRVVSLLLGAALLCGHGAFCCRVSQGKVCFADEFKHPCKMAYFHELSSRVSFQEARLACESE
GGVLLSLENEAEQKLIESMLQNLTKEPGTGISDGDWGLWRNGDGTSGACPDLYQWSDGSNSQ
YRNWYTDEPSCGSEKCVVMYHQPTANPGLGGPYLYQWNDDRCNMKHNICKYEPEINPTAPVEK
PYLTNQPGDTHQNVVVTEAGIIPNLIYVVIPTIPLLLLILVAFGTCCFQMLHKSQGRKTKTSPNQ
STLWISKSTRKESGMEV

Important features of the protein:

Signal peptide:

amino acids 1-21

Transmembrane domain:

amino acids 214-235

N-glycosylation sites.

amino acids 86-89, 255-258

cAMP- and cGMP-dependent protein kinase phosphorylation site.

amino acids 266-269

N-myristoylation sites.

amino acids 27-32, 66-71, 91-96, 93-98, 102-107, 109-114, 140-
145, 212-217

**SECRETED AND TRANSMEMBRANE
POLYPEPTIDES AND NUCLEIC ACIDS
ENCODING THE SAME**

**CROSS REFERENCE TO RELATED
APPLICATIONS**

[0001] This is a continuation application claiming priority under 35 USC § 120 to U.S. Ser. No. 10/006,867 Filed Dec. 6, 2001, and which claims priority under 35 USC § 119 to U.S. provisional serial Nos. 60/063,435 Filed Oct. 29, 1997; 60/064,215 Filed Oct. 29, 1997; 60/082,797 Filed Apr. 22, 1998; 60/083,495 Filed Apr. 29, 1998; 60/085,579 Filed May 15, 1998; 60/087,759 Filed Jun. 2, 1998; 60/088,021 Filed Jun. 4, 1998; 60/088,029 Filed Jun. 4, 1998; 60/088,030 Filed Jun. 4, 1998; 60/088,734 Filed Jun. 10, 1998; 60/088,740 Filed Jun. 10, 1998; 60/088,811 Filed Jun. 10, 1998; 60/088,824 Filed Jun. 10, 1998; 60/088,825 Filed Jun. 10, 1998; 60/088,863 Filed Jun. 11, 1998; 60/089,105 Filed Jun. 12, 1998; 60/089,514 Filed Jun. 16, 1998; 60/089,653 Filed Jun. 17, 1998; 60/089,952 Filed Jun. 19, 1998; 60/090,246 Filed Jun. 22, 1998; 60/090,444 Filed Jun. 24, 1998; 60/090,688 Filed Jun. 25, 1998; 60/090,696 Filed Jun. 25, 1998; 60/090,862 Filed Jun. 26, 1998; 60/091,628 Filed Jul. 2, 1998; 60/096,012 Filed Aug. 10, 1998; 60/096,757 Filed Aug. 17, 1998; 60/096,949 Filed Aug. 18, 1998; 60/096,959 Filed Aug. 18, 1998; 60/097,954 Filed Aug. 26, 1998; 60/097,971 Filed Aug. 26, 1998; 60/097,979 Filed Aug. 26, 1998; 60/098,749 Filed Sep. 1, 1998; 60/099,741 Filed Sep. 10, 1998; 60/099,763 Filed Sep. 10, 1998; 60/099,792 Filed Sep. 10, 1998; 60/099,812 Filed Sep. 10, 1998; 60/099,815 Filed Sep. 10, 1998; 60/100,627 Filed Sep. 16, 1998; 60/100,662 Filed Sep. 16, 1998; 60/100,683 Filed Sep. 17, 1998; 60/100,684 Filed Sep. 17, 1998; 60/100,930 Filed Sep. 17, 1998; 60/101,279 Filed Sep. 22, 1998; 60/101,475 Filed Sep. 23, 1998; 60/101,738 Filed Sep. 24, 1998; 60/101,743 Filed Sep. 24, 1998; 60/101,916 Filed Sep. 24, 1998; 60/102,570 Filed Sep. 30, 1998; 60/103,449 Filed Oct. 6, 1998; 60/103,678 Filed Oct. 8, 1998; 60/103,679 Filed Oct. 8, 1998; 60/103,711 Filed Oct. 8, 1998; 60/105,000 Filed Oct. 20, 1998; 60/105,002 Filed Oct. 20, 1998; 60/105,881 Filed Oct. 27, 1998; 60/106,030 Filed Oct. 28, 1998; 60/106,464 Filed Oct. 30, 1998; 60/106,856 Filed Nov. 3, 1998; 60/108,807 Filed Nov. 17, 1998; 60/112,419 Filed Dec. 15, 1998; 60/112,422 Filed Dec. 15, 1998; 60/112,853 Filed Dec. 16, 1998; 60/113,011 Filed Dec. 16, 1998; 60/112,854 Filed Dec. 16, 1998; 60/113,300 Filed Dec. 22, 1998; 60/113,408 Filed Dec. 22, 1998; 60/113,430 Filed Dec. 23, 1998; 60/113,621 Filed Dec. 23, 1998; 60/114,223 Filed Dec. 30, 1998; 60/115,614 Filed Jan. 12, 1999; 60/116,527 Filed Jan. 20, 1999; 60/116,843 Filed Jan. 22, 1999; 60/119,285 Filed Feb. 9, 1999; 60/119,287 Filed Feb. 9, 1999; 60/119,525 Filed Feb. 10, 1999; 60/119,549 Filed Feb. 10, 1999; 60/120,014 Filed Feb. 11, 1999; 60/129,122 Filed Apr. 13, 1999; 60/129,674 Filed Apr. 16, 1999; 60/131,291 Filed Apr. 27, 1999; 60/138,387 Filed Jun. 9, 1999; 60/144,791 Filed Jul. 20, 1999; 60/169,495 Filed Dec. 7, 1999; 60/175,481 Filed Jan. 11, 2000; 60/191,007 Filed Mar. 21, 2000; 60/199,397 Filed Apr. 25, 2000 and which claims priority under 35 USC § 120 to U.S. patent applications and PCT International Patent Application Ser. Nos. 09/380,139 Filed Aug. 25, 1999; 09/311,832 Filed May 14, 1999; 09/380,137 Filed Aug. 25, 1999, now abandoned; 09/380,138 Filed Aug. 25, 1999, now abandoned; 09/380,142 Filed Aug. 25, 1999, now abandoned; 09/397,342 Filed

Sep. 15, 1999; 09/403,297 Filed Oct. 18, 1999, now abandoned; 09/423,844 Filed Nov. 12, 1999, now abandoned; 09/644,848 Filed Aug. 22, 2000; 09/665,350 Filed Sep. 18, 2000; 09/664,610 Filed Sep. 18, 2000, now abandoned; 09/709,238 Filed Nov. 8, 2000; 09/747,259 Filed Dec. 20, 2000; 09/816,744 Filed Mar. 22, 2001; 09/854,208 Filed May 10, 2001; 09/854,280 Filed May 10, 2001; 09/870,574 Filed May 30, 2001; 09/874,503 Filed Jun. 5, 2001; 09/908,827 Filed Jul. 18, 2001; 09/869,566 Filed Feb. 19, 2002; PCT/US98/19330 Filed Sep. 16, 1998; PCT/US99/05028 Filed Mar. 8, 1999; PCT/US99/10733 Filed May 14, 1999; PCT/US99/12252 Filed Jun. 2, 1999; PCT/US99/20111 Filed Sep. 1, 1999; PCT/US99/21090 Filed Sep. 15, 1999; PCT/US99/21194 Filed Sep. 15, 1999; PCT/US99/30720 Filed Dec. 22, 1999; PCT/US00/04341 Filed Feb. 18, 2000; PCT/US00/04342 Filed Feb. 18, 2000; PCT/US00/04414 Filed Feb. 22, 2000; PCT/US00/05601 Filed Mar. 1, 2000; PCT/US00/08439 Filed Mar. 30, 2000; PCT/US00/14042 Filed May 22, 2000; PCT/US00/15264 Filed Jun. 2, 2000; PCT/US00/23522 Filed Aug. 23, 2000; PCT/US00/23328 Filed Aug. 24, 2000; PCT/US00/30873 Filed Nov. 10, 2000; PCT/US00/32678 Filed Dec. 1, 2000; PCT/US00/34956 Filed Dec. 20, 2000; PCT/US01/06520 Filed Feb. 28, 2001; PCT/US01/06666 Filed Mar. 1, 2001; PCT/US01/17443 Filed May 30, 2001; PCT/US01/17800 Filed Jun. 1, 2001; PCT/US01/19692 Filed Jun. 20, 2001; PCT/US01/21066 Filed Jun. 29, 2001; PCT/US01/21735 Filed Jul. 9, 2001, the entire disclosures of which are hereby incorporated by reference.

BACKGROUND OF THE INVENTION

[0002] The present invention relates generally to the identification and isolation of novel DNA and to the recombinant production of novel polypeptides.

[0003] Extracellular proteins play important roles in, among other things, the formation, differentiation and maintenance of multicellular organisms. The fate of many individual cells, e.g., proliferation, migration, differentiation, or interaction with other cells, is typically governed by information received from other cells and/or the immediate environment. This information is often transmitted by secreted polypeptides (for instance, mitogenic factors, survival factors, cytotoxic factors, differentiation factors, neuropeptides, and hormones) which are, in turn, received and interpreted by diverse cell receptors or membrane-bound proteins. These secreted polypeptides or signaling molecules normally pass through the cellular secretory pathway to reach their site of action in the extracellular environment.

[0004] Secreted proteins have various industrial applications, including as pharmaceuticals, diagnostics, biosensors and bioreactors. Most protein drugs available at present, such as thrombolytic agents, interferons, interleukins, erythropoietins, colony stimulating factors, and various other cytokines, are secretory proteins. Their receptors, which are membrane proteins, also have potential as therapeutic or diagnostic agents. Efforts are being undertaken by both industry and academia to identify new, native secreted proteins. Many efforts are focused on the screening of mammalian recombinant DNA libraries to identify the coding sequences for novel secreted proteins. Examples of screening methods and techniques are described in the literature [see, for example, Klein et al., Proc. Natl. Acad. Sci. 93:7108-7113 (1996); U.S. Pat. No. 5,536,637].

[0005] Membrane-bound proteins and receptors can play important roles in, among other things, the formation, differentiation and maintenance of multicellular organisms. The fate of many individual cells, e.g., proliferation, migration, differentiation, or interaction with other cells, is typically governed by information received from other cells and/or the immediate environment. This information is often transmitted by secreted polypeptides (for instance, mitogenic factors, survival factors, cytotoxic factors, differentiation factors, neuropeptides, and hormones) which are, in turn, received and interpreted by diverse cell receptors or membrane-bound proteins. Such membrane-bound proteins and cell receptors include, but are not limited to, cytokine receptors, receptor kinases, receptor phosphatases, receptors involved in cell-cell interactions, and cellular adhesion molecules like selectins and integrins. For instance, transduction of signals that regulate cell growth and differentiation is regulated in part by phosphorylation of various cellular proteins. Protein tyrosine kinases, enzymes that catalyze that process, can also act as growth factor receptors. Examples include fibroblast growth factor receptor and nerve growth factor receptor.

[0006] Membrane-bound proteins and receptor molecules have various industrial applications, including as pharmaceutical and diagnostic agents. Receptor immunoadhesins, for instance, can be employed as therapeutic agents to block receptor-ligand interactions. The membrane-bound proteins can also be employed screening of potential peptide or small molecule inhibitors of the relevant receptor/ligand interaction.

[0007] Efforts are being undertaken by both industry and academia to identify new, native receptor or membrane-bound proteins. Many efforts are focused on the screening of mammalian recombinant DNA libraries to identify the coding sequences for novel receptor or membrane-bound proteins.

SUMMARY OF INVENTION

[0008] In one embodiment, the invention provides an isolated nucleic acid molecule comprising a nucleotide sequence that encodes a PRO polypeptide.

[0009] In one aspect, the isolated nucleic acid molecule comprises a nucleotide sequence having at least about 80% nucleic acid sequence identity, alternatively at least about 81% nucleic acid sequence identity, alternatively at least about 82% nucleic acid sequence identity, alternatively at least about 83% nucleic acid sequence identity, alternatively at least about 84% nucleic acid sequence identity, alternatively at least about 85% nucleic acid sequence identity, alternatively at least about 86% nucleic acid sequence identity, alternatively at least about 87% nucleic acid sequence identity, alternatively at least about 88% nucleic acid sequence identity, alternatively at least about 89% nucleic acid sequence identity, alternatively at least about 90% nucleic acid sequence identity, alternatively at least about 91% nucleic acid sequence identity, alternatively at least about 92% nucleic acid sequence identity, alternatively at least about 93% nucleic acid sequence identity, alternatively at least about 94% nucleic acid sequence identity, alternatively at least about 95% nucleic acid sequence identity, alternatively at least about 96% nucleic acid sequence identity, alternatively at least about 97% nucleic

acid sequence identity, alternatively at least about 98% nucleic acid sequence identity and alternatively at least about 99% nucleic acid sequence identity to (a) a DNA molecule encoding a PRO polypeptide having a full-length amino acid sequence as disclosed herein, an amino acid sequence lacking the signal peptide as disclosed herein, an extracellular domain of a transmembrane protein, with or without the signal peptide, as disclosed herein or any other specifically defined fragment of the full-length amino acid sequence as disclosed herein, or (b) the complement of the DNA molecule of (a).

[0010] In other aspects, the isolated nucleic acid molecule comprises a nucleotide sequence having at least about 80% nucleic acid sequence identity, alternatively at least about 81% nucleic acid sequence identity, alternatively at least about 82% nucleic acid sequence identity, alternatively at least about 83% nucleic acid sequence identity, alternatively at least about 84% nucleic acid sequence identity, alternatively at least about 85% nucleic acid sequence identity, alternatively at least about 86% nucleic acid sequence identity, alternatively at least about 87% nucleic acid sequence identity, alternatively at least about 88% nucleic acid sequence identity, alternatively at least about 89% nucleic acid sequence identity, alternatively at least about 90% nucleic acid sequence identity, alternatively at least about 91% nucleic acid sequence identity, alternatively at least about 92% nucleic acid sequence identity, alternatively at least about 93% nucleic acid sequence identity, alternatively at least about 94% nucleic acid sequence identity, alternatively at least about 95% nucleic acid sequence identity, alternatively at least about 96% nucleic acid sequence identity, alternatively at least about 97% nucleic acid sequence identity, alternatively at least about 98% nucleic acid sequence identity and alternatively at least about 99% nucleic acid sequence identity to (a) a DNA molecule comprising the coding sequence of a full-length PRO polypeptide cDNA as disclosed herein, the coding sequence of a PRO polypeptide lacking the signal peptide as disclosed herein, the coding sequence of an extracellular domain of a transmembrane PRO polypeptide, with or without the signal peptide, as disclosed herein or the coding sequence of any other specifically defined fragment of the full-length amino acid sequence as disclosed herein, or (b) the complement of the DNA molecule of (a).

[0011] In a further aspect, the invention concerns an isolated nucleic acid molecule comprising a nucleotide sequence having at least about 80% nucleic acid sequence identity, alternatively at least about 81% nucleic acid sequence identity, alternatively at least about 82% nucleic acid sequence identity, alternatively at least about 83% nucleic acid sequence identity, alternatively at least about 84% nucleic acid sequence identity, alternatively at least about 85% nucleic acid sequence identity, alternatively at least about 86% nucleic acid sequence identity, alternatively at least about 87% nucleic acid sequence identity, alternatively at least about 88% nucleic acid sequence identity, alternatively at least about 89% nucleic acid sequence identity, alternatively at least about 90% nucleic acid sequence identity, alternatively at least about 91% nucleic acid sequence identity, alternatively at least about 92% nucleic acid sequence identity, alternatively at least about 93% nucleic acid sequence identity, alternatively at least about 94% nucleic acid sequence identity, alternatively at least about 95% nucleic acid sequence identity, alternatively

at least about 96% nucleic acid sequence identity, alternatively at least about 97% nucleic acid sequence identity, alternatively at least about 98% nucleic acid sequence identity and alternatively at least about 99% nucleic acid sequence identity to (a) a DNA molecule that encodes the same mature polypeptide encoded by any of the human protein cDNAs deposited with the ATCC as disclosed herein, or (b) the complement of the DNA molecule of (a).

[0012] Another aspect the invention provides an isolated nucleic acid molecule comprising a nucleotide sequence encoding a PRO polypeptide which is either transmembrane domain-deleted or transmembrane domain-inactivated, or is complementary to such encoding nucleotide sequence, wherein the transmembrane domain(s) of such polypeptide are disclosed herein. Therefore, soluble extracellular domains of the herein described PRO polypeptides are contemplated.

[0013] Another embodiment is directed to fragments of a PRO polypeptide coding sequence, or the complement thereof, that may find use as, for example, hybridization probes, for encoding fragments of a PRO polypeptide that may optionally encode a polypeptide comprising a binding site for an anti-PRO antibody or as antisense oligonucleotide probes. Such nucleic acid fragments are usually at least about 20 nucleotides in length, alternatively at least about 30 nucleotides in length, alternatively at least about 40 nucleotides in length, alternatively at least about 50 nucleotides in length, alternatively at least about 60 nucleotides in length, alternatively at least about 70 nucleotides in length, alternatively at least about 80 nucleotides in length, alternatively at least about 90 nucleotides in length, alternatively at least about 100 nucleotides in length, alternatively at least about 110 nucleotides in length, alternatively at least about 120 nucleotides in length, alternatively at least about 130 nucleotides in length, alternatively at least about 140 nucleotides in length, alternatively at least about 150 nucleotides in length, alternatively at least about 160 nucleotides in length, alternatively at least about 170 nucleotides in length, alternatively at least about 180 nucleotides in length, alternatively at least about 190 nucleotides in length, alternatively at least about 200 nucleotides in length, alternatively at least about 250 nucleotides in length, alternatively at least about 300 nucleotides in length, alternatively at least about 350 nucleotides in length, alternatively at least about 400 nucleotides in length, alternatively at least about 450 nucleotides in length, alternatively at least about 500 nucleotides in length, alternatively at least about 600 nucleotides in length, alternatively at least about 700 nucleotides in length, alternatively at least about 800 nucleotides in length, alternatively at least about 900 nucleotides in length and alternatively at least about 1000 nucleotides in length, wherein in this context the term "about" means the referenced nucleotide sequence length plus or minus 10% of that referenced length. It is noted that novel fragments of a PRO polypeptide-encoding nucleotide sequence may be determined in a routine manner by aligning the PRO polypeptide-encoding nucleotide sequence with other known nucleotide sequences using any of a number of well known sequence alignment programs and determining which PRO polypeptide-encoding nucleotide sequence fragment(s) are novel. All of such PRO polypeptide-encoding nucleotide sequences are contemplated herein. Also contemplated are the PRO polypeptide fragments encoded by these nucleotide molecule frag-

ments, preferably those PRO polypeptide fragments that comprise a binding site for an anti-PRO antibody.

[0014] In another embodiment, the invention provides isolated PRO polypeptide encoded by any of the isolated nucleic acid sequences hereinabove identified.

[0015] In a certain aspect, the invention concerns an isolated PRO polypeptide, comprising an amino acid sequence having at least about 80% amino acid sequence identity, alternatively at least about 81% amino acid sequence identity, alternatively at least about 82% amino acid sequence identity, alternatively at least about 83% amino acid sequence identity, alternatively at least about 84% amino acid sequence identity, alternatively at least about 85% amino acid sequence identity, alternatively at least about 86% amino acid sequence identity, alternatively at least about 87% amino acid sequence identity, alternatively at least about 88% amino acid sequence identity, alternatively at least about 89% amino acid sequence identity, alternatively at least about 90% amino acid sequence identity, alternatively at least about 91% amino acid sequence identity, alternatively at least about 92% amino acid sequence identity, alternatively at least about 93% amino acid sequence identity, alternatively at least about 94% amino acid sequence identity, alternatively at least about 95% amino acid sequence identity, alternatively at least about 96% amino acid sequence identity, alternatively at least about 97% amino acid sequence identity, alternatively at least about 98% amino acid sequence identity and alternatively at least about 99% amino acid sequence identity to a PRO polypeptide having a full-length amino acid sequence as disclosed herein, an amino acid sequence lacking the signal peptide as disclosed herein, an extracellular domain of a transmembrane protein, with or without the peptide, as disclosed herein or any other specifically defined fragment of the full-length amino acid sequence as disclosed herein.

[0016] In a further aspect, the invention concerns an isolated PRO polypeptide comprising an amino acid sequence having at least about 80% amino acid sequence identity, alternatively at least about 81% amino acid sequence identity, alternatively at least about 82% amino acid sequence identity, alternatively at least about 83% amino acid sequence identity, alternatively at least about 84% amino acid sequence identity, alternatively at least about 85% amino acid sequence identity, alternatively at least about 86% amino acid sequence identity, alternatively at least about 87% amino acid sequence identity, alternatively at least about 88% amino acid sequence identity, alternatively at least about 89% amino acid sequence identity, alternatively at least about 90% amino acid sequence identity, alternatively at least about 91% amino acid sequence identity, alternatively at least about 92% amino acid sequence identity, alternatively at least about 93% amino acid sequence identity, alternatively at least about 94% amino acid sequence identity, alternatively at least about 95% amino acid sequence identity, alternatively at least about 96% amino acid sequence identity, alternatively at least about 97% amino acid sequence identity, alternatively at least about 98% amino acid sequence identity and alternatively at least about 99% amino acid sequence identity to an amino acid sequence encoded by any of the human protein cDNAs deposited with the ATCC as disclosed herein.

[0017] In a specific aspect, the invention provides an isolated PRO polypeptide without the N-terminal signal sequence and/or the initiating methionine and is encoded by a nucleotide sequence that encodes such an amino acid sequence as hereinbefore described. Processes for producing the same are also herein described, wherein those processes comprise culturing a host cell comprising a vector which comprises the appropriate encoding nucleic acid molecule under conditions suitable for expression of the PRO polypeptide and recovering the PRO polypeptide from the cell culture.

[0018] Another aspect the invention provides an isolated PRO polypeptide which is either transmembrane domain-deleted or transmembrane domain-inactivated. Processes for producing the same are also herein described, wherein those processes comprise culturing a host cell comprising a vector which comprises the appropriate encoding nucleic acid molecule under conditions suitable for expression of the PRO polypeptide and recovering the PRO polypeptide from the cell culture.

[0019] In yet another embodiment, the invention concerns agonists and antagonists of a native PRO polypeptide as defined herein. In a particular embodiment, the agonist or antagonist is an anti-PRO antibody or a small molecule.

[0020] In a further embodiment, the invention concerns a method of identifying agonists or antagonists to a PRO polypeptide which comprise contacting the PRO polypeptide with a candidate molecule and monitoring a biological activity mediated by said PRO polypeptide. Preferably, the PRO polypeptide is a native PRO polypeptide.

[0021] In a still further embodiment, the invention concerns a composition of matter comprising a PRO polypeptide, or an agonist or antagonist of a PRO polypeptide as herein described, or an anti-PRO antibody, in combination with a carrier. Optionally, the carrier is a pharmaceutically acceptable carrier.

[0022] Another embodiment of the present invention is directed to the use of a PRO polypeptide, or an agonist or antagonist thereof as hereinbefore described, or an anti-PRO antibody, for the preparation of a medicament useful in the treatment of a condition which is responsive to the PRO polypeptide, an agonist or antagonist thereof or an anti-PRO antibody.

[0023] In other embodiments of the present invention, the invention provides vectors comprising DNA encoding any of the herein described polypeptides. Host cell comprising any such vector are also provided. By way of example, the host cells may be CHO cells, *E. coli*, or yeast. A process for producing any of the herein described polypeptides is further provided and comprises culturing host cells under conditions suitable for expression of the desired polypeptide and recovering the desired polypeptide from the cell culture.

[0024] In other embodiments, the invention provides chimeric molecules comprising any of the herein described polypeptides fused to a heterologous polypeptide or amino acid sequence. Example of such chimeric molecules comprise any of the herein described polypeptides fused to an epitope tag sequence or a Fc region of an immunoglobulin.

[0025] In another embodiment, the invention provides an antibody which binds, preferably specifically, to any of the

above or below described polypeptides. Optionally, the antibody is a monoclonal antibody, humanized antibody, antibody fragment or single-chain antibody.

[0026] In yet other embodiments, the invention provides oligonucleotide probes useful for isolating genomic and cDNA nucleotide sequences or as antisense probes, wherein those probes may be derived from any of the above or below described nucleotide sequences.

[0027] In yet other embodiments, the present invention is directed to methods of using the PRO polypeptides of the present invention for a variety of uses based upon the functional biological assay data presented in the Examples below.

BRIEF DESCRIPTION OF DRAWINGS

[0028] FIG. 1 shows a nucleotide sequence (SEQ ID NO:1) of a native sequence PRO180 cDNA, wherein SEQ ID NO:1 is a clone designated herein as "DNA26843-1389".

[0029] FIG. 2 shows the amino acid sequence (SEQ ID NO:2) derived from the coding sequence of SEQ ID NO:1 shown in FIG. 1.

[0030] FIG. 3 shows a nucleotide sequence (SEQ ID NO:3) of a native sequence PRO218 cDNA, wherein SEQ ID NO:3 is a clone designated herein as "DNA30867-1335".

[0031] FIG. 4 shows the amino acid sequence (SEQ ID NO:4) derived from the coding sequence of SEQ ID NO:3 shown in FIG. 3.

[0032] FIG. 5 shows a nucleotide sequence (SEQ ID NO:5) of a native sequence PRO263 cDNA, wherein SEQ ID NO:5 is a clone designated herein as "DNA34431-1177".

[0033] FIG. 6 shows the amino acid sequence (SEQ ID NO:6) derived from the coding sequence of SEQ ID NO:5 shown in FIG. 5.

[0034] FIG. 7 shows a nucleotide sequence (SEQ ID NO:7) of a native sequence PRO295 cDNA, wherein SEQ ID NO:7 is a clone designated herein as "DNA38268-1188".

[0035] FIG. 8 shows the amino acid sequence (SEQ ID NO:8) derived from the coding sequence of SEQ ID NO:7 shown in FIG. 7.

[0036] FIG. 9 shows a nucleotide sequence (SEQ ID NO:9) of a native sequence PRO874 cDNA, wherein SEQ ID NO:9 is a clone designated herein as "DNA40621-1440".

[0037] FIG. 10 shows the amino acid sequence (SEQ ID NO:10) derived from the coding sequence of SEQ ID NO:9 shown in FIG. 9.

[0038] FIG. 11 shows a nucleotide sequence (SEQ ID NO:11) of a native sequence PRO300 cDNA, wherein SEQ ID NO:11 is a clone designated herein as "DNA40625-1189".

[0039] FIG. 12 shows the amino acid sequence (SEQ ID NO:12) derived from the coding sequence of SEQ ID NO:11 shown in FIG. 11.

[0040] FIG. 13 shows a nucleotide sequence (SEQ ID NO:13) of a native sequence PRO1864 cDNA, wherein SEQ ID NO:13 is a clone designated herein as "DNA45409-2511".

[0041] FIG. 14 shows the amino acid sequence (SEQ ID NO:14) derived from the coding sequence of SEQ ID NO:13 shown in FIG. 13.

[0042] FIG. 15 shows a nucleotide sequence (SEQ ID NO:15) of a native sequence PRO1282 cDNA, wherein SEQ ID NO:15 is a clone designated herein as "DNA45495-1550".

[0043] FIG. 16 shows the amino acid sequence (SEQ ID NO:16) derived from the coding sequence of SEQ ID NO:15 shown in FIG. 15.

[0044] FIG. 17 shows a nucleotide sequence (SEQ ID NO:17) of a native sequence PRO1063 cDNA, wherein SEQ ID NO:17 is a clone designated herein as "DNA49820-1427".

[0045] FIG. 18 shows the amino acid sequence (SEQ ID NO:18) derived from the coding sequence of SEQ ID NO:17 shown in FIG. 17.

[0046] FIG. 19 shows a nucleotide sequence (SEQ ID NO:19) of a native sequence PRO1773 cDNA, wherein SEQ ID NO:19 is a clone designated herein as "DNA56406-1704".

[0047] FIG. 20 shows the amino acid sequence (SEQ ID NO:20) derived from the coding sequence of SEQ ID NO:19 shown in FIG. 19.

[0048] FIG. 21 shows a nucleotide sequence (SEQ ID NO:21) of a native sequence PRO1013 cDNA, wherein SEQ ID NO:21 is a clone designated herein as "DNA56410-1414".

[0049] FIG. 22 shows the amino acid sequence (SEQ ID NO:22) derived from the coding sequence of SEQ ID NO:21 shown in FIG. 21.

[0050] FIG. 23 shows a nucleotide sequence (SEQ ID NO:23) of a native sequence PRO937 cDNA, wherein SEQ ID NO:23 is a clone designated herein as "DNA56436-1448".

[0051] FIG. 24 shows the amino acid sequence (SEQ ID NO:24) derived from the coding sequence of SEQ ID NO:23 shown in FIG. 23.

[0052] FIG. 25 shows a nucleotide sequence (SEQ ID NO:25) of a native sequence PRO842 cDNA, wherein SEQ ID NO:25 is a clone designated herein as "DNA56855-1447".

[0053] FIG. 26 shows the amino acid sequence (SEQ ID NO:26) derived from the coding sequence of SEQ ID NO:25 shown in FIG. 25.

[0054] FIG. 27 shows a nucleotide sequence (SEQ ID NO:27) of a native sequence PRO1180 cDNA, wherein SEQ ID NO:27 is a clone designated herein as "DNA56860-1510".

[0055] FIG. 28 shows the amino acid sequence (SEQ ID NO:28) derived from the coding sequence of SEQ ID NO:27 shown in FIG. 27.

[0056] FIG. 29 shows a nucleotide sequence (SEQ ID NO:29) of a native sequence PRO831 cDNA, wherein SEQ ID NO:29 is a clone designated herein as "DNA56862-1343".

[0057] FIG. 30 shows the amino acid sequence (SEQ ID NO:30) derived from the coding sequence of SEQ ID NO:29 shown in FIG. 29.

[0058] FIG. 31 shows a nucleotide sequence (SEQ ID NO:31) of a native sequence PRO1115 cDNA, wherein SEQ ID NO:31 is a clone designated herein as "DNA56868-1478".

[0059] FIG. 32 shows the amino acid sequence (SEQ ID NO:32) derived from the coding sequence of SEQ ID NO:31 shown in FIG. 31.

[0060] FIG. 33 shows a nucleotide sequence (SEQ ID NO:33) of a native sequence PRO1277 cDNA, wherein SEQ ID NO:33 is a clone designated herein as "DNA56869-1545".

[0061] FIG. 34 shows the amino acid sequence (SEQ ID NO:34) derived from the coding sequence of SEQ ID NO:33 shown in FIG. 33.

[0062] FIG. 35 shows a nucleotide sequence (SEQ ID NO:35) of a native sequence PRO1074 cDNA, wherein SEQ ID NO:35 is a clone designated herein as "DNA57704-1452".

[0063] FIG. 36 shows the amino acid sequence (SEQ ID NO:36) derived from the coding sequence of SEQ ID NO:35 shown in FIG. 35.

[0064] FIG. 37 shows a nucleotide sequence (SEQ ID NO:37) of a native sequence PRO1344 cDNA, wherein SEQ ID NO:37 is a clone designated herein as "DNA58723-1588".

[0065] FIG. 38 shows the amino acid sequence (SEQ ID NO:38) derived from the coding sequence of SEQ ID NO:37 shown in FIG. 37.

[0066] FIG. 39 shows a nucleotide sequence (SEQ ID NO:39) of a native sequence PRO1136 cDNA, wherein SEQ ID NO:39 is a clone designated herein as "DNA57827-1493".

[0067] FIG. 40 shows the amino acid sequence (SEQ ID NO:40) derived from the coding sequence of SEQ ID NO:39 shown in FIG. 39.

[0068] FIG. 41 shows a nucleotide sequence (SEQ ID NO:41) of a native sequence PRO1109 cDNA, wherein SEQ ID NO:41 is a clone designated herein as "DNA58737-1473".

[0069] FIG. 42 shows the amino acid sequence (SEQ ID NO:42) derived from the coding sequence of SEQ ID NO:41 shown in FIG. 41.

[0070] FIG. 43 shows a nucleotide sequence (SEQ ID NO:43) of a native sequence PRO1003 cDNA, wherein SEQ ID NO:43 is a clone designated herein as "DNA58846-1409".

[0071] FIG. 44 shows the amino acid sequence (SEQ ID NO:44) derived from the coding sequence of SEQ ID NO:43 shown in FIG. 43.

[0072] FIG. 45 shows a nucleotide sequence (SEQ ID NO:45) of a native sequence PRO1138 cDNA, wherein SEQ ID NO:45 is a clone designated herein as "DNA58850-1495".

[0073] FIG. 46 shows the amino acid sequence (SEQ ID NO:46) derived from the coding sequence of SEQ ID NO:45 shown in FIG. 45.

[0074] FIG. 47 shows a nucleotide sequence (SEQ ID NO:47) of a native sequence PRO994 cDNA, wherein SEQ ID NO:47 is a clone designated herein as "DNA58855-1422".

[0075] FIG. 48 shows the amino acid sequence (SEQ ID NO:48) derived from the coding sequence of SEQ ID NO:47 shown in FIG. 47.

[0076] FIG. 49 shows a nucleotide sequence (SEQ ID NO:49) of a native sequence PRO1069 cDNA, wherein SEQ ID NO:49 is a clone designated herein as "DNA59211-1450".

[0077] FIG. 50 shows the amino acid sequence (SEQ ID NO:50) derived from the coding sequence of SEQ ID NO:49 shown in FIG. 49.

[0078] FIG. 51 shows a nucleotide sequence (SEQ ID NO:51) of a native sequence PRO1411 cDNA, wherein SEQ ID NO:51 is a clone designated herein as "DNA59212-1627".

[0079] FIG. 52 shows the amino acid sequence (SEQ ID NO:52) derived from the coding sequence of SEQ ID NO:51 shown in FIG. 51.

[0080] FIG. 53 shows a nucleotide sequence (SEQ ID NO:53) of a native sequence PRO1129 cDNA, wherein SEQ ID NO:53 is a clone designated herein as "DNA59213-1487".

[0081] FIG. 54 shows the amino acid sequence (SEQ ID NO:54) derived from the coding sequence of SEQ ID NO:53 shown in FIG. 53.

[0082] FIG. 55 shows a nucleotide sequence (SEQ ID NO:55) of a native sequence PRO1027 cDNA, wherein SEQ ID NO:55 is a clone designated herein as "DNA59605-1418".

[0083] FIG. 56 shows the amino acid sequence (SEQ ID NO:56) derived from the coding sequence of SEQ ID NO:55 shown in FIG. 55.

[0084] FIG. 57 shows a nucleotide sequence (SEQ ID NO:57) of a native sequence PRO1106 cDNA, wherein SEQ ID NO:57 is a clone designated herein as "DNA59609-1470".

[0085] FIG. 58 shows the amino acid sequence (SEQ ID NO:58) derived from the coding sequence of SEQ ID NO:57 shown in FIG. 57.

[0086] FIG. 59 shows a nucleotide sequence (SEQ ID NO:59) of a native sequence PRO1291 cDNA, wherein SEQ ID NO:59 is a clone designated herein as "DNA59610-1556".

[0087] FIG. 60 shows the amino acid sequence (SEQ ID NO:60) derived from the coding sequence of SEQ ID NO:59 shown in FIG. 59.

[0088] FIG. 61 shows a nucleotide sequence (SEQ ID NO:61) of a native sequence PRO3573 cDNA, wherein SEQ ID NO:61 is a clone designated herein as "DNA59837-2545".

[0089] FIG. 62 shows the amino acid sequence (SEQ ID NO:62) derived from the coding sequence of SEQ ID NO:61 shown in FIG. 61.

[0090] FIG. 63 shows a nucleotide sequence (SEQ ID NO:63) of a native sequence PRO3566 cDNA, wherein SEQ ID NO:63 is a clone designated herein as "DNA59844-2542".

[0091] FIG. 64 shows the amino acid sequence (SEQ ID NO:64) derived from the coding sequence of SEQ ID NO:63 shown in FIG. 63.

[0092] FIG. 65 shows a nucleotide sequence (SEQ ID NO:65) of a native sequence PRO1098 cDNA, wherein SEQ ID NO:65 is a clone designated herein as "DNA59854-1459".

[0093] FIG. 66 shows the amino acid sequence (SEQ ID NO:66) derived from the coding sequence of SEQ ID NO:65 shown in FIG. 65.

[0094] FIG. 67 shows a nucleotide sequence (SEQ ID NO:67) of a native sequence PRO1158 cDNA, wherein SEQ ID NO:67 is a clone designated herein as "DNA60625-1507".

[0095] FIG. 68 shows the amino acid sequence (SEQ ID NO:68) derived from the coding sequence of SEQ ID NO:67 shown in FIG. 67.

[0096] FIG. 69 shows a nucleotide sequence (SEQ ID NO:69) of a native sequence PRO1124 cDNA, wherein SEQ ID NO:69 is a clone designated herein as "DNA60629-1481".

[0097] FIG. 70 shows the amino acid sequence (SEQ ID NO:70) derived from the coding sequence of SEQ ID NO:69 shown in FIG. 69.

[0098] FIG. 71 shows a nucleotide sequence (SEQ ID NO:71) of a native sequence PRO1287 cDNA, wherein SEQ ID NO:71 is a clone designated herein as "DNA61755-1554".

[0099] FIG. 72 shows the amino acid sequence (SEQ ID NO:72) derived from the coding sequence of SEQ ID NO:71 shown in FIG. 71.

[0100] FIG. 73 shows a nucleotide sequence (SEQ ID NO:73) of a native sequence PRO1335 cDNA, wherein SEQ ID NO:73 is a clone designated herein as "DNA62812-1594".

[0101] FIG. 74 shows the amino acid sequence (SEQ ID NO:74) derived from the coding sequence of SEQ ID NO:73 shown in FIG. 73.

[0102] FIG. 75 shows a nucleotide sequence (SEQ ID NO:75) of a native sequence PRO1315 cDNA, wherein SEQ ID NO:75 is a clone designated herein as "DNA62815-1576".

[0103] FIG. 76 shows the amino acid sequence (SEQ ID NO:76) derived from the coding sequence of SEQ ID NO:75 shown in FIG. 75.

[0104] FIG. 77 shows a nucleotide sequence (SEQ ID NO:77) of a native sequence PRO1357 cDNA, wherein SEQ ID NO:77 is a clone designated herein as "DNA64881-1602".

[0105] FIG. 78 shows the amino acid sequence (SEQ ID NO:78) derived from the coding sequence of SEQ ID NO:77 shown in FIG. 77.

[0106] FIG. 79 shows a nucleotide sequence (SEQ ID NO:79) of a native sequence PRO1356 cDNA, wherein SEQ ID NO:79 is a clone designated herein as "DNA64886-1601".

[0107] FIG. 80 shows the amino acid sequence (SEQ ID NO:80) derived from the coding sequence of SEQ ID NO:79 shown in FIG. 79.

[0108] FIG. 81 shows a nucleotide sequence (SEQ ID NO:81) of a native sequence PRO1557 cDNA, wherein SEQ ID NO:81 is a clone designated herein as "DNA64902-1667".

[0109] FIG. 82 shows the amino acid sequence (SEQ ID NO:82) derived from the coding sequence of SEQ ID NO:81 shown in FIG. 81.

[0110] FIG. 83 shows a nucleotide sequence (SEQ ID NO:83) of a native sequence PRO1347 cDNA, wherein SEQ ID NO:83 is a clone designated herein as "DNA64950-1590".

[0111] FIG. 84 shows the amino acid sequence (SEQ ID NO:84) derived from the coding sequence of SEQ ID NO:83 shown in FIG. 83.

[0112] FIG. 85 shows a nucleotide sequence (SEQ ID NO:85) of a native sequence PRO1302 cDNA, wherein SEQ ID NO:85 is a clone designated herein as "DNA65403-1565".

[0113] FIG. 86 shows the amino acid sequence (SEQ ID NO:86) derived from the coding sequence of SEQ ID NO:85 shown in FIG. 85.

[0114] FIG. 87 shows a nucleotide sequence (SEQ ID NO:87) of a native sequence PRO1270 cDNA, wherein SEQ ID NO:87 is a clone designated herein as "DNA66308-1537".

[0115] FIG. 88 shows the amino acid sequence (SEQ ID NO:88) derived from the coding sequence of SEQ ID NO:87 shown in FIG. 87.

[0116] FIG. 89 shows a nucleotide sequence (SEQ ID NO:89) of a native sequence PRO1268 cDNA, wherein SEQ ID NO:89 is a clone designated herein as "DNA66519-1535".

[0117] FIG. 90 shows the amino acid sequence (SEQ ID NO:90) derived from the coding sequence of SEQ ID NO:89 shown in FIG. 89.

[0118] FIG. 91 shows a nucleotide sequence (SEQ ID NO:91) of a native sequence PRO1327 cDNA, wherein SEQ ID NO:91 is a clone designated herein as "DNA66521-1583".

[0119] FIG. 92 shows the amino acid sequence (SEQ ID NO:92) derived from the coding sequence of SEQ ID NO:91 shown in FIG. 91.

[0120] FIG. 93 shows a nucleotide sequence (SEQ ID NO:93) of a native sequence PRO1328 cDNA, wherein SEQ ID NO:93 is a clone designated herein as "DNA66658-1584".

[0121] FIG. 94 shows the amino acid sequence (SEQ ID NO:94) derived from the coding sequence of SEQ ID NO:93 shown in FIG. 93.

[0122] FIG. 95 shows a nucleotide sequence (SEQ ID NO:95) of a native sequence PRO1329 cDNA, wherein SEQ ID NO:95 is a clone designated herein as "DNA66660-1585".

[0123] FIG. 96 shows the amino acid sequence (SEQ ID NO:96) derived from the coding sequence of SEQ ID NO:95 shown in FIG. 95.

[0124] FIG. 97 shows a nucleotide sequence (SEQ ID NO:97) of a native sequence PRO1340 cDNA, wherein SEQ ID NO:97 is a clone designated herein as "DNA66663-1598".

[0125] FIG. 98 shows the amino acid sequence (SEQ ID NO:98) derived from the coding sequence of SEQ ID NO:97 shown in FIG. 97.

[0126] FIG. 99 shows a nucleotide sequence (SEQ ID NO:99) of a native sequence PRO1342 cDNA, wherein SEQ ID NO:99 is a clone designated herein as "DNA66674-1599".

[0127] FIG. 100 shows the amino acid sequence (SEQ ID NO:100) derived from the coding sequence of SEQ ID NO:99 shown in FIG. 99.

[0128] FIG. 101 shows a nucleotide sequence (SEQ ID NO:101) of a native sequence PRO3579 cDNA, wherein SEQ ID NO:101 is a clone designated herein as "DNA68862-2546".

[0129] FIG. 102 shows the amino acid sequence (SEQ ID NO:102) derived from the coding sequence of SEQ ID NO:101 shown in FIG. 101.

[0130] FIG. 103 shows a nucleotide sequence (SEQ ID NO:103) of a native sequence PRO1472 cDNA, wherein SEQ ID NO:103 is a clone designated herein as "DNA68866-1644".

[0131] FIG. 104 shows the amino acid sequence (SEQ ID NO:104) derived from the coding sequence of SEQ ID NO:103 shown in FIG. 103.

[0132] FIG. 105 shows a nucleotide sequence (SEQ ID NO:105) of a native sequence PRO1461 cDNA, wherein SEQ ID NO:105 is a clone designated herein as "DNA68871-1638".

[0133] FIG. 106 shows the amino acid sequence (SEQ ID NO:106) derived from the coding sequence of SEQ ID NO:105 shown in FIG. 105.

[0134] FIG. 107 shows a nucleotide sequence (SEQ ID NO:107) of a native sequence PRO1568 cDNA, wherein SEQ ID NO:107 is a clone designated herein as "DNA68880-1676".

[0135] FIG. 108 shows the amino acid sequence (SEQ ID NO:108) derived from the coding sequence of SEQ ID NO:107 shown in FIG. 107.

[0136] FIG. 109 shows a nucleotide sequence (SEQ ID NO:109) of a native sequence PRO1753 cDNA, wherein SEQ ID NO:109 is a clone designated herein as "DNA68883-1691".

[0137] FIG. 110 shows the amino acid sequence (SEQ ID NO:110) derived from the coding sequence of SEQ ID NO:109 shown in FIG. 109.

[0138] FIG. 111 shows a nucleotide sequence (SEQ ID NO:111) of a native sequence PRO1570 cDNA, wherein SEQ ID NO:111 is a clone designated herein as "DNA68885-1678".

[0139] FIG. 112 shows the amino acid sequence (SEQ ID NO:112) derived from the coding sequence of SEQ ID NO:111 shown in FIG. 111.

[0140] FIG. 113 shows a nucleotide sequence (SEQ ID NO:113) of a native sequence PRO1446 cDNA, wherein SEQ ID NO:113 is a clone designated herein as "DNA71277-1636".

[0141] FIG. 114 shows the amino acid sequence (SEQ ID NO:114) derived from the coding sequence of SEQ ID NO:113 shown in FIG. 113.

[0142] FIG. 115 shows a nucleotide sequence (SEQ ID NO:115) of a native sequence PRO1565 cDNA, wherein SEQ ID NO:115 is a clone designated herein as "DNA73727-1673".

[0143] FIG. 116 shows the amino acid sequence (SEQ ID NO:116) derived from the coding sequence of SEQ ID NO:115 shown in FIG. 115.

[0144] FIG. 117 shows a nucleotide sequence (SEQ ID NO:117) of a native sequence PRO1572 cDNA, wherein SEQ ID NO:117 is a clone designated herein as "DNA73734-1680".

[0145] FIG. 118 shows the amino acid sequence (SEQ ID NO:118) derived from the coding sequence of SEQ ID NO:117 shown in FIG. 117.

[0146] FIG. 119 shows a nucleotide sequence (SEQ ID NO:119) of a native sequence PRO1573 cDNA, wherein SEQ ID NO:119 is a clone designated herein as "DNA73735-1681".

[0147] FIG. 120 shows the amino acid sequence (SEQ ID NO:120) derived from the coding sequence of SEQ ID NO:119 shown in FIG. 119.

[0148] FIG. 121 shows a nucleotide sequence (SEQ ID NO:121) of a native sequence PRO1550 cDNA, wherein SEQ ID NO:121 is a clone designated herein as "DNA76393-1664".

[0149] FIG. 122 shows the amino acid sequence (SEQ ID NO:122) derived from the coding sequence of SEQ ID NO:121 shown in FIG. 121.

[0150] FIG. 123 shows a nucleotide sequence (SEQ ID NO:123) of a native sequence PRO1693 cDNA, wherein SEQ ID NO:123 is a clone designated herein as "DNA77301-1708".

[0151] FIG. 124 shows the amino acid sequence (SEQ ID NO:124) derived from the coding sequence of SEQ ID NO:123 shown in FIG. 123.

[0152] FIG. 125 shows a nucleotide sequence (SEQ ID NO:125) of a native sequence PRO1566 cDNA, wherein SEQ ID NO:125 is a clone designated herein as "DNA77568-1626".

[0153] FIG. 126 shows the amino acid sequence (SEQ ID NO:126) derived from the coding sequence of SEQ ID NO:125 shown in FIG. 125.

[0154] FIG. 127 shows a nucleotide sequence (SEQ ID NO:127) of a native sequence PRO1774 cDNA, wherein SEQ ID NO:127 is a clone designated herein as "DNA77626-1705".

[0155] FIG. 128 shows the amino acid sequence (SEQ ID NO:128) derived from the coding sequence of SEQ ID NO:127 shown in FIG. 127.

[0156] FIG. 129 shows a nucleotide sequence (SEQ ID NO:129) of a native sequence PRO1928 cDNA, wherein SEQ ID NO:129 is a clone designated herein as "DNA81754-2532".

[0157] FIG. 130 shows the amino acid sequence (SEQ ID NO:130) derived from the coding sequence of SEQ ID NO:129 shown in FIG. 129.

[0158] FIG. 131 shows a nucleotide sequence (SEQ ID NO:131) of a native sequence PRO1865 cDNA, wherein SEQ ID NO:131 is a clone designated herein as "DNA81757-2512".

[0159] FIG. 132 shows the amino acid sequence (SEQ ID NO:132) derived from the coding sequence of SEQ ID NO:131 shown in FIG. 131.

[0160] FIG. 133 shows a nucleotide sequence (SEQ ID NO:133) of a native sequence PRO1925 cDNA, wherein SEQ ID NO:133 is a clone designated herein as "DNA82302-2529".

[0161] FIG. 134 shows the amino acid sequence (SEQ ID NO:134) derived from the coding sequence of SEQ ID NO:133 shown in FIG. 133.

[0162] FIG. 135 shows a nucleotide sequence (SEQ ID NO:135) of a native sequence PRO1926 cDNA, wherein SEQ ID NO:135 is a clone designated herein as "DNA82340-2530".

[0163] FIG. 136 shows the amino acid sequence (SEQ ID NO:136) derived from the coding sequence of SEQ ID NO:135 shown in FIG. 135.

[0164] FIG. 137 shows a nucleotide sequence (SEQ ID NO:137) of a native sequence PRO1801 cDNA, wherein SEQ ID NO:137 is a clone designated herein as "DNA83500-2506".

[0165] FIG. 138 shows the amino acid sequence (SEQ ID NO:138) derived from the coding sequence of SEQ ID NO:137 shown in FIG. 137.

[0166] FIG. 139 shows a nucleotide sequence (SEQ ID NO:139) of a native sequence PRO4405 cDNA, wherein SEQ ID NO:139 is a clone designated herein as "DNA84920-2614".

[0167] FIG. 140 shows the amino acid sequence (SEQ ID NO:140) derived from the coding sequence of SEQ ID NO:139 shown in FIG. 139.

[0168] FIG. 141 shows a nucleotide sequence (SEQ ID NO:141) of a native sequence PRO3435 cDNA, wherein SEQ ID NO:141 is a clone designated herein as "DNA85066-2534".

[0169] FIG. 142 shows the amino acid sequence (SEQ ID NO:142) derived from the coding sequence of SEQ ID NO:141 shown in FIG. 141.

[0170] FIG. 143 shows a nucleotide sequence (SEQ ID NO:143) of a native sequence PRO3543 cDNA, wherein SEQ ID NO:143 is a clone designated herein as "DNA86571-2551".

[0171] FIG. 144 shows the amino acid sequence (SEQ ID NO:144) derived from the coding sequence of SEQ ID NO:143 shown in FIG. 143.

[0172] FIG. 145 shows a nucleotide sequence (SEQ ID NO:145) of a native sequence PRO3443 cDNA, wherein SEQ ID NO:145 is a clone designated herein as "DNA87991-2540".

[0173] FIG. 146 shows the amino acid sequence (SEQ ID NO:146) derived from the coding sequence of SEQ ID NO:145 shown in FIG. 145.

[0174] FIG. 147 shows a nucleotide sequence (SEQ ID NO:147) of a native sequence PRO3442 cDNA, wherein SEQ ID NO:147 is a clone designated herein as "DNA92238-2539".

[0175] FIG. 148 shows the amino acid sequence (SEQ ID NO:148) derived from the coding sequence of SEQ ID NO:147 shown in FIG. 147.

[0176] FIG. 149 shows a nucleotide sequence (SEQ ID NO:149) of a native sequence PRO5990 cDNA, wherein SEQ ID NO:149 is a clone designated herein as "DNA96042-2682".

[0177] FIG. 150 shows the amino acid sequence (SEQ ID NO:150) derived from the coding sequence of SEQ ID NO:149 shown in FIG. 149.

[0178] FIG. 151 shows a nucleotide sequence (SEQ ID NO:151) of a native sequence PRO4342 cDNA, wherein SEQ ID NO:151 is a clone designated herein as "DNA96787-2534".

[0179] FIG. 152 shows the amino acid sequence (SEQ ID NO:152) derived from the coding sequence of SEQ ID NO:151 shown in FIG. 151.

[0180] FIG. 153 shows a nucleotide sequence (SEQ ID NO:153) of a native sequence PRO10096 cDNA, wherein SEQ ID NO:153 is a clone designated herein as "DNA125185-2806".

[0181] FIG. 154 shows the amino acid sequence (SEQ ID NO:154) derived from the coding sequence of SEQ ID NO:153 shown in FIG. 153.

[0182] FIG. 155 shows a nucleotide sequence (SEQ ID NO:155) of a native sequence PRO10272 cDNA, wherein SEQ ID NO:155 is a clone designated herein as "DNA147531-2821".

[0183] FIG. 156 shows the amino acid sequence (SEQ ID NO:156) derived from the coding sequence of SEQ ID NO:155 shown in FIG. 155.

[0184] FIG. 157 shows a nucleotide sequence (SEQ ID NO:157) of a native sequence PRO5801 cDNA, wherein SEQ ID NO:157 is a clone designated herein as "DNA115291-2681".

[0185] FIG. 158 shows the amino acid sequence (SEQ ID NO:158) derived from the coding sequence of SEQ ID NO:157 shown in FIG. 157.

[0186] FIG. 159 shows a nucleotide sequence (SEQ ID NO:159) of a native sequence PRO20110 cDNA, wherein SEQ ID NO:159 is a clone designated herein as "DNA166819".

[0187] FIG. 160 shows the amino acid sequence (SEQ ID NO:160) derived from the coding sequence of SEQ ID NO:159 shown in FIG. 159.

[0188] FIG. 161 shows a nucleotide sequence (SEQ ID NO:161) of a native sequence PRO20040 cDNA, wherein SEQ ID NO:161 is a clone designated herein as "DNA164625-2890".

[0189] FIG. 162 shows the amino acid sequence (SEQ ID NO:162) derived from the coding sequence of SEQ ID NO:161 shown in FIG. 161.

[0190] FIG. 163 shows a nucleotide sequence (SEQ ID NO:163) of a native sequence PRO20233 cDNA, wherein SEQ ID NO:163 is a clone designated herein as "DNA165608".

[0191] FIG. 164 shows the amino acid sequence (SEQ ID NO:164) derived from the coding sequence of SEQ ID NO:163 shown in FIG. 163.

[0192] FIG. 165 shows a nucleotide sequence (SEQ ID NO:165) of a native sequence PRO19670 cDNA, wherein SEQ ID NO:165 is a clone designated herein as "DNA131639-2874".

[0193] FIG. 166 shows the amino acid sequence (SEQ ID NO:166) derived from the coding sequence of SEQ ID NO:165 shown in FIG. 165.

[0194] FIG. 167 shows a nucleotide sequence (SEQ ID NO:167) of a native sequence PRO1890 cDNA, wherein SEQ ID NO:167 is a clone designated herein as "DNA79230-2525".

[0195] FIG. 168 shows the amino acid sequence (SEQ ID NO:168) derived from the coding sequence of SEQ ID NO:167 shown in FIG. 167.

DETAILED DESCRIPTION

1. Definitions

[0196] The terms "PRO polypeptide" and "PRO" as used herein and when immediately followed by a numerical designation refer to various polypeptides, wherein the complete designation (i.e., PRO/number) refers to specific polypeptide sequences as described herein. The terms "PRO/number polypeptide" and "PRO/number" wherein the term "number" is provided as an actual numerical designation as used herein encompass native sequence polypeptides and polypeptide variants (which are further defined herein). The PRO polypeptides described herein may be isolated from a variety of sources, such as from human tissue types or from another source, or prepared by recombinant or synthetic methods. The term "PRO polypeptide" refers to each individual PRO/number polypeptide disclosed herein. All disclosures in this specification which refer to the "PRO polypeptide" refer to each of the polypeptides individually as well as jointly. For example, descriptions of the prepa-

ration of, purification of, derivation of, formation of antibodies to or against, administration of, compositions containing, treatment of a disease with, etc., pertain to each polypeptide of the invention individually. The term "PRO polypeptide" also includes variants of the PRO/number polypeptides disclosed herein.

[0197] A "native sequence PRO polypeptide" comprises a polypeptide having the same amino acid sequence as the corresponding PRO polypeptide derived from nature. Such native sequence PRO polypeptides can be isolated from nature or can be produced by recombinant or synthetic means. The term "native sequence PRO polypeptide" specifically encompasses naturally-occurring truncated or secreted forms of the specific PRO polypeptide (e.g., an extracellular domain sequence), naturally-occurring variant forms (e.g., alternatively spliced forms) and naturally-occurring allelic variants of the polypeptide. In various embodiments of the invention, the native sequence PRO polypeptides disclosed herein are mature or full-length native sequence polypeptides comprising the full-length amino acids sequences shown in the accompanying figures. Start and stop codons are shown in bold font and underlined in the figures. However, while the PRO polypeptide disclosed in the accompanying figures are shown to begin with methionine residues designated herein as amino acid position 1 in the figures, it is conceivable and possible that other methionine residues located either upstream or downstream from the amino acid position 1 in the figures may be employed as the starting amino acid residue for the PRO polypeptides.

[0198] The PRO polypeptide "extracellular domain" or "ECD" refers to a form of the PRO polypeptide which is essentially free of the transmembrane and cytoplasmic domains. Ordinarily, a PRO polypeptide ECD will have less than 1% of such transmembrane and/or cytoplasmic domains and preferably, will have less than 0.5% of such domains. It will be understood that any transmembrane domains identified for the PRO polypeptides of the present invention are identified pursuant to criteria routinely employed in the art for identifying that type of hydrophobic domain. The exact boundaries of a transmembrane domain may vary but most likely by no more than about 5 amino acids at either end of the domain as initially identified herein. Optionally, therefore, an extracellular domain of a PRO polypeptide may contain from about 5 or fewer amino acids on either side of the transmembrane domain/extracellular domain boundary as identified in the Examples or specification and such polypeptides, with or without the associated signal peptide, and nucleic acid encoding them, are contemplated by the present invention.

[0199] The approximate location of the "signal peptides" of the various PRO polypeptides disclosed herein are shown in the present specification and/or the accompanying figures. It is noted, however, that the C-terminal boundary of a signal peptide may vary, but most likely by no more than about 5 amino acids on either side of the signal peptide C-terminal boundary as initially identified herein, wherein the C-terminal boundary of the signal peptide may be identified pursuant to criteria routinely employed in the art for identifying that type of amino acid sequence element (e.g., Nielsen et al., *Prot. Eng.* 10:1-6 (1997) and von Heinje et al., *Nucl. Acids. Res.* 14:4683-4690 (1986)). Moreover, it is also recognized that, in some cases, cleavage of a signal

sequence from a secreted polypeptide is not entirely uniform, resulting in more than one secreted species. These mature polypeptides, where the signal peptide is cleaved within no more than about 5 amino acids on either side of the C-terminal boundary of the signal peptide as identified herein, and the polynucleotides encoding them, are contemplated by the present invention.

[0200] "PRO polypeptide variant" means an active PRO polypeptide as defined above or below having at least about 80% amino acid sequence identity with a full-length native sequence PRO polypeptide sequence as disclosed herein, a PRO polypeptide sequence lacking the signal peptide as disclosed herein, an extracellular domain of a PRO polypeptide, with or without the signal peptide, as disclosed herein or any other fragment of a full-length PRO polypeptide sequence as disclosed herein. Such PRO polypeptide variants include, for instance, PRO polypeptides wherein one or more amino acid residues are added, or deleted, at the N- or C-terminus of the full-length native amino acid sequence. Ordinarily, a PRO polypeptide variant will have at least about 80% amino acid sequence identity, alternatively at least about 81% amino acid sequence identity, alternatively at least about 82% amino acid sequence identity, alternatively at least about 83% amino acid sequence identity, alternatively at least about 84% amino acid sequence identity, alternatively at least about 85% amino acid sequence identity, alternatively at least about 86% amino acid sequence identity, alternatively at least about 87% amino acid sequence identity, alternatively at least about 88% amino acid sequence identity, alternatively at least about 89% amino acid sequence identity, alternatively at least about 90% amino acid sequence identity, alternatively at least about 91% amino acid sequence identity, alternatively at least about 92% amino acid sequence identity, alternatively at least about 93% amino acid sequence identity, alternatively at least about 94% amino acid sequence identity, alternatively at least about 95% amino acid sequence identity, alternatively at least about 96% amino acid sequence identity, alternatively at least about 97% amino acid sequence identity, alternatively at least about 98% amino acid sequence identity and alternatively at least about 99% amino acid sequence identity to a full-length native sequence PRO polypeptide sequence as disclosed herein, a PRO polypeptide sequence lacking the signal peptide as disclosed herein, an extracellular domain of a PRO polypeptide, with or without the signal peptide, as disclosed herein or any other specifically defined fragment of a full-length PRO polypeptide sequence as disclosed herein. Ordinarily, PRO variant polypeptides are at least about 10 amino acids in length, alternatively at least about 20 amino acids in length, alternatively at least about 30 amino acids in length, alternatively at least about 40 amino acids in length, alternatively at least about 50 amino acids in length, alternatively at least about 60 amino acids in length, alternatively at least about 70 amino acids in length, alternatively at least about 80 amino acids in length, alternatively at least about 90 amino acids in length, alternatively at least about 100 amino acids in length, alternatively at least about 150 amino acids in length, alternatively at least about 200 amino acids in length, alternatively at least about 300 amino acids in length, or more.

[0201] "Percent (%) amino acid sequence identity" with respect to the PRO polypeptide sequences identified herein is defined as the percentage of amino acid residues in a

candidate sequence that are identical with the amino acid residues in the specific PRO polypeptide sequence, after aligning the sequences and introducing gaps, if necessary, to achieve the maximum percent sequence identity, and not considering any conservative substitutions as part of the sequence identity. Alignment for purposes of determining percent amino acid sequence identity can be achieved in various ways that are within the skill in the art, for instance, using publicly available computer software such as BLAST, BLAST-2, ALIGN or Megalign (DNASTAR) software. Those skilled in the art can determine appropriate parameters for measuring alignment, including any algorithms needed to achieve maximal alignment over the full length of the sequences being compared. For purposes herein, however, % amino acid sequence identity values are generated using the sequence comparison computer program ALIGN-2, wherein the complete source code for the ALIGN-2 program is provided in Table 1 below. The ALIGN-2 sequence comparison computer program was authored by Genentech, Inc. and the source code shown in Table 1 below has been filed with user documentation in the U.S. Copyright Office, Washington D.C., 20559, where it is registered under U.S. Copyright Registration No. TXU510087. The ALIGN-2 program is publicly available through Genentech, Inc., South San Francisco, Calif. or may be compiled from the source code provided in Table 1 below. The ALIGN-2 program should be compiled for use on a UNIX operating system, preferably digital UNIX V4.0D. All sequence comparison parameters are set by the ALIGN-2 program and do not vary.

[0202] In situations where ALIGN-2 is employed for amino acid sequence comparisons, the % amino acid sequence identity of a given amino acid sequence A to, with, or against a given amino acid sequence B (which can alternatively be phrased as a given amino acid sequence A that has or comprises a certain % amino acid sequence identity to, with, or against a given amino acid sequence B) is calculated as follows:

100 times the fraction X/Y

[0203] where X is the number of amino acid residues scored as identical matches by the sequence alignment program ALIGN-2 in that program's alignment of A and B, and where Y is the total number of amino acid residues in B. It will be appreciated that where the length of amino acid sequence A is not equal to the length of amino acid sequence B, the % amino acid sequence identity of A to B will not equal the % amino acid sequence identity of B to A. As examples of % amino acid sequence identity calculations using this method, Tables 2 and 3 demonstrate how to calculate the % amino acid sequence identity of the amino acid sequence designated "Comparison Protein" to the amino acid sequence designated "PRO", wherein "PRO" represents the amino acid sequence of a hypothetical PRO polypeptide of interest, "Comparison Protein" represents the amino acid sequence of a polypeptide against which the "PRO" polypeptide of interest is being compared, and "X", "Y" and "Z" each represent different hypothetical amino acid residues.

[0204] Unless specifically stated otherwise, all % amino acid sequence identity values used herein are obtained as described in the immediately preceding paragraph using the ALIGN-2 computer program. However, % amino acid sequence identity values may also be obtained as described

below by using the WU-BLAST-2 computer program (Altschul et al., *Methods in Enzymology* 266:460-480 (1996)). Most of the WU-BLAST-2 search parameters are set to the default values. Those not set to default values, i.e., the adjustable parameters, are set with the following values: overlap span=1, overlap fraction=0.125, word threshold (T)=11, and scoring matrix=BLOSUM62. When WU-BLAST-2 is employed, a % amino acid sequence identity value is determined by dividing (a) the number of matching identical amino acid residues between the amino acid sequence of the PRO polypeptide of interest having a sequence derived from the native PRO polypeptide and the comparison amino acid sequence of interest (i.e., the sequence against which the PRO polypeptide of interest is being compared which may be a PRO variant polypeptide) as determined by WU-BLAST-2 by (b) the total number of amino acid residues of the PRO polypeptide of interest. For example, in the statement "a polypeptide comprising an the amino acid sequence A which has or having at least 80% amino acid sequence identity to the amino acid sequence B", the amino acid sequence A is the comparison amino acid sequence of interest and the amino acid sequence B is the amino acid sequence of the PRO polypeptide of interest.

[0205] Percent amino acid sequence identity may also be determined using the sequence comparison program NCBI-BLAST2 (Altschul et al., *Nucleic Acids Res.* 25:3389-3402 (1997)). The NCBI-BLAST2 sequence comparison program may be downloaded from <http://www.ncbi.nlm.nih.gov> or otherwise obtained from the National Institute of Health, Bethesda, Md. NCBI-BLAST2 uses several search parameters, wherein all of those search parameters are set to default values including, for example, unmask=yes, strand=all, expected occurrences=10, minimum low complexity length=15/5, multi-pass e-value=0.01, constant for multi-pass=25, dropoff for final gapped alignment=25 and scoring matrix=BLOSUM62.

[0206] In situations where NCBI-BLAST2 is employed for amino acid sequence comparisons, the % amino acid sequence identity of a given amino acid sequence A to, with, or against a given amino acid sequence B (which can alternatively be phrased as a given amino acid sequence A that has or comprises a certain % amino acid sequence identity to, with, or against a given amino acid sequence B) is calculated as follows:

100 times the fraction X/Y

[0207] where X is the number of amino acid residues scored as identical matches by the sequence alignment program NCBI-BLAST2 in that program's alignment of A and B, and where Y is the total number of amino acid residues in B. It will be appreciated that where the length of amino acid sequence A is not equal to the length of amino acid sequence B, the % amino acid sequence identity of A to B will not equal the % amino acid sequence identity of B to A.

[0208] "PRO variant polynucleotide" or "PRO variant nucleic acid sequence" means a nucleic acid molecule which encodes an active PRO polypeptide as defined below and which has at least about 80% nucleic acid sequence identity with a nucleotide acid sequence encoding a full-length native sequence PRO polypeptide sequence as disclosed herein, a full-length native sequence PRO polypeptide sequence lacking the signal peptide as disclosed herein, an

extracellular domain of a PRO polypeptide, with or without the signal peptide, as disclosed herein or any other fragment of a full-length PRO polypeptide sequence as disclosed herein. Ordinarily, a PRO variant polynucleotide will have at least about 80% nucleic acid sequence identity, alternatively at least about 81% nucleic acid sequence identity, alternatively at least about 82% nucleic acid sequence identity, alternatively at least about 83% nucleic acid sequence identity, alternatively at least about 84% nucleic acid sequence identity, alternatively at least about 85% nucleic acid sequence identity, alternatively at least about 86% nucleic acid sequence identity, alternatively at least about 87% nucleic acid sequence identity, alternatively at least about 88% nucleic acid sequence identity, alternatively at least about 89% nucleic acid sequence identity, alternatively at least about 90% nucleic acid sequence identity, alternatively at least about 91% nucleic acid sequence identity, alternatively at least about 92% nucleic acid sequence identity, alternatively at least about 93% nucleic acid sequence identity, alternatively at least about 94% nucleic acid sequence identity, alternatively at least about 95% nucleic acid sequence identity, alternatively at least about 96% nucleic acid sequence identity, alternatively at least about 97% nucleic acid sequence identity, alternatively at least about 98% nucleic acid sequence identity and alternatively at least about 99% nucleic acid sequence identity with a nucleic acid sequence encoding a full-length native sequence PRO polypeptide sequence as disclosed herein, a full-length native sequence PRO polypeptide sequence lacking the signal peptide as disclosed herein, an extracellular domain of a PRO polypeptide, with or without the signal sequence, as disclosed herein or any other fragment of a full-length PRO polypeptide sequence as disclosed herein. Variants do not encompass the native nucleotide sequence.

[0209] Ordinarily, PRO variant polynucleotides are at least about 30 nucleotides in length, alternatively at least about 60 nucleotides in length, alternatively at least about 90 nucleotides in length, alternatively at least about 120 nucleotides in length, alternatively at least about 150 nucleotides in length, alternatively at least about 180 nucleotides in length, alternatively at least about 210 nucleotides in length, alternatively at least about 240 nucleotides in length, alternatively at least about 270 nucleotides in length, alternatively at least about 300 nucleotides in length, alternatively at least about 450 nucleotides in length, alternatively at least about 600 nucleotides in length, alternatively at least about 900 nucleotides in length, or more.

[0210] "Percent (%) nucleic acid sequence identity" with respect to PRO-encoding nucleic acid sequences identified herein is defined as the percentage of nucleotides in a candidate sequence that are identical with the nucleotides in the PRO nucleic acid sequence of interest, after aligning the sequences and introducing gaps, if necessary, to achieve the maximum percent sequence identity. Alignment for purposes of determining percent nucleic acid sequence identity can be achieved in various ways that are within the skill in the art, for instance, using publicly available computer software such as BLAST, BLAST-2, ALIGN or Megalign (DNASTAR) software. For purposes herein, however, % nucleic acid sequence identity values are generated using the sequence comparison computer program ALIGN-2, wherein the complete source code for the ALIGN-2 program is provided in Table 1 below. The ALIGN-2 sequence comparison computer program was authored by Genentech, Inc.

and the source code shown in Table 1 below has been filed with user documentation in the U.S. Copyright Office, Washington D.C., 20559, where it is registered under U.S. Copyright Registration No. TXU510087. The ALIGN-2 program is publicly available through Genentech, Inc., South San Francisco, Calif. or may be compiled from the source code provided in Table 1 below. The ALIGN-2 program should be compiled for use on a UNIX operating system, preferably digital UNIX V4.0D. All sequence comparison parameters are set by the ALIGN-2 program and do not vary.

[0211] In situations where ALIGN-2 is employed for nucleic acid sequence comparisons, the % nucleic acid sequence identity of a given nucleic acid sequence C to, with, or against a given nucleic acid sequence D (which can alternatively be phrased as a given nucleic acid sequence C that has or comprises a certain % nucleic acid sequence identity to, with, or against a given nucleic acid sequence D) is calculated as follows:

100 times the fraction W/Z

[0212] where W is the number of nucleotides scored as identical matches by the sequence alignment program ALIGN-2 in that program's alignment of C and D, and where Z is the total number of nucleotides in D. It will be appreciated that where the length of nucleic acid sequence C is not equal to the length of nucleic acid sequence D, the % nucleic acid sequence identity of C to D will not equal the % nucleic acid sequence identity of D to C. As examples of % nucleic acid sequence identity calculations, Tables 4 and 5, demonstrate how to calculate the % nucleic acid sequence identity of the nucleic acid sequence designated "Comparison DNA" to the nucleic acid sequence designated "PRO-DNA", wherein "PRO-DNA" represents a hypothetical PRO-encoding nucleic acid sequence of interest, "Comparison DNA" represents the nucleotide sequence of a nucleic acid molecule against which the "PRO-DNA" nucleic acid molecule of interest is being compared, and "N", "L" and "V" each represent different hypothetical nucleotides.

[0213] Unless specifically stated otherwise, all % nucleic acid sequence identity values used herein are obtained as described in the immediately preceding paragraph using the ALIGN-2 computer program. However, % nucleic acid sequence identity values may also be obtained as described below by using the WU-BLAST-2 computer program (Altschul et al., Methods in Enzymology 266:460-480 (1996)). Most of the WU-BLAST-2 search parameters are set to the default values. Those not set to default values, i.e., the adjustable parameters, are set with the following values: overlap span=1, overlap fraction=0.125, word threshold (T)=11, and scoring matrix=BLOSUM62. When WU-BLAST-2 is employed, a % nucleic acid sequence identity value is determined by dividing (a) the number of matching identical nucleotides between the nucleic acid sequence of the PRO polypeptide-encoding nucleic acid molecule of interest having a sequence derived from the native sequence PRO polypeptide-encoding nucleic acid and the comparison nucleic acid molecule of interest (i.e., the sequence against which the PRO polypeptide-encoding nucleic acid molecule of interest is being compared which may be a variant PRO polynucleotide) as determined by WU-BLAST-2 by (b) the total number of nucleotides of the PRO polypeptide-encoding nucleic acid molecule of interest. For example, in the statement "an isolated nucleic acid

molecule comprising a nucleic acid sequence A which has or having at least 80% nucleic acid sequence identity to the nucleic acid sequence B", the nucleic acid sequence A is the comparison nucleic acid molecule of interest and the nucleic acid sequence B is the nucleic acid sequence of the PRO polypeptide-encoding nucleic acid molecule of interest.

[0214] Percent nucleic acid sequence identity may also be determined using the sequence comparison program NCBI-BLAST2 (Altschul et al., *Nucleic Acids Res.* 25:3389-3402 (1997)). The NCBI-BLAST2 sequence comparison program may be downloaded from <http://www.ncbi.nlm.nih.gov> or otherwise obtained from the National Institute of Health, Bethesda, Md. NCBI-BLAST2 uses several search parameters, wherein all of those search parameters are set to default values including, for example, unmask=yes, strand=all, expected occurrences=10, minimum low complexity length=15/5, multi-pass e-value=0.01, constant for multi-pass=25, dropoff for final gapped alignment=25 and scoring matrix=BLOSUM62.

[0215] In situations where NCBI-BLAST2 is employed for sequence comparisons, the % nucleic acid sequence identity of a given nucleic acid sequence C to, with, or against a given nucleic acid sequence D (which can alternatively be phrased as a given nucleic acid sequence C that has or comprises a certain % nucleic acid sequence identity to, with, or against a given nucleic acid sequence D) is calculated as follows:

$$100 \text{ times the fraction } W/Z$$

[0216] where W is the number of nucleotides scored as identical matches by the sequence alignment program NCBI-BLAST2 in that program's alignment of C and D, and where Z is the total number of nucleotides in D. It will be appreciated that where the length of nucleic acid sequence C is not equal to the length of nucleic acid sequence D, the % nucleic acid sequence identity of C to D will not equal the % nucleic acid sequence identity of D to C.

[0217] In other embodiments, PRO variant polynucleotides are nucleic acid molecules that encode an active PRO polypeptide and which are capable of hybridizing, preferably under stringent hybridization and wash conditions, to nucleotide sequences encoding a full-length PRO polypeptide as disclosed herein. PRO variant polypeptides may be those that are encoded by a PRO variant polynucleotide.

[0218] "Isolated," when used to describe the various polypeptides disclosed herein, means polypeptide that has been identified and separated and/or recovered from a component of its natural environment. Contaminant components of its natural environment are materials that would typically interfere with diagnostic or therapeutic uses for the polypeptide, and may include enzymes, hormones, and other proteinaceous or non-proteinaceous solutes. In preferred embodiments, the polypeptide will be purified (1) to a degree sufficient to obtain at least 15 residues of N-terminal or internal amino acid sequence by use of a spinning cup sequenator, or (2) to homogeneity by SDS-PAGE under non-reducing or reducing conditions using Coomassie blue or, preferably, silver stain. Isolated polypeptide includes polypeptide in situ within recombinant cells, since at least one component of the PRO polypeptide natural environment will not be present. Ordinarily, however, isolated polypeptide will be prepared by at least one purification step.

[0219] An "isolated" PRO polypeptide-encoding nucleic acid or other polypeptide-encoding nucleic acid is a nucleic acid molecule that is identified and separated from at least one contaminant nucleic acid molecule with which it is ordinarily associated in the natural source of the polypeptide-encoding nucleic acid. An isolated polypeptide-encoding nucleic acid molecule is other than in the form or setting in which it is found in nature. Isolated polypeptide-encoding nucleic acid molecules therefore are distinguished from the specific polypeptide-encoding nucleic acid molecule as it exists in natural cells. However, an isolated polypeptide-encoding nucleic acid molecule includes polypeptide-encoding nucleic acid molecules contained in cells that ordinarily express the polypeptide where, for example, the nucleic acid molecule is in a chromosomal location different from that of natural cells.

[0220] The term "control sequences" refers to DNA sequences necessary for the expression of an operably linked coding sequence in a particular host organism. The control sequences that are suitable for prokaryotes, for example, include a promoter, optionally an operator sequence, and a ribosome binding site. Eukaryotic cells are known to utilize promoters, polyadenylation signals, and enhancers.

[0221] Nucleic acid is "operably linked" when it is placed into a functional relationship with another nucleic acid sequence. For example, DNA for a presequence or secretory leader is operably linked to DNA for a polypeptide if it is expressed as a preprotein that participates in the secretion of the polypeptide; a promoter or enhancer is operably linked to a coding sequence if it affects the transcription of the sequence; or a ribosome binding site is operably linked to a coding sequence if it is positioned so as to facilitate translation. Generally, "operably linked" means that the DNA sequences being linked are contiguous, and, in the case of a secretory leader, contiguous and in reading phase. However, enhancers do not have to be contiguous. Linking is accomplished by ligation at convenient restriction sites. If such sites do not exist, the synthetic oligonucleotide adaptors or linkers are used in accordance with conventional practice.

[0222] The term "antibody" is used in the broadest sense and specifically covers, for example, single anti-PRO monoclonal antibodies (including agonist, antagonist, and neutralizing antibodies), anti-PRO antibody compositions with polypeptidic specificity, single chain anti-PRO antibodies, and fragments of anti-PRO antibodies (see below). The term "monoclonal antibody" as used herein refers to an antibody obtained from a population of substantially homogeneous antibodies, i.e., the individual antibodies comprising the population are identical except for possible naturally-occurring mutations that may be present in minor amounts.

[0223] "Stringency" of hybridization reactions is readily determinable by one of ordinary skill in the art, and generally is an empirical calculation dependent upon probe length, washing temperature, and salt concentration. In general, longer probes require higher temperatures for proper annealing, while shorter probes need lower temperatures. Hybridization generally depends on the ability of denatured DNA to reanneal when complementary strands are present in an environment below their melting temperature. The higher the degree of desired homology between the probe and hybridizable sequence, the higher the relative temperature which can be used. As a result, it follows that

higher relative temperatures would tend to make the reaction conditions more stringent, while lower temperatures less so. For additional details and explanation of stringency of hybridization reactions, see Ausubel et al., *Current Protocols in Molecular Biology*, Wiley Interscience Publishers, (1995).

[0224] “Stringent conditions” or “high stringency conditions”, as defined herein, may be identified by those that: (1) employ low ionic strength and high temperature for washing, for example 0.015 M sodium chloride/0.0015 M sodium citrate/0.1% sodium dodecyl sulfate at 50° C.; (2) employ during hybridization a denaturing agent, such as formamide, for example, 50% (v/v) formamide with 0.1% bovine serum albumin/0.1% Ficoll/0.1% polyvinylpyrrolidone/50 mM sodium phosphate buffer at pH 6.5 with 750 mM sodium chloride, 75 mM sodium citrate at 42° C.; or (3) employ 50% formamide, 5×SSC (0.75 M NaCl, 0.075 M sodium citrate), 50 mM sodium phosphate (pH 6.8), 0.1% sodium pyrophosphate, 5× Denhardt’s solution, sonicated salmon sperm DNA (50 µg/ml), 0.1% SDS, and 10% dextran sulfate at 42° C., with washes at 42° C. in 0.2×SSC (sodium chloride/sodium citrate) and 50% formamide at 55° C., followed by a high-stringency wash consisting of 0.1×SSC containing EDTA at 55° C.

[0225] “Moderately stringent conditions” may be identified as described by Sambrook et al., *Molecular Cloning: A Laboratory Manual*, New York: Cold Spring Harbor Press, 1989, and include the use of washing solution and hybridization conditions (e.g., temperature, ionic strength and % SDS) less stringent than those described above. An example of moderately stringent conditions is overnight incubation at 37° C. in a solution comprising: 20% formamide, 5×SSC (150 mM NaCl, 15 mM trisodium citrate), 50 mM sodium phosphate (pH 7.6), 5× Denhardt’s solution, 10% dextran sulfate, and 20 mg/ml denatured sheared salmon sperm DNA, followed by washing the filters in 1×SSC at about 37-50° C. The skilled artisan will recognize how to adjust the temperature, ionic strength, etc. as necessary to accommodate factors such as probe length and the like.

[0226] The term “epitope tagged” when used herein refers to a chimeric polypeptide comprising a PRO polypeptide fused to a “tag polypeptide”. The tag polypeptide has enough residues to provide an epitope against which an antibody can be made, yet is short enough such that it does not interfere with activity of the polypeptide to which it is fused. The tag polypeptide preferably also is fairly unique so that the antibody does not substantially cross-react with other epitopes. Suitable tag polypeptides generally have at least six amino acid residues and usually between about 8 and 50 amino acid residues (preferably, between about 10 and 20 amino acid residues).

[0227] As used herein, the term “immunoadhesin” designates antibody-like molecules which combine the binding specificity of a heterologous protein (an “adhesin”) with the effector functions of immunoglobulin constant domains. Structurally, the immunoadhesins comprise a fusion of an amino acid sequence with the desired binding specificity which is other than the antigen recognition and binding site of an antibody (i.e., is “heterologous”), and an immunoglobulin constant domain sequence. The adhesin part of an immunoadhesin molecule typically is a contiguous amino acid sequence comprising at least the binding site of a receptor or a ligand. The immunoglobulin constant domain

sequence in the immunoadhesin may be obtained from any immunoglobulin, such as IgG-1, IgG-2, IgG-3, or IgG-4 subtypes, IgA (including IgA-1 and IgA-2), IgE, IgD or IgM.

[0228] “Active” or “activity” for the purposes herein refers to form(s) of a PRO polypeptide which retain a biological and/or an immunological activity of native or naturally-occurring PRO, wherein “biological” activity refers to a biological function (either inhibitory or stimulatory) caused by a native or naturally-occurring PRO other than the ability to induce the production of an antibody against an antigenic epitope possessed by a native or naturally-occurring PRO and an “immunological” activity refers to the ability to induce the production of an antibody against an antigenic epitope possessed by a native or naturally-occurring PRO.

[0229] The term “antagonist” is used in the broadest sense, and includes any molecule that partially or fully blocks, inhibits, or neutralizes a biological activity of a native PRO polypeptide disclosed herein. In a similar manner, the term “agonist” is used in the broadest sense and includes any molecule that mimics a biological activity of a native PRO polypeptide disclosed herein. Suitable agonist or antagonist molecules specifically include agonist or antagonist antibodies or antibody fragments, fragments or amino acid sequence variants of native PRO polypeptides, peptides, antisense oligonucleotides, small organic molecules, etc. Methods for identifying agonists or antagonists of a PRO polypeptide may comprise contacting a PRO polypeptide with a candidate agonist or antagonist molecule and measuring a detectable change in one or more biological activities normally associated with the PRO polypeptide.

[0230] “Treatment” refers to both therapeutic treatment and prophylactic or preventative measures, wherein the object is to prevent or slow down (lessen) the targeted pathologic condition or disorder. Those in need of treatment include those already with the disorder as well as those prone to have the disorder or those in whom the disorder is to be prevented.

[0231] “Chronic” administration refers to administration of the agent(s) in a continuous mode as opposed to an acute mode, so as to maintain the initial therapeutic effect (activity) for an extended period of time. “Intermittent” administration is treatment that is not consecutively done without interruption, but rather is cyclic in nature.

[0232] “Mammal” for purposes of treatment refers to any animal classified as a mammal, including humans, domestic and farm animals, and zoo, sports, or pet animals, such as dogs, cats, cattle, horses, sheep, pigs, goats, rabbits, etc. Preferably, the mammal is human.

[0233] Administration “in combination with” one or more further therapeutic agents includes simultaneous (concurrent) and consecutive administration in any order.

[0234] “Carriers” as used herein include pharmaceutically acceptable carriers, excipients, or stabilizers which are non-toxic to the cell or mammal being exposed thereto at the dosages and concentrations employed. Often the physiologically acceptable carrier is an aqueous pH buffered solution. Examples of physiologically acceptable carriers include buffers such as phosphate, citrate, and other organic acids; antioxidants including ascorbic acid; low molecular weight

(less than about 10 residues) polypeptide; proteins, such as serum albumin, gelatin, or immunoglobulins; hydrophilic polymers such as polyvinylpyrrolidone; amino acids such as glycine, glutamine, asparagine, arginine or lysine; monosaccharides, disaccharides, and other carbohydrates including glucose, mannose, or dextrans; chelating agents such as EDTA; sugar alcohols such as mannitol or sorbitol; salt-forming counterions such as sodium; and/or nonionic surfactants such as TWEENTM, polyethylene glycol (PEG), and PLURONICSTM.

[0235] "Antibody fragments" comprise a portion of an intact antibody, preferably the antigen binding or variable region of the intact antibody. Examples of antibody fragments include Fab, Fab', F(ab')₂, and Fv fragments; diabodies; linear antibodies (Zapata et al., Protein Eng. 8(10): 1057-1062 [1995]); single-chain antibody molecules; and multispecific antibodies formed from antibody fragments.

[0236] Papain digestion of antibodies produces two identical antigen-binding fragments, called "Fab" fragments, each with a single antigen-binding site, and a residual "Fc" fragment, a designation reflecting the ability to crystallize readily. Pepsin treatment yields an F(ab')₂ fragment that has two antigen-combining sites and is still capable of cross-linking antigen.

[0237] "Fv" is the minimum antibody fragment which contains a complete antigen-recognition and -binding site. This region consists of a dimer of one heavy- and one light-chain variable domain in tight, non-covalent association. It is in this configuration that the three CDRs of each variable domain interact to define an antigen-binding site on the surface of the V_H-V_L dimer. Collectively, the six CDRs confer antigen-binding specificity to the antibody. However, even a single variable domain (or half of an Fv comprising only three CDRs specific for an antigen) has the ability to recognize and bind antigen, although at a lower affinity than the entire binding site.

[0238] The Fab fragment also contains the constant domain of the light chain and the first constant domain (CH1) of the heavy chain. Fab fragments differ from Fab' fragments by the addition of a few residues at the carboxy terminus of the heavy chain CH1 domain including one or more cysteines from the antibody hinge region. Fab'-SH is the designation herein for Fab' in which the cysteine residue(s) of the constant domains bear a free thiol group. F(ab')₂ antibody fragments originally were produced as pairs of Fab' fragments which have hinge cysteines between them. Other chemical couplings of antibody fragments are also known.

[0239] The "light chains" of antibodies (immunoglobulins) from any vertebrate species can be assigned to one of two clearly distinct types, called kappa and lambda, based on the amino acid sequences of their constant domains.

[0240] Depending on the amino acid sequence of the constant domain of their heavy chains, immunoglobulins can be assigned to different classes. There are five major classes of immunoglobulins: IgA, IgD, IgE, IgG, and IgM, and several of these may be further divided into subclasses (isotypes), e.g., IgG1, IgG2, IgG3, IgG4, IgA, and IgA2.

[0241] "Single-chain Fv" or "sFv" antibody fragments comprise the V_H and V_L domains of antibody, wherein these domains are present in a single polypeptide chain. Prefer-

ably, the Fv polypeptide further comprises a polypeptide linker between the V_H and V_L domains which enables the sFv to form the desired structure for antigen binding. For a review of sFv, see Pluckthun in *The Pharmacology of Monoclonal Antibodies*, vol.113, Rosenberg and Moore eds., Springer-Verlag, New York, pp. 269-315 (1994).

[0242] The term "diabodies" refers to small antibody fragments with two antigen-binding sites, which fragments comprise a heavy-chain variable domain (V_H) connected to a light-chain variable domain (V_L) in the same polypeptide chain (V_H-V_L). By using a linker that is too short to allow pairing between the two domains on the same chain, the domains are forced to pair with the complementary domains of another chain and create two antigen-binding sites. Diabodies are described more fully in, for example, EP 404,097; WO 93/11161; and Hollinger et al., Proc. Natl. Acad. Sci. USA, 90:6444-6448 (1993).

[0243] An "isolated" antibody is one which has been identified and separated and/or recovered from a component of its natural environment. Contaminant components of its natural environment are materials which would interfere with diagnostic or therapeutic uses for the antibody, and may include enzymes, hormones, and other proteinaceous or nonproteinaceous solutes. In preferred embodiments, the antibody will be purified (1) to greater than 95% by weight of antibody as determined by the Lowry method, and most preferably more than 99% by weight, (2) to a degree sufficient to obtain at least 15 residues of N-terminal or internal amino acid sequence by use of a spinning cup sequenator, or (3) to homogeneity by SDS-PAGE under reducing or nonreducing conditions using Coomassie blue or, preferably, silver stain. Isolated antibody includes the antibody in situ within recombinant cells since at least one component of the antibody's natural environment will not be present. Ordinarily, however, isolated antibody will be prepared by at least one purification step.

[0244] An antibody that "specifically binds to" or is "specific for" a particular polypeptide or an epitope on a particular polypeptide is one that binds to that particular polypeptide or epitope on a particular polypeptide without substantially binding to any other polypeptide or polypeptide epitope.

[0245] The word "label" when used herein refers to a detectable compound or composition which is conjugated directly or indirectly to the antibody so as to generate a "labeled" antibody. The label may be detectable by itself (e.g. radioisotope labels or fluorescent labels) or, in the case of an enzymatic label, may catalyze chemical alteration of a substrate compound or composition which is detectable.

[0246] By "solid phase" is meant a non-aqueous matrix to which the antibody of the present invention can adhere. Examples of solid phases encompassed herein include those formed partially or entirely of glass (e.g., controlled pore glass), polysaccharides (e.g., agarose), polyacrylamides, polystyrene, polyvinyl alcohol and silicones. In certain embodiments, depending on the context, the solid phase can comprise the well of an assay plate; in others it is a purification column (e.g., an affinity chromatography column). This term also includes a discontinuous solid phase of discrete particles, such as those described in U.S. Pat. No. 4,275,149.

[0247] A “liposome” is a small vesicle composed of various types of lipids, phospholipids and/or surfactant which is useful for delivery of a drug (such as a PRO polypeptide or antibody thereto) to a mammal. The components of the liposome are commonly arranged in a bilayer

formation, similar to the lipid arrangement of biological membranes.

[0248] A “small molecule” is defined herein to have a molecular weight below about 500 Daltons

TABLE 1

```

/*
 *
 * C—C increased from 12 to 15
 * Z is average of EQ
 * B is average of ND
 * match with stop is _M; stop—stop = 0; J (joker) match = 0
 */
#define _M -8 /* value of a match with a stop */
int
_day[26][26] = {
/* A B C D E F G H I J K L M N O P Q R S T U V W X Y Z */
/* A */ {2, 0, -2, 0, 0, -4, 1, -1, -1, 0, -1, -2, -1, 0, _M, 1, 0, -2, 1, 1, 0, 0, -6, 0, -3, 0},
/* B */ {0, 3, -4, 3, 2, -5, 0, 1, -2, 0, 0, -3, -2, 2, _M, -1, 1, 0, 0, 0, -2, -5, 0, -3, 1},
/* C */ {-2, -4, 15, -5, -5, -4, -3, -3, -2, 0, -5, -6, -5, -4, _M, -3, -5, -4, 0, -2, 0, -2, -8, 0, 0, -5},
/* D */ {0, 3, -5, 4, 3, -6, 1, 1, -2, 0, 0, -4, -3, 2, _M, -1, 2, -1, 0, 0, 0, -2, -7, 0, -4, 2},
/* E */ {0, 2, -5, 3, 4, -5, 0, 1, -2, 0, 0, -3, -2, 1, _M, -1, 2, -1, 0, 0, 0, -2, -7, 0, -4, 3},
/* F */ {-4, -5, -4, -6, -5, 9, -5, -2, 1, 0, -5, 2, 0, -4, _M, -5, -5, -4, -3, -3, 0, -1, 0, 0, 7, -5},
/* G */ {1, 0, -3, 1, 0, -5, 5, -2, -3, 0, -2, -4, -3, 0, _M, -1, -1, -3, 1, 0, 0, -1, -7, 0, -5, 0},
/* H */ {-1, 1, -3, 1, 1, -2, -2, 6, -2, 0, 0, -2, -2, 2, _M, 0, 3, 2, -1, -1, 0, -2, -3, 0, 0, 2},
/* I */ {-1, -2, -2, -2, -2, 1, -3, -2, 5, 0, -2, 2, 2, -2, _M, -2, -2, -2, -1, 0, 0, 4, -5, 0, -1, -2},
/* J */ {0, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0, _M, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0},
/* K */ {-1, 0, -5, 0, 0, -5, -2, 0, -2, 0, 5, -3, 0, 1, _M, -1, 1, 3, 0, 0, 0, -2, -3, 0, -4, 0},
/* L */ {-2, -3, -6, -4, -3, 2, -4, -2, 2, 0, -3, 6, 4, -3, _M, -3, -2, -3, -3, -1, 0, 2, -2, 0, -1, -2},
/* M */ {-1, -2, -5, -3, -2, 0, -3, -2, 2, 0, 0, 4, 6, -2, _M, -2, -1, 0, -2, -1, 0, 2, -4, 0, -2, -1},
/* N */ {0, 2, -4, 2, 1, -4, 0, 2, -2, 0, 1, -3, -2, 2, _M, -1, 1, 0, 1, 0, 0, -2, -4, 0, -2, 1},
/* O */ {_M, _M, _M, _M, _M, _M, _M, _M, _M, _M, _M, _M, _M, _M, _M, _M, _M, _M, _M, _M, _M, _M, _M, _M},
/* P */ {1, -1, -3, -1, -1, -5, -1, 0, -2, 0, -1, -3, -2, -1, _M, 6, 0, 0, 1, 0, 0, -1, -6, 0, -5, 0},
/* Q */ {0, 1, -5, 2, 2, -5, -1, 3, -2, 0, 1, -2, -1, 1, _M, 0, 4, 1, -1, -1, 0, -2, -5, 0, -4, 3},
/* R */ {-2, 0, -4, -1, -1, -4, -3, 2, -2, 0, 3, -3, 0, 0, _M, 0, 1, 6, 0, -1, 0, -2, 2, 0, -4, 0},
/* S */ {1, 0, 0, 0, -3, 1, -1, -1, 0, 0, -3, -2, 1, _M, 1, -1, 0, 2, 1, 0, -1, -2, 0, -3, 0},
/* T */ {1, 0, -2, 0, 0, -3, 0, -1, 0, 0, 0, -1, -1, 0, _M, 0, -1, -1, 1, 3, 0, 0, -5, 0, -3, 0},
/* U */ {0, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0, _M, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0},
/* V */ {0, -2, -2, -2, -2, -1, -1, -2, 4, 0, -2, 2, 2, -2, _M, -1, -2, -2, -1, 0, 0, 4, -6, 0, -2, -2},
/* W */ {-6, -5, -8, -7, -7, 0, -7, -3, -5, 0, -3, -2, -4, -4, _M, -6, -5, 2, -2, -5, 0, -6, 17, 0, 0, -6},
/* X */ {0, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0, _M, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0},
/* Y */ {-3, -3, 0, -4, -4, 7, -5, 0, -1, 0, -4, -1, -2, -2, _M, -5, -4, -4, -3, -3, 0, -2, 0, 0, 10, -4},
/* Z */ {0, 1, -5, 2, 3, -5, 0, 2, -2, 0, 0, -2, -1, 1, _M, 0, 3, 0, 0, 0, -2, -6, 0, -4, 4},
};
/*
 */
#include <stdio.h>
#include <ctype.h>
#define MAXJMP 16 /* max jumps in a diag */
#define MAXGAP 24 /* don't continue to penalize gaps larger than this */
#define JMPS 1024 /* max jmps in an path */
#define MX 4 /* save if there's at least MX-1 bases since last jmp */
#define DMAT 3 /* value of matching bases */
#define DMIS 0 /* penalty for mismatched bases */
#define DINS0 8 /* penalty for a gap */
#define DINS1 1 /* penalty per base */
#define PINS0 8 /* penalty for a gap */
#define PINS1 4 /* penalty per residue */
struct jmp {
    short n[MAXJMP]; /* size of jmp (neg for dely) */
    unsigned short x[MAXJMP]; /* base no. of jmp in seq x */
    /* limits seq to 2^16 -1 */
};
struct diag {
    int score; /* score at last jmp */
    long offset; /* offset of prev block */
    short ijmp; /* current jmp index */
    struct jmp jp; /* list of jmps */
};
struct path {
    int spc; /* number of leading spaces */
    short n[JMPS]; /* size of jmp (gap) */
    int x[JMPS]; /* loc of jmp (last elem before gap) */
}

```

TABLE 1-continued

```

};
char      *ofile;           /* output file name */
char      *namex[2];        /* seq names: getseqs() */
char      *prog;            /* prog name for err msgs */
char      *seqx[2];         /* seqs: getseqs() */
int       dmax;             /* best diag: nw() */
int       dmax0;            /* final diag */
int       dna;              /* set if dna: main() */
int       endgaps;          /* set if penalizing end gaps */
int       gapx, gapy;        /* total gaps in seqs */
int       len0, len1;        /* seq lens */
int       ngapx, ngapy;      /* total size of gaps */
int       smax;             /* max score: nw() */
int       *xbm;             /* bitmap for matching */
long      offset;          /* current offset in jmp file */
struct    diag *dx;         /* holds diagonals */
struct    path pp[2];       /* holds path for seqs */
char      *calloc(), *malloc(), *index(), *strcpy();
char      *getseq(), *g_calloc();
/* Needleman-Wunsch alignment program
 *
 * usage: progs file1 file2
 * where file1 and file2 are two dna or two protein sequences.
 * The sequences can be in upper- or lower-case and may contain ambiguity
 * Any lines beginning with ';', '>' or '<' are ignored
 * Max file length is 65535 (limited by unsigned short x in the jmp struct)
 * A sequence with 1/3 or more of its elements ACGTU is assumed to be DNA
 * Output is in the file "align.out"
 *
 * The program may create a tmp file in /tmp to hold info about traceback.
 * Original version developed under BSD 4.3 on a vax 8650
 */
#include "nw.h"
#include "day.h"
static  __dbval[26] = {
    1,14,2,13,0,0,4,11,0,0,12,0,3,15,0,0,0,5,6,8,8,7,9,0,10,0
};
static  __pbval[26] = {
    1, 2|(1<<('D'-'A'))|(1<<('N'-'A')), 4, 8, 16, 32, 64,
    128, 256, 0xFFFFFFFF, 1<<10, 1<<11, 1<<12, 1<<13, 1<<14,
    1<<15, 1<<16, 1<<17, 1<<18, 1<<19, 1<<20, 1<<21, 1<<22,
    1<<23, 1<<24, 1<<25|(1<<('E'-'A'))|(1<<('Q'-'A'))
};
main(ac, av)
int      ac;
char     *av[];
{
    prog = av[0];
    if(ac != 3) {
        fprintf(stderr, "usage: %s file1 file2\n", prog);
        fprintf(stderr, "where file1 and file2 are two dna or two protein sequences.\n");
        fprintf(stderr, "The sequences can be in upper- or lower-case\n");
        fprintf(stderr, "Any lines beginning with ';', '>' or '<' are ignored\n");
        fprintf(stderr, "Output is in the file \"align.out\"\n");
        exit(1);
    }
    namex[0] = av[1];
    namex[1] = av[2];
    seqx[0] = getseq(namex[0], &len0);
    seqx[1] = getseq(namex[1], &len1);
    xbm = (dna)? __dbval : __pbval;
    endgaps = 0;          /* 1 to penalize endgaps */
    ofile = "align.out";  /* output file */
    nw();                /* fill in the matrix, get the possible jmps */
    readjumps();          /* get the actual jmps */
    print();              /* print stats, alignment */
    cleanup(0);           /* unlink any tmp files */
}
/* do the alignment, return best score: main()
 * dna: values in Fitch and Smith, PNAS, 80, 1382-1386, 1983
 * pro: PAM 250 values
 * When scores are equal, we prefer mismatches to any gap, prefer
 * a new gap to extending an ongoing gap, and prefer a gap in seqx
 * to a gap in seq y.
 */
nw()

```

main

nw

TABLE 1-continued

```

{
    char    *px, *py;        /* seqs and ptrs */
    int      *ndely, *dely;   /* keep track of dely */
    int      ndelx, delx;     /* keep track of delx */
    int      *tmp;           /* for swapping row0, row1 */
    int      mis;            /* score for each type */
    int      ins0, ins1;      /* insertion penalties */
    register id;             /* diagonal index */
    register ij;            /* jmp index */
    register *col0, *col1;    /* score for curr, last row */
    register xx, yy;         /* index into seqs */
    dx = (struct diag *)g_calloc("to get diags", len0+len1+1, sizeof(struct diag));
    ndely = (int *)g_calloc("to get ndely", len1+1, sizeof(int));
    dely = (int *)g_calloc("to get dely", len1+1, sizeof(int));
    col0 = (int *)g_calloc("to get col0", len1+1, sizeof(int));
    col1 = (int *)g_calloc("to get col1", len1+1, sizeof(int));
    ins0 = (dna)? DINS0 : PINS0;
    ins1 = (dna)? DINS1 : PINS1;
    smax = -10000;
    if (endgaps) {
        for (col0[0] = dely[0] = -ins0, yy = 1; yy <= len1; yy++) {
            col0[yy] = dely[yy] = col0[yy-1] - ins1;
            ndely[yy] = yy;
        }
        col0[0] = 0;        /* Waterman Bull Math Biol 84 */
    }
    else
        for (yy = 1; yy <= len1; yy++)
            dely[yy] = -ins0;
    /* fill in match matrix
    */
    for (px = seqx[0], xx = 1; xx <= len0; px++, xx++) {
        /* initialize first entry in col
        */
        if (endgaps) {
            if (xx == 1)
                col1[0] = delx = -(ins0+ins1);
            else
                col1[0] = delx = col0[0]-ins1;
            ndelx = xx;
        }
        else {
            col1[0] = 0;
            delx = -ins0;
            ndelx = 0;
        }
    }

    for (py = seqx[1], yy = 1; yy <= len1; py++, yy++) {
        mis = col0[yy-1];
        if (dna)
            mis += (xbm[*px-'A']&xbm[*py-'A'])? DMAT : DMIS;
        else
            mis += _day[*px-'A'][*py-'A'];
        /* update penalty for del in x seq;
        * favor new del over ongong del
        * ignore MAXGAP if weighting endgaps
        */
        if (endgaps || ndely[yy] < MAXGAP) {
            if (col0[yy] - ins0 >= dely[yy]) {
                dely[yy] = col0[yy] - (ins0+ins1);
                ndely[yy] = 1;
            } else {
                dely[yy] -= ins1;
                ndely[yy]++;
            }
        } else {
            if (col0[yy] - (ins0+ins1) >= dely[yy]) {
                dely[yy] = col0[yy] - (ins0+ins1);
                ndely[yy] = 1;
            } else
                ndely[yy]++;
        }
        /* update penalty for del in y seq;
        * favor new del over ongong del
        */
        if (endgaps || ndelx < MAXGAP) {

```

TABLE 1-continued

```

        if(col1[yy-1] - ins0 >= delx) {
            delx = col1[yy-1] - (ins0+ins1);
            ndelx = 1;
        } else {
            delx -= ins1;
            ndelx++;
        }
    } else {
        if (col1[yy-1] - (ins0+ins1) >= delx) {
            delx = col1[yy-1] - (ins0+ins1);
            ndelx = 1;
        } else
            ndelx++;
    }
    /* pick the maximum score; we're favoring
    * mis over any del and delx over dely
    */

    id = xx - yy + len1 - 1;
    if (mis >= delx && mis >= dely[yy])
        col1[yy] = mis;
    else if (delx >= dely[yy]) {
        col1[yy] = delx;
        ij = dx[id].ijmp;
        if (dx[id].jp.n[0] && (!dna || (ndelx >= MAXJMP
        && xx > dx[id].jp.x[ij]+MX) || mis > dx[id].score+DINS0)) {
            dx[id].ijmp++;
            if (++ij >= MAXJMP) {
                writeimps(id);
                ij = dx[id].ijmp = 0;
                dx[id].offset = offset;
                offset += sizeof(struct jmp) + sizeof(offset);
            }
        }
        dx[id].jp.n[ij] = ndelx;
        dx[id].jp.x[ij] = xx;
        dx[id].score = delx;
    }
    else {
        col1[yy] = dely[yy];
        ij = dx[id].ijmp;
        if (dx[id].jp.n[0] && (!dna || (ndely[yy] >= MAXJMP
        && xx > dx[id].jp.x[ij]+MX) || mis > dx[id].score+DINS0)) {
            dx[id].ijmp++;
            if (++ij >= MAXJMP) {
                writeimps(id);
                ij = dx[id].ijmp = 0;
                dx[id].offset = offset;
                offset += sizeof(struct jmp) + sizeof(offset);
            }
        }
        dx[id].jp.n[ij] = ndely[yy];
        dx[id].jp.x[ij] = xx;
        dx[id].score = dely[yy];
    }
    if (xx == len0 && yy < len1) {
        /* last col
        */
        if (endgaps)
            col1[yy] -= ins0+ins1*(len1-yy);
        if (col1[yy] > smax) {
            smax = col1[yy];
            dmax = id;
        }
    }
}
if (endgaps && xx < len0)
    col1[yy-1] -= ins0+ins1*(len0-xx);
if (col1[yy-1] > smax) {
    smax = col1[yy-1];
    dmax = id;
}
tmp = col0; col0 = col1; col1 = tmp;
}
(void) free((char *)ndely);
(void) free((char *)dely);

```

...nw

TABLE 1-continued

```

(void) free((char *)col0);
(void) free((char *)col1);
    }
/*
 *
 * print() -- only routine visible outside this module
 *
 * static:
 * getmat() -- trace back best path, count matches: print()
 * pr_align() -- print alignment of described in array p[]: print()
 * dumpblock() -- dump a block of lines with numbers, stars: pr_align()
 * nums() -- put out a number line: dumpblock()
 * putline() -- put out a line (name, [num], seq, [num]): dumpblock()
 * stars() -- put a line of stars: dumpblock()
 * stripname() -- strip any path and prefix from a seqname
 */
#include "nw.h"
#define SPC 3
#define P_LINE 256 /* maximum output line */
#define P_SPC 3 /* space between name or num and seq */
extern _day[26][26];
int olen; /* set output line length */
FILE *fx; /* output file */
print()
{
    int lx, ly, firstgap, lastgap; /* overlap */
    if ((fx = fopen(ofile, "w")) == 0) {
        fprintf(stderr, "%s: can't write %s\n", prog, ofile);
        cleanup(1);
    }
    fprintf(fx, "<first sequence: %s (length = %d)\n", namex[0], len0);
    fprintf(fx, "<second sequence: %s (length = %d)\n", namex[1], len1);
    olen = 60;
    lx = len0;
    ly = len1;
    firstgap = lastgap = 0;
    if (dmax < len1 - 1) { /* leading gap in x */
        pp[0].spc = firstgap = len1 - dmax - 1;
        ly -= pp[0].spc;
    }
    else if (dmax > len1 - 1) { /* leading gap in y */
        pp[1].spc = firstgap = dmax - (len1 - 1);
        lx -= pp[1].spc;
    }
    if (dmax0 < len0 - 1) { /* trailing gap in x */
        lastgap = len0 - dmax0 - 1;
        lx -= lastgap;
    }
    else if (dmax0 > len0 - 1) { /* trailing gap in y */
        lastgap = dmax0 - (len0 - 1);
        ly -= lastgap;
    }
    getmat(lx, ly, firstgap, lastgap);
    pr_align();
}
/*
 * trace back the best path, count matches
 */
static
getmat(lx, ly, firstgap, lastgap)
{
    int lx, ly; /* "core" (minus endgaps) */
    int firstgap, lastgap; /* leading trailing overlap */
    {
        int nm, i0, i1, siz0, siz1;
        char outx[32];
        double pct;
        register n0, n1;
        register char *p0, *p1;
        /* get total matches, score
        */
        i0 = i1 = siz0 = siz1 = 0;
        p0 = seqx[0] + pp[1].spc;
        p1 = seqx[1] + pp[0].spc;
        n0 = pp[1].spc + 1;
        n1 = pp[0].spc + 1;
        nm = 0;

```

TABLE 1-continued

```

while ( *p0 && *p1 ) {
    if (siz0) {
        p1++;
        n1++;
        siz0--;
    }
    else if (siz1) {
        p0++;
        n0++;
        siz1--;
    }
    else {
        if (xbm[*p0-'A']&xbm[*p1-'A'])
            nm++;
        if (n0++ == pp[0].x[i0])
            siz0 = pp[0].n[i0++];
        if (n1++ == pp[1].x[i1])
            siz1 = pp[1].n[i1++];
        p0++;
        p1++;
    }
}
/* pct homology:
 * if penalizing endgaps, base is the shorter seq
 * else, knock off overhangs and take shorter core
 */
if (endgaps)
    lx = (len0 < len1)? len0 : len1;
else
    lx = (lx < ly)? lx : ly;
pct = 100.*(double)nm/(double)lx;
fprintf(fx, "\n");
fprintf(fx, "<end match%s in an overlap of %d: %.2f percent similarity\n",
        nm, (nm == 1)? "" : "es", lx, pct);
fprintf(fx, "<gaps in first sequence: %d", gapx);
if (gapx) {
    (void) sprintf(outx, "(%d %s%s)",
        ngapx, (dna)? "base": "residue", (ngapx == 1)? "" : "s");
    fprintf(fx, "%s", outx);
}
fprintf(fx, ", gaps in second sequence: %d", gapy);
if (gapy) {
    (void) sprintf(outx, "(%d %s%s)",
        ngapy, (dna)? "base": "residue", (ngapy == 1)? "" : "s");
    fprintf(fx, "%s", outx);
}
if (dna)
    fprintf(fx,
        "\n<score: %d (match = %d, mismatch = %d, gap penalty = %d + %d per base)\n",
        smax, DMAT, DMIS, DINS0, DINS1);
else
    fprintf(fx,
        "\n<score: %d (Dayhoff PAM 250 matrix, gap penalty = %d + %d per residue)\n",
        smax, PINS0, PINS1);
if (endgaps)
    fprintf(fx,
        "<endgaps penalized. left endgap: %d %s%s, right endgap: %d %s%s\n",
        firstgap, (dna)? "base": "residue", (firstgap == 1)? "" : "s",
        lastgap, (dna)? "base": "residue", (lastgap == 1)? "" : "s");
else
    fprintf(fx, "<endgaps not penalized\n");
}
static      nm;          /* matches in core -- for checking */
static      lmax;        /* lengths of stripped file names */
static      ij[2];       /* jmp index for a path */
static      nc[2];       /* number at start of current line */
static      ni[2];       /* current elem number -- for gapping */
static      siz[2];
static char *ps[2];      /* ptr to current element */
static char *po[2];      /* ptr to next output char slot */
static char out[2][P_LINE]; /* output line */
static char star[P_LINE]; /* set by stars() */
/*
 * print alignment of described in struct path pp[]
 */
static
pr_align()

```

...getmat

pr_align

TABLE 1-continued

```
{
  int      nn;          /* char count */
  int      more;
  register i;
  for (i = 0, lmax = 0; i < 2; i++) {
    nn = stripname(name[x[i]]);
    if (nn > lmax)
      lmax = nn;
    nc[i] = 1;
    ni[i] = 1;
    siz[i] = ij[i] = 0;
    ps[i] = seq[x[i]];
    po[i] = out[i];
  }
  for (nn = nm = 0, more = 1; more;) {
    for (i = more = 0; i < 2; i++) {
      /*
       * do we have more of this sequence?
       */
      if (!*ps[i])
        continue;
      more++;
      if (pp[i].spc) {
        /* leading space */
        *po[i]++ = ' ';
        pp[i].spc--;
      }
      else if (siz[i]) {
        /* in a gap */
        *po[i]++ = '-';
        siz[i]--;
      }
      else {
        /* we're putting a seq element
         */
        *po[i] = *ps[i];
        if (islower(*ps[i]))
          *ps[i] = toupper(*ps[i]);
        po[i]++;
        ps[i]++;
        /*
         * are we at next gap for this seq?
         */
        if (ni[i] == pp[i].x[ij[i]]) {
          /*
           * we need to merge all gaps
           * at this location
           */
          siz[i] == pp[i].n[ij[i]++];
          while (ni[i] == pp[i].x[ij[i]])
            siz[i] += pp[i].n[ij[i]++];
        }
        ni[i]++;
      }
    }
    if (++nn == olen || !more && nn) {
      dumpblock();
      for (i = 0; i < 2; i++)
        po[i] = out[i];
      nn = 0;
    }
  }
}
/*
 * dump a block of lines, including numbers, stars: pr_align()
 */
static
dumpblock()
{
  register i;
  for(i = 0; i < 2; i++)
    *po[i]-- = '\0';

  (void) puts("\n", fx);
  for (i = 0; i < 2; i++) {
    if (*out[i] && (*out[i] != ' ' || *(po[i]) != ' ')) {
      if (i == 0)
        nums(i);
      if (i == 0 && *out[1])
        ...dumpblock
    }
  }
}
```

TABLE 1-continued

<pre> stars(); putline(i); if (i == 0 && *out[1]) fprintf(fx, star); if (i == 1) nums(i); } } } } /* * put out a number line: dumpblock() */ static nums(ix) int ix; /* index in out[] holding seq line */ { char nline[P_LINE]; register i, j; register char *pn, *px, *py; for(pn = nline, i = 0; i < lmax+P_SPC; i++, pn++) *pn = ' '; for (i = nc[ix], py = out[ix]; *py; py++, pn++) { if (*py == ' ' *py == '-') *pn = ' '; else { if (i%10 == 0 (i == 1 && nc[ix] != 1)) { j = (i < 0)? -i : i; for (px = pn; j; j/= 10, px--) *px = j%10 + '0'; if (i < 0) *px = '-'; } else *pn = ' '; i++; } } *pn = '\0'; nc[ix] = i; for (pn = nline; *pn; pn++) (void) putc(*pn, fx); (void) putc('\n', fx); } /* * put out a line (name, [num], seq. [num]): dumpblock() */ static putline(ix) int ix; { int i; register char *px; for (px = namex[ix], i = 0; *px && *px != '.'; px++, i++) (void) putc(*px, fx); for (i < lmax+P_SPC; i++) (void) putc(' ', fx); /* these count from 1: * ni[] is current element (from 1) * nc[] is number at start of current line */ for (px = out[ix]; *px; px++) (void) putc(*px&0x7F, fx); (void) putc('\n', fx); } /* * put a line of stars (seqs always in out[0], out[1]): dumpblock() */ static stars() { int i; register char *p0, *p1, cx, *px; if (!(*out[0] (*out[0] == ' ' && *(p0[0]) == ' ') !*out[1] (*out[1] == ' ' && *(p0[1]) == ' ')) return; px = star; </pre>	nums putline ...putline stars
--	---

TABLE 1-continued

<pre>for (i = lmax+P__SPC; i; i--) *px++ = ' '; for (p0 = out[0], p1 = out[1]; *p0 && *p1; p0++, p1++) { if (isalpha(*p0) && isalpha(*p1)) { if (xbm[*p0-'A']&xbm[*p1-'A']) { cx = '*'; nm++; } else if (!dna && __day[*p0-'A'][*p1-'A'] > 0) cx = '.'; else cx = ' '; } else cx = ' '; *px++ = cx; } *px++ = '\n'; *px = '\0'; } /* * strip path or prefix from pn, return len: pr__align() */ static stripname(pn) char *pn; /* file name (may be path) */ { register char *px, *py; py = 0; for (px = pn; *px; px++) if (*px == '/') py = px + 1; if (py) (void) strcpy(pn, py); return(strlen(pn)); } /* * cleanup() -- cleanup any tmp file * getseq() -- read in seq, set dna, len, maxlen * g__calloc() -- calloc() with error checkin * readjumps() -- get the good jumps, from tmp file if necessary * writejumps() -- write a filled array of jumps to a tmp file: nw() */ #include "nw.h" #include <sys/file.h> char *jname = "/tmp/homgXXXXXX"; /* tmp file for jumps */ FILE *fj; int cleanup(); /* cleanup tmp file */ long lseek(); /* * remove any tmp file if we blow */ cleanup(i) int i; { if (fj) (void) unlink(jname); exit(i); } /* * read, return ptr to seq, set dna, len, maxlen * skip lines starting with ';', '<', or '>' * seq in upper or lower case */ char * getseq(file, len) char *file; /* file name */ int *len; /* seq len */ { char line[1024], *pseq; register char *px, *py; int natgc, tlen; FILE *fp; if ((fp = fopen(file, "r")) == 0) { fprintf(stderr, "%s: can't read %s\n", prog, file); exit(1); } </pre>		
		stripname
		cleanup
		getseq

TABLE 1-continued

```

    }
    tlen = natgc = 0;
    while (fgets(line, 1024, fp)) {
        if (*line == ';' || *line == '<' || *line == '>')
            continue;
        for (px = line; *px != '\n'; px++)
            if (isupper(*px) || islower(*px))
                tlen++;
    }
    if ((pseq = malloc((unsigned)(tlen+6))) == 0) {
        fprintf(stderr, "%s: malloc() failed to get %d bytes for %s\n", prog, tlen+6, file);
        exit(1);
    }
    pseq[0] = pseq[1] = pseq[2] = pseq[3] = '\0';
    ...getseq

    py = pseq + 4;
    *len = tlen;
    rewind(fp);
    while (fgets(line, 1024, fp)) {
        if (*line == ';' || *line == '<' || *line == '>')
            continue;
        for (px = line; *px != '\n'; px++) {
            if (isupper(*px))
                *py++ = *px;
            else if (islower(*px))
                *py++ = toupper(*px);
            if (index("ATGCU", *(py-1)))
                natgc++;
        }
    }
    *py++ = '\0';
    *py = '\0';
    (void) fclose(fp);
    dna = natgc > (tlen/3);
    return(pseq+4);
}
char *
g__alloc(msg, nx, sz)
char *msg;          /* program, calling routine */
int nx, sz;          /* number and size of elements */
{
    char *px, *calloc();
    if ((px = calloc((unsigned)nx, (unsigned)sz)) == 0) {
        if (*msg) {
            fprintf(stderr, "%s: g__alloc() failed %s (n= %d, sz= %d)\n", prog, msg, nx, sz);
            exit(1);
        }
    }
    return(px);
}
/*
 * get final jmps from dx[] or tmp file, set pp[], reset dmax: main()
 */
readjmps()
{
    int fd = -1;
    int siz, i0, i1;
    register i, j, xx;
    if (fj) {
        (void) fclose(fj);
        if ((fd = open(jname, O_RDONLY, 0)) < 0) {
            fprintf(stderr, "%s: can't open() %s\n", prog, jname);
            cleanup(1);
        }
    }
    for (i = i0 = i1 = 0, dmax0 = dmax, xx = len0; i++) {
        while (1) {
            for (j = dx[dmax].ijmp; j >= 0 && dx[dmax].jp.x[j] >= xx; j--)
                ;
            ...readjmps

            if (j < 0 && dx[dmax].offset && fj) {
                (void) lseek(fd, dx[dmax].offset, 0);
                (void) read(fd, (char *)&dx[dmax].jp, sizeof(struct jmp));
                (void) read(fd, (char *)&dx[dmax].offset, sizeof(dx[dmax].offset));
                dx[dmax].ijmp = MAXJMP-1;
            }
        }
    }
}

```

TABLE 1-continued

```

    }
    else
        break;
    }
    if (i >= JMPS) {
        fprintf(stderr, "%s: too many gaps in alignment\n", prog);
        cleanup(1);
    }
    if (j >= 0) {
        siz = dx[dmax].jp.n[j];
        xx = dx[dmax].jp.x[j];
        dmax += siz;
        if (siz < 0) { /* gap in second seq */
            pp[1].n[i1] = -siz;
            xx += siz;
            /* id = xx - yy + len1 - 1
            */
            pp[1].x[i1] = xx - dmax + len1 - 1;
            gapy++;
            ngapy -= siz;
/* ignore MAXGAP when doing endgaps */
            siz = (-siz < MAXGAP || endgaps)? -siz : MAXGAP;
            i1++;
        }
        else if (siz > 0) { /* gap in first seq */
            pp[0].n[i0] = siz;
            pp[0].x[i0] = xx;
            gapx++;
            ngapx += siz;
/* ignore MAXGAP when doing endgaps */
            siz = (siz < MAXGAP || endgaps)? siz : MAXGAP;
            i0++;
        }
    }
    else
        break;
}
/* reverse the order of jmps
*/
for (j = 0, i0--; j < i0; j++, i0--) {
    i = pp[0].n[j]; pp[0].n[j] = pp[0].n[i0]; pp[0].n[i0] = i;
    i = pp[0].x[j]; pp[0].x[j] = pp[0].x[i0]; pp[0].x[i0] = i;
}
for (j = 0, i1--; j < i1; j++, i1--) {
    i = pp[1].n[j]; pp[1].n[j] = pp[1].n[i1]; pp[1].n[i1] = i;
    i = pp[1].x[j]; pp[1].x[j] = pp[1].x[i1]; pp[1].x[i1] = i;
}
if (fd >= 0)
    (void) close(fd);
if (fj) {
    (void) unlink(jname);
    fj = 0;
    offset = 0;
}
}

/*
 * write a filled jmp struct offset of the prev one (if any): nw()
 */
writejumps(ix)
int ix;
{
    char *mktemp();
    if (!fj) {
        if (mktemp(jname) < 0) {
            fprintf(stderr, "%s: can't mktemp() %s\n", prog, jname);
            cleanup(1);
        }
        if ((fj = fopen(jname, "w")) == 0) {
            fprintf(stderr, "%s: can't write %s\n", prog, jname);
            exit(1);
        }
    }
    (void) fwrite((char *)&dx[ix].jp, sizeof(struct jmp), 1, fj);
    (void) fwrite((char *)&dx[ix].offset, sizeof(dx[ix].offset), 1, fj);
}

```

[0249]

TABLE 2

PRO	XXXXXXXXXXXXXXXXX	(Length = 15 amino acids)
Comparison Protein	XXXXXXXXYYYYYYY	(Length = 12 amino acids)

% amino acid sequence identity = (the number of identically matching amino acid residues between the two polypeptide sequences as determined by ALIGN-2) divided by (the total number of amino acid residues of the PRO polypeptide) = 5 divided by 15 = 33.3%

[0250]

TABLE 3

PRO	XXXXXXXXXX	(Length = 10 amino acids)
Comparison Protein	XXXXXXXXYYYZZYZ	(Length = 15 amino acids)

% amino acid sequence identity = (the number of identically matching amino acid residues between the two polypeptide sequences as determined by ALIGN-2) divided by (the total number of amino acid residues of the PRO polypeptide) = 5 divided by 10 = 50%

[0251]

TABLE 4

PRO-DNA	NNNNNNNNNNNN	(Length = 14 nucleotides)
Comparison DNA	NNNNNLLLLLLLL	(Length = 16 nucleotides)

% nucleic acid sequence identity = (the number of identically matching nucleotides between the two nucleic acid sequences as determined by ALIGN-2) divided by (the total number of nucleotides of the PRO-DNA nucleic acid sequence) = 6 divided by 14 = 42.9%

[0252]

TABLE 5

PRO-DNA	NNNNNNNNNNNN	(Length = 12 nucleotides)
Comparison DNA	NNNLLLVV	(Length = 9 nucleotides)

% nucleic acid sequence identity = (the number of identically matching nucleotides between the two nucleic acid sequences as determined by ALIGN-2) divided by (the total number of nucleotides of the PRO-DNA nucleic acid sequence) = 4 divided by 12 = 33.3%

II. Compositions and Methods of the Invention

A. Full-Length PRO Polypeptides

[0253] The present invention provides newly identified and isolated nucleotide sequences encoding polypeptides referred to in the present application as PRO polypeptides. In particular, cDNAs encoding various PRO polypeptides have been identified and isolated, as disclosed in further detail in the Examples below. It is noted that proteins produced in separate expression rounds may be given different PRO numbers but the UNQ number is unique for any given DNA and the encoded protein, and will not be changed. However, for sake of simplicity, in the present specification the protein encoded by the full length native nucleic acid molecules disclosed herein as well as all further native homologues and variants included in the foregoing definition of PRO, will be referred to as "PRO/number", regardless of their origin or mode of preparation.

[0254] As disclosed in the Examples below, various cDNA clones have been deposited with the ATCC. The actual nucleotide sequences of those clones can readily be determined by the skilled artisan by sequencing of the deposited clone using routine methods in the art. The predicted amino acid sequence can be determined from the nucleotide sequence using routine skill. For the PRO polypeptides and encoding nucleic acids described herein, Applicants have identified what is believed to be the reading frame best identifiable with the sequence information available at the time.

B. PRO Polypeptide Variants

[0255] In addition to the full-length native sequence PRO polypeptides described herein, it is contemplated that PRO variants can be prepared. PRO variants can be prepared by introducing appropriate nucleotide changes into the PRO DNA, and/or by synthesis of the desired PRO polypeptide. Those skilled in the art will appreciate that amino acid changes may alter post-translational processes of the PRO, such as changing the number or position of glycosylation sites or altering the membrane anchoring characteristics.

[0256] Variations in the native full-length sequence PRO or in various domains of the PRO described herein, can be made, for example, using any of the techniques and guidelines for conservative and non-conservative mutations set forth, for instance, in U.S. Pat. No. 5,364,934. Variations may be a substitution, deletion or insertion of one or more codons encoding the PRO that results in a change in the amino acid sequence of the PRO as compared with the native sequence PRO. Optionally the variation is by substitution of at least one amino acid with any other amino acid in one or more of the domains of the PRO. Guidance in determining which amino acid residue may be inserted, substituted or deleted without adversely affecting the desired activity may be found by comparing the sequence of the PRO with that of homologous known protein molecules and minimizing the number of amino acid sequence changes made in regions of high homology. Amino acid substitutions can be the result of replacing one amino acid with another amino acid having similar structural and/or chemical properties, such as the replacement of a leucine with a serine, i.e., conservative amino acid replacements. Insertions or deletions may optionally be in the range of about 1 to 5 amino acids. The variation allowed may be determined by systematically making insertions, deletions or substitutions of amino acids in the sequence and testing the resulting variants for activity exhibited by the full-length or mature native sequence.

[0257] PRO polypeptide fragments are provided herein. Such fragments may be truncated at the N-terminus or C-terminus, or may lack internal residues, for example, when compared with a full length native protein. Certain fragments lack amino acid residues that are not essential for a desired biological activity of the PRO polypeptide.

[0258] PRO fragments may be prepared by any of a number of conventional techniques. Desired peptide fragments may be chemically synthesized. An alternative approach involves generating PRO fragments by enzymatic digestion, e.g., by treating the protein with an enzyme known to cleave proteins at sites defined by particular amino acid residues, or by digesting the DNA with suitable restric-

tion enzymes and isolating the desired fragment. Yet another suitable technique involves isolating and amplifying a DNA fragment encoding a desired polypeptide fragment, by polymerase chain reaction (PCR). Oligonucleotides that define the desired termini of the DNA fragment are employed at the 5' and 3' primers in the PCR. Preferably, PRO polypeptide fragments share at least one biological and/or immunological activity with the native PRO polypeptide disclosed herein.

[0259] In particular embodiments, conservative substitutions of interest are shown in Table 6 under the heading of preferred substitutions. If such substitutions result in a change in biological activity, then more substantial changes, denominated exemplary substitutions in Table 6, or as further described below in reference to amino acid classes, are introduced and the products screened.

TABLE 6

Original Residue	Exemplary Substitutions	Preferred Substitutions
Ala (A)	val; leu; ile	val
Arg (R)	lys; gln; asn	lys
Asn (N)	gln; his; lys; arg	gln
Asp (D)	glu	glu
Cys (C)	ser	ser
Gln (Q)	asn	asn
Glu (E)	asp	asp
Gly (G)	pro; ala	ala
His (H)	asn; gln; lys; arg	arg
Ile (I)	leu; val; met; ala; phe; norleucine	leu
Leu (L)	norleucine; ile; val; met; ala; phe	ile
Lys (K)	arg; gln; asn	arg
Met (M)	leu; phe; ile	leu
Phe (F)	leu; val; ile; ala; tyr	leu
Pro (P)	ala	ala
Ser (S)	thr	thr
Thr (T)	ser	ser
Trp (W)	tyr; phe	tyr
Tyr (Y)	trp; phe; thr; ser	phe
Val (V)	ile; leu; met; phe; ala; norleucine	leu

[0260] Substantial modifications in function or immunological identity of the PRO polypeptide are accomplished by selecting substitutions that differ significantly in their effect on maintaining (a) the structure of the polypeptide backbone in the area of the substitution, for example, as a sheet or helical conformation, (b) the charge or hydrophobicity of the molecule at the target site, or (c) the bulk of the side chain. Naturally occurring residues are divided into groups based on common side-chain properties:

- [0261] (1) hydrophobic: norleucine, met, ala, val, leu, ile;
- [0262] (2) neutral hydrophilic: cys, ser, thr;
- [0263] (3) acidic: asp, glu;
- [0264] (4) basic: asn, gln, his, lys, arg;
- [0265] (5) residues that influence chain orientation: gly, pro; and
- [0266] (6) aromatic: trp, tyr, phe.

[0267] Non-conservative substitutions will entail exchanging a member of one of these classes for another

class. Such substituted residues also may be introduced into the conservative substitution sites or, more preferably, into the remaining (non-conserved) sites.

[0268] The variations can be made using methods known in the art such as oligonucleotide-mediated (site-directed) mutagenesis, alanine scanning, and PCR mutagenesis. Site-directed mutagenesis [Carter et al., Nucl. Acids Res., 13:4331 (1986); Zoller et al., Nucl. Acids Res., 10:6487 (1987)], cassette mutagenesis [Wells et al., Gene, 34:315 (1985)], restriction selection mutagenesis [Wells et al., Philos. Trans. R. Soc. London SerA, 317:415 (1986)] or other known techniques can be performed on the cloned DNA to produce the PRO variant DNA.

[0269] Scanning amino acid analysis can also be employed to identify one or more amino acids along a contiguous sequence. Among the preferred scanning amino acids are relatively small, neutral amino acids. Such amino acids include alanine, glycine, serine, and cysteine. Alanine is typically a preferred scanning amino acid among this group because it eliminates the side-chain beyond the beta-carbon and is less likely to alter the main-chain conformation of the variant [Cunningham and Wells, Science, 244: 1081-1085 (1989)]. Alanine is also typically preferred because it is the most common amino acid. Further, it is frequently found in both buried and exposed positions [Creighton, The Proteins, (W. H. Freeman & Co., N.Y.); Chothia, J. Mol. Biol., 150:1 (1976)]. If alanine substitution does not yield adequate amounts of variant, an isoteric amino acid can be used.

C. Modifications of PRO

[0270] Covalent modifications of PRO are included within the scope of this invention. One type of covalent modification includes reacting targeted amino acid residues of a PRO polypeptide with an organic derivatizing agent that is capable of reacting with selected side chains or the N- or C-terminal residues of the PRO. Derivatization with bifunctional agents is useful, for instance, for crosslinking PRO to a water-insoluble support matrix or surface for use in the method for purifying anti-PRO antibodies, and vice-versa. Commonly used crosslinking agents include, e.g., 1,1-bis (diazooacetyl)-2-phenylethane, glutaraldehyde, N-hydroxysuccinimide esters, for example, esters with 4-azidosalicylic acid, homobifunctional imidoesters, including disuccinimidyl esters such as 3,3'-dithiobis(succinimidylpropionate), bifunctional maleimides such as bis-N-maleimido-1,8-octane and agents such as methyl-3-[(p-azidophenyl)dithio]propioimide.

[0271] Other modifications include deamidation of glutamyl and asparaginy residues to the corresponding glutamyl and aspartyl residues, respectively, hydroxylation of proline and lysine, phosphorylation of hydroxyl groups of seryl or threonyl residues, methylation of the α -amino groups of lysine, arginine, and histidine side chains [T. E. Creighton, Proteins: Structure and Molecular Properties, W. H. Freeman & Co., San Francisco, pp. 79-86 (1983)], acetylation of the N-terminal amine, and amidation of any C-terminal carboxyl group.

[0272] Another type of covalent modification of the PRO polypeptide included within the scope of this invention comprises altering the native glycosylation pattern of the polypeptide. "Altering the native glycosylation pattern" is

intended for purposes herein to mean deleting one or more carbohydrate moieties found in native sequence PRO (either by removing the underlying glycosylation site or by deleting the glycosylation by chemical and/or enzymatic means), and/or adding one or more glycosylation sites that are not present in the native sequence PRO. In addition, the phrase includes qualitative changes in the glycosylation of the native proteins, involving a change in the nature and proportions of the various carbohydrate moieties present.

[0273] Addition of glycosylation sites to the PRO polypeptide may be accomplished by altering the amino acid sequence. The alteration may be made, for example, by the addition of, or substitution by, one or more serine or threonine residues to the native sequence PRO (for O-linked glycosylation sites). The PRO amino acid sequence may optionally be altered through changes at the DNA level, particularly by mutating the DNA encoding the PRO polypeptide at preselected bases such that codons are generated that will translate into the desired amino acids.

[0274] Another means of increasing the number of carbohydrate moieties on the PRO polypeptide is by chemical or enzymatic coupling of glycosides to the polypeptide. Such methods are described in the art, e.g., in WO 87/05330 published Sep. 11, 1987, and in Aplin and Wriston, *CRC Crit. Rev. Biochem.*, pp. 259-306 (1981).

[0275] Removal of carbohydrate moieties present on the PRO polypeptide may be accomplished chemically or enzymatically or by mutational substitution of codons encoding for amino acid residues that serve as targets for glycosylation. Chemical deglycosylation techniques are known in the art and described, for instance, by Hakimuddin, et al., *Arch. Biochem. Biophys.*, 259:52 (1987) and by Edge et al., *Anal.*

[0276] *Biochem.*, 118:131 (1981). Enzymatic cleavage of carbohydrate moieties on polypeptides can be achieved by the use of a variety of endo- and exo-glycosidases as described by Thotakura et al., *Meth. Enzymol.*, 138:350 (1987).

[0277] Another type of covalent modification of PRO comprises linking the PRO polypeptide to one of a variety of nonproteinaceous polymers, e.g., polyethylene glycol (PEG), polypropylene glycol, or polyoxyalkylenes, in the manner set forth in U.S. Pat. Nos. 4,640,835; 4,496,689; 4,301,144; 4,670,417; 4,791,192 or 4,179,337.

[0278] The PRO of the present invention may also be modified in a way to form a chimeric molecule comprising PRO fused to another, heterologous polypeptide or amino acid sequence.

[0279] In one embodiment, such a chimeric molecule comprises a fusion of the PRO with a tag polypeptide which provides an epitope to which an anti-tag antibody can selectively bind. The epitope tag is generally placed at the amino- or carboxyl-terminus of the PRO. The presence of such epitope-tagged forms of the PRO can be detected using an antibody against the tag polypeptide. Also, provision of the epitope tag enables the PRO to be readily purified by affinity purification using an anti-tag antibody or another type of affinity matrix that binds to the epitope tag. Various tag polypeptides and their respective antibodies are well known in the art. Examples include poly-histidine (poly-his) or poly-histidine-glycine (poly-his-gly) tags; the flu HA tag polypeptide and its antibody 12CA5 [Field et al., *Mol. Cell.*

Biol., 8:2159-2165 (1988)]; the c-myc tag and the 8F9, 3C7, 6E10, G4, B7 and 9E10 antibodies thereto [Evan et al., *Molecular and Cellular Biology*, 5:3610-3616 (1985)]; and the Herpes Simplex virus glycoprotein D (gD) tag and its antibody [Paborsky et al., *Protein Engineering*, 3(6):547-553 (1990)]. Other tag polypeptides include the Flag-peptide [Hopp et al., *BioTechnology*, 6:1204-1210 (1988)]; the KT3 epitope peptide [Martin et al., *Science*, 255:192-194 (1992)]; an α -tubulin epitope peptide [Skinner et al., *J. Biol. Chem.*, 266:15163-15166 (1991)]; and the T7 gene 10 protein peptide tag [Lutz-Freyermuth et al., *Proc. Natl. Acad. Sci. USA*, 87:6393-6397 (1990)].

[0280] In an alternative embodiment, the chimeric molecule may comprise a fusion of the PRO with an immunoglobulin or a particular region of an immunoglobulin. For a bivalent form of the chimeric molecule (also referred to as an "immunoadhesin"), such a fusion could be to the Fc region of an IgG molecule. The Ig fusions preferably include the substitution of a soluble (transmembrane domain deleted or inactivated) form of a PRO polypeptide in place of at least one variable region within an Ig molecule. In a particularly preferred embodiment, the immunoglobulin fusion includes the hinge, CH2 and CH3, or the hinge, CH1, CH2 and CH3 regions of an IgG1 molecule. For the production of immunoglobulin fusions see also U.S. Pat. No. 5,428,130 issued Jun. 27, 1995.

D. Preparation of PRO

[0281] The description below relates primarily to production of PRO by culturing cells transformed or transfected with a vector containing PRO nucleic acid. It is, of course, contemplated that alternative methods, which are well known in the art, may be employed to prepare PRO. For instance, the PRO sequence, or portions thereof, may be produced by direct peptide synthesis using solid-phase techniques [see, e.g., Stewart et al., *Solid-Phase Peptide Synthesis*, W. H. Freeman Co., San Francisco, Calif. (1969); Merrifield, *J. Am. Chem. Soc.*, 85:2149-2154 (1963)]. In vitro protein synthesis may be performed using manual techniques or by automation. Automated synthesis may be accomplished, for instance, using an Applied Biosystems Peptide Synthesizer (Foster City, Calif.) using manufacturer's instructions. Various portions of the PRO may be chemically synthesized separately and combined using chemical or enzymatic methods to produce the full-length PRO.

1. Isolation of DNA Encoding PRO

[0282] DNA encoding PRO may be obtained from a cDNA library prepared from tissue believed to possess the PRO mRNA and to express it at a detectable level. Accordingly, human PRO DNA can be conveniently obtained from a cDNA library prepared from human tissue, such as described in the Examples. The PRO-encoding gene may also be obtained from a genomic library or by known synthetic procedures (e.g., automated nucleic acid synthesis).

[0283] Libraries can be screened with probes (such as antibodies to the PRO or oligonucleotides of at least about 20-80 bases) designed to identify the gene of interest or the protein encoded by it. Screening the cDNA or genomic library with the selected probe may be conducted using

standard procedures, such as described in Sambrook et al., *Molecular Cloning: A Laboratory Manual* (New York: Cold Spring Harbor Laboratory Press, 1989). An alternative means to isolate the gene encoding PRO is to use PCR methodology [Sambrook et al., supra; Dieffenbach et al., *PCR Primer: A Laboratory Manual* (Cold Spring Harbor Laboratory Press, 1995)].

[0284] The Examples below describe techniques for screening a cDNA library. The oligonucleotide sequences selected as probes should be of sufficient length and sufficiently unambiguous that false positives are minimized. The oligonucleotide is preferably labeled such that it can be detected upon hybridization to DNA in the library being screened. Methods of labeling are well known in the art, and include the use of radiolabels like ^{32}P -labeled ATP, biotinylation or enzyme labeling. Hybridization conditions, including moderate stringency and high stringency, are provided in Sambrook et al., supra.

[0285] Sequences identified in such library screening methods can be compared and aligned to other known sequences deposited and available in public databases such as GenBank or other private sequence databases. Sequence identity (at either the amino acid or nucleotide level) within defined regions of the molecule or across the full-length sequence can be determined using methods known in the art and as described herein.

[0286] Nucleic acid having protein coding sequence may be obtained by screening selected cDNA or genomic libraries using the deduced amino acid sequence disclosed herein for the first time, and, if necessary, using conventional primer extension procedures as described in Sambrook et al., supra, to detect precursors and processing intermediates of mRNA that may not have been reverse-transcribed into cDNA.

2. Selection and Transformation of Host Cells

[0287] Host cells are transfected or transformed with expression or cloning vectors described herein for PRO production and cultured in conventional nutrient media modified as appropriate for inducing promoters, selecting transformants, or amplifying the genes encoding the desired sequences. The culture conditions, such as media, temperature, pH and the like, can be selected by the skilled artisan without undue experimentation. In general, principles, protocols, and practical techniques for maximizing the productivity of cell cultures can be found in *Mammalian Cell Biotechnology: a Practical Approach*, M. Butler, ed. (IRL Press, 1991) and Sambrook et al., supra.

[0288] Methods of eukaryotic cell transfection and prokaryotic cell transformation are known to the ordinarily skilled artisan, for example, CaCl_2 , CaPO_4 , liposome-mediated and electroporation. Depending on the host cell used, transformation is performed using standard techniques appropriate to such cells. The calcium treatment employing calcium chloride, as described in Sambrook et al., supra, or electroporation is generally used for prokaryotes. Infection with *Agrobacterium tumefaciens* is used for transformation of certain plant cells, as described by Shaw et al., *Gene*, 23:315 (1983) and WO 89/05859 published Jun. 29, 1989. For mammalian cells without such cell walls, the calcium phosphate precipitation method of Graham and van der Eb, *Virology*, 52:456-457 (1978) can be employed. General

aspects of mammalian cell host system transfections have been described in U.S. Pat. No. 4,399,216. Transformations into yeast are typically carried out according to the method of Van Solingen et al., *J. Bact.*, 130:946 (1977) and Hsiao et al., *Proc. Natl. Acad. Sci. (USA)*, 76:3829 (1979). However, other methods for introducing DNA into cells, such as by nuclear microinjection, electroporation, bacterial protoplast fusion with intact cells, or polycations, e.g., polybrene, polyornithine, may also be used. For various techniques for transforming mammalian cells, see Keown et al., *Methods in Enzymology*, 185:527-537 (1990) and Mansour et al., *Nature*, 336:348-352 (1988).

[0289] Suitable host cells for cloning or expressing the DNA in the vectors herein include prokaryote, yeast, or higher eukaryote cells. Suitable prokaryotes include but are not limited to eubacteria, such as Gram-negative or Gram-positive organisms, for example, Enterobacteriaceae such as *E. coli*. Various *E. coli* strains are publicly available, such as *E. coli* K12 strain MM294 (ATCC 31,446); *E. coli* X1776 (ATCC 31,537); *E. coli* strain W3110 (ATCC 27,325) and K5 772 (ATCC 53,635). Other suitable prokaryotic host cells include Enterobacteriaceae such as Escherichia, e.g., *E. coli*, Enterobacter, Erwinia, Klebsiella, Proteus, Salmonella, e.g., *Salmonella typhimurium*, Serratia, e.g., *Serratia marcescans*, and Shigella, as well as Bacilli such as *B. subtilis* and *B. licheniformis* (e.g., *B. licheniformis* 41P disclosed in DD 266,710 published Apr. 12, 1989), Pseudomonas such as *P. aeruginosa*, and Streptomyces. These examples are illustrative rather than limiting. Strain W3110 is one particularly preferred host or parent host because it is a common host strain for recombinant DNA product fermentations. Preferably, the host cell secretes minimal amounts of proteolytic enzymes. For example, strain W3110 may be modified to effect a genetic mutation in the genes encoding proteins endogenous to the host, with examples of such hosts including *E. coli* W3110 strain 1A2, which has the complete genotype tonA; *E. coli* W3110 strain 9E4, which has the complete genotype tonA ptr3; *E. coli* W3110 strain 27C7 (ATCC 55,244), which has the complete genotype tonA ptr3 phoA E15 (argF-lac)169 degP ompT kan^r; *E. coli* W3110 strain 37D6, which has the complete genotype tonA ptr3 phoA E15 (argF-lac)169 degP ompT rbs7 ilvG kan^r; *E. coli* W3110 strain 40B4, which is strain 37D6 with a non-kanamycin resistant degP deletion mutation; and an *E. coli* strain having mutant periplasmic protease disclosed in U.S. Pat. No. 4,946,783 issued Aug. 7, 1990. Alternatively, in vitro methods of cloning, e.g., PCR or other nucleic acid polymerase reactions, are suitable.

[0290] In addition to prokaryotes, eukaryotic microbes such as filamentous fungi or yeast are suitable cloning or expression hosts for PRO-encoding vectors. *Saccharomyces cerevisiae* is a commonly used lower eukaryotic host microorganism. Others include *Schizosaccharomyces pombe* (Beach and Nurse, *Nature*, 290: 140 [1981]; EP 139,383 published May 2, 1985); Kluyveromyces hosts (U.S. Pat. No. 4,943,529; Fleer et al., *Bio/Technology*, 9:968-975 (1991)) such as, e.g., *K. lactis* (MW98-8C, CBS683, CBS4574; Louvencourt et al., *J. Bacteriol.*, 154(2):737-742 [1983]), *K. fragilis* (ATCC 12,424), *K. bulgaricus* (ATCC 16,045), *K. wickerhamii* (ATCC 24,178), *K. waltii* (ATCC 56,500), *K. drosophilum* (ATCC 36,906; Van den Berg et al., *Bio/Technology*, 8:135 (1990)), *K. thermotolerans*, and *K. marxianus*; yarrowia (EP 402,226); *Pichia pastoris* (EP 183,070; Sreekrishna et al., *J. Basic Microbiol.*, 28:265-278

[1988]); *Candida*; *Trichoderma reesia* (EP 244,234); *Neurospora crassa* (Case et al., Proc. Natl. Acad. Sci. USA, 76:5259-5263 [1979]); Schwanniomycetes such as *Schwanniomycetes occidentalis* (EP 394,538 published Oct. 31, 1990); and filamentous fungi such as, e.g., *Neurospora*, *Penicillium*, *Tolypocladium* (WO 91/000357 published Jan. 10, 1991), and *Aspergillus* hosts such as *A. nidulans* (Ballance et al., Biochem. Biophys. Res. Commun., 112:284-289 [1983]; Tilburn et al., Gene, 26:205-221 [1983]; Yelton et al., Proc. Natl. Acad. Sci. USA, 81: 1470-1474 [1984]) and *A. niger* (Kelly and Hynes, EMBO J., 4:475-479 [1985]). Methylophilic yeasts are suitable herein and include, but are not limited to, yeast capable of growth on methanol selected from the genera consisting of *Hansenula*, *Candida*, *Kloeckera*, *Pichia*, *Saccharomyces*, *Torulopsis*, and *Rhodotorula*. A list of specific species that are exemplary of this class of yeasts may be found in C. Anthony, The Biochemistry of Methylophilic yeasts, 269 (1982).

[0291] Suitable host cells for the expression of glycosylated PRO are derived from multicellular organisms. Examples of invertebrate cells include insect cells such as *Drosophila* S2 and *Spodoptera* Sf9, as well as plant cells. Examples of useful mammalian host cell lines include Chinese hamster ovary (CHO) and COS cells. More specific examples include monkey kidney CV1 line transformed by SV40 (COS-7, ATCC CRL 1651); human embryonic kidney line (293 or 293 cells subcloned for growth in suspension culture, Graham et al., J. Gen. Virol., 36:59 (1977)); Chinese hamster ovary cells/DHFR (CHO, Urlaub and Chasin, Proc. Natl. Acad. Sci. USA, 77:4216 (1980)); mouse sertoli cells (TM4, Mather, Biol. Reprod., 23:243-251 (1980)); human lung cells (W138, ATCC CCL 75); human liver cells (Hep G2, HB 8065); and mouse mammary tumor (MMT 060562, ATCC CCL51). The selection of the appropriate host cell is deemed to be within the skill in the art.

3. Selection and Use of a Replicable Vector

[0292] The nucleic acid (e.g., cDNA or genomic DNA) encoding PRO may be inserted into a replicable vector for cloning (amplification of the DNA) or for expression. Various vectors are publicly available. The vector may, for example, be in the form of a plasmid, cosmid, viral particle, or phage. The appropriate nucleic acid sequence may be inserted into the vector by a variety of procedures. In general, DNA is inserted into an appropriate restriction endonuclease site(s) using techniques known in the art. Vector components generally include, but are not limited to, one or more of a signal sequence, an origin of replication, one or more marker genes, an enhancer element, a promoter, and a transcription termination sequence. Construction of suitable vectors containing one or more of these components employs standard ligation techniques which are known to the skilled artisan.

[0293] The PRO may be produced recombinantly not only directly, but also as a fusion polypeptide with a heterologous polypeptide, which may be a signal sequence or other polypeptide having a specific cleavage site at the N-terminus of the mature protein or polypeptide. In general, the signal sequence may be a component of the vector, or it may be a part of the PRO-encoding DNA that is inserted into the vector. The signal sequence may be a prokaryotic signal sequence selected, for example, from the group of the alkaline phosphatase, penicillinase, lpp, or heat-stable

enterotoxin II leaders. For yeast secretion the signal sequence may be, e.g., the yeast invertase leader, alpha factor leader (including *Saccharomyces* and *Kluyveromyces* α -factor leaders, the latter described in U.S. Pat. No. 5,010, 182), or acid phosphatase leader, the *C. albicans* glucoamylase leader (EP 362,179 published Apr. 4, 1990), or the signal described in WO 90/13646 published Nov. 15, 1990. In mammalian cell expression, mammalian signal sequences may be used to direct secretion of the protein, such as signal sequences from secreted polypeptides of the same or related species, as well as viral secretory leaders.

[0294] Both expression and cloning vectors contain a nucleic acid sequence that enables the vector to replicate in one or more selected host cells. Such sequences are well known for a variety of bacteria, yeast, and viruses. The origin of replication from the plasmid pBR322 is suitable for most Gram-negative bacteria, the 2μ plasmid origin is suitable for yeast, and various viral origins (SV40, polyoma, adenovirus, VSV or BPV) are useful for cloning vectors in mammalian cells.

[0295] Expression and cloning vectors will typically contain a selection gene, also termed a selectable marker. Typical selection genes encode proteins that (a) confer resistance to antibiotics or other toxins, e.g., ampicillin, neomycin, methotrexate, or tetracycline, (b) complement auxotrophic deficiencies, or (c) supply critical nutrients not available from complex media, e.g., the gene encoding D-alanine racemase for *Bacilli*.

[0296] An example of suitable selectable markers for mammalian cells are those that enable the identification of cells competent to take up the PRO-encoding nucleic acid, such as DHFR or thymidine kinase. An appropriate host cell when wild-type DHFR is employed is the CHO cell line deficient in DHFR activity, prepared and propagated as described by Urlaub et al., Proc. Natl. Acad. Sci. USA, 77:4216 (1980). A suitable selection gene for use in yeast is the trp 1 gene present in the yeast plasmid YRp7 [Stinchcomb et al., Nature, 282:39 (1979); Kingsman et al., Gene, 7:141 (1979); Tschemper et al., Gene, 10:157 (1980)]. The trp 1 gene provides a selection marker for a mutant strain of yeast lacking the ability to grow in tryptophan, for example, ATCC No. 44076 or PEP4-1 [Jones, Genetics, 85:12 (1977)].

[0297] Expression and cloning vectors usually contain a promoter operably linked to the PRO-encoding nucleic acid sequence to direct mRNA synthesis. Promoters recognized by a variety of potential host cells are well known. Promoters suitable for use with prokaryotic hosts include the β -lactamase and lactose promoter systems [Chang et al., Nature, 275:615 (1978); Goeddel et al., Nature, 281:544 (1979)], alkaline phosphatase, a tryptophan (trp) promoter system [Goeddel, Nucleic Acids Res., 8:4057 (1980); EP 36,776], and hybrid promoters such as the tac promoter [deBoer et al., Proc. Natl. Acad. Sci. USA, 80:21-25 (1983)]. Promoters for use in bacterial systems also will contain a Shine-Dalgarno (S.D.) sequence operably linked to the DNA encoding PRO.

[0298] Examples of suitable promoting sequences for use with yeast hosts include the promoters for 3-phosphoglycerate kinase [Hitzeman et al., J. Biol. Chem., 255:2073 (1980)] or other glycolytic enzymes [Hess et al., J. Adv. Enzyme Reg., 7:149 (1968); Holland, Biochemistry,

17:4900 (1978)], such as enolase, glyceraldehyde-3-phosphate dehydrogenase, hexokinase, pyruvate decarboxylase, phosphofructokinase, glucose-6-phosphate isomerase, 3-phosphoglycerate mutase, pyruvate kinase, triosephosphate isomerase, phosphoglucose isomerase, and glucokinase.

[0299] Other yeast promoters, which are inducible promoters having the additional advantage of transcription controlled by growth conditions, are the promoter regions for alcohol dehydrogenase 2, isocytochrome C, acid phosphatase, degradative enzymes associated with nitrogen metabolism, metallothionein, glyceraldehyde-3-phosphate dehydrogenase, and enzymes responsible for maltose and galactose utilization. Suitable vectors and promoters for use in yeast expression are further described in EP 73,657.

[0300] PRO transcription from vectors in mammalian host cells is controlled, for example, by promoters obtained from the genomes of viruses such as polyoma virus, fowlpox virus (UK 2,211,504 published Jul. 5, 1989), adenovirus (such as Adenovirus 2), bovine papilloma virus, avian sarcoma virus, cytomegalovirus, a retrovirus, hepatitis-B virus and Simian Virus 40 (SV40), from heterologous mammalian promoters, e.g., the actin promoter or an immunoglobulin promoter, and from heat-shock promoters, provided such promoters are compatible with the host cell systems.

[0301] Transcription of a DNA encoding the PRO by higher eukaryotes may be increased by inserting an enhancer sequence into the vector. Enhancers are cis-acting elements of DNA, usually about from 10 to 300 bp, that act on a promoter to increase its transcription. Many enhancer sequences are now known from mammalian genes (globin, elastase, albumin, α -fetoprotein, and insulin). Typically, however, one will use an enhancer from a eukaryotic cell virus. Examples include the SV40 enhancer on the late side of the replication origin (bp 100-270), the cytomegalovirus early promoter enhancer, the polyoma enhancer on the late side of the replication origin, and adenovirus enhancers. The enhancer may be spliced into the vector at a position 5' or 3' to the PRO coding sequence, but is preferably located at a site 5' from the promoter.

[0302] Expression vectors used in eukaryotic host cells (yeast, fungi, insect, plant, animal, human, or nucleated cells from other multicellular organisms) will also contain sequences necessary for the termination of transcription and for stabilizing the mRNA. Such sequences are commonly available from the 5' and, occasionally 3', untranslated regions of eukaryotic or viral DNAs or cDNAs. These regions contain nucleotide segments transcribed as polyadenylated fragments in the untranslated portion of the mRNA encoding PRO.

[0303] Still other methods, vectors, and host cells suitable for adaptation to the synthesis of PRO in recombinant vertebrate cell culture are described in Gething et al., *Nature*, 293:620-625 (1981); Mantei et al., *Nature*, 281:40-46 (1979); EP 117,060; and EP 117,058.

4. Detecting Gene Amplification/Expression

[0304] Gene amplification and/or expression may be measured in a sample directly, for example, by conventional Southern blotting, Northern blotting to quantitate the transcription of mRNA [Thomas, *Proc. Natl. Acad. Sci. USA*,

77:5201(1980)], dot blotting (DNA analysis), or in situ hybridization, using an appropriately labeled probe, based on the sequences provided herein. Alternatively, antibodies may be employed that can recognize specific duplexes, including DNA duplexes, RNA duplexes, and DNA-RNA hybrid duplexes or DNA-protein duplexes. The antibodies in turn may be labeled and the assay may be carried out where the duplex is bound to a surface, so that upon the formation of duplex on the surface, the presence of antibody bound to the duplex can be detected.

[0305] Gene expression, alternatively, may be measured by immunological methods, such as immunohistochemical staining of cells or tissue sections and assay of cell culture or body fluids, to quantitate directly the expression of gene product. Antibodies useful for immunohistochemical staining and/or assay of sample fluids may be either monoclonal or polyclonal, and may be prepared in any mammal. Conveniently, the antibodies may be prepared against a native sequence PRO polypeptide or against a synthetic peptide based on the DNA sequences provided herein or against exogenous sequence fused to PRO DNA and encoding a specific antibody epitope.

5. Purification of Polypeptide

[0306] Forms of PRO may be recovered from culture medium or from host cell lysates. If membrane-bound, it can be released from the membrane using a suitable detergent solution (e.g. Triton-X 100) or by enzymatic cleavage. Cells employed in expression of PRO can be disrupted by various physical or chemical means, such as freeze-thaw cycling, sonication, mechanical disruption, or cell lysing agents.

[0307] It may be desired to purify PRO from recombinant cell proteins or polypeptides. The following procedures are exemplary of suitable purification procedures: by fractionation on an ion-exchange column; ethanol precipitation; reverse phase HPLC; chromatography on silica or on a cation-exchange resin such as DEAE; chromatofocusing; SDS-PAGE; ammonium sulfate precipitation; gel filtration using, for example, Sephadex G-75; protein A Sepharose columns to remove contaminants such as IgG; and metal chelating columns to bind epitope-tagged forms of the PRO. Various methods of protein purification may be employed and such methods are known in the art and described for example in Deutscher, *Methods in Enzymology*, 182 (1990); Scopes, *Protein Purification: Principles and Practice*, Springer-Verlag, New York (1982). The purification step(s) selected will depend, for example, on the nature of the production process used and the particular PRO produced.

E. Uses for PRO

[0308] Nucleotide sequences (or their complement) encoding PRO have various applications in the art of molecular biology, including uses as hybridization probes, in chromosome and gene mapping and in the generation of anti-sense RNA and DNA. PRO nucleic acid will also be useful for the preparation of PRO polypeptides by the recombinant techniques described herein.

[0309] The full-length native sequence PRO gene, or portions thereof, may be used as hybridization probes for a cDNA library to isolate the full-length PRO cDNA or to isolate still other cDNAs (for instance, those encoding naturally-occurring variants of PRO or PRO from other

species) which have a desired sequence identity to the native PRO sequence disclosed herein. Optionally, the length of the probes will be about 20 to about 50 bases. The hybridization probes may be derived from at least partially novel regions of the full length native nucleotide sequence wherein those regions may be determined without undue experimentation or from genomic sequences including promoters, enhancer elements and introns of native sequence PRO. By way of example, a screening method will comprise isolating the coding region of the PRO gene using the known DNA sequence to synthesize a selected probe of about 40 bases. Hybridization probes may be labeled by a variety of labels, including radionucleotides such as ^{32}P or ^{35}S , or enzymatic labels such as alkaline phosphatase coupled to the probe via avidin/biotin coupling systems. Labeled probes having a sequence complementary to that of the PRO gene of the present invention can be used to screen libraries of human cDNA, genomic DNA or mRNA to determine which members of such libraries the probe hybridizes to. Hybridization techniques are described in further detail in the Examples below.

[0310] Any EST sequences disclosed in the present application may similarly be employed as probes, using the methods disclosed herein.

[0311] Other useful fragments of the PRO nucleic acids include antisense or sense oligonucleotides comprising a single-stranded nucleic acid sequence (either RNA or DNA) capable of binding to target PRO mRNA (sense) or PRO DNA (antisense) sequences. Antisense or sense oligonucleotides, according to the present invention, comprise a fragment of the coding region of PRO DNA. Such a fragment generally comprises at least about 14 nucleotides, preferably from about 14 to 30 nucleotides. The ability to derive an antisense or a sense oligonucleotide, based upon a cDNA sequence encoding a given protein is described in, for example, Stein and Cohen (Cancer Res. 48:2659, 1988) and van der Krol et al. (BioTechniques 6:958, 1988).

[0312] Binding of antisense or sense oligonucleotides to target nucleic acid sequences results in the formation of duplexes that block transcription or translation of the target sequence by one of several means, including enhanced degradation of the duplexes, premature termination of transcription or translation, or by other means. The antisense oligonucleotides thus may be used to block expression of PRO proteins. Antisense or sense oligonucleotides further comprise oligonucleotides having modified sugar-phosphodiester backbones (or other sugar linkages, such as those described in WO 91/06629) and wherein such sugar linkages are resistant to endogenous nucleases. Such oligonucleotides with resistant sugar linkages are stable in vivo (i.e., capable of resisting enzymatic degradation) but retain sequence specificity to be able to bind to target nucleotide sequences.

[0313] Other examples of sense or antisense oligonucleotides include those oligonucleotides which are covalently linked to organic moieties, such as those described in WO 90/10048, and other moieties that increases affinity of the oligonucleotide for a target nucleic acid sequence, such as poly-(L-lysine). Further still, intercalating agents, such as ellipticine, and alkylating agents or metal may be attached to sense or antisense oligonucleotides to modify binding specificities of the antisense or sense oligonucleotide for the target nucleotide sequence.

[0314] Antisense or sense oligonucleotides may be introduced into a cell containing the target nucleic acid sequence by any gene transfer method, including, for example, CaPO_4 -mediated DNA transfection, electroporation, or by using gene transfer vectors such as Epstein-Barr virus. In a preferred procedure, an antisense or sense oligonucleotide is inserted into a suitable retroviral vector. A cell containing the target nucleic acid sequence is contacted with the recombinant retroviral vector, either in vivo or ex vivo. Suitable retroviral vectors include, but are not limited to, those derived from the murine retrovirus M-MuLV, N2 (a retrovirus derived from M-MuLV), or the double copy vectors designated DCT5A, DCT5B and DCT5C (see WO 90/13641).

[0315] Sense or antisense oligonucleotides also may be introduced into a cell containing the target nucleotide sequence by formation of a conjugate with a ligand binding molecule, as described in WO 91/04753. Suitable ligand binding molecules include, but are not limited to, cell surface receptors, growth factors, other cytokines, or other ligands that bind to cell surface receptors. Preferably, conjugation of the ligand binding molecule does not substantially interfere with the ability of the ligand binding molecule to bind to its corresponding molecule or receptor, or block entry of the sense or antisense oligonucleotide or its conjugated version into the cell.

[0316] Alternatively, a sense or an antisense oligonucleotide may be introduced into a containing the target nucleic acid sequence by formation of an oligonucleotide-lipid complex, as described in WO 90/10448. The sense or antisense oligonucleotide-lipid complex is preferably dissociated within the cell by an endogenous lipase.

[0317] Antisense or sense RNA or DNA molecules are generally at least about 5 bases in length, about 10 bases in length, about 15 bases in length, about 20 bases in length, about 25 bases in length, about 30 bases in length, about 35 bases in length, about 40 bases in length, about 45 bases in length, about 50 bases in length, about 55 bases in length, about 60 bases in length, about 65 bases in length, about 70 bases in length, about 75 bases in length, about 80 bases in length, about 85 bases in length, about 90 bases in length, about 95 bases in length, about 100 bases in length, or more.

[0318] The probes may also be employed in PCR techniques to generate a pool of sequences for identification of closely related PRO coding sequences.

[0319] Nucleotide sequences encoding a PRO can also be used to construct hybridization probes for mapping the gene which encodes that PRO and for the genetic analysis of individuals with genetic disorders. The nucleotide sequences provided herein may be mapped to a chromosome and specific regions of a chromosome using known techniques, such as in situ hybridization, linkage analysis against known chromosomal markers, and hybridization screening with libraries.

[0320] When the coding sequences for PRO encode a protein which binds to another protein (example, where the PRO is a receptor), the PRO can be used in assays to identify the other proteins or molecules involved in the binding interaction. By such methods, inhibitors of the receptor/ligand binding interaction can be identified. Proteins involved in such binding interactions can also be used to

screen for peptide or small molecule inhibitors or agonists of the binding interaction. Also, the receptor PRO can be used to isolate correlative ligand(s). Screening assays can be designed to find lead compounds that mimic the biological activity of a native PRO or a receptor for PRO. Such screening assays will include assays amenable to high-throughput screening of chemical libraries, making them particularly suitable for identifying small molecule drug candidates. Small molecules contemplated include synthetic organic or inorganic compounds. The assays can be performed in a variety of formats, including protein-protein binding assays, biochemical screening assays, immunoassays and cell based assays, which are well characterized in the art.

[0321] Nucleic acids which encode PRO or its modified forms can also be used to generate either transgenic animals or “knock out” animals which, in turn, are useful in the development and screening of therapeutically useful reagents. A transgenic animal (e.g., a mouse or rat) is an animal having cells that contain a transgene, which transgene was introduced into the animal or an ancestor of the animal at a prenatal, e.g., an embryonic stage. A transgene is a DNA which is integrated into the genome of a cell from which a transgenic animal develops. In one embodiment, cDNA encoding PRO can be used to clone genomic DNA encoding PRO in accordance with established techniques and the genomic sequences used to generate transgenic animals that contain cells which express DNA encoding PRO. Methods for generating transgenic animals, particularly animals such as mice or rats, have become conventional in the art and are described, for example, in U.S. Pat. Nos. 4,736,866 and 4,870,009. Typically, particular cells would be targeted for PRO transgene incorporation with tissue-specific enhancers. Transgenic animals that include a copy of a transgene encoding PRO introduced into the germ line of the animal at an embryonic stage can be used to examine the effect of increased expression of DNA encoding PRO. Such animals can be used as tester animals for reagents thought to confer protection from, for example, pathological conditions associated with its overexpression. In accordance with this facet of the invention, an animal is treated with the reagent and a reduced incidence of the pathological condition, compared to untreated animals bearing the transgene, would indicate a potential therapeutic intervention for the pathological condition.

[0322] Alternatively, non-human homologues of PRO can be used to construct a PRO “knock out” animal which has a defective or altered gene encoding PRO as a result of homologous recombination between the endogenous gene encoding PRO and altered genomic DNA encoding PRO introduced into an embryonic stem cell of the animal. For example, cDNA encoding PRO can be used to clone genomic DNA encoding PRO in accordance with established techniques. A portion of the genomic DNA encoding PRO can be deleted or replaced with another gene, such as a gene encoding a selectable marker which can be used to monitor integration. Typically, several kilobases of unaltered flanking DNA (both at the 5' and 3' ends) are included in the vector [see e.g., Thomas and Capecchi, *Cell*, 51:503 (1987) for a description of homologous recombination vectors]. The vector is introduced into an embryonic stem cell line by electroporation and cells in which the introduced DNA has homologously recombined with the endogenous DNA are selected [see e.g., Li et al., *Cell*, 69:915 (1992)]. The

selected cells are then injected into a blastocyst of an animal (e.g., a mouse or rat) to form aggregation chimeras [see e.g., Bradley, in *Teratocarcinomas and Embryonic Stem Cells: A Practical Approach*, E. J. Robertson, ed. (IRL, Oxford, 1987), pp. 113-152]. A chimeric embryo can then be implanted into a suitable pseudopregnant female foster animal and the embryo brought to term to create a “knock out” animal. Progeny harboring the homologously recombined DNA in their germ cells can be identified by standard techniques and used to breed animals in which all cells of the animal contain the homologously recombined DNA. Knock-out animals can be characterized for instance, for their ability to defend against certain pathological conditions and for their development of pathological conditions due to absence of the PRO polypeptide.

[0323] Nucleic acid encoding the PRO polypeptides may also be used in gene therapy. In gene therapy applications, genes are introduced into cells in order to achieve in vivo synthesis of a therapeutically effective genetic product, for example for replacement of a defective gene. “Gene therapy” includes both conventional gene therapy where a lasting effect is achieved by a single treatment, and the administration of gene therapeutic agents, which involves the one time or repeated administration of a therapeutically effective DNA or mRNA. Antisense RNAs and DNAs can be used as therapeutic agents for blocking the expression of certain genes in vivo. It has already been shown that short antisense oligonucleotides can be imported into cells where they act as inhibitors, despite their low intracellular concentrations caused by their restricted uptake by the cell membrane. (Zamecnik et al., *Proc. Natl. Acad. Sci. USA* 83:4143-4146 [1986]). The oligonucleotides can be modified to enhance their uptake, e.g. by substituting their negatively charged phosphodiester groups by uncharged groups.

[0324] There are a variety of techniques available for introducing nucleic acids into viable cells. The techniques vary depending upon whether the nucleic acid is transferred into cultured cells in vitro, or in vivo in the cells of the intended host. Techniques suitable for the transfer of nucleic acid into mammalian cells in vitro include the use of liposomes, electroporation, microinjection, cell fusion, DEAE-dextran, the calcium phosphate precipitation method, etc. The currently preferred in vivo gene transfer techniques include transfection with viral (typically retroviral) vectors and viral coat protein-liposome mediated transfection (Dzau et al., *Trends in Biotechnology* 11, 205-210 [1993]). In some situations it is desirable to provide the nucleic acid source with an agent that targets the target cells, such as an antibody specific for a cell surface membrane protein or the target cell, a ligand for a receptor on the target cell, etc. Where liposomes are employed, proteins which bind to a cell surface membrane protein associated with endocytosis may be used for targeting and/or to facilitate uptake, e.g. capsid proteins or fragments thereof tropic for a particular cell type, antibodies for proteins which undergo internalization in cycling, proteins that target intracellular localization and enhance intracellular half-life. The technique of receptor-mediated endocytosis is described, for example, by Wu et al., *J. Biol. Chem.* 262, 4429-4432 (1987); and Wagner et al., *Proc. Natl. Acad. Sci. USA* 87, 3410-3414 (1990). For review of gene marking and gene therapy protocols see Anderson et al., *Science* 256, 808-813 (1992).

[0325] The PRO polypeptides described herein may also be employed as molecular weight markers for protein electrophoresis purposes and the isolated nucleic acid sequences may be used for recombinantly expressing those markers.

[0326] The nucleic acid molecules encoding the PRO polypeptides or fragments thereof described herein are useful for chromosome identification. In this regard, there exists an ongoing need to identify new chromosome markers, since relatively few chromosome marking reagents, based upon actual sequence data are presently available. Each PRO nucleic acid molecule of the present invention can be used as a chromosome marker.

[0327] The PRO polypeptides and nucleic acid molecules of the present invention may also be used diagnostically for tissue typing, wherein the PRO polypeptides of the present invention may be differentially expressed in one tissue as compared to another, preferably in a diseased tissue as compared to a normal tissue of the same tissue type. PRO nucleic acid molecules will find use for generating probes for PCR, Northern analysis, Southern analysis and Western analysis.

[0328] The PRO polypeptides described herein may also be employed as therapeutic agents. The PRO polypeptides of the present invention can be formulated according to known methods to prepare pharmaceutically useful compositions, whereby the PRO product hereof is combined in admixture with a pharmaceutically acceptable carrier vehicle. Therapeutic formulations are prepared for storage by mixing the active ingredient having the desired degree of purity with optional physiologically acceptable carriers, excipients or stabilizers (Remington's Pharmaceutical Sciences 16th edition, Osol, A. Ed. (1980)), in the form of lyophilized formulations or aqueous solutions. Acceptable carriers, excipients or stabilizers are nontoxic to recipients at the dosages and concentrations employed, and include buffers such as phosphate, citrate and other organic acids; antioxidants including ascorbic acid; low molecular weight (less than about 10 residues) polypeptides; proteins, such as serum albumin, gelatin or immunoglobulins; hydrophilic polymers such as polyvinylpyrrolidone, amino acids such as glycine, glutamine, asparagine, arginine or lysine; monosaccharides, disaccharides and other carbohydrates including glucose, mannose, or dextrans; chelating agents such as EDTA; sugar alcohols such as mannitol or sorbitol; salt-forming counterions such as sodium; and/or nonionic surfactants such as TWEENTM, PLURONICSTM or PEG.

[0329] The formulations to be used for in vivo administration must be sterile. This is readily accomplished by filtration through sterile filtration membranes, prior to or following lyophilization and reconstitution.

[0330] Therapeutic compositions herein generally are placed into a container having a sterile access port, for example, an intravenous solution bag or vial having a stopper pierceable by a hypodermic injection needle.

[0331] The route of administration is in accord with known methods, e.g. injection or infusion by intravenous, intraperitoneal, intracerebral, intramuscular, intraocular, intraarterial or intralesional routes, topical administration, or by sustained release systems.

[0332] Dosages and desired drug concentrations of pharmaceutical compositions of the present invention may vary

depending on the particular use envisioned. The determination of the appropriate dosage or route of administration is well within the skill of an ordinary physician. Animal experiments provide reliable guidance for the determination of effective doses for human therapy. Interspecies scaling of effective doses can be performed following the principles laid down by Mordenti, J. and Chappell, W. "The use of interspecies scaling in toxicokinetics" In *Toxicokinetics and New Drug Development*, Yacobi et al., Eds., Pergamon Press, New York 1989, pp. 42-96.

[0333] When in vivo administration of a PRO polypeptide or agonist or antagonist thereof is employed, normal dosage amounts may vary from about 10 ng/kg to up to 100 mg/kg of mammal body weight or more per day, preferably about 1 μ g/kg/day to 10 mg/kg/day, depending upon the route of administration. Guidance as to particular dosages and methods of delivery is provided in the literature; see, for example, U.S. Pat. Nos. 4,657,760; 5,206,344; or 5,225,212. It is anticipated that different formulations will be effective for different treatment compounds and different disorders, that administration targeting one organ or tissue, for example, may necessitate delivery in a manner different from that to another organ or tissue.

[0334] Where sustained-release administration of a PRO polypeptide is desired in a formulation with release characteristics suitable for the treatment of any disease or disorder requiring administration of the PRO polypeptide, microencapsulation of the PRO polypeptide is contemplated. Microencapsulation of recombinant proteins for sustained release has been successfully performed with human growth hormone (rhGH), interferon- (rhIFN-), interleukin-2, and MN rgp120. Johnson et al., *Nat. Med.*, 2:795-799 (1996); Yasuda, *Biomed. Ther.*, 27:1221-1223 (1993); Hora et al., *Bio/Technology*, 8:755-758 (1990); Cleland, "Design and Production of Single Immunization Vaccines Using Polylactide Polyglycolide Microsphere Systems," in *Vaccine Design: The Subunit and Adjuvant Approach*, Powell and Newman, eds, (Plenum Press: New York, 1995), pp. 439-462; WO 97/03692, WO 96/40072, WO 96/07399; and U.S. Pat. No. 5,654,010.

[0335] The sustained-release formulations of these proteins were developed using polylactic-coglycolic acid (PLGA) polymer due to its biocompatibility and wide range of biodegradable properties. The degradation products of PLGA, lactic and glycolic acids, can be cleared quickly within the human body. Moreover, the degradability of this polymer can be adjusted from months to years depending on its molecular weight and composition. Lewis, "Controlled release of bioactive agents from lactide/glycolide polymer," in: M. Chasin and R. Langer (Eds.), *Biodegradable Polymers as Drug Delivery Systems* (Marcel Dekker: New York, 1990), pp.1-41.

[0336] This invention encompasses methods of screening compounds to identify those that mimic the PRO polypeptide (agonists) or prevent the effect of the PRO polypeptide (antagonists). Screening assays for antagonist drug candidates are designed to identify compounds that bind or complex with the PRO polypeptides encoded by the genes identified herein, or otherwise interfere with the interaction of the encoded polypeptides with other cellular proteins. Such screening assays will include assays amenable to

high-throughput screening of chemical libraries, making them particularly suitable for identifying small molecule drug candidates.

[0337] The assays can be performed in a variety of formats, including protein-protein binding assays, biochemical screening assays, immunoassays, and cell-based assays, which are well characterized in the art.

[0338] All assays for antagonists are common in that they call for contacting the drug candidate with a PRO polypeptide encoded by a nucleic acid identified herein under conditions and for a time sufficient to allow these two components to interact.

[0339] In binding assays, the interaction is binding and the complex formed can be isolated or detected in the reaction mixture. In a particular embodiment, the PRO polypeptide encoded by the gene identified herein or the drug candidate is immobilized on a solid phase, e.g., on a microtiter plate, by covalent or non-covalent attachments. Non-covalent attachment generally is accomplished by coating the solid surface with a solution of the PRO polypeptide and drying. Alternatively, an immobilized antibody, e.g., a monoclonal antibody, specific for the PRO polypeptide to be immobilized can be used to anchor it to a solid surface. The assay is performed by adding the non-immobilized component, which may be labeled by a detectable label, to the immobilized component, e.g., the coated surface containing the anchored component. When the reaction is complete, the non-reacted components are removed, e.g., by washing, and complexes anchored on the solid surface are detected. When the originally non-immobilized component carries a detectable label, the detection of label immobilized on the surface indicates that complexing occurred. Where the originally non-immobilized component does not carry a label, complexing can be detected, for example, by using a labeled antibody specifically binding the immobilized complex.

[0340] If the candidate compound interacts with but does not bind to a particular PRO polypeptide encoded by a gene identified herein, its interaction with that polypeptide can be assayed by methods well known for detecting protein-protein interactions. Such assays include traditional approaches, such as, e.g., cross-linking, co-immunoprecipitation, and co-purification through gradients or chromatographic columns. In addition, protein-protein interactions can be monitored by using a yeast-based genetic system described by Fields and co-workers (Fields and Song, *Nature* (London), 340:245-246 (1989); Chien et al., *Proc. Natl. Acad. Sci. USA*, 88:9578-9582 (1991)) as disclosed by Chevray and Nathans, *Proc. Natl. Acad. Sci. USA*, 89: 5789-5793 (1991). Many transcriptional activators, such as yeast GAL4, consist of two physically discrete modular domains, one acting as the DNA-binding domain, the other one functioning as the transcription-activation domain. The yeast expression system described in the foregoing publications (generally referred to as the "two-hybrid system") takes advantage of this property, and employs two hybrid proteins, one in which the target protein is fused to the DNA-binding domain of GAL4, and another, in which candidate activating proteins are fused to the activation domain. The expression of a GAL1-lacZ reporter gene under control of a GAL4-activated promoter depends on reconstitution of GAL4 activity via protein-protein interaction. Colonies containing interacting polypeptides are detected

with a chromogenic substrate for β -galactosidase. A complete kit (MATCHMAKER™) for identifying protein-protein interactions between two specific proteins using the two-hybrid technique is commercially available from Clontech. This system can also be extended to map protein domains involved in specific protein interactions as well as to pinpoint amino acid residues that are crucial for these interactions.

[0341] Compounds that interfere with the interaction of a gene encoding a PRO polypeptide identified herein and other intra- or extracellular components can be tested as follows: usually a reaction mixture is prepared containing the product of the gene and the intra- or extracellular component under conditions and for a time allowing for the interaction and binding of the two products. To test the ability of a candidate compound to inhibit binding, the reaction is run in the absence and in the presence of the test compound. In addition, a placebo may be added to a third reaction mixture, to serve as positive control. The binding (complex formation) between the test compound and the intra- or extracellular component present in the mixture is monitored as described hereinabove. The formation of a complex in the control reaction(s) but not in the reaction mixture containing the test compound indicates that the test compound interferes with the interaction of the test compound and its reaction partner.

[0342] To assay for antagonists, the PRO polypeptide may be added to a cell along with the compound to be screened for a particular activity and the ability of the compound to inhibit the activity of interest in the presence of the PRO polypeptide indicates that the compound is an antagonist to the PRO polypeptide. Alternatively, antagonists may be detected by combining the PRO polypeptide and a potential antagonist with membrane-bound PRO polypeptide receptors or recombinant receptors under appropriate conditions for a competitive inhibition assay. The PRO polypeptide can be labeled, such as by radioactivity, such that the number of PRO polypeptide molecules bound to the receptor can be used to determine the effectiveness of the potential antagonist. The gene encoding the receptor can be identified by numerous methods known to those of skill in the art, for example, ligand panning and FACS sorting. Coligan et al., *Current Protocols in Immun.*, 1(2): Chapter 5 (1991). Preferably, expression cloning is employed wherein polyadenylated RNA is prepared from a cell responsive to the PRO polypeptide and a cDNA library created from this RNA is divided into pools and used to transfect COS cells or other cells that are not responsive to the PRO polypeptide. Transfected cells that are grown on glass slides are exposed to labeled PRO polypeptide. The PRO polypeptide can be labeled by a variety of means including iodination or inclusion of a recognition site for a site-specific protein kinase. Following fixation and incubation, the slides are subjected to autoradiographic analysis. Positive pools are identified and sub-pools are prepared and re-transfected using an interactive sub-pooling and re-screening process, eventually yielding a single clone that encodes the putative receptor.

[0343] As an alternative approach for receptor identification, labeled PRO polypeptide can be photoaffinity-linked with cell membrane or extract preparations that express the receptor molecule. Cross-linked material is resolved by PAGE and exposed to X-ray film. The labeled complex containing the receptor can be excised, resolved into peptide

fragments, and subjected to protein micro-sequencing. The amino acid sequence obtained from micro-sequencing would be used to design a set of degenerate oligonucleotide probes to screen a cDNA library to identify the gene encoding the putative receptor.

[0344] In another assay for antagonists, mammalian cells or a membrane preparation expressing the receptor would be incubated with labeled PRO polypeptide in the presence of the candidate compound. The ability of the compound to enhance or block this interaction could then be measured.

[0345] More specific examples of potential antagonists include an oligonucleotide that binds to the fusions of immunoglobulin with PRO polypeptide, and, in particular, antibodies including, without limitation, poly- and monoclonal antibodies and antibody fragments, single-chain antibodies, anti-idiotypic antibodies, and chimeric or humanized versions of such antibodies or fragments, as well as human antibodies and antibody fragments. Alternatively, a potential antagonist may be a closely related protein, for example, a mutated form of the PRO polypeptide that recognizes the receptor but imparts no effect, thereby competitively inhibiting the action of the PRO polypeptide.

[0346] Another potential PRO polypeptide antagonist is an antisense RNA or DNA construct prepared using antisense technology, where, e.g., an antisense RNA or DNA molecule acts to block directly the translation of mRNA by hybridizing to targeted mRNA and preventing protein translation. Antisense technology can be used to gene expression through triple-helix formation or antisense DNA or RNA, both of which methods are based on binding of a polynucleotide to DNA or RNA. For example, the 5' coding portion of the polynucleotide sequence, which encodes the mature PRO polypeptides herein, is used to design an antisense RNA oligonucleotide of from about 10 to 40 base pairs in length. A DNA oligonucleotide is designed to be complementary to a region of the gene involved in transcription (triple helix—see Lee et al., *Nucl. Acids Res.*, 6:3073 (1979); Cooney et al., *Science*, 241: 456 (1988); Dervan et al., *Science*, 251:1360 (1991)), thereby preventing transcription and the production of the PRO polypeptide. The antisense RNA oligonucleotide hybridizes to the mRNA in vivo and blocks translation of the mRNA molecule into the PRO polypeptide (antisense—Okano, *Neurochem.*, 56:560 (1991); Oligodeoxynucleotides as Antisense Inhibitors of Gene Expression (CRC Press: Boca Raton, Fla., 1988). The oligonucleotides described above can also be delivered to cells such that the antisense RNA or DNA may be expressed in vivo to inhibit production of the PRO polypeptide. When antisense DNA is used, oligodeoxyribonucleotides derived from the translation-initiation site, e.g., between about -10 and +10 positions of the target gene nucleotide sequence, are preferred.

[0347] Potential antagonists include small molecules that bind to the active site, the receptor binding site, or growth factor or other relevant binding site of the PRO polypeptide, thereby blocking the normal biological activity of the PRO polypeptide. Examples of small molecules include, but are not limited to, small peptides or peptide-like molecules, preferably soluble peptides, and synthetic non-peptidyl organic or inorganic compounds.

[0348] Ribozymes are enzymatic RNA molecules capable of catalyzing the specific cleavage of RNA. Ribozymes act

by sequence-specific hybridization to the complementary target RNA, followed by endonucleolytic cleavage. Specific ribozyme cleavage sites within a potential RNA target can be identified by known techniques. For further details see, e.g., Rossi, *Current Biology*, 4:469-471 (1994), and PCT publication No. WO 97/33551 (published Sep. 18, 1997).

[0349] Nucleic acid molecules in triple-helix formation used to inhibit transcription should be single-stranded and composed of deoxynucleotides. The base composition of these oligonucleotides is designed such that it promotes triple-helix formation via Hoogsteen base-pairing rules, which generally require sizeable stretches of purines or pyrimidines on one strand of a duplex. For further details see, e.g., PCT publication No. WO 97/33551, *supra*.

[0350] These small molecules can be identified by any one or more of the screening assays discussed hereinabove and/or by any other screening techniques well known for those skilled in the art.

[0351] Diagnostic and therapeutic uses of the herein disclosed molecules may also be based upon the positive functional assay hits disclosed and described below.

F. Anti-PRO Antibodies

[0352] The present invention further provides anti-PRO antibodies. Exemplary antibodies include polyclonal, monoclonal, humanized, bispecific, and heteroconjugate antibodies.

1. Polyclonal Antibodies

[0353] The anti-PRO antibodies may comprise polyclonal antibodies. Methods of preparing polyclonal antibodies are known to the skilled artisan. Polyclonal antibodies can be raised in a mammal, for example, by one or more injections of an immunizing agent and, if desired, an adjuvant. Typically, the immunizing agent and/or adjuvant will be injected in the mammal by multiple subcutaneous or intraperitoneal injections. The immunizing agent may include the PRO polypeptide or a fusion protein thereof. It may be useful to conjugate the immunizing agent to a protein known to be immunogenic in the mammal being immunized. Examples of such immunogenic proteins include but are not limited to keyhole limpet hemocyanin, serum albumin, bovine thyroglobulin, and soybean trypsin inhibitor. Examples of adjuvants which may be employed include Freund's complete adjuvant and MPL-TDM adjuvant (monophosphoryl Lipid A, synthetic trehalose dicorynomycolate). The immunization protocol may be selected by one skilled in the art without undue experimentation.

2. Monoclonal Antibodies

[0354] The anti-PRO antibodies may, alternatively, be monoclonal antibodies. Monoclonal antibodies may be prepared using hybridoma methods, such as those described by Kohler and Milstein, *Nature*, 256:495 (1975). In a hybridoma method, a mouse, hamster, or other appropriate host animal, is typically immunized with an immunizing agent to elicit lymphocytes that produce or are capable of producing antibodies that will specifically bind to the immunizing agent. Alternatively, the lymphocytes may be immunized in vitro.

[0355] The immunizing agent will typically include the PRO polypeptide or a fusion protein thereof. Generally,

either peripheral blood lymphocytes ("PBLs") are used if cells of human origin are desired, or spleen cells or lymph node cells are used if non-human mammalian sources are desired. The lymphocytes are then fused with an immortalized cell line using a suitable fusing agent, such as polyethylene glycol, to form a hybridoma cell [Goding, *Monoclonal Antibodies: Principles and Practice*, Academic Press, (1986) pp. 59-103]. Immortalized cell lines are usually transformed mammalian cells, particularly myeloma cells of rodent, bovine and human origin. Usually, rat or mouse myeloma cell lines are employed. The hybridoma cells may be cultured in a suitable culture medium that preferably contains one or more substances that inhibit the growth or survival of the unfused, immortalized cells. For example, if the parental cells lack the enzyme hypoxanthine guanine phosphoribosyl transferase (HGPRT or HPRT), the culture medium for the hybridomas typically will include hypoxanthine, aminopterin, and thymidine ("HAT medium"), which substances prevent the growth of HGPRT-deficient cells.

[0356] Preferred immortalized cell lines are those that fuse efficiently, support stable high level expression of antibody by the selected antibody-producing cells, and are sensitive to a medium such as HAT medium. More preferred immortalized cell lines are murine myeloma lines, which can be obtained, for instance, from the Salk Institute Cell Distribution Center, San Diego, Calif. and the American Type Culture Collection, Manassas, Va. Human myeloma and mouse-human heteromyeloma cell lines also have been described for the production of human monoclonal antibodies [Kozbor, *J. Immunol.*, 133:3001 (1984); Brodeur et al., *Monoclonal Antibody Production Techniques and Applications*, Marcel Dekker, Inc., New York, (1987) pp. 51-63].

[0357] The culture medium in which the hybridoma cells are cultured can then be assayed for the presence of monoclonal antibodies directed against PRO. Preferably, the binding specificity of monoclonal antibodies produced by the hybridoma cells is determined by immunoprecipitation or by an in vitro binding assay, such as radioimmunoassay (RIA) or enzyme-linked immunoabsorbent assay (ELISA). Such techniques and assays are known in the art. The binding affinity of the monoclonal antibody can, for example, be determined by the Scatchard analysis of Munson and Pollard, *Anal. Biochem.*, 107:220 (1980).

[0358] After the desired hybridoma cells are identified, the clones may be subcloned by limiting dilution procedures and grown by standard methods [Goding, *supra*]. Suitable culture media for this purpose include, for example, Dulbecco's Modified Eagle's Medium and RPMI-1640 medium. Alternatively, the hybridoma cells may be grown in vivo as ascites in a mammal.

[0359] The monoclonal antibodies secreted by the subclones may be isolated or purified from the culture medium or ascites fluid by conventional immunoglobulin purification procedures such as, for example, protein A-Sepharose, hydroxylapatite chromatography, gel electrophoresis, dialysis, or affinity chromatography.

[0360] The monoclonal antibodies may also be made by recombinant DNA methods, such as those described in U.S. Pat. No. 4,816,567. DNA encoding the monoclonal antibodies of the invention can be readily isolated and sequenced using conventional procedures (e.g., by using oligonucleotide probes that are capable of binding specifically to genes

encoding the heavy and light chains of murine antibodies). The hybridoma cells of the invention serve as a preferred source of such DNA. Once isolated, the DNA may be placed into expression vectors, which are then transfected into host cells such as simian COS cells, Chinese hamster ovary (CHO) cells, or myeloma cells that do not otherwise produce immunoglobulin protein, to obtain the synthesis of monoclonal antibodies in the recombinant host cells. The DNA also may be modified, for example, by substituting the coding sequence for human heavy and light chain constant domains in place of the homologous murine sequences [U.S. Pat. No. 4,816,567; Morrison et al., *supra*] or by covalently joining to the immunoglobulin coding sequence all or part of the coding sequence for a non-immunoglobulin polypeptide. Such a non-immunoglobulin polypeptide can be substituted for the constant domains of an antibody of the invention, or can be substituted for the variable domains of one antigen-combining site of an antibody of the invention to create a chimeric bivalent antibody.

[0361] The antibodies may be monovalent antibodies. Methods for preparing monovalent antibodies are well known in the art. For example, one method involves recombinant expression of immunoglobulin light chain and modified heavy chain. The heavy chain is truncated generally at any point in the Fc region so as to prevent heavy chain crosslinking. Alternatively, the relevant cysteine residues are substituted with another amino acid residue or are deleted so as to prevent crosslinking.

[0362] In vitro methods are also suitable for preparing monovalent antibodies. Digestion of antibodies to produce fragments thereof, particularly, Fab fragments, can be accomplished using routine techniques known in the art.

3. Human and Humanized Antibodies

[0363] The anti-PRO antibodies of the invention may further comprise humanized antibodies or human antibodies. Humanized forms of non-human (e.g., murine) antibodies are chimeric immunoglobulins, immunoglobulin chains or fragments thereof (such as Fv, Fab, Fab', F(ab')₂ or other antigen-binding subsequences of antibodies) which contain minimal sequence derived from non-human immunoglobulin. Humanized antibodies include human immunoglobulins (recipient antibody) in which residues from a complementary determining region (CDR) of the recipient are replaced by residues from a CDR of a non-human species (donor antibody) such as mouse, rat or rabbit having the desired specificity, affinity and capacity. In some instances, Fv framework residues of the human immunoglobulin are replaced by corresponding non-human residues. Humanized antibodies may also comprise residues which are found neither in the recipient antibody nor in the imported CDR or framework sequences. In general, the humanized antibody will comprise substantially all of at least one, and typically two, variable domains, in which all or substantially all of the CDR regions correspond to those of a non-human immunoglobulin and all or substantially all of the FR regions are those of a human immunoglobulin consensus sequence. The humanized antibody optimally also will comprise at least a portion of an immunoglobulin constant region (Fc), typically that of a human immunoglobulin [Jones et al., *Nature*, 321:522-525 (1986); Riechmann et al., *Nature*, 332:323-329 (1988); and Presta, *Curr. Op. Struct. Biol.*, 2:593-596 (1992)].

[0364] Methods for humanizing non-human antibodies are well known in the art. Generally, a humanized antibody has one or more amino acid residues introduced into it from a source which is non-human. These non-human amino acid residues are often referred to as "import" residues, which are typically taken from an "import" variable domain. Humanization can be essentially performed following the method of Winter and co-workers [Jones et al., *Nature*, 321:522-525 (1986); Riechmann et al., *Nature*, 332:323-327 (1988); Verhoeven et al., *Science*, 239:1534-1536 (1988)], by substituting rodent CDRs or CDR sequences for the corresponding sequences of a human antibody. Accordingly, such "humanized" antibodies are chimeric antibodies (U.S. Pat. No. 4,816,567), wherein substantially less than an intact human variable domain has been substituted by the corresponding sequence from a non-human species. In practice, humanized antibodies are typically human antibodies in which some CDR residues and possibly some FR residues are substituted by residues from analogous sites in rodent antibodies.

[0365] Human antibodies can also be produced using various techniques known in the art, including phage display libraries [Hoogenboom and Winter, *J. Mol. Biol.*, 227:381 (1991); Marks et al., *J. Mol. Biol.*, 222:581 (1991)]. The techniques of Cole et al. and Boerner et al. are also available for the preparation of human monoclonal antibodies (Cole et al., *Monoclonal Antibodies and Cancer Therapy*, Alan R. Liss, p. 77 (1985) and Boerner et al., *J. Immunol.*, 147(1):86-95 (1991)]. Similarly, human antibodies can be made by introducing of human immunoglobulin loci into transgenic animals, e.g., mice in which the endogenous immunoglobulin genes have been partially or completely inactivated. Upon challenge, human antibody production is observed, which closely resembles that seen in humans in all respects, including gene rearrangement, assembly, and antibody repertoire. This approach is described, for example, in U.S. Pat. Nos. 5,545,807; 5,545,806; 5,569,825; 5,625,126; 5,633,425; 5,661,016, and in the following scientific publications: Marks et al., *Bio/Technology* 10, 779-783 (1992); Lonberg et al., *Nature* 368 856-859 (1994); Morrison, *Nature* 368, 812-13 (1994); Fishwild et al., *Nature Biotechnology* 14, 845-51 (1996); Neuberger, *Nature Biotechnology* 14, 826 (1996); Lonberg and Huszar, *Intern. Rev. Immunol.* 13 65-93 (1995).

[0366] The antibodies may also be affinity matured using known selection and/or mutagenesis methods as described above. Preferred affinity matured antibodies have an affinity which is five times, more preferably 10 times, even more preferably 20 or 30 times greater than the starting antibody (generally murine, humanized or human) from which the matured antibody is prepared.

4. Bispecific Antibodies

[0367] Bispecific antibodies are monoclonal, preferably human or humanized, antibodies that have binding specificities for at least two different antigens. In the present case, one of the binding specificities is for the PRO, the other one is for any other antigen, and preferably for a cell-surface protein or receptor or receptor subunit.

[0368] Methods for making bispecific antibodies are known in the art. Traditionally, the recombinant production of bispecific antibodies is based on the co-expression of two

immunoglobulin heavy-chain/light-chain pairs, where the two heavy chains have different specificities [Milstein and Cuello, *Nature*, 305:537-539 (1983)]. Because of the random assortment of immunoglobulin heavy and light chains, these hybridomas (quadromas) produce a potential mixture of ten different antibody molecules, of which only one has the correct bispecific structure. The purification of the correct molecule is usually accomplished by affinity chromatography steps. Similar procedures are disclosed in WO 93/08829, published May 13, 1993, and in Traunecker et al., *EMBO J.*, 10:3655-3659 (1991).

[0369] Antibody variable domains with the desired binding specificities (antibody-antigen combining sites) can be fused to immunoglobulin constant domain sequences. The fusion preferably is with an immunoglobulin heavy-chain constant domain, comprising at least part of the hinge, CH2, and CH3 regions. It is preferred to have first heavy-chain constant region (CH1) containing the site necessary for light-chain binding present in at least one of the fusions. DNAs encoding the immunoglobulin heavy-chain fusions and, if desired, the immunoglobulin light chain, are inserted into separate expression vectors, and are co-transfected into a suitable host organism. For further details of generating bispecific antibodies see, for example, Suresh et al., *Methods in Enzymology*, 121:210 (1986).

[0370] According to another approach described in WO 96/27011, the interface between a pair of antibody molecules can be engineered to maximize the percentage of heterodimers which are recovered from recombinant cell culture. The preferred interface comprises at least a part of the CH3 region of an antibody constant domain. In this method, one or more small amino acid side chains from the interface of the first antibody molecule are replaced with larger side chains (e.g. tyrosine or tryptophan). Compensatory "cavities" of identical or similar size to the large side chain(s) are created on the interface of the second antibody molecule by replacing large amino acid side chains with smaller ones (e.g. alanine or threonine). This provides a mechanism for increasing the yield of the heterodimer over other unwanted end-products such as homodimers.

[0371] Bispecific antibodies can be prepared as full length antibodies or antibody fragments (e.g. F(ab')₂ bispecific antibodies). Techniques for generating bispecific antibodies from antibody fragments have been described in the literature. For example, bispecific antibodies can be prepared using chemical linkage. Brennan et al., *Science* 229:81 (1985) describe a procedure wherein intact antibodies are proteolytically cleaved to generate F(ab')₂ fragments. These fragments are reduced in the presence of the dithiol complexing agent sodium arsenite to stabilize vicinal dithiols and prevent intermolecular disulfide formation. The Fab' fragments generated are then converted to thionitrobenzoate (TNB) derivatives. One of the Fab'-TNB derivatives is then reconverted to the Fab'-thiol by reduction with mercaptoethylamine and is mixed with an equimolar amount of the other Fab'-TNB derivative to form the bispecific antibody. The bispecific antibodies produced can be used as agents for the selective immobilization of enzymes.

[0372] Fab' fragments may be directly recovered from *E. coli* and chemically coupled to form bispecific antibodies. Shalaby et al., *J. Exp. Med.* 175:217 (1992) describe the production of a fully humanized bispecific antibody F(ab')₂

molecule. Each Fab' fragment was separately secreted from *E. coli* and subjected to directed chemical coupling in vitro to form the bispecific antibody. The bispecific antibody thus formed was able to bind to cells overexpressing the ErbB2 receptor and normal human T cells, as well as trigger the lytic activity of human cytotoxic lymphocytes against human breast tumor targets.

[0373] Various technique for making and isolating bispecific antibody fragments directly from recombinant cell culture have also been described. For example, bispecific antibodies have been produced using leucine zippers. Kostelný et al., *J. Immunol.* 148 (5):1547-1553 (1992). The leucine zipper peptides from the Fos and Jun proteins were linked to the Fab' portions of two different antibodies by gene fusion. The antibody homodimers were reduced at the hinge region to form monomers and then re-oxidized to form the antibody heterodimers. This method can also be utilized for the production of antibody homodimers. The "diabody" technology described by Hollinger et al., *Proc. Natl. Acad. Sci. USA* 90:6444-6448 (1993) has provided an alternative mechanism for making bispecific antibody fragments. The fragments comprise a heavy-chain variable domain (V_H) connected to a light-chain variable domain (V_L) by a linker which is too short to allow pairing between the two domains on the same chain. Accordingly, the V_H and V_L domains of one fragment are forced to pair with the complementary V_L and V_H domains of another fragment, thereby forming two antigen-binding sites. Another strategy for making bispecific antibody fragments by the use of single-chain Fv (sFv) dimers has also been reported. See, Gruber et al., *J. Immunol.* 152:5368 (1994). Antibodies with more than two valencies are contemplated. For example, trispecific antibodies can be prepared. Tutt et al., *J. Immunol.* 147:60 (1991).

[0374] Exemplary bispecific antibodies may bind to two different epitopes on a given PRO polypeptide herein. Alternatively, an anti-PRO polypeptide arm may be combined with an arm which binds to a triggering molecule on a leukocyte such as a T-cell receptor molecule (e.g. CD2, CD3, CD28, or B7), or Fc receptors for IgG (Fc γ R), such as Fc γ RI (CD64), Fc γ RII (CD32) and Fc γ RIII (CD16) so as to focus cellular defense mechanisms to the cell expressing the particular PRO polypeptide. Bispecific antibodies may also be used to localize cytotoxic agents to cells which express a particular PRO polypeptide. These antibodies possess a PRO-binding arm and an arm which binds a cytotoxic agent or a radionuclide chelator, such as EOTUBE, DPTA, DOTA, or TETA. Another bispecific antibody of interest binds the PRO polypeptide and further binds tissue factor (TF).

5. Heteroconjugate Antibodies

[0375] Heteroconjugate antibodies are also within the scope of the present invention. Heteroconjugate antibodies are composed of two covalently joined antibodies. Such antibodies have, for example, been proposed to target immune system cells to unwanted cells [U.S. Pat. No. 4,676,980], and for treatment of HIV infection [WO 91/00360; WO 92/200373; EP 03089]. It is contemplated that the antibodies may be prepared in vitro using known methods in synthetic protein chemistry, including those involving crosslinking agents. For example, immunotoxins may be constructed using a disulfide exchange reaction or by forming a thioether bond. Examples of suitable reagents for

this purpose include iminothiolate and methyl-4-mercaptobutyrimidate and those disclosed, for example, in U.S. Pat. No. 4,676,980.

6. Effector Function Engineering

[0376] It may be desirable to modify the antibody of the invention with respect to effector function, so as to enhance, e.g., the effectiveness of the antibody in treating cancer. For example, cysteine residue(s) may be introduced into the Fc region, thereby allowing interchain disulfide bond formation in this region. The homodimeric antibody thus generated may have improved internalization capability and/or increased complement-mediated cell killing and antibody-dependent cellular cytotoxicity (ADCC). See Caron et al., *J. Exp. Med.*, 176: 1191-1195 (1992) and Shopes, *J. Immunol.*, 148: 2918-2922 (1992). Homodimeric antibodies with enhanced anti-tumor activity may also be prepared using heterobifunctional cross-linkers as described in Wolff et al. *Cancer Research*, 53: 2560-2565 (1993). Alternatively, an antibody can be engineered that has dual Fc regions and may thereby have enhanced complement lysis and ADCC capabilities. See Stevenson et al., *Anti-Cancer Drug Design*, 3: 219-230 (1989).

7. Immunoconjugates

[0377] The invention also pertains to immunoconjugates comprising an antibody conjugated to a cytotoxic agent such as a chemotherapeutic agent, toxin (e.g., an enzymatically active toxin of bacterial, fungal, plant, or animal origin, or fragments thereof), or a radioactive isotope (i.e., a radioconjugate).

[0378] Chemotherapeutic agents useful in the generation of such immunoconjugates have been described above. Enzymatically active toxins and fragments thereof that can be used include diphtheria A chain, nonbinding active fragments of diphtheria toxin, exotoxin A chain (from *Pseudomonas aeruginosa*), ricin A chain, abrin A chain, modeccin A chain, alpha-sarcin, *Aleurites fordii* proteins, dianthin proteins, *Phytolacca americana* proteins (PAPI, PAPII, and PAP-S), *momordica charantia* inhibitor, curcin, crotin, *sapaonaria officinalis* inhibitor, gelonin, mitogellin, restrictocin, phenomycin, enomycin, and the tricothecenes. A variety of radionuclides are available for the production of radioconjugated antibodies. Examples include ^{212}Bi , ^{131}I , ^{131}In , ^{90}Y , and ^{186}Re . Conjugates of the antibody and cytotoxic agent are made using a variety of bifunctional protein-coupling agents such as N-succinimidyl-3-(2-pyridyldithiol) propionate (SPDP), iminothiolane (IT), bifunctional derivatives of imidoesters (such as dimethyl adipimide HCL), active esters (such as disuccinimidyl suberate), aldehydes (such as glutaraldehyde), bis-azido compounds (such as bis (p-azidobenzoyl) hexanediamine), bis-diazonium derivatives (such as bis-(p-diazoniumbenzoyl)-ethylenediamine), diisocyanates (such as tolyene 2,6-diisocyanate), and bis-active fluorine compounds (such as 1,5-difluoro-2,4-dinitrobenzene). For example, a ricin immunotoxin can be prepared as described in Vitetta et al., *Science*, 238: 1098 (1987). Carbon-14-labeled 1-isothiocyanatobenzyl-3-methyldiethylene triaminepentaacetic acid (MX-DTPA) is an exemplary chelating agent for conjugation of radionuclide to the antibody. See WO 94/11026.

[0379] In another embodiment, the antibody may be conjugated to a "receptor" (such streptavidin) for utilization in

tumor pretargeting wherein the antibody-receptor conjugate is administered to the patient, followed by removal of unbound conjugate from the circulation using a clearing agent and then administration of a "ligand" (e.g., avidin) that is conjugated to a cytotoxic agent (e.g., a radionucleotide).

8. Immunoliposomes

[0380] The antibodies disclosed herein may also be formulated as immunoliposomes. Liposomes containing the antibody are prepared by methods known in the art, such as described in Epstein et al., *Proc. Natl. Acad. Sci. USA*, 82: 3688 (1985); Hwang et al., *Proc. Natl. Acad. Sci. USA*, 77: 4030 (1980); and U.S. Pat. Nos. 4,485,045 and 4,544,545. Liposomes with enhanced circulation time are disclosed in U.S. Pat. No. 5,013,556.

[0381] Particularly useful liposomes can be generated by the reverse-phase evaporation method with a lipid composition comprising phosphatidylcholine, cholesterol, and PEG-derivatized phosphatidylethanolamine (PEG-PE). Liposomes are extruded through filters of defined pore size to yield liposomes with the desired diameter. Fab' fragments of the antibody of the present invention can be conjugated to the liposomes as described in Martin et al., *J. Biol. Chem.*, 257: 286-288 (1982) via a disulfide-interchange reaction. A chemotherapeutic agent (such as Doxorubicin) is optionally contained within the liposome. See Gabizon et al., *J. National Cancer Inst.*, 81(19): 1484 (1989).

9. Pharmaceutical Compositions of Antibodies

[0382] Antibodies specifically binding a PRO polypeptide identified herein, as well as other molecules identified by the screening assays disclosed hereinbefore, can be administered for the treatment of various disorders in the form of pharmaceutical compositions.

[0383] If the PRO polypeptide is intracellular and whole antibodies are used as inhibitors, internalizing antibodies are preferred. However, lipofections or liposomes can also be used to deliver the antibody, or an antibody fragment, into cells. Where antibody fragments are used, the smallest inhibitory fragment that specifically binds to the binding domain of the target protein is preferred. For example, based upon the variable-region sequences of an antibody, peptide molecules can be designed that retain the ability to bind the target protein sequence. Such peptides can be synthesized chemically and/or produced by recombinant DNA technology. See, e.g., Marasco et al., *Proc. Natl. Acad. Sci. USA*, 90: 7889-7893 (1993). The formulation herein may also contain more than one active compound as necessary for the particular indication being treated, preferably those with complementary activities that do not adversely affect each other. Alternatively, or in addition, the composition may comprise an agent that enhances its function, such as, for example, a cytotoxic agent, cytokine, chemotherapeutic agent, or growth-inhibitory agent. Such molecules are suitably present in combination in amounts that are effective for the purpose intended.

[0384] The active ingredients may also be entrapped in microcapsules prepared, for example, by coacervation techniques or by interfacial polymerization, for example, hydroxymethylcellulose or gelatin-microcapsules and poly-(methylmethacrylate) microcapsules, respectively, in colloidal drug delivery systems (for example, liposomes, albumin

microspheres, microemulsions, nano-particles, and nano-capsules) or in macroemulsions. Such techniques are disclosed in Remington's *Pharmaceutical Sciences*, supra.

[0385] The formulations to be used for in vivo administration must be sterile. This is readily accomplished by filtration through sterile filtration membranes.

[0386] Sustained-release preparations may be prepared. Suitable examples of sustained-release preparations include semipermeable matrices of solid hydrophobic polymers containing the antibody, which matrices are in the form of shaped articles, e.g., films, or microcapsules. Examples of sustained-release matrices include polyesters, hydrogels (for example, poly(2-hydroxyethyl-methacrylate), or poly(vinylalcohol)), polylactides (U.S. Pat. No. 3,773,919), copolymers of L-glutamic acid and 65 ethyl-L-glutamate, non-degradable ethylene-vinyl acetate, degradable lactic acid-glycolic acid copolymers such as the LUPRON DEPOT™ (injectable microspheres composed of lactic acid-glycolic acid copolymer and leuprolide acetate), and poly-D-(-)-3-hydroxybutyric acid. While polymers such as ethylene-vinyl acetate and lactic acid-glycolic acid enable release of molecules for over 100 days, certain hydrogels release proteins for shorter time periods. When encapsulated antibodies remain in the body for a long time, they may denature or aggregate as a result of exposure to moisture at 37° C., resulting in a loss of biological activity and possible changes in immunogenicity. Rational strategies can be devised for stabilization depending on the mechanism involved. For example, if the aggregation mechanism is discovered to be intermolecular S—S bond formation through thio-disulfide interchange, stabilization may be achieved by modifying sulfhydryl residues, lyophilizing from acidic solutions, controlling moisture content, using appropriate additives, and developing specific polymer matrix compositions.

C. Uses for Anti-PRO Antibodies

[0387] The anti-PRO antibodies of the invention have various utilities. For example, anti-PRO antibodies may be used in diagnostic assays for PRO, e.g., detecting its expression (and in some cases, differential expression) in specific cells, tissues, or serum. Various diagnostic assay techniques known in the art may be used, such as competitive binding assays, direct or indirect sandwich assays and immunoprecipitation assays conducted in either heterogeneous or homogeneous phases [Zola, *Monoclonal Antibodies: A Manual of Techniques*, CRC Press, Inc. (1987) pp.147-158]. The antibodies used in the diagnostic assays can be labeled with a detectable moiety. The detectable moiety should be capable of producing, either directly or indirectly, a detectable signal. For example, the detectable moiety may be a radioisotope, such as ³H, ¹⁴C, ³²P, ³⁵S, or ¹²⁵I, a fluorescent or chemiluminescent compound, such as fluorescein isothiocyanate, rhodamine, or luciferin, or an enzyme, such as alkaline phosphatase, beta-galactosidase or horseradish peroxidase. Any method known in the art for conjugating the antibody to the detectable moiety may be employed, including those methods described by Hunter et al., *Nature*, 144:945 (1962); David et al., *Biochemistry*, 13:1014 (1974); Pain et al., *J. Immunol. Meth.*, 40:219 (1981); and Nygren, *J. Histochem. and Cytochem.*, 30:407 (1982).

[0388] Anti-PRO antibodies also are useful for the affinity purification of PRO from recombinant cell culture or natural

sources. In this process, the antibodies against PRO are immobilized on a suitable support, such as Sephadex resin or filter paper, using methods well known in the art. The immobilized antibody then is contacted with a sample containing the PRO to be purified, and thereafter the support is washed with a suitable solvent that will remove substantially all the material in the sample except the PRO, which is bound to the immobilized antibody. Finally, the support is washed with another suitable solvent that will release the PRO from the antibody.

[0389] The following examples are offered for illustrative purposes only, and are not intended to limit the scope of the present invention in any way.

[0390] All patent and literature references cited in the present specification are hereby incorporated by reference in their entirety.

EXAMPLES

[0391] Commercially available reagents referred to in the examples were used according to manufacturer's instructions unless otherwise indicated. The source of those cells identified in the following examples, and throughout the specification, by ATCC accession numbers is the American Type Culture Collection, Manassas, Va.

Example 1

Extracellular Domain Homology Screening to Identify Novel Polypeptides and cDNA Encoding Therefor

[0392] The extracellular domain (ECD) sequences (including the secretion signal sequence, if any) from about 950 known secreted proteins from the Swiss-Prot public database were used to search EST databases. The EST databases included public databases (e.g., Dayhoff, GenBank), and proprietary databases (e.g., LIFESEQ™, Incyte Pharmaceuticals, Palo Alto, Calif.). The search was performed using the computer program BLAST or BLAST-2 (Altschul et al., *Methods in Enzymology* 266:460-480 (1996)) as a comparison of the ECD protein sequences to a 6 frame translation of the EST sequences. Those comparisons with a BLAST score of 70 (or in some cases 90) or greater that did not encode known proteins were clustered and assembled into consensus DNA sequences with the program "phrap" (Phil Green, University of Washington, Seattle, Wash.).

[0393] Using this extracellular domain homology screen, consensus DNA sequences were assembled relative to the other identified EST sequences using phrap. In addition, the consensus DNA sequences obtained were often (but not always) extended using repeated cycles of BLAST or BLAST-2 and phrap to extend the consensus sequence as far as possible using the sources of EST sequences discussed above.

[0394] Based upon the consensus sequences obtained as described above, oligonucleotides were then synthesized and used to identify by PCR a cDNA library that contained the sequence of interest and for use as probes to isolate a clone of the full-length coding sequence for a PRO polypeptide. Forward and reverse PCR primers generally range from 20 to 30 nucleotides and are often designed to give a PCR

product of about 100-1000 bp in length. The probe sequences are typically 40-55 bp in length. In some cases, additional oligonucleotides are synthesized when the consensus sequence is greater than about 1-1.5 kbp. In order to screen several libraries for a full-length clone, DNA from the libraries was screened by PCR amplification, as per Ausubel et al., *Current Protocols in Molecular Biology*, with the PCR primer pair. A positive library was then used to isolate clones encoding the gene of interest using the probe oligonucleotide and one of the primer pairs.

[0395] The cDNA libraries used to isolate the cDNA clones were constructed by standard methods using commercially available reagents such as those from Invitrogen, San Diego, Calif. The cDNA was primed with oligo dT containing a NotI site, linked with blunt to SalI hemikinased adaptors, cleaved with NotI, sized appropriately by gel electrophoresis, and cloned in a defined orientation into a suitable cloning vector (such as pRKB or pRKD; pRK5B is a precursor of pRK5D that does not contain the SfiI site; see, Holmes et al., *Science*, 253:1278-1280 (1991)) in the unique XhoI and NotI sites.

Example 2

Isolation of cDNA Clones by Amylase Screening

[0396] 1. Preparation of oligo dT primed cDNA library

[0397] mRNA was isolated from a human tissue of interest using reagents and protocols from Invitrogen, San Diego, Calif. (Fast Track 2). This RNA was used to generate an oligo dT primed cDNA library in the vector pRK5D using reagents and protocols from Life Technologies, Gaithersburg, Md. (Super Script Plasmid System). In this procedure, the double stranded cDNA was sized to greater than 1000 bp and the SalI/NotI linked cDNA was cloned into XhoI/NotI cleaved vector. pRK5D is a cloning vector that has an sp6 transcription initiation site followed by an SfiI restriction enzyme site preceding the XhoI/NotI cDNA cloning sites.

[0398] 2. Preparation of random primed cDNA library

[0399] A secondary cDNA library was generated in order to preferentially represent the 5' ends of the primary cDNA clones. Sp6 RNA was generated from the primary library (described above), and this RNA was used to generate a random primed cDNA library in the vector pSST-AMY.0 using reagents and protocols from Life Technologies (Super Script Plasmid System, referenced above). In this procedure the double stranded cDNA was sized to 500-1000 bp, linked with blunt to NotI adaptors, cleaved with SfiI, and cloned into SfiI/NotI cleaved vector. pSST-AMY.0 is a cloning vector that has a yeast alcohol dehydrogenase promoter preceding the cDNA cloning sites and the mouse amylase sequence (the mature sequence without the secretion signal) followed by the yeast alcohol dehydrogenase terminator, after the cloning sites. Thus, cDNAs cloned into this vector that are fused in frame with amylase sequence will lead to the secretion of amylase from appropriately transfected yeast colonies.

[0400] 3. Transformation and Detection

[0401] DNA from the library described in paragraph 2 above was chilled on ice to which was added electrocompetent DH10B bacteria (Life Technologies, 20 ml). The bacteria and vector mixture was then electroporated as

recommended by the manufacturer. Subsequently, SOC media (Life Technologies, 1 ml) was added and the mixture was incubated at 37° C. for 30 minutes. The transformants were then plated onto 20 standard 150 mm LB plates containing ampicillin and incubated for 16 hours (37° C.). Positive colonies were scraped off the plates and the DNA was isolated from the bacterial pellet using standard protocols, e.g. CsCl-gradient. The purified DNA was then carried on to the yeast protocols below.

[0402] The yeast methods were divided into three categories: (1) Transformation of yeast with the plasmid/cDNA combined vector; (2) Detection and isolation of yeast clones secreting amylase; and (3) PCR amplification of the insert directly from the yeast colony and purification of the DNA for sequencing and further analysis.

[0403] The yeast strain used was HD56-5A (ATCC-90785). This strain has the following genotype: MAT alpha, ura3-52, leu2-3, leu2-112, his3-11, his3-15, MAL⁺, SUC⁺, GAL⁺. Preferably, yeast mutants can be employed that have deficient post-translational pathways. Such mutants may have translocation deficient alleles in sec 71, sec 72, sec 62, with truncated sec 71 being most preferred. Alternatively, antagonists (including antisense nucleotides and/or ligands) which interfere with the normal operation of these genes, other proteins implicated in this post translation pathway (e.g., SEC61p, SEC72p, SEC62p, SEC63p, TDJ1p or SSA1p-4p) or the complex formation of these proteins may also be preferably employed in combination with the amylase-expressing yeast.

[0404] Transformation was performed based on the protocol outlined by Gietz et al., Nucl. Acid. Res., 20:1425 (1992). Transformed cells were then inoculated from agar into YEPD complex media broth (100 ml) and grown overnight at 30° C. The YEPD broth was prepared as described in Kaiser et al., Methods in Yeast Genetics, Cold Spring Harbor Press, Cold Spring Harbor, N.Y., p. 207 (1994). The overnight culture was then diluted to about 2x10⁶ cells/ml (approx. OD₆₀₀=0.1) into fresh YEPD broth (500 ml) and regrown to 1x10⁷ cells/ml (approx. OD₆₀₀=0.4-0.5).

[0405] The cells were then harvested and prepared for transformation by transfer into GS3 rotor bottles in a Sorval GS3 rotor at 5,000 rpm for 5 minutes, the supernatant discarded, and then resuspended into sterile water, and centrifuged again in 50 ml falcon tubes at 3,500 rpm in a Beckman GS-6KR centrifuge. The supernatant was discarded and the cells were subsequently washed with LiAc/TE (10 ml, 10 mM Tris-HCl, 1 mM EDTA pH 7.5, 100 mM Li₂OOCCH₃), and resuspended into LiAc/TE (2.5 ml).

[0406] Transformation took place by mixing the prepared cells (100 μ l) with freshly denatured single stranded salmon testes DNA (Lofstrand Labs, Gaithersburg, Md.) and transforming DNA (1 μ g, vol.<10 μ l) in microfuge tubes. The mixture was mixed briefly by vortexing, then 40% PEG/TE (600 μ l, 40% polyethylene glycol-4000, 10 mM Tris-HCl, 1 mM EDTA, 100 mM Li₂OOCCH₃, pH 7.5) was added. This mixture was gently mixed and incubated at 30° C. while agitating for 30 minutes. The cells were then heat shocked at 42° C. for 15 minutes, and the reaction vessel centrifuged in a microfuge at 12,000 rpm for 5-10 seconds, decanted and resuspended into TE (500 μ l, 10 mM Tris-HCl, 1 mM EDTA pH 7.5) followed by recentrifugation. The cells were then

diluted into TE (1 ml) and aliquots (200 μ l) were spread onto the selective media previously prepared in 150 mm growth plates (VWR).

[0407] Alternatively, instead of multiple small reactions, the transformation was performed using a single, large scale reaction, wherein reagent amounts were scaled up accordingly.

[0408] The selective media used was a synthetic complete dextrose agar lacking uracil (SCD-Ura) prepared as described in Kaiser et al., Methods in Yeast Genetics, Cold Spring Harbor Press, Cold Spring Harbor, N.Y., p. 208-210 (1994). Transformants were grown at 30° C. for 2-3 days.

[0409] The detection of colonies secreting amylase was performed by including red starch in the selective growth media. Starch was coupled to the red dye (Reactive Red-120, Sigma) as per the procedure described by Biely et al., Anal. Biochem., 172:176-179 (1988). The coupled starch was incorporated into the SCD-Ura agar plates at a final concentration of 0.15% (w/v), and was buffered with potassium phosphate to a pH of 7.0 (50-100 mM final concentration).

[0410] The positive colonies were picked and streaked across fresh selective media (onto 150 mm plates) in order to obtain well isolated and identifiable single colonies. Well isolated single colonies positive for amylase secretion were detected by direct incorporation of red starch into buffered SCD-Ura agar. Positive colonies were determined by their ability to break down starch resulting in a clear halo around the positive colony visualized directly.

[0411] 4. Isolation of DNA by PCR Amplification

[0412] When a positive colony was isolated, a portion of it was picked by a toothpick and diluted into sterile water (30 μ l) in a 96 well plate. At this time, the positive colonies were either frozen and stored for subsequent analysis or immediately amplified. An aliquot of cells (5 μ l) was used as a template for the PCR reaction in a 25 μ l volume containing: 0.5 μ l KlenTaq (Clontech, Palo Alto, Calif.); 4.0 μ l 10 mM dNTP's (Perkin Elmer-Cetus); 2.5 μ l Kentaq buffer (Clontech); 0.25 μ l forward oligo 1; 0.25 μ l reverse oligo 2; 12.5 μ l distilled water. The sequence of the forward oligonucleotide 1 was:

[0413] 5'-TGTAACGACGGCCAGTTAAATA-GACCTGCAATTATTAATCT-3' (SEQ ID NO:169)

[0414] The sequence of reverse oligonucleotide 2 was:

[0415] 5'-CAGGAAACAGCTATGACCACCTG-CACACCTGCAAATCCATT-3' (SEQ ID NO:170)

[0416] PCR was then performed as follows:

a.	Denature	92° C.,	5 minutes
b.	3 cycles of:	Denature	92° C., 30 seconds
		Anneal	59° C., 30 seconds
		Extend	72° C., 60 seconds
c.	3 cycles of:	Denature	92° C., 30 seconds
		Anneal	57° C., 30 seconds
		Extend	72° C., 60 seconds
d.	25 cycles of:	Denature	92° C., 30 seconds
		Anneal	55° C., 30 seconds
		Extend	72° C., 60 seconds
e.	Hold	4° C.	

[0417] The underlined regions of the oligonucleotides annealed to the ADH promoter region and the amylase region, respectively, and amplified a 307 bp region from vector pSST-AMY.0 when no insert was present. Typically, the first 18 nucleotides of the 5' end of these oligonucleotides contained annealing sites for the sequencing primers. Thus, the total product of the PCR reaction from an empty vector was 343 bp. However, signal sequence-fused cDNA resulted in considerably longer nucleotide sequences.

[0418] Following the PCR, an aliquot of the reaction (5 µl) was examined by agarose gel electrophoresis in a 1% agarose gel using a Tris-Borate-EDTA (TBE) buffering system as described by Sambrook et al., supra. Clones resulting in a single strong PCR product larger than 400 bp were further analyzed by DNA sequencing after purification with a 96 Qiaquick PCR clean-up column (Qiagen Inc., Chatsworth, Calif.).

Example 3

Isolation of cDNA Clones Using Signal Algorithm Analysis

[0419] Various polypeptide-encoding nucleic acid sequences were identified by applying a proprietary signal sequence finding algorithm developed by Genentech, Inc. (South San Francisco, Calif.) upon ESTs as well as clustered and assembled EST fragments from public (e.g., GenBank) and/or private (LIFESeq®, Incyte Pharmaceuticals, Inc., Palo Alto, Calif.) databases. The signal sequence algorithm computes a secretion signal score based on the character of the DNA nucleotides surrounding the first and optionally second methionine codon(s) (ATG) at the 5'-end of the sequence or sequence fragment under consideration. The nucleotides following the first ATG must code for at least 35 unambiguous amino acids without any stop codons. If the first ATG has the required amino acids, the second is not examined. If neither meets the requirement, the candidate sequence is not scored. In order to determine whether the EST sequence contains an authentic signal sequence, the DNA and corresponding amino acid sequences surrounding the ATG codon are scored using a set of seven sensors (evaluation parameters) known to be associated with secretion signals. Use of this algorithm resulted in the identification of numerous polypeptide-encoding nucleic acid sequences.

Example 4

Isolation of cDNA Clones Encoding Human PRO Polypeptides

[0420] Using the techniques described in Examples 1 to 3 above, numerous full-length cDNA clones were identified as encoding PRO polypeptides as disclosed herein. These cDNAs were then deposited under the terms of the Budapest Treaty with the American Type Culture Collection, 10801 University Blvd., Manassas, Va. 20110-2209, USA (ATCC) as shown in Tabl 7 below.

TABLE 7

Material	ATCC Dep. No.	Deposit Date
DNA26843-1389	203099	Aug. 4, 1998
DNA30867-1335	209807	Apr. 28, 1998
DNA34431-1177	209399	Oct. 17, 1997

TABLE 7-continued

Material	ATCC Dep. No.	Deposit Date
DNA38268-1188	209421	Oct. 28, 1997
DNA40621-1440	209922	Jun. 2, 1998
DNA40625-1189	209788	Apr. 21, 1998
DNA45409-2511	203579	Jan. 12, 1999
DNA45495-1550	203156	Aug. 25, 1998
DNA49820-1427	209932	Jun. 2, 1998
DNA56406-1704	203478	Nov. 17, 1998
DNA56410-1414	209923	Jun. 2, 1998
DNA56436-1448	209902	May 27, 1998
DNA56855-1447	203004	Jun. 23, 1998
DNA56860-1510	209952	Jun. 9, 1998
DNA56862-1343	203174	Sep. 1, 1998
DNA56868-1478	203024	Jun. 23, 1998
DNA56869-1545	203161	Aug. 25, 1998
DNA57704-1452	209953	Jun. 9, 1998
DNA58723-1588	203133	Aug. 18, 1998
DNA57827-1493	203045	Jul. 1, 1998
DNA58737-1473	203136	Aug. 18, 1998
DNA58846-1409	209957	Jun. 9, 1998
DNA58850-1495	209956	Jun. 9, 1998
DNA58855-1422	203018	Jun. 23, 1998
DNA59211-1450	209960	Jun. 9, 1998
DNA59212-1627	203245	Sep. 9, 1998
DNA59213-1487	209959	Jun. 9, 1998
DNA59605-1418	203005	Jun. 23, 1998
DNA59609-1470	209963	Jun. 9, 1998
DNA59610-1556	209990	Jun. 16, 1998
DNA59837-2545	203658	Feb. 9, 1999
DNA59844-2542	203650	Feb. 9, 1999
DNA59854-1459	209974	Jun. 16, 1998
DNA60625-1507	209975	Jun. 16, 1998
DNA60629-1481	209979	Jun. 16, 1998
DNA61755-1554	203112	Aug. 11, 1998
DNA62812-1594	203248	Sep. 9, 1998
DNA62815-1576	203247	Sep. 9, 1998
DNA64881-1602	203240	Sep. 9, 1998
DNA64886-1601	203241	Sep. 9, 1998
DNA64902-1667	203317	Oct. 6, 1998
DNA64950-1590	203224	Sep. 15, 1998
DNA65403-1565	203230	Sep. 15, 1998
DNA66308-1537	203159	Aug. 25, 1998
DNA66519-1535	203236	Sep. 15, 1998
DNA66521-1583	203225	Sep. 15, 1998
DNA66658-1584	203229	Sep. 15, 1998
DNA66660-1585	203279	Sep. 22, 1998
DNA66663-1598	203268	Sep. 22, 1998
DNA66674-1599	203281	Sep. 22, 1998
DNA68862-2546	203652	Feb. 9, 1999
DNA68866-1644	203283	Sep. 22, 1998
DNA68871-1638	203280	Sep. 22, 1998
DNA68880-1676	203319	Oct. 6, 1998
DNA68883-1691	203535	Dec. 15, 1998
DNA68885-1678	203311	Oct. 6, 1998
DNA71277-1636	203285	Sep. 22, 1998
DNA73727-1673	203459	Nov. 3, 1998
DNA73734-1680	203363	Oct. 20, 1998
DNA73735-1681	203356	Oct. 20, 1998
DNA76393-1664	203323	Oct. 6, 1998
DNA77301-1708	203407	Oct. 27, 1998
DNA77568-1626	203134	Aug. 18, 1998
DNA77626-1705	203536	Dec. 15, 1998
DNA81754-2532	203542	Dec. 15, 1998
DNA81757-2512	203543	Dec. 15, 1998
DNA82302-2529	203534	Dec. 15, 1998
DNA82340-2530	203547	Dec. 22, 1998
DNA83500-2506	203391	Oct. 29, 1998
DNA84920-2614	203966	Apr. 27, 1999
DNA85066-2534	203588	Jan. 12, 1999
DNA86571-2551	203660	Feb. 9, 1999
DNA87991-2540	203656	Feb. 9, 1999
DNA92238-2539	203602	Jan. 20, 1999

TABLE 7-continued

Material	ATCC Dep. No.	Deposit Date
DNA96042-2682	PTA-382	Jul. 20, 1999
DNA96787-2534	203589	Jan. 12, 1999
DNA125185-2806	PTA-1031	Dec. 7, 1999
DNA147531-2821	PTA-1185	Jan. 11, 2000
DNA115291-2681	PTA-202	Jun. 8, 1999
DNA164625-28890	PTA-1535	Mar. 21, 2000
DNA131639-2874	PTA-1784	Apr. 25, 2000
DNA79230-2525	203549	Dec. 22, 1998

[0421] These deposits were made under the provisions of the Budapest Treaty on the International Recognition of the Deposit of Microorganisms for the Purpose of Patent Procedure and the Regulations thereunder (Budapest Treaty). This assures maintenance of a viable culture of the deposit for 30 years from the date of deposit. The deposits will be made available by ATCC under the terms of the Budapest Treaty, and subject to an agreement between Genentech, Inc. and ATCC, which assures permanent and unrestricted availability of the progeny of the culture of the deposit to the public upon issuance of the pertinent U.S. patent or upon laying open to the public of any U.S. or foreign patent application, whichever comes first, and assures availability of the progeny to one determined by the U.S. Commissioner of Patents and Trademarks to be entitled thereto according to 35 USC § 122 and the Commissioner's rules pursuant thereto (including 37 CFR § 1.14 with particular reference to 886 OG 638).

[0422] The assignee of the present application has agreed that if a culture of the materials on deposit should die or be lost or destroyed when cultivated under suitable conditions, the materials will be promptly replaced on notification with another of the same. Availability of the deposited material is not to be construed as a license to practice the invention in contravention of the rights granted under the authority of any government in accordance with its patent laws.

Example 3

Use of PRO as a Hybridization Probe

[0423] The following method describes use of a nucleotide sequence encoding PRO as a hybridization probe.

[0424] DNA comprising the coding sequence of full-length or mature PRO as disclosed herein is employed as a probe to screen for homologous DNAs (such as those encoding naturally-occurring variants of PRO) in human tissue cDNA libraries or human tissue genomic libraries.

[0425] Hybridization and washing of filters containing either library DNAs is performed under the following high stringency conditions. Hybridization of radiolabeled PRO-derived probe to the filters is performed in a solution of 50% formamide, 5×SSC, 0.1% SDS, 0.1% sodium pyrophosphate, 50 mM sodium phosphate, pH 6.8, 2× Denhardt's solution, and 10% dextran sulfate at 42° C. for 20 hours. Washing of the filters is performed in an aqueous solution of 0.1×SSC and 0.1% SDS at 42° C.

[0426] DNAs having a desired sequence identity with the DNA encoding full-length native sequence PRO can then be identified using standard techniques known in the art.

Example 6

Expression of PRO in *E. Coli*

[0427] This example illustrates preparation of an unglycosylated form of PRO by recombinant expression in *E. coli*.

[0428] The DNA sequence encoding PRO is initially amplified using selected PCR primers. The primers should contain restriction enzyme sites which correspond to the restriction enzyme sites on the selected expression vector. A variety of expression vectors may be employed. An example of a suitable vector is pBR322 (derived from *E. coli*; see Bolivar et al., Gene, 2:95 (1977)) which contains genes for ampicillin and tetracycline resistance. The vector is digested with restriction enzyme and dephosphorylated. The PCR amplified sequences are then ligated into the vector. The vector will preferably include sequences which encode for an antibiotic resistance gene, a trp promoter, a polyhis leader (including the first six STII codons, polyhis sequence, and enterokinase cleavage site), the PRO coding region, lambda transcriptional terminator, and an argU gene.

[0429] The ligation mixture is then used to transform a selected *E. coli* strain using the methods described in Sambrook et al., supra. Transformants are identified by their ability to grow on LB plates and antibiotic resistant colonies are then selected. Plasmid DNA can be isolated and confirmed by restriction analysis and DNA sequencing.

[0430] Selected clones can be grown overnight in liquid culture medium such as LB broth supplemented with antibiotics. The overnight culture may subsequently be used to inoculate a larger scale culture. The cells are then grown to a desired optical density, during which the expression promoter is turned on.

[0431] After culturing the cells for several more hours, the cells can be harvested by centrifugation. The cell pellet obtained by the centrifugation can be solubilized using various agents known in the art, and the solubilized PRO protein can then be purified using a metal chelating column under conditions that allow tight binding of the protein.

[0432] PRO may be expressed in *E. coli* in a poly-His tagged form, using the following procedure. The DNA encoding PRO is initially amplified using selected PCR primers. The primers will contain restriction enzyme sites which correspond to the restriction enzyme sites on the selected expression vector, and other useful sequences providing for efficient and reliable translation initiation, rapid purification on a metal chelation column, and proteolytic removal with enterokinase. The PCR-amplified, poly-His tagged sequences are then ligated into an expression vector, which is used to transform an *E. coli* host based on strain 52 (W3110 fuhA(tonA) Ion galE rpoHts (htpRts) clpP(lacIq). Transformants are first grown in LB containing 50 mg/ml carbenicillin at 30° C. with shaking until an O.D.600 of 3-5 is reached. Cultures are then diluted 50-100 fold into CRAP media (prepared by mixing 3.57 g (NH₄)₂SO₄, 0.71 g sodium citrate-2H₂O, 1.07 g KCl, 5.36 g Difco yeast extract, 5.36 g Sheffield hycase SF in 500 mL water, as well as 110 mM MPOS, pH 7.3, 0.55% (w/v) glucose and 7 mM MgSO₄) and grown for approximately 20-30 hours at 30° C. with shaking. Samples are removed to verify expression by SDS-page analysis, and the bulk culture is centrifuged to pellet the cells. Cell pellets are frozen until purification and refolding.

[0433] *E. coli* paste from 0.5 to 1 L fermentations (6-10 g pellets) is resuspended in 10 volumes (w/v) in 7 M guanidine, 20 mM Tris, pH 8 buffer. Solid sodium sulfite and sodium tetrathionate is added to make final concentrations of 0.1M and 0.02 M, respectively, and the solution is stirred overnight at 4° C. This step results in a denatured protein with all cysteine residues blocked by sulfitolization. The solution is centrifuged at 40,000 rpm in a Beckman Ultracentrifuge for 30 min. The supernatant is diluted with 3-5 volumes of metal chelate column buffer (6 M guanidine, 20 mM Tris, pH 7.4) and filtered through 0.22 micron filters to clarify. The clarified extract is loaded onto a 5 ml Qiagen Ni-NTA metal chelate column equilibrated in the metal chelate column buffer. The column is washed with additional buffer containing 50 mM imidazole (Calbiochem, Utrol grade), pH 7.4. The protein is eluted with buffer containing 250 mM imidazole. Fractions containing the desired protein are pooled and stored at 4° C. Protein concentration is estimated by its absorbance at 280 nm using the calculated extinction coefficient based on its amino acid sequence.

[0434] The proteins are refolded by diluting the sample slowly into freshly prepared refolding buffer consisting of: 20 mM Tris, pH 8.6, 0.3 M NaCl, 2.5 M urea, 5 mM cysteine, 20 mM glycine and 1 mM EDTA. Refolding volumes are chosen so that the final protein concentration is between 50 to 100 micrograms/ml. The refolding solution is stirred gently at 4° C. for 12-36 hours. The refolding reaction is quenched by the addition of TFA to a final concentration of 0.4% (pH of approximately 3). Before further purification of the protein, the solution is filtered through a 0.22 micron filter and acetonitrile is added to 2-10% final concentration. The refolded protein is chromatographed on a Poros R1/H reversed phase column using a mobile buffer of 0.1% TFA with elution with a gradient of acetonitrile from 10 to 80%. Aliquots of fractions with A280 absorbance are analyzed on SDS polyacrylamide gels and fractions containing homogeneous refolded protein are pooled. Generally, the properly refolded species of most proteins are eluted at the lowest concentrations of acetonitrile since those species are the most compact with their hydrophobic interiors shielded from interaction with the reversed phase resin. Aggregated species are usually eluted at higher acetonitrile concentrations. In addition to resolving misfolded forms of from the desired form, the reversed phase step also removes endotoxin from the samples.

[0435] Fractions containing the desired folded PRO polypeptide are pooled and the acetonitrile removed using a gentle stream of nitrogen directed at the solution. Proteins are formulated into 20 mM Hepes, pH 6.8 with 0.14 M sodium chloride and 4% mannitol by dialysis or by gel filtration using G25 Superfine (Pharmacia) resins equilibrated in the formulation buffer and sterile filtered.

[0436] Many of the PRO polypeptides disclosed herein were successfully expressed as described above.

Example 7

Expression of PRO in Mammalian Cells

[0437] This example illustrates preparation of a potentially glycosylated form of PRO by recombinant expression in mammalian cells.

[0438] The vector, pRK5 (see EP 307,247, published Mar. 15, 1989), is employed as the expression vector. Optionally,

the PRO DNA is ligated into pRK5 with selected restriction enzymes to allow insertion of the PRO DNA using ligation methods such as described in Sambrook et al., supra. The resulting vector is called pRK5-PRO.

[0439] In one embodiment, the selected host cells may be 293 cells. Human 293 cells (ATCC CCL 1573) are grown to confluence in tissue culture plates in medium such as DMEM supplemented with fetal calf serum and optionally, nutrient components and/or antibiotics. About 10 µg pRK5-PRO DNA is mixed with about 1 µg DNA encoding the VA RNA gene [Thimmappaya et al., Cell, 31:543 (1982)] and dissolved in 500 µl of 1 mM Tris-HCl, 0.1 mM EDTA, 0.227 M CaCl₂. To this mixture is added, dropwise, 500 µl of 50 mM HEPES (pH 7.35), 280 mM NaCl, 1.5 mM NaPO₄, and a precipitate is allowed to form for 10 minutes at 25° C. The precipitate is suspended and added to the 293 cells and allowed to settle for about four hours at 37° C. The culture medium is aspirated off and 2 ml of 20% glycerol in PBS is added for 30 seconds. The 293 cells are then washed with serum free medium, fresh medium is added and the cells are incubated for about 5 days.

[0440] Approximately 24 hours after the transfections, the culture medium is removed and replaced with culture medium (alone) or culture medium containing 200 µCi/ml ³⁵S-cysteine and 200 µCi/ml ³⁵S-methionine. After a 12 hour incubation, the conditioned medium is collected, concentrated on a spin filter, and loaded onto a 15% SDS gel. The processed gel may be dried and exposed to film for a selected period of time to reveal the presence of PRO polypeptide. The cultures containing transfected cells may undergo further incubation (in serum free medium) and the medium is tested in selected bioassays.

[0441] In an alternative technique, PRO may be introduced into 293 cells transiently using the dextran sulfate method described by Somparyrac et al., Proc. Natl. Acad. Sci., 12:7575 (1981). 293 cells are grown to maximal density in a spinner flask and 700 µg pRK5-PRO DNA is added. The cells are first concentrated from the spinner flask by centrifugation and washed with PBS. The DNA-dextran precipitate is incubated on the cell pellet for four hours. The cells are treated with 20% glycerol for 90 seconds, washed with tissue culture medium, and re-introduced into the spinner flask containing tissue culture medium, 5 µg/ml bovine insulin and 0.1 µg/ml bovine transferrin. After about four days, the conditioned media is centrifuged and filtered to remove cells and debris. The sample containing expressed PRO can then be concentrated and purified by any selected method, such as dialysis and/or column chromatography.

[0442] In another embodiment, PRO can be expressed in CHO cells. The pRK5-PRO can be transfected into CHO cells using known reagents such as CaPO₄ or DEAE-dextran. As described above, the cell cultures can be incubated, and the medium replaced with culture medium (alone) or medium containing a radiolabel such as ³⁵S-methionine. After determining the presence of PRO polypeptide, the culture medium may be replaced with serum free medium. Preferably, the cultures are incubated for about 6 days, and then the conditioned medium is harvested. The medium containing the expressed PRO can then be concentrated and purified by any selected method.

[0443] Epitope-tagged PRO may also be expressed in host CHO cells. The PRO may be subcloned out of the pRK5

vector. The subclone insert can undergo PCR to fuse in frame with a selected epitope tag such as a poly-his tag into a Baculovirus expression vector. The poly-his tagged PRO insert can then be subcloned into a SV40 driven vector containing a selection marker such as DHFR for selection of stable clones. Finally, the CHO cells can be transfected (as described above) with the SV40 driven vector. Labeling may be performed, as described above, to verify expression. The culture medium containing the expressed poly-His tagged PRO can then be concentrated and purified by any selected method, such as by Ni^{2+} -chelate affinity chromatography.

[0444] PRO may also be expressed in CHO and/or COS cells by a transient expression procedure or in CHO cells by another stable expression procedure.

[0445] Stable expression in CHO cells is performed using the following procedure. The proteins are expressed as an IgG construct (immunoadhesin), in which the coding sequences for the soluble forms (e.g. extracellular domains) of the respective proteins are fused to an IgG1 constant region sequence containing the hinge, CH2 and CH2 domains and/or is a poly-His tagged form.

[0446] Following PCR amplification, the respective DNAs are subcloned in a CHO expression vector using standard techniques as described in Ausubel et al., *Current Protocols of Molecular Biology*, Unit 3.16, John Wiley and Sons (1997). CHO expression vectors are constructed to have compatible restriction sites 5' and 3' of the DNA of interest to allow the convenient shuttling of cDNA's. The vector used expression in CHO cells is as described in Lucas et al., *Nucl. Acids Res.* 24:9 (1774-1779) (1996), and uses the SV40 early promoter/enhancer to drive expression of the cDNA of interest and dihydrofolate reductase (DHFR). DHFR expression permits selection for stable maintenance of the plasmid following transfection.

[0447] Twelve micrograms of the desired plasmid DNA is introduced into approximately 10 million CHO cells using commercially available transfection reagents Superfect® (Quiagen), Dospere® or Eugene® (Boehringer Mannheim). The cells are grown as described in Lucas et al., supra. Approximately 3×10^{-7} cells are frozen in an ampule for further growth and production as described below.

[0448] The ampules containing the plasmid DNA are thawed by placement into water bath and mixed by vortexing. The contents are pipetted into a centrifuge tube containing 10 mLs of media and centrifuged at 1000 rpm for 5 minutes. The supernatant is aspirated and the cells are resuspended in 10 mL of selective media (0.2 μm filtered PS20 with 5% 0.2 μm diafiltered fetal bovine serum). The cells are then aliquoted into a 100 mL spinner containing 90 mL of selective media. After 1-2 days, the cells are transferred into a 250 mL spinner filled with 150 mL selective growth medium and incubated at 37° C. After another 2-3 days, 250 mL, 500 mL and 2000 mL spinners are seeded with 3×10^5 cells/mL. The cell media is exchanged with fresh media by centrifugation and resuspension in production medium. Although any suitable CHO media may be employed, a production medium described in U.S. Pat. No. 5,122,469, issued Jun. 16, 1992 may actually be used. A 3L production spinner is seeded at 1.2×10^6 cells/mL. On day 0, the cell number pH is determined. On day 1, the spinner is sampled and sparging with filtered air is commenced. On day 2, the spinner is sampled, the temperature shifted to 33°

C., and 30 mL of 500 g/L glucose and 0.6 mL of 10% antifoam (e.g., 35% polydimethylsiloxane emulsion, Dow Corning 365 Medical Grade Emulsion) taken. Throughout the production, the pH is adjusted as necessary to keep it at around 7.2. After 10 days, or until the viability dropped below 70%, the cell culture is harvested by centrifugation and filtering through a 0.22 μm filter. The filtrate was either stored at 4° C. or immediately loaded onto columns for purification.

[0449] For the poly-His tagged constructs, the proteins are purified using a Ni-NTA column (Qiagen). Before purification, imidazole is added to the conditioned media to a concentration of 5 mM. The conditioned media is pumped onto a 6 ml Ni-NTA column equilibrated in 20 mM Hepes, pH 7.4, buffer containing 0.3 M NaCl and 5 mM imidazole at a flow rate of 4-5 ml/min. at 4° C. After loading, the column is washed with additional equilibration buffer and the protein eluted with equilibration buffer containing 0.25 M imidazole. The highly purified protein is subsequently desalted into a storage buffer containing 10 mM Hepes, 0.14 M NaCl and 4% mannitol, pH 6.8, with a 25 ml G25 Superfine (Pharmacia) column and stored at -80° C.

[0450] Immunoadhesin (Fc-containing) constructs are purified from the conditioned media as follows. The conditioned medium is pumped onto a 5 ml Protein A column (Pharmacia) which had been equilibrated in 20 mM Na phosphate buffer, pH 6.8. After loading, the column is washed extensively with equilibration buffer before elution with 100 mM citric acid, pH 3.5. The eluted protein is immediately neutralized by collecting 1 ml fractions into tubes containing 275 μL of 1 M Tris buffer, pH 9. The highly purified protein is subsequently desalted into storage buffer as described above for the poly-His tagged proteins. The homogeneity is assessed by SDS polyacrylamide gels and by N-terminal amino acid sequencing by Edman degradation.

[0451] Many of the PRO polypeptides disclosed herein were successfully expressed as described above.

Example 8

Expression of PRO in Yeast

[0452] The following method describes recombinant expression of PRO in yeast.

[0453] First, yeast expression vectors are constructed for intracellular production or secretion of PRO from the ADH2/GAPDH promoter. DNA encoding PRO and the promoter is inserted into suitable restriction enzyme sites in the selected plasmid to direct intracellular expression of PRO. For secretion, DNA encoding PRO can be cloned into the selected plasmid, together with DNA encoding the ADH2/GAPDH promoter, a native PRO signal peptide or other mammalian signal peptide, or, for example, a yeast alpha-factor or invertase secretory signal/leader sequence, and linker sequences (if needed) for expression of PRO.

[0454] Yeast cells, such as yeast strain AB110, can then be transformed with the expression plasmids described above and cultured in selected fermentation media. The transformed yeast supernatants can be analyzed by precipitation with 10% trichloroacetic acid and separation by SDS-PAGE, followed by staining of the gels with Coomassie Blue stain.

[0455] Recombinant PRO can subsequently be isolated and purified by removing the cells from the fermentation

medium by centrifugation and then concentrating the medium using selected cartridge filters. The concentrate containing PRO may further be purified using selected column chromatography resins.

[0456] Many of the PRO polypeptides disclosed herein were successfully expressed as described above.

Example 9

Expression of PRO in Baculovirus-Infected Insect Cells

[0457] The following method describes recombinant expression of PRO in Baculovirus-infected insect cells.

[0458] The sequence coding for PRO is fused upstream of an epitope tag contained a baculovirus expression vector. Such epitope tags include poly-his tags and immunoglobulin tags (like Fc regions of IgG). A variety of plasmids may be employed, including plasmids derived from commercially available plasmids such as pVL1393 (Novagen). Briefly, the sequence encoding PRO or the desired portion of the coding sequence of PRO such as the sequence encoding the extracellular domain of a transmembrane protein or the sequence encoding the mature protein if the protein is extracellular is amplified by PCR with primers complementary to the 5' and 3' regions. The 5' primer may incorporate flanking (selected) restriction enzyme sites. The product is then digested with those selected restriction enzymes and subcloned into the expression vector.

[0459] Recombinant baculovirus is generated by co-transfecting the above plasmid and BaculoGold™ virus DNA (Pharmingen) into *Spodoptera frugiperda* ("Sf9") cells (ATCC CRL 1711) using lipofectin (commercially available from GIBCO-BRL). After 4-5 days of incubation at 28° C., the released viruses are harvested and used for further amplifications. Viral infection and protein expression are performed as described by O'Reilley et al., Baculovirus expression vectors: A Laboratory Manual, Oxford: Oxford University Press (1994).

[0460] Expressed poly-his tagged PRO can then be purified, for example, by Ni²⁺-chelate affinity chromatography as follows. Extracts are prepared from recombinant virus-infected Sf9 cells as described by Rupert et al., Nature, 362:175-179 (1993). Briefly, Sf9 cells are washed, resuspended in sonication buffer (25 mL Hepes, pH 7.9; 12.5 mM MgCl₂; 0.1 mM EDTA; 10% glycerol; 0.1% NP-40; 0.4 M KCl), and sonicated twice for 20 seconds on ice. The sonicates are cleared by centrifugation, and the supernatant is diluted 50-fold in loading buffer (50 mM phosphate, 300 mM NaCl, 10% glycerol, pH 7.8) and filtered through a 0.45 μ m filter. A Ni²⁺-NTA agarose column (commercially available from Qiagen) is prepared with a bed volume of 5 mL, washed with 25 mL of water and equilibrated with 25 mL of loading buffer. The filtered cell extract is loaded onto the column at 0.5 mL per minute. The column is washed to baseline A₂₈₀ with loading buffer, at which point fraction collection is started. Next, the column is washed with a secondary wash buffer (50 mM phosphate; 300 mM NaCl, 10% glycerol, pH 6.0), which elutes nonspecifically bound protein. After reaching A₂₈₀ baseline again, the column is developed with a 0 to 500 mM Imidazole gradient in the secondary wash buffer. One mL fractions are collected and analyzed by SDS-PAGE and silver staining or Western blot

with Ni²⁺-NTA-conjugated to alkaline phosphatase (Qiagen). Fractions containing the eluted His₁₀-tagged PRO are pooled and dialyzed against loading buffer.

[0461] Alternatively, purification of the IgG tagged (or Fc tagged) PRO can be performed using known chromatography techniques, including for instance, Protein A or protein G column chromatography.

[0462] Many of the PRO polypeptides disclosed herein were successfully expressed as described above.

Example 10

Preparation of Antibodies That Bind PRO

[0463] This example illustrates preparation of monoclonal antibodies which can specifically bind PRO.

[0464] Techniques for producing the monoclonal antibodies are known in the art and are described, for instance, in Goding, supra. Immunogens that may be employed include purified PRO, fusion proteins containing PRO, and cells expressing recombinant PRO on the cell surface. Selection of the immunogen can be made by the skilled artisan without undue experimentation.

[0465] Mice, such as Balb/c, are immunized with the PRO immunogen emulsified in complete Freund's adjuvant and injected subcutaneously or intraperitoneally in an amount from 1-100 micrograms. Alternatively, the immunogen is emulsified in MPL-TDM adjuvant (Ribi Immunochemical Research, Hamilton, Mont.) and injected into the animal's hind foot pads. The immunized mice are then boosted 10 to 12 days later with additional immunogen emulsified in the selected adjuvant. Thereafter, for several weeks, the mice may also be boosted with additional immunization injections. Serum samples may be periodically obtained from the mice by retro-orbital bleeding for testing in ELISA assays to detect anti-PRO antibodies.

[0466] After a suitable antibody titer has been detected, the animals "positive" for antibodies can be injected with a final intravenous injection of PRO. Three to four days later, the mice are sacrificed and the spleen cells are harvested. The spleen cells are then fused (using 35% polyethylene glycol) to a selected murine myeloma cell line such as P3X63AgU.1, available from ATCC, No. CRL 1597. The fusions generate hybridoma cells which can then be plated in 96 well tissue culture plates containing HAT (hypoxanthine, aminopterin, and thymidine) medium to inhibit proliferation of non-fused cells, myeloma hybrids, and spleen cell hybrids.

[0467] The hybridoma cells will be screened in an ELISA for reactivity against PRO. Determination of "positive" hybridoma cells secreting the desired monoclonal antibodies against PRO is within the skill in the art.

[0468] The positive hybridoma cells can be injected intraperitoneally into syngeneic Balb/c mice to produce ascites containing the anti-PRO monoclonal antibodies. Alternatively, the hybridoma cells can be grown in tissue culture flasks or roller bottles. Purification of the monoclonal antibodies produced in the ascites can be accomplished using ammonium sulfate precipitation, followed by gel exclusion chromatography. Alternatively, affinity chromatography based upon binding of antibody to protein A or protein G can be employed.

Example 11

Purification of PRO Polypeptides Using Specific Antibodies

[0469] Native or recombinant PRO polypeptides may be purified by a variety of standard techniques in the art of protein purification. For example, pro-PRO polypeptide, mature PRO polypeptide, or pre-PRO polypeptide is purified by immunoaffinity chromatography using antibodies specific for the PRO polypeptide of interest. In general, an immunoaffinity column is constructed by covalently coupling the anti-PRO polypeptide antibody to an activated chromatographic resin.

[0470] Polyclonal immunoglobulins are prepared from immune sera either by precipitation with ammonium sulfate or by purification on immobilized Protein A (Pharmacia LKB Biotechnology, Piscataway, N.J.). Likewise, monoclonal antibodies are prepared from mouse ascites fluid by ammonium sulfate precipitation or chromatography on immobilized Protein A. Partially purified immunoglobulin is covalently attached to a chromatographic resin such as CnBr-activated SEPHAROSE™ (Pharmacia LKB Biotechnology). The antibody is coupled to the resin, the resin is blocked, and the derivative resin is washed according to the manufacturer's instructions.

[0471] Such an immunoaffinity column is utilized in the purification of PRO polypeptide by preparing a fraction from cells containing PRO polypeptide in a soluble form. This preparation is derived by solubilization of the whole cell or of a subcellular fraction obtained via differential centrifugation by the addition of detergent or by other methods well known in the art. Alternatively, soluble PRO polypeptide containing a signal sequence may be secreted in useful quantity into the medium in which the cells are grown.

[0472] A soluble PRO polypeptide-containing preparation is passed over the immunoaffinity column, and the column is washed under conditions that allow the preferential absorbance of PRO polypeptide (e.g., high ionic strength buffers in the presence of detergent). Then, the column is eluted under conditions that disrupt antibody/PRO polypeptide binding (e.g., a low pH buffer such as approximately pH 2-3, or a high concentration of a chaotrope such as urea or thiocyanate ion), and PRO polypeptide is collected.

Example 12

Drug Screening

[0473] This invention is particularly useful for screening compounds by using PRO polypeptides or binding fragment thereof in any of a variety of drug screening techniques. The PRO polypeptide or fragment employed in such a test may either be free in solution, affixed to a solid support, borne on a cell surface, or located intracellularly. One method of drug screening utilizes eukaryotic or prokaryotic host cells which are stably transformed with recombinant nucleic acids expressing the PRO polypeptide or fragment. Drugs are screened against such transformed cells in competitive binding assays. Such cells, either in viable or fixed form, can be used for standard binding assays. One may measure, for example, the formation of complexes between PRO polypeptide or a fragment and the agent being tested.

Alternatively, one can examine the diminution in complex formation between the PRO polypeptide and its target cell or target receptors caused by the agent being tested.

[0474] Thus, the present invention provides methods of screening for drugs or any other agents which can affect a PRO polypeptide-associated disease or disorder. These methods comprise contacting such an agent with a PRO polypeptide or fragment thereof and assaying (i) for the presence of a complex between the agent and the PRO polypeptide or fragment, or (ii) for the presence of a complex between the PRO polypeptide or fragment and the cell, by methods well known in the art. In such competitive binding assays, the PRO polypeptide or fragment is typically labeled. After suitable incubation, free PRO polypeptide or fragment is separated from that present in bound form, and the amount of free or uncomplexed label is a measure of the ability of the particular agent to bind to PRO polypeptide or to interfere with the PRO polypeptide/cell complex.

[0475] Another technique for drug screening provides high throughput screening for compounds having suitable binding affinity to a polypeptide and is described in detail in WO 84/03564, published on Sep. 13, 1984. Briefly stated, large numbers of different small peptide test compounds are synthesized on a solid substrate, such as plastic pins or some other surface. As applied to a PRO polypeptide, the peptide test compounds are reacted with PRO polypeptide and washed. Bound PRO polypeptide is detected by methods well known in the art. Purified PRO polypeptide can also be coated directly onto plates for use in the aforementioned drug screening techniques. In addition, non-neutralizing antibodies can be used to capture the peptide and immobilize it on the solid support.

[0476] This invention also contemplates the use of competitive drug screening assays in which neutralizing antibodies capable of binding PRO polypeptide specifically compete with a test compound for binding to PRO polypeptide or fragments thereof. In this manner, the antibodies can be used to detect the presence of any peptide which shares one or more antigenic determinants with PRO polypeptide.

Example 13

Rational Drug Design

[0477] The goal of rational drug design is to produce structural analogs of biologically active polypeptide of interest (i.e., a PRO polypeptide) or of small molecules with which they interact, e.g., agonists, antagonists, or inhibitors. Any of these examples can be used to fashion drugs which are more active or stable forms of the PRO polypeptide or which enhance or interfere with the function of the PRO polypeptide in vivo (c.f., Hodgson, Bio/Technology, 9: 19-21(1991)).

[0478] In one approach, the three-dimensional structure of the PRO polypeptide, or of a PRO polypeptide-inhibitor complex, is determined by X-ray crystallography, by computer modeling or, most typically, by a combination of the two approaches. Both the shape and charges of the PRO polypeptide must be ascertained to elucidate the structure and to determine active site(s) of the molecule. Less often, useful information regarding the structure of the PRO polypeptide may be gained by modeling based on the structure of homologous proteins. In both cases, relevant

structural information is used to design analogous PRO polypeptide-like molecules or to identify efficient inhibitors. Useful examples of rational drug design may include molecules which have improved activity or stability as shown by Braxton and Wells, *Biochemistry*, 31:7796-7801 (1992) or which act as inhibitors, agonists, or antagonists of native peptides as shown by Athauda et al., *J. Biochem.*, 113:742-746(1993).

[0479] It is also possible to isolate a target-specific antibody, selected by functional assay, as described above, and then to solve its crystal structure. This approach, in principle, yields a pharmacore upon which subsequent drug design can be based. It is possible to bypass protein crystallography altogether by generating anti-idiotypic antibodies (anti-ids) to a functional, pharmacologically active antibody. As a mirror image of a mirror image, the binding site of the anti-ids would be expected to be an analog of the original receptor. The anti-id could then be used to identify and isolate peptides from banks of chemically or biologically produced peptides. The isolated peptides would then act as the pharmacore.

[0480] By virtue of the present invention, sufficient amounts of the PRO polypeptide may be made available to perform such analytical studies as X-ray crystallography. In addition, knowledge of the PRO polypeptide amino acid sequence provided herein will provide guidance to those employing computer modeling techniques in place of or in addition to X-ray crystallography.

Example 14

Pericyte c-Fos Induction (Assay 93)

[0481] This assay shows that certain polypeptides of the invention act to induce the expression of c-fos in pericyte cells and, therefore, are useful not only as diagnostic markers for particular types of pericyte-associated tumors but also for giving rise to antagonists which would be expected to be useful for the therapeutic treatment of pericyte-associated tumors. Induction of c-fos expression in pericytes is also indicative of the induction of angiogenesis and, as such, PRO polypeptides capable of inducing the expression of c-fos would be expected to be useful for the treatment of conditions where induced angiogenesis would be beneficial including, for example, wound healing, and the like. Specifically, on day 1, pericytes are received from VEC Technologies and all but 5 ml of media is removed from flask. On day 2, the pericytes are trypsinized, washed, spun and then plated onto 96 well plates. On day 7, the media is removed and the pericytes are treated with 100 μ l of PRO polypeptide test samples and controls (positive control=DME+5% serum+PDGF at 500 ng/ml; negative control=protein 32). Replicates are averaged and SD/CV are determined. Fold increase over Protein 32 (buffer control) value indicated by chemiluminescence units (RLU) luminometer reading verses frequency is plotted on a histogram. Two-fold above Protein 32 value is considered positive for the assay. ASY Matrix: Growth media=low glucose DMEM=20% FBS+1 $\times$ pen strep+1 $\times$ fungizone. Assay Media=low glucose DMEM+5% FBS.

[0482] The following polypeptides tested positive in this assay: PRO1347 and PRO1340.

Example 15

Ability of PRO Polypeptides to Stimulate the Release of Proteoglycans from Cartilage (Assay 97)

[0483] The ability of various PRO polypeptides to stimulate the release of proteoglycans from cartilage tissue was tested as follows.

[0484] The metacarpophalangeal joint of 4-6 month old pigs was aseptically dissected, and articular cartilage was removed by free hand slicing being careful to avoid the underlying bone. The cartilage was minced and cultured in bulk for 24 hours in a humidified atmosphere of 95% air, 5% CO₂ in serum free (SF) media (DME/F12 1:1) with 0.1% BSA and 100 U/ml penicillin and 100 μ g/ml streptomycin. After washing three times, approximately 100 mg of articular cartilage was aliquoted into micronics tubes and incubated for an additional 24 hours in the above SF media. PRO polypeptides were then added at 1% either alone or in combination with 18 ng/ml interleukin-1 α , a known stimulator of proteoglycan release from cartilage tissue. The supernatant was then harvested and assayed for the amount of proteoglycans using the 1,9-dimethyl-methylene blue (DMB) colorimetric assay (Farndale and Buttle, *Biochem. Biophys. Acta* 883:173-177 (1985)). A positive result in this assay indicates that the test polypeptide will find use, for example, in the treatment of sports-related joint problems, articular cartilage defects, osteoarthritis or rheumatoid arthritis.

[0485] When various PRO polypeptides were tested in the above assay, the polypeptides demonstrated a marked ability to stimulate release of proteoglycans from cartilage tissue both basally and after stimulation with interleukin-1 α and at 24 and 72 hours after treatment, thereby indicating that these PRO polypeptides are useful for stimulating proteoglycan release from cartilage tissue. As such, these PRO polypeptides are useful for the treatment of sports-related joint problems, articular cartilage defects, osteoarthritis or rheumatoid arthritis. The polypeptides testing positive in this assay are: PRO1565, PRO1693, PRO1801 and PRO10096.

Example 16

Detection of Polypeptides That Affect Glucose or FFA Uptake in Skeletal Muscle (Assay 106)

[0486] This assay is designed to determine whether PRO polypeptides show the ability to affect glucose or FFA uptake by skeletal muscle cells. PRO polypeptides testing positive in this assay would be expected to be useful for the therapeutic treatment of disorders where either the stimulation or inhibition of glucose uptake by skeletal muscle would be beneficial including, for example, diabetes or hyper- or hypo-insulinemia.

[0487] In a 96 well format, PRO polypeptides to be assayed are added to primary rat differentiated skeletal muscle, and allowed to incubate overnight. Then fresh media with the PRO polypeptide and $\pm$ insulin are added to the wells. The sample media is then monitored to determine glucose and FFA uptake by the skeletal muscle cells. The insulin will stimulate glucose and FFA uptake by the skeletal muscle, and insulin in media without the PRO polypeptide is used as a positive control, and a limit for scoring. As the PRO polypeptide being tested may either stimulate or inhibit

glucose and FFA uptake, results are scored as positive in the assay if greater than 1.5 times or less than 0.5 times the insulin control.

[0488] The following PRO polypeptides tested positive as either stimulators or inhibitors of glucose and/or FFA uptake in this assay: PRO4405.

Example 17

Identification of PRO Polypeptides That Stimulate TNF- α Release In Human Blood (Assay 128)

[0489] This assay shows that certain PRO polypeptides of the present invention act to stimulate the release of TNF- α in human blood. PRO polypeptides testing positive in this assay are useful for, among other things, research purposes where stimulation of the release of TNF- α would be desired and for the therapeutic treatment of conditions wherein enhanced TNF- α release would be beneficial. Specifically, 200 μ l of human blood supplemented with 50 mM Hepes buffer (pH 7.2) is aliquotted per well in a 96 well test plate. To each well is then added 300 μ l of either the test PRO polypeptide in 50 mM Hepes buffer (at various concentrations) or 50 mM Hepes buffer alone (negative control) and the plates are incubated at 37° C. for 6 hours. The samples are then centrifuged and 50 μ l of plasma is collected from each well and tested for the presence of TNF- α by ELISA assay. A positive in the assay is a higher amount of TNF- α in the PRO polypeptide treated samples as compared to the negative control samples.

[0490] The following PRO polypeptides tested positive in this assay: PRO263, PRO295, PRO1282, PRO1063, PRO1356, PRO3543, and PRO5990.

Example 18

Tumor Versus Normal Differential Tissue Expression Distribution

[0491] Oligonucleotide probes were constructed from some of the PRO polypeptide-encoding nucleotide sequences shown in the accompanying figures for use in quantitative PCR amplification reactions. The oligonucleotide probes were chosen so as to give an approximately 200-600 base pair amplified fragment from the 3' end of its associated template in a standard PCR reaction. The oligonucleotide probes were employed in standard quantitative PCR amplification reactions with cDNA libraries isolated from different human tumor and normal human tissue samples and analyzed by agarose gel electrophoresis so as to obtain a quantitative determination of the level of expression of the PRO polypeptide-encoding nucleic acid in the various tumor and normal tissues tested. β -actin was used as a control to assure that equivalent amounts of nucleic acid was used in each reaction. Identification of the differential expression of the PRO polypeptide-encoding nucleic acid in one or more tumor tissues as compared to one or more normal tissues of the same tissue type renders the molecule useful diagnostically for the determination of the presence or absence of tumor in a subject suspected of possessing a tumor as well as therapeutically as a target for the treatment of a tumor in a subject possessing such a tumor. These assays provided the following results.

Molecule	is more highly expressed in:	as compared to:
DNA26843-1389	normal lung rectum tumor	lung tumor normal rectum
DNA30867-1335	normal kidney	kidney tumor
DNA40621-1440	normal lung	lung tumor
DNA40625-1189	normal lung	lung tumor
DNA45409-2511	melanoma tumor	normal akin
DNA56406-1704	kidney tumor normal skin	normal kidney melanoma tumor
DNA56410-1414	normal stomach	stomach tumor
DNA56436-1448	normal skin	melanoma tumor
DNA56855-1447	normal esophagus rectum tumor	esophageal tumor normal rectum
DNA56860-1510	normal kidney rectum tumor	kidney tumor normal rectum
DNA56862-1343	kidney tumor normal lung	normal kidney lung tumor
DNA56868-1478	normal stomach normal lung	stomach tumor lung tumor
DNA56869-1545	normal esophagus normal skin	esophageal tumor melanoma tumor
DNA57704-1452	normal stomach rectum tumor	stomach tumor normal rectum
DNA58723-1588	normal stomach kidney tumor normal akin	stomach tumor normal kidney melanoma tumor
DNA57827-1493	normal stomach normal skin	stomach tumor melanoma tumor
DNA58737-1473	esophageal tumor normal stomach	normal esophagus stomach tumor
DNA58846-1409	lung tumor	normal lung
DNA58850-1495	esophageal tumor kidney tumor	normal esophagus normal kidney
DNA58855-1422	normal stomach rectum tumor	stomach tumor normal rectum
DNA59211-1450	normal kidney	kidney tumor
DNA59212-1627	normal skin	melanoma tumor
DNA59213-1487	normal stomach normal skin	stomach tumor melanoma tumor
DNA59605-1418	melanoma tumor	normal skin
DNA59609-1470	esophageal tumor	normal esophagus
DNA59610-1556	esophageal tumor lung tumor normal skin	normal esophagus normal lung melanoma tumor
DNA59837-2545	normal skin	melanoma tumor
DNA59844-2542	normal skin esophageal tumor	melanoma tumor normal esophagus
DNA59854-1459	normal esophagus stomach tumor normal lung	esophageal tumor normal stomach lung tumor
DNA60625-1507	normal lung	lung tumor
DNA60629-1481	normal esophagus normal rectum	esophageal tumor rectum tumor
DNA61755-1554	normal stomach kidney tumor	stomach tumor normal kidney
DNA62812-1594	normal stomach normal lung normal rectum normal skin	stomach tumor lung tumor rectum tumor melanoma tumor
DNA62815-1576	esophageal tumor	normal esophagus
DNA64881-1602	normal stomach normal lung	stomach tumor lung tumor
DNA64902-1667	esophageal tumor kidney tumor	normal esophagus normal kidney
DNA65403-1565	normal esophagus	esophageal tumor
DNA66308-1537	normal lung	lung tumor
DNA66519-1535	kidney tumor	normal kidney
DNA66521-1583	normal esophagus normal stomach normal lung normal rectum normal skin	esophageal tumor stomach tumor lung tumor rectum tumor melanoma tumor
DNA66658-1584	normal lung melanoma tumor	lung tumor normal skin
DNA66660-1585	lung tumor	normal lung

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Molecule	is more highly expressed in:	as compared to:
DNA66674-1599	kidney tumor	normal kidney
	normal lung	lung tumor
DNA68862-2546	melanoma tumor	normal skin
DNA68866-1644	normal stomach	stomach tumor
DNA68871-1638	lung tumor	normal lung
	normal skin	melanoma tumor
DNA68880-1676	normal lung	lung tumor
	normal skin	melanoma tumor
DNA68883-1691	esophageal tumor	normal esophagus
DNA68885-1678	lung tumor	normal lung
DNA71277-1636	normal stomach	stomach tumor
DNA73734-1680	normal lung	lung tumor
DNA73735-1681	esophageal tumor	normal esophagus
	normal kidney	kidney tumor
	lung tumor	normal lung
	normal skin	melanoma tumor
DNA76393-1664	esophageal tumor	normal esophagus
	stomach tumor	normal stomach
	lung tumor	normal lung
	rectum tumor	normal rectum
DNA77568-1626	normal stomach	stomach tumor
	lung tumor	normal lung
DNA77626-1705	normal rectum	rectum tumor
DNA81754-2532	normal skin	melanoma tumor
DNA81757-2512	esophageal tumor	normal esophagus
	normal stomach	stomach tumor
	melanoma tumor	normal skin
DNA82302-2529	normal stomach	stomach tumor
	normal lung	lung tumor
DNA82340-2530	normal esophagus	esophageal tumor
DNA85066-2534	lung tumor	normal lung
	normal skin	melanoma tumor
DNA87991-2540	esophageal tumor	normal esophagus
DNA92238-2539	normal skin	melanoma tumor
DNA96787-2534	normal kidney	kidney tumor

Example 19

Identification of Receptor/Ligand Interactions

[0492] In this assay, various PRO polypeptides are tested for ability to bind to a panel of potential receptor or ligand molecules for the purpose of identifying receptor/ligand interactions. The identification of a ligand for a known receptor, a receptor for a known ligand or a novel receptor/ligand pair is useful for a variety of indications including, for example, targeting bioactive molecules (linked to the ligand or receptor) to a cell known to express the receptor or ligand, use of the receptor or ligand as a reagent to detect the presence of the ligand or receptor in a composition suspected of containing the same, wherein the composition may comprise cells suspected of expressing the ligand or receptor, modulating the growth of or another biological or immunological activity of a cell known to express or respond to the receptor or ligand, modulating the immune response of cells or toward cells that express the receptor or ligand, allowing the preparation of agonists, antagonists and/or antibodies directed against the receptor or ligand which will modulate the growth of or a biological or immunological activity of a cell expressing the receptor or ligand, and various other indications which will be readily apparent to the ordinarily skilled artisan.

[0493] The assay is performed as follows. A PRO polypeptide of the present invention suspected of being a ligand for a receptor is expressed as a fusion protein containing the Fc domain of human IgG (an immunoadhesin).

Receptor-ligand binding is detected by allowing interaction of the immunoadhesin polypeptide with cells (e.g. Cos cells) expressing candidate PRO polypeptide receptors and visualization of bound immunoadhesin with fluorescent reagents directed toward the Fc fusion domain and examination by microscope. Cells expressing candidate receptors are produced by transient transfection, in parallel, of defined subsets of a library of cDNA expression vectors encoding PRO polypeptides that may function as receptor molecules. Cells are then incubated for 1 hour in the presence of the PRO polypeptide immunoadhesin being tested for possible receptor binding. The cells are then washed and fixed with paraformaldehyde. The cells are then incubated with fluorescent conjugated antibody directed against the Fc portion of the PRO polypeptide immunoadhesin (e.g. FITC conjugated goat anti-human-Fc antibody). The cells are then washed again and examined by microscope. A positive interaction is judged by the presence of fluorescent labeling of cells transfected with cDNA encoding a particular PRO polypeptide receptor or pool of receptors and an absence of similar fluorescent labeling of similarly prepared cells that have been transfected with other cDNA or pools of cDNA. If a defined pool of cDNA expression vectors is judged to be positive for interaction with a PRO polypeptide immunoadhesin, the individual cDNA species that comprise the pool are tested individually (the pool is "broken down") to determine the specific cDNA that encodes a receptor able to interact with the PRO polypeptide immunoadhesin.

[0494] In another embodiment of this assay, an epitope-tagged potential ligand PRO polypeptide (e.g. 8 histidine "His" tag) is allowed to interact with a panel of potential receptor PRO polypeptide molecules that have been expressed as fusions with the Fc domain of human IgG (immunoadhesins). Following a 1 hour co-incubation with the epitope tagged PRO polypeptide, the candidate receptors are each immunoprecipitated with protein A beads and the beads are washed. Potential ligand interaction is determined by western blot analysis of the immunoprecipitated complexes with antibody directed towards the epitope tag. An interaction is judged to occur if a band of the anticipated molecular weight of the epitope tagged protein is observed in the western blot analysis with a candidate receptor, but is not observed to occur with the other members of the panel of potential receptors.

[0495] Using these assays, the following receptor/ligand interactions have been herein identified:

- [0496] (1) PRO10272 binds to PRO5801.
- [0497] (2) PRO20110 binds to the human IL-17 receptor (Yao et al., *Cytokine* 9(11):794-800 (1997); also herein designated as PRO1) and to PRO20040.
- [0498] (3) PRO10096 binds to PRO20233.
- [0499] (4) PRO19670 binds to PRO1890.

[0500] The foregoing written specification is considered to be sufficient to enable one skilled in the art to practice the invention. The present invention is not to be limited in scope by the construct deposited, since the deposited embodiment is intended as a single illustration of certain aspects of the invention and any constructs that are functionally equivalent are within the scope of this invention. The deposit of material herein does not constitute an admission that the

written description herein contained is inadequate to enable the practice of any aspect of the invention, including the best mode thereof, nor is it to be construed as limiting the scope of the claims to the specific illustrations that it represents.

Indeed, various modifications of the invention in addition to those shown and described herein will become apparent to those skilled in the art from the foregoing description and fall within the scope of the appended claims.

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Tyr Ile Arg Asp Phe Gln Lys Ile Ser Leu Arg Val Glu Ala Asn
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gcgaaccagc agctgaattt cacagaagct aaggaggcct gtaggctgct	350
gggactaagt ttggccggca aggaccaagt tgaaacagcc ttgaaagcta	400
gctttgaaac ttgcagctat ggctgggttg gagatggatt cgtggctatc	450
tctaggatta gcccaaacc ccaagtgtgg aaaaatggg tggggtgtcct	500
gatttggaag gttccagtga gccgacagtt tgcagcctat tgttacaact	550
catctgatac ttggactaac tcgtgcattc cagaaattat caccaccaa	600
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cagtgcagct acctactcgg tggcatcccc ttactctaca atacctgccc	700
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aagaaaactg ataaaaacc agaagagtcc aagagtcaa gcaaaactac	1100
cgtgcgatgc ctggaagctg aagtttagat gagacagaaa tgaggagaca	1150
cacctgaggg ttggtttctt catgctcctt accctgcccc agctggggaa	1200
atcaaaaggg ccaaagaacc aaagaagaaa gtccaccctt ggttcctaac	1250
tggaatcagc tcaggactgc cattggacta tggagtgcac caaagagaat	1300
gcccttctoc ttattgtaac cctgtctgga tctatcctc ctacctccaa	1350

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agcttccac ggcttttcta gcctggctat gtcctaataa tatccactg      1400
ggagaaagga gttttgcaaa gtgcaaggac ctaaaacatc tcatcagtat      1450
ccagtggtaa aaaggcctcc tggctgtctg aggctagggtg ggttgaaagc      1500
caaggagtca ctgagaccaa ggctttctct actgattccg cagctcagac      1550
cctttcttca gctctgaaag agaaacacgt atcccacctg acatgtcctt      1600
ctgagcccgg taagagcaaa agaatggcag aaaagttag ccctgaaag      1650
ccatggagat ttcataact tgagacctaa tctctgtaaa gctaaaataa      1700
agaaatagaa caaggctgag gatacgacag tacactgtca gcagggactg      1750
taaacacaga cagggtcaaa gtgttttctc tgaacacatt gagttggaat      1800
cactgttagt aacacacaca ctacttttt ctggtctcta cactgctga      1850
tattttctct aggaaatata cttttacaag taacaaaaat aaaaactctt      1900
ataaatttct atttttatct gagttacaga aatgattact aaggaagatt      1950
actcagtaat ttgtttaaaa agtaataaaa ttcaacaaac atttgctgaa      2000
tagctactat atgtcaagtg ctgtgcaagg tattacactc tgtaattgaa      2050
tattattcct caaaaaattg cacatagtag aacgctatct gggaagctat      2100
ttttttcagt ttgatatatt ctagcttctc tacttccaaa ctaattttta      2150
tttttgctga gactaatctt attcatittc tctaatatgg caaccattat      2200
aaccttaatt tattattaac atacctaaga agtacattgt tacctctata      2250
taccaaagca cattttaaaa gtgccattaa caaatgtatc actagccctc      2300
ctttttccaa caagaaggga ctgagagatg cagaaatatt tgtgacaaaa      2350
aattaaagca tttagaaaaa tt                                     2372

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<210> SEQ ID NO 6

<211> LENGTH: 322

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 6

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Met Ala Arg Cys Phe Ser Leu Val Leu Leu Thr Ser Ile Trp
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Thr Thr Arg Leu Leu Val Gln Gly Ser Leu Arg Ala Glu Glu Leu
20          25          30

Ser Ile Gln Val Ser Cys Arg Ile Met Gly Ile Thr Leu Val Ser
35          40          45

Lys Lys Ala Asn Gln Gln Leu Asn Phe Thr Glu Ala Lys Glu Ala
50          55          60

Cys Arg Leu Leu Gly Leu Ser Leu Ala Gly Lys Asp Gln Val Glu
65          70          75

Thr Ala Leu Lys Ala Ser Phe Glu Thr Cys Ser Tyr Gly Trp Val
80          85          90

Gly Asp Gly Phe Val Val Ile Ser Arg Ile Ser Pro Asn Pro Lys
95          100         105

Cys Gly Lys Asn Gly Val Gly Val Leu Ile Trp Lys Val Pro Val
110         115         120

Ser Arg Gln Phe Ala Ala Tyr Cys Tyr Asn Ser Ser Asp Thr Trp

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	125		130		135
Thr Asn Ser Cys Ile Pro Glu Ile Ile Thr Thr Lys Asp Pro Ile					
	140		145		150
Phe Asn Thr Gln Thr Ala Thr Gln Thr Thr Glu Phe Ile Val Ser					
	155		160		165
Asp Ser Thr Tyr Ser Val Ala Ser Pro Tyr Ser Thr Ile Pro Ala					
	170		175		180
Pro Thr Thr Thr Pro Pro Ala Pro Ala Ser Thr Ser Ile Pro Arg					
	185		190		195
Arg Lys Lys Leu Ile Cys Val Thr Glu Val Phe Met Glu Thr Ser					
	200		205		210
Thr Met Ser Thr Glu Thr Glu Pro Phe Val Glu Asn Lys Ala Ala					
	215		220		225
Phe Lys Asn Glu Ala Ala Gly Phe Gly Gly Val Pro Thr Ala Leu					
	230		235		240
Leu Val Leu Ala Leu Leu Phe Phe Gly Ala Ala Ala Gly Leu Gly					
	245		250		255
Phe Cys Tyr Val Lys Arg Tyr Val Lys Ala Phe Pro Phe Thr Asn					
	260		265		270
Lys Asn Gln Gln Lys Glu Met Ile Glu Thr Lys Val Val Lys Glu					
	275		280		285
Glu Lys Ala Asn Asp Ser Asn Pro Asn Glu Glu Ser Lys Lys Thr					
	290		295		300
Asp Lys Asn Pro Glu Glu Ser Lys Ser Pro Ser Lys Thr Thr Val					
	305		310		315
Arg Cys Leu Glu Ala Glu Val					
	320				

<210> SEQ ID NO 7
<211> LENGTH: 2586
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 7

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ccgcagcgca actcgggtcca gtcggggcgg cggtcgggg cgagagcg	150
agatgcagcg gcttggggcc accctgctgt gcctgctgct ggcggcggcg	200
gtcccccacg ccccgcgcc cgctccgacg gcgacctcg ctccagtcaa	250
gcccggcccc gctctcagct acccgcagga ggaggccacc ctcaatgaga	300
tgttccgcga ggttgaggaa ctgatggagg acacgcagca caaattgcgc	350
agcgcggtgg aagagatgga ggcagaagaa gctgctgcta aagcatcatc	400
agaagtgaac ctggcaaact tacctcccag ctatcacaat gagaccaaca	450
cagacacgaa ggttggaat aataccatcc atgtgcaccg agaaattcac	500
aagataacca acaaccagac tggacaaatg gtcttttcag agacagttat	550
cacatctgtg ggagacgaag aaggcagaag gagccacgag tgcatcatcg	600
acgaggactg tgggccacgc atgtactgcc agtttgccag cttccagtac	650
acctgccagc catgccgggg ccagaggatg ctctgcaccc gggacagtga	700

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gtgctgtgga gaccagctgt gtgtctgggg tcaactgcacc aaaatggcca	750
ccaggggcag caatgggacc atctgtgaca accagaggga ctgccagccg	800
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cctgcccgtg gagggcgagc tttgccatga ccccgccagc cggcttcttg	900
acctcatcac ctgggagcta gagcctgatg gagccttgga ccgatgccct	950
tgtgccagtg gcctcctctg ccagccccac agccacagcc tgggtgatgt	1000
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tgcccagaga ggtccccgat gagtatgaag ttggcagctt catggaggag	1100
gtgccccagg agctggagga cctgggaggg agcctgactg aagagatggc	1150
gctgggggag cctgcggctg ccgccgctgc actgctggga ggggaagaga	1200
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tatttcccca ggtgtgtgct ttaggcgtgg gctgaccagg cttcttctca	1300
catcttcttc ccagtaagtt tcccctctgg cttgacagca tgagggtgtg	1350
tgcatattgt cagctcccc aggtgtttct ccaggcttca cagtctggtg	1400
cttgggagag tcaggcaggg ttaaactgca ggagcagttt gccaccctg	1450
tccagattat tggctgcttt gcctctacca gttggcagac agccgtttgt	1500
tctacatggc tttgataatt gtttgagggg aggagatgga aacaatgtgg	1550
agtctccctc tgatttggtt tggggaaatg tggagaagag tgccctgctt	1600
tgcaaacatc aacctggcaa aaatgcaaca aatgaatttt ccacgcagtt	1650
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gaaggagaaat gggatttttc ttgaggcctg cacatctgga attaaaggtca	2050
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gttctcacag tgtggggcag ccgtccttct aatgaagaca atgatattga	2150
cactgtccct ctttggcagt tgcattagta actttgaaag gtatatgact	2200
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caattatcaa ccacgtggag aaaatcaaac cgagcagggc tgtgtgaaac	2350
atggttgtaa tatgcgactg cgaacactga actctacgcc actccacaaa	2400
tgatgttttc aggtgtcatg gactgttgcc accatgtatt catccagagt	2450
tcttaaagtt taaagttgca catgattgta taagcatgct ttctttgagt	2500
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<210> SEQ ID NO 8

<211> LENGTH: 350

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 8

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

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	335	340	345
Gly Gly Glu Glu Ile			
350			
<210> SEQ ID NO 9			
<211> LENGTH: 1395			
<212> TYPE: DNA			
<213> ORGANISM: Homo Sapien			
<400> SEQUENCE: 9			
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tgtgcgtctt ccagggctac tcatccaaag gcctaatacca acgttctgtc			150
ttcaatctgc aaatctatgg ggtcctgggg ctcttctgga cccttaactg			200
ggtactggcc ctgggccaat gcgtcctcgc tggagccttt gcctccttct			250
actgggcctt ccacaagccc caggacatcc ctaccttccc cttaatctct			300
gccttcaccc gcacactccg ttaccacact gggtcattgg catttgagc			350
cctcatcctg acccttgtgc agatagcccg ggtcatcttg gatatattg			400
accacaagct cagaggagtg cagaaccctg tagcccgctg catcatgtgc			450
tgtttcaagt gctgcctctg gtgtctggaa aaatttatca agttcctaaa			500
ccgcaatgoa tacatcatga tcgccatcta cgggaagaat ttctgtgtct			550
cagccaaaaa tgcgttcatg ctactcatgc gaaacattgt cagggtggtc			600
gtcctggaca aagtcacaga cctgctgtg ttctttggga agctgctggt			650
ggtcggaggc gtgggggtcc tgtccttctt ttttttctcc ggtgcaccc			700
cggggctggg taaagacttt aagagccccc acctcaacta ttactggctg			750
cccacatgca cctccatcct gggggcctat gtcacgcga gcggcttctt			800
cagcgttttc ggcattgtgt tggacacgct ctctctctgc ttcttggaag			850
acctggagcg gaacaacggc tccctggacc ggccctacta catgtccaag			900
agccttctaa agattctggg caagaagaac gaggcgcccc cggacaacaa			950
gaagagggaag aagtgcacgc tccggccctg atccaggact gcacccacc			1000
cccaccgtcc agccatccaa cctcacttcg ccttacagggt ctccattttg			1050
tggtaaaaaa aggttttagg ccaggcgccg tggctcacgc ctgtaatcca			1100
acacttttag aggctgaggc gggcgatca cctgagtcag gatttcgaga			1150
ccagcctggc caacatggtg aaacctccgt ctctattaaa aatacaaaaa			1200
ttagccgaga gtggtggcat gcacctgtca tcccagctac tcgggaggct			1250
gaggcaggag aatcgcttga acccgggagg cagaggttgc agtgagccga			1300
gatcgcgcca ctgcactcca acctgggtga cagactctgt ctccaaaaca			1350
aaacaaacaa acaaaaagat ttattataag atattttggt aactc			1395
<210> SEQ ID NO 10			
<211> LENGTH: 321			
<212> TYPE: PRT			
<213> ORGANISM: Homo Sapien			
<400> SEQUENCE: 10			

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

<210> SEQ ID NO 11
<211> LENGTH: 1901
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 11

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ccgtgagccg cctcatcttc acgtttcttc tcttctctggg ggtgctggtg	200
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ggcctgtggc ctgatgttca tgtactacac tgagccagc ggctgccacg	750
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caggctcctg cagagcccca tcccccgcc acaccacac ggtggagctg	1650
cctcttctct cccctcctcc ctggtgcccc tactcagcat ctcggtgaa	1700
agggctccct tgtcctcagg ctccacggga gcggggctgc tggagagagc	1750
ggggaactcc caccacagtg gggcatccgg cactgaagcc ctggtgttcc	1800
tggtcacgtc cccagggga ccctgcccc ttctggact tcgtgcotta	1850
ctgagtctct aagacttttt ctaataaaca agccagtgcg tgtaaaaaaa	1900
a	1901

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<210> SEQ ID NO 12

<211> LENGTH: 457

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 12

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Met Gly Ala Cys Leu Gly Ala Cys Ser Leu Leu Ser Cys Ala Ser
 1           5           10          15

Cys Leu Cys Gly Ser Ala Pro Cys Ile Leu Cys Ser Cys Cys Pro
 20          25          30

Ala Ser Arg Asn Ser Thr Val Ser Arg Leu Ile Phe Thr Phe Phe
 35          40          45

Leu Phe Leu Gly Val Leu Val Ser Ile Ile Met Leu Ser Pro Gly
 50          55          60

Val Glu Ser Gln Leu Tyr Lys Leu Pro Trp Val Cys Glu Glu Gly
 65          70          75

Ala Gly Ile Pro Thr Val Leu Gln Gly His Ile Asp Cys Gly Ser
 80          85          90

Leu Leu Gly Tyr Arg Ala Val Tyr Arg Met Cys Phe Ala Thr Ala
 95          100         105

Ala Phe Phe Phe Phe Phe Phe Thr Leu Leu Met Leu Cys Val Ser
 110         115         120

Ser Ser Arg Asp Pro Arg Ala Ala Ile Gln Asn Gly Phe Trp Phe
 125         130         135

Phe Lys Phe Leu Ile Leu Val Gly Leu Thr Val Gly Ala Phe Tyr
 140         145         150

Ile Pro Asp Gly Ser Phe Thr Asn Ile Trp Phe Tyr Phe Gly Val
 155         160         165

Val Gly Ser Phe Leu Phe Ile Leu Ile Gln Leu Val Leu Leu Ile
 170         175         180

Asp Phe Ala His Ser Trp Asn Gln Arg Trp Leu Gly Lys Ala Glu
 185         190         195

Glu Cys Asp Ser Arg Ala Trp Tyr Ala Gly Leu Phe Phe Phe Thr
 200         205         210

Leu Leu Phe Tyr Leu Leu Ser Ile Ala Ala Val Ala Leu Met Phe
 215         220         225

Met Tyr Tyr Thr Glu Pro Ser Gly Cys His Glu Gly Lys Val Phe
 230         235         240

Ile Ser Leu Asn Leu Thr Phe Cys Val Cys Val Ser Ile Ala Ala
 245         250         255

Val Leu Pro Lys Val Gln Asp Ala Gln Pro Asn Ser Gly Leu Leu
 260         265         270

Gln Ala Ser Val Ile Thr Leu Tyr Thr Met Phe Val Thr Trp Ser
 275         280         285

Ala Leu Ser Ser Ile Pro Glu Gln Lys Cys Asn Pro His Leu Pro
 290         295         300

Thr Gln Leu Gly Asn Glu Thr Val Val Ala Gly Pro Glu Gly Tyr
 305         310         315

Glu Thr Gln Trp Trp Asp Ala Pro Ser Ile Val Gly Leu Ile Ile
 320         325         330

Phe Leu Leu Cys Thr Leu Phe Ile Ser Leu Arg Ser Ser Asp His

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335					340					345				
Arg	Gln	Val	Asn	Ser	Leu	Met	Gln	Thr	Glu	Glu	Cys	Pro	Pro	Met
350					355					360				
Leu	Asp	Ala	Thr	Gln	Gln	Gln	Gln	Gln	Gln	Val	Ala	Ala	Cys	Glu
365					370					375				
Gly	Arg	Ala	Phe	Asp	Asn	Glu	Gln	Asp	Gly	Val	Thr	Tyr	Ser	Tyr
380					385					390				
Ser	Phe	Phe	His	Phe	Cys	Leu	Val	Leu	Ala	Ser	Leu	His	Val	Met
395					400					405				
Met	Thr	Leu	Thr	Asn	Trp	Tyr	Lys	Pro	Gly	Glu	Thr	Arg	Lys	Met
410					415					420				
Ile	Ser	Thr	Trp	Thr	Ala	Val	Trp	Val	Lys	Ile	Cys	Ala	Ser	Trp
425					430					435				
Ala	Gly	Leu	Leu	Leu	Tyr	Leu	Trp	Thr	Leu	Val	Ala	Pro	Leu	Leu
440					445					450				
Leu	Arg	Asn	Arg	Asp	Phe	Ser								
455														

<210> SEQ ID NO 13
<211> LENGTH: 1572
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 13

cgggccagcc	tggggcggcc	ggccaggaac	caccggttaa	ggtgtcttct	50
ctttagggat	ggtgaggttg	gaaaagact	cctgtaacct	tcctccagga	100
tgaaccacct	gccagaagac	atggagaacg	ctctaccg	gagccagagc	150
tcccatgctt	ctctgcgcaa	tatccattcc	atcaaccca	cacaactcat	200
ggccaggatt	gagtcctatg	aaggaaggga	aaagaaggc	atatctgatg	250
tcaggaggac	tttctgtttg	ttgtcacct	ttgacctctt	attcgtaaca	300
ttactgtgga	taatagagtt	aaatgtgaat	ggaggcattg	agaacacatt	350
agagaaggag	gtgatgcagt	atgactacta	ttcttcatat	tttgatatat	400
ttcttctggc	agtttttcga	tttaaagtg	taatactg	atatgctgtg	450
tgcagactgc	gccattggtg	ggcaatagcg	ttgacaacgg	cagtgaccag	500
tgccctttta	ctagcaaaag	tgatccttcc	gaagcttttc	tctcaagggg	550
cttttggtta	tgtgctgccc	atcatttcat	tcacacctgc	ctggattgag	600
acgtggttcc	tggaattcaa	agtgttacct	caagaagcag	aagaagaaaa	650
cagactcctg	atagttcagg	atgcttcaga	gagggcagca	cttataacctg	700
gtggtctttc	tgatggtcag	ttttattccc	ctcctgaatc	cgaagcagga	750
tctgaagaag	ctgaagaaaa	acaggacagt	gagaaaccac	ttttagaact	800
atgagtacta	cttttggttaa	atgtgaaaaa	ccctcacaga	aagtcacgca	850
ggcaaaaaga	ggcaggcagt	ggagtctccc	tgtcgacagt	aaagttgaaa	900
tggtgacgtc	cactgctggc	tttattgaac	agctaataaa	gattttattta	950
ttgtaatacc	tcacaaacgt	tgtaccatat	ccatgcacat	ttagttgcct	1000
gcctgtggct	ggtaaggtaa	tgtcatgatt	catcctctct	tcagtgagac	1050

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tgagcctgat gtgtaacaa ataggatgaag aaagtcttgt gctgtattcc	1100
taatcaaaag acttaataata ttgaagtaac acttttttag taagcaagat	1150
acctttttat ttcaattcac agaattggaat ttttttggtt catgtctcag	1200
atattttttg tattttctttt ttaacactct acatttcctt tgttttttaa	1250
ctcatgcaca tgtgctcttt gtacagtttt aaaaagtga ataaaatctg	1300
acatgtcaat gtggctagtt ttatttttct tgttttgcatt tatgtgtatg	1350
gcctgaagtgt ttggacttgc aaaaggggaa gaaaggaatt gcgaatacat	1400
gtaaaatgtc accagacatt tgtattattt ttatcatgaa atcatgtttt	1450
tctctgattg ttctgaaatg ttctaaatac tcttattttg aatgcacaaa	1500
atgacttaaa ccattcatat catgtttcct ttgcgttcag ccaatttcaa	1550
ttaaaatgaa ctaaattaaa aa	1572

<210> SEQ ID NO 14

<211> LENGTH: 234

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 14

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

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230

<210> SEQ ID NO 15

<211> LENGTH: 2768

<212> TYPE: DNA

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 15

actcgaacgc agttgcttcg ggacccagga cccctcggg cccgacccgc	50
caggaaagac tgagggccgc gcctgccccg cccggctccc tgcgcgcgcg	100
ccgcctcccg ggacagaaga tgtgtccag ggtccctctg ctgctgccgc	150
tgctcctgct actggccctg gggcctggg tgagggctg cccatccggc	200
tgccagtgcg gccagccaca gacagtcttc tgcactgcc gccaggggac	250
cacggtgccc cgagacgtgc caccgcacac ggtggggctg tacgtctttg	300
agaacggcat caccatgctc gacgcaggca gctttgccg cctgccgggc	350
ctgcagctcc tggacctgtc acagaaccag atcgccagcc tgcccagcgg	400
ggtcttccag ccactcgcca acctcagcaa cctggacctg acggccaaca	450
ggctgcata aatcaccaat gagaccttcc gtggcctgcg gcgcctcgag	500
cgccctctacc tgggcaagaa ccgcctccgc cacatccagc ctggtgcctt	550
cgacacgctc gaccgcctcc tggagctcaa gctgcaggac aacgagctgc	600
gggcactgcc cccgctgcgc ctgccccgcc tgcgtctgct ggacctcagc	650
cacaacagcc tcctggccct ggagcccgcc atcctggaca ctgccaacgt	700
ggaggcgctg cggctggctg gtctggggct gcagcagctg gacgaggggc	750
tcttcagccg cttgcgcaac ctccacgacc tggatgtgtc cgacaaccag	800
ctggagcgag tgccacctgt gatccgaggc ctccggggcc tgacgcgcct	850
gcggctggcc ggcaacaccc gcattgcccc gctgcggccc gaggacctgg	900
ccggcctggc tgccctgcag gagctggatg tgagcaacct aagcctgcag	950
gccctgcctg gcgacctctc gggctctctc ccccgctgc ggctgctggc	1000
agctgccccg aacccttca actgcgtgtg cccctgagc tggtttgccc	1050
cctgggtgcg cgagagccac gtcacactgg ccagccctga ggagacgcgc	1100
tgccacttcc cgccaagaa cgttgcccg gcgtccttg agcttgacta	1150
cgccgacttt ggctgcccag ccaccaccac cacagccaca gtgcccacca	1200
cgaggcccggt ggtgcgggag cccacagcct tgtcttctag cttggtcct	1250
acctggctta gcccacagc gccggccact gagggcccca gccgcacctc	1300
cactgccccg ccgactgtag ggcctgtccc ccagccccag gactgcccac	1350
cgctccacctg cctcaatggg ggcacatgcc acctggggac acggcaccac	1400
ctggcgctgt tgtgcccga aggttccag ggcctgtact gtgagagcca	1450
gatggggcag gggacacggc ccagccctac accagtcacg ccgaggccac	1500
cacggtccct gaccctgggc atcgagccgg tgagccccac ctccctgcgc	1550
gtggggctgc agcgctacct ccaggggagc tccgtgcagc tcaggagcct	1600
ccgtctcacc tatcgcaacc tatcggggcc tgataagcgg ctggtgacgc	1650

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tgcgactgcc tgcctcgctc gctgagtaca cggtcaccca gctgcggccc	1700
aacgccactt actccgtctg tgtcatgcct ttggggcccg ggcgggtgcc	1750
ggagggcgag gaggcctgcg gggaggccca tacaccccca gccgtccact	1800
ccaaccacgc ccagtcacc caggcccgcg agggcaacct gccgtcctc	1850
attgcgcccg ccctggccgc ggtgtcctg gccgcgtgg ctgcggtggg	1900
ggcagcctac tgtgtgcggc gggggcgggc catggcagca ggcgctcagg	1950
acaaagggca ggtggggcca ggggctgggc ccctggaact ggagggagtg	2000
aaggtcccct tggagccagg cccgaaggca acagaggcg gtggagaggc	2050
cctgcccagc gggctctagt gtgaggtgcc actcatgggc ttcccagggc	2100
ctggcctcca gtcacccctc cagcacaagc cctacatcta agccagagag	2150
agacagggca gctggggccg ggctctcagc cagtgagatg gccagccccc	2200
tctctgtgcc acaccacgta agttctcagt cccaacctcg gggatgtgtg	2250
cagacagggc tgtgtgacca cagctggggc ctgttccctc tggacctcg	2300
ttctctcatc tgtgagatgc tgtggccag ctgacgagcc ctaacgtccc	2350
cagaaccgag tgcctatgag gacagtgtcc gccctgccct ccgcaacctg	2400
cagtccttgg gcacggcggg ccctgccatg tgtgtgtaac gcatgcctgg	2450
gtcctgtctg gctctccac tccaggcggc ccctgggggc cagtgaagga	2500
agctcccgga aagagcagag ggagagcggg taggcggctg tgtgactcta	2550
gtcttggccc caggaagcga aggaacaaaa gaaactggaa aggaagatgc	2600
tttaggaaca tgttttgctt ttttaaaata tatatattta taagagatcc	2650
tttcccattt attctgggaa gatgtttttc aaactcagag acaaggactt	2700
tggtttttgt aagacaaacg atgatatgaa ggccttttgt aagaaaaaat	2750
aaaagatgaa gtgtgaaa	2768

<210> SEQ ID NO 16

<211> LENGTH: 673

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 16

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

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110					115					120				
Arg	Gly	Leu	Arg	Arg	Leu	Glu	Arg	Leu	Tyr	Leu	Gly	Lys	Asn	Arg
				125					130					135
Ile	Arg	His	Ile	Gln	Pro	Gly	Ala	Phe	Asp	Thr	Leu	Asp	Arg	Leu
				140					145					150
Leu	Glu	Leu	Lys	Leu	Gln	Asp	Asn	Glu	Leu	Arg	Ala	Leu	Pro	Pro
				155					160					165
Leu	Arg	Leu	Pro	Arg	Leu	Leu	Leu	Leu	Asp	Leu	Ser	His	Asn	Ser
				170					175					180
Leu	Leu	Ala	Leu	Glu	Pro	Gly	Ile	Leu	Asp	Thr	Ala	Asn	Val	Glu
				185					190					195
Ala	Leu	Arg	Leu	Ala	Gly	Leu	Gly	Leu	Gln	Gln	Leu	Asp	Glu	Gly
				200					205					210
Leu	Phe	Ser	Arg	Leu	Arg	Asn	Leu	His	Asp	Leu	Asp	Val	Ser	Asp
				215					220					225
Asn	Gln	Leu	Glu	Arg	Val	Pro	Pro	Val	Ile	Arg	Gly	Leu	Arg	Gly
				230					235					240
Leu	Thr	Arg	Leu	Arg	Leu	Ala	Gly	Asn	Thr	Arg	Ile	Ala	Gln	Leu
				245					250					255
Arg	Pro	Glu	Asp	Leu	Ala	Gly	Leu	Ala	Ala	Leu	Gln	Glu	Leu	Asp
				260					265					270
Val	Ser	Asn	Leu	Ser	Leu	Gln	Ala	Leu	Pro	Gly	Asp	Leu	Ser	Gly
				275					280					285
Leu	Phe	Pro	Arg	Leu	Arg	Leu	Leu	Ala	Ala	Ala	Arg	Asn	Pro	Phe
				290					295					300
Asn	Cys	Val	Cys	Pro	Leu	Ser	Trp	Phe	Gly	Pro	Trp	Val	Arg	Glu
				305					310					315
Ser	His	Val	Thr	Leu	Ala	Ser	Pro	Glu	Glu	Thr	Arg	Cys	His	Phe
				320					325					330
Pro	Pro	Lys	Asn	Ala	Gly	Arg	Leu	Leu	Leu	Glu	Leu	Asp	Tyr	Ala
				335					340					345
Asp	Phe	Gly	Cys	Pro	Ala	Thr	Thr	Thr	Thr	Ala	Thr	Val	Pro	Thr
				350					355					360
Thr	Arg	Pro	Val	Val	Arg	Glu	Pro	Thr	Ala	Leu	Ser	Ser	Ser	Leu
				365					370					375
Ala	Pro	Thr	Trp	Leu	Ser	Pro	Thr	Ala	Pro	Ala	Thr	Glu	Ala	Pro
				380					385					390
Ser	Pro	Pro	Ser	Thr	Ala	Pro	Pro	Thr	Val	Gly	Pro	Val	Pro	Gln
				395					400					405
Pro	Gln	Asp	Cys	Pro	Pro	Ser	Thr	Cys	Leu	Asn	Gly	Gly	Thr	Cys
				410					415					420
His	Leu	Gly	Thr	Arg	His	His	Leu	Ala	Cys	Leu	Cys	Pro	Glu	Gly
				425					430					435
Phe	Thr	Gly	Leu	Tyr	Cys	Glu	Ser	Gln	Met	Gly	Gln	Gly	Thr	Arg
				440					445					450
Pro	Ser	Pro	Thr	Pro	Val	Thr	Pro	Arg	Pro	Pro	Arg	Ser	Leu	Thr
				455					460					465
Leu	Gly	Ile	Glu	Pro	Val	Ser	Pro	Thr	Ser	Leu	Arg	Val	Gly	Leu
				470					475					480
Gln	Arg	Tyr	Leu	Gln	Gly	Ser	Ser	Val	Gln	Leu	Arg	Ser	Leu	Arg
				485					490					495

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Leu	Thr	Tyr	Arg	Asn	Leu	Ser	Gly	Pro	Asp	Lys	Arg	Leu	Val	Thr
				500					505					510
Leu	Arg	Leu	Pro	Ala	Ser	Leu	Ala	Glu	Tyr	Thr	Val	Thr	Gln	Leu
				515					520					525
Arg	Pro	Asn	Ala	Thr	Tyr	Ser	Val	Cys	Val	Met	Pro	Leu	Gly	Pro
				530					535					540
Gly	Arg	Val	Pro	Glu	Gly	Glu	Glu	Ala	Cys	Gly	Glu	Ala	His	Thr
				545					550					555
Pro	Pro	Ala	Val	His	Ser	Asn	His	Ala	Pro	Val	Thr	Gln	Ala	Arg
				560					565					570
Glu	Gly	Asn	Leu	Pro	Leu	Leu	Ile	Ala	Pro	Ala	Leu	Ala	Ala	Val
				575					580					585
Leu	Leu	Ala	Ala	Leu	Ala	Ala	Val	Gly	Ala	Ala	Tyr	Cys	Val	Arg
				590					595					600
Arg	Gly	Arg	Ala	Met	Ala	Ala	Ala	Ala	Gln	Asp	Lys	Gly	Gln	Val
				605					610					615
Gly	Pro	Gly	Ala	Gly	Pro	Leu	Glu	Leu	Glu	Gly	Val	Lys	Val	Pro
				620					625					630
Leu	Glu	Pro	Gly	Pro	Lys	Ala	Thr	Glu	Gly	Gly	Gly	Glu	Ala	Leu
				635					640					645
Pro	Ser	Gly	Ser	Glu	Cys	Glu	Val	Pro	Leu	Met	Gly	Phe	Pro	Gly
				650					655					660
Pro	Gly	Leu	Gln	Ser	Pro	Leu	His	Ala	Lys	Pro	Tyr	Ile		
				665					670					

<210> SEQ ID NO 17
<211> LENGTH: 1672
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 17

gcagcggcga ggcggcggtg gtggctgagt ccgtggtggc agaggcgaag	50
gcgacagctc atgcgggtcc ggatagggct gacgctgctg ctgtgtgcgg	100
tgctgctgag cttggcctcg gcgtcctcgg atgaagaagg cagccaggat	150
gaatccttag attccaagac tactttgaca tcagatgagt cagtaaagga	200
ccatactact gcaggcagag tagttgctgg tcaaatatctt cttgattcag	250
aagaatctga attagaatcc tctattcaag aagaggaaga cagcctcaag	300
agccaagagg gggaaagtgt cacagaagat atcagctttc tagagtctcc	350
aatccagaa aacaaggact atgaagagcc aaagaaagta cggaaaccag	400
ctttgaccgc cattgaaggc acagcacatg gggagccctg ccacttcct	450
tttcttttcc tagataagga gtatgatgaa tgtacatcag atgggagggg	500
agatggcaga ctgtggtgtg ctacaacctg tgactacaaa gcagatgaaa	550
agtggggctt ttgtgaaact gaagaagagg ctgctaagag acggcagatg	600
caggaagcag aaatgatgta tcaaactgga atgaaatcc ttaatggaag	650
caataagaaa agccaaaaaa gagaagcata tcggtatctc caaaaggcag	700
caagcatgaa ccataccaaa gccctggaga gagtgtcata tgctctttta	750
tttggtgatt acttgccaca gaatatccag gcagcgagag agatgtttga	800

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gaagctgact gaggaaggct ctccaaggg acagactgct cttggctttc      850
tgtatgcctc tggacttggt gttaattcaa gtcaggcaaa ggctcttgta      900
tattatacat ttggagctct tgggggcaat ctaatagccc acatggtttt      950
ggtaagtaga ctttagtgga aggctaataa tattaacatc agaagaattt     1000
gtggtttata gcggccacaa ctttttcagc tttcatgac cagatttgct     1050
tgtattaaga ccaaataatt agttgaactt ccttcaaatt cttgttaatg     1100
gatataacac atggaatcta catgtaaatg aaagttggtg gagtccacaa     1150
tttttcttta aaatgattag ttggctgat tgcccctaaa aagagagatc     1200
tgataaatgg ctctttttaa attttctctg agttggaatt gtcagaatca     1250
ttttttacat tagattatca taattttaaa aatttttctt tagtttttca     1300
aaattttgta aatggtggct atagaaaaac aacatgaaat attatacaat     1350
attttgcaac aatgccctaa gaattgttaa aattcatgga gttatttggt     1400
cagaatgact ccagagagct ctactttctg ttttttactt ttcatgattg     1450
gctgtcttcc catttattct ggtcatttat tgctagtac actgtgcttg     1500
cttccagtag tctcattttc cctattttgc taatttgta ctttttcttt     1550
gctaatttgg aagattaact catttttaat aaaattatgt ctaagattaa     1600
aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa     1650
aaaaaaaaaa aaaaaaaaaa aa                                           1672

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<210> SEQ ID NO 18

<211> LENGTH: 301

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 18

```

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

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	155		160		165
Gly Phe Cys Glu Thr Glu Glu Glu Ala	Ala Lys Arg Arg Gln Met				
170	175				180
Gln Glu Ala Glu Met Met Tyr Gln Thr	Gly Met Lys Ile Leu Asn				
185	190				195
Gly Ser Asn Lys Lys Ser Gln Lys Arg	Glu Ala Tyr Arg Tyr Leu				
200	205				210
Gln Lys Ala Ala Ser Met Asn His Thr	Lys Ala Leu Glu Arg Val				
215	220				225
Ser Tyr Ala Leu Leu Phe Gly Asp Tyr	Leu Pro Gln Asn Ile Gln				
230	235				240
Ala Ala Arg Glu Met Phe Glu Lys Leu	Thr Glu Glu Gly Ser Pro				
245	250				255
Lys Gly Gln Thr Ala Leu Gly Phe Leu	Tyr Ala Ser Gly Leu Gly				
260	265				270
Val Asn Ser Ser Gln Ala Lys Ala Leu	Val Tyr Tyr Thr Phe Gly				
275	280				285
Ala Leu Gly Gly Asn Leu Ile Ala His	Met Val Leu Val Ser Arg				
290	295				300

Leu

<210> SEQ ID NO 19
<211> LENGTH: 1508
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 19

aattcagatt ttaagcccat tctgcagtgg aatttcatga actagcaaga	50
ggacaccatc ttcttgtatt atacaagaaa ggagtgtacc tatcacacac	100
agggggaaaa atgctctttt ggggtgctagg cctcctaadc ctctgtggtt	150
ttctgtggac tcgtaaagga aaactaaaga ttgaagacat cactgataag	200
tacattttta tcaactggatg tgactcgggc tttggaaact tggcagccag	250
aacttttgat aaaaagggat ttcatgtaat cgctgcctgt ctgactgaat	300
caggatcaac agctttaaag gcagaaacct cagagagact tcgtactgtg	350
cttctggatg tgaccgaccc agagaatgac aagaggactg cccagtgggt	400
gaagaaccaa gttggggaga aaggtctctg gggctctgac aataatgctg	450
gtgttcccgg cgtgctggct cccactgact ggctgacact agaggactac	500
agagaacctt ttgaagtga cctgtttgga ctcatcagtg tgacactaaa	550
tatgcttcct ttggtcaaga aagctcaagg gagagttatt aatgtctcca	600
gtgttgagg tcgccttgca atcggtggag ggggctatac tccatccaaa	650
tatgcagtgg aaggtttcaa tgacagctta agacgggaca tgaagcttt	700
tggtgtgcac gtctcatgca ttgaaccagg attgttcaaa aaaaacttgg	750
cagatccagt aaaggttaatt gaaaaaaac tcgccatttg ggagcagctg	800
tctccagaca tcaaacaaca atatggagaa gggttacattg aaaaaagtct	850
agacaaactg aaaggcaata aatcctatgt gaacatggac ctctctccgg	900
tggtagagtg catggaccac gctctaacaa gtctcttccc taagactcat	950

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tatgccgctg gaaaagatgc caaaattttc tggatacctc tgtctcacat	1000
gccagcagct ttgcaagact ttttattgtt gaaacagaaa gcagagctgg	1050
ctaattccaa ggcagtgtga ctacagtaac cacaatgtc tcctccaggc	1100
tatgaaattg gccgatttca agaacacatc tccttttcaa cccatttcct	1150
tatctgctcc aacctggact catttagatc gtgcttattt ggattgcaaa	1200
agggagtccc accatcgctg gtggtatccc agggtcctg ctcaagtttt	1250
ctttgaaaaa gagggctgga atggtacatc acataggcaa gtccctgccct	1300
gtatttaggc tttgcctgct tgggtgtgatg taagggaat tgaagactt	1350
gccattcaa aatgatcttt accgtggcct gcccattgct tatggtcccc	1400
agcatttaca gtaacttgct aatgtaagt atcatctctt atctaaatat	1450
taaaagataa gtcaacccaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa	1500
aaaaaaaa	1508

<210> SEQ ID NO 20

<211> LENGTH: 319

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 20

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

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215										220										225									
Ile	Trp	Glu	Gln	Leu	Ser	Pro	Asp	Ile	Lys	Gln	Gln	Tyr	Gly	Glu															
230										235										240									
Gly	Tyr	Ile	Glu	Lys	Ser	Leu	Asp	Lys	Leu	Lys	Gly	Asn	Lys	Ser															
245										250										255									
Tyr	Val	Asn	Met	Asp	Leu	Ser	Pro	Val	Val	Glu	Cys	Met	Asp	His															
260										265										270									
Ala	Leu	Thr	Ser	Leu	Phe	Pro	Lys	Thr	His	Tyr	Ala	Ala	Gly	Lys															
275										280										285									
Asp	Ala	Lys	Ile	Phe	Trp	Ile	Pro	Leu	Ser	His	Met	Pro	Ala	Ala															
290										295										300									
Leu	Gln	Asp	Phe	Leu	Leu	Leu	Lys	Gln	Lys	Ala	Glu	Leu	Ala	Asn															
305										310										315									

Pro Lys Ala Val

<210> SEQ ID NO 21
<211> LENGTH: 1849
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 21

ctgagggcggc ggtagcatgg agggggagag tacgtcggcg gtgctctcgg	50
gctttgtgct cggcgcactc gctttccagc acctcaacac ggactcggac	100
acggaaggtt ttcttcttgg ggaagtaaaa ggtgaagcca agaacagcat	150
tactgattcc caaatggatg atgttgaagt tgtttataca attgacattc	200
agaaatatat tccatgctat cagcttttta gcttttataa ttcttcaggc	250
gaagtaaatg agcaagcact gaagaaaata ttatcaaatg tcaaaaagaa	300
tggtgtaggt tggtacaaat tccgtcgtca ttcagatcag atcatgacgt	350
ttagagagag gctgcttcac aaaaacttgc aggagcattt ttcaaaccaa	400
gacctgtttt ttctgctatt aacaccaagt ataataacag aaagctgctc	450
tactcatcga ctggaacatt ccttatataa acctcaaaaa ggactttttc	500
acaggggtacc tttagtgggt gccaatctgg gcatgtctga acaactgggt	550
tataaaactg tatcaggttc ctgtatgtcc actggtttta gccgagcagt	600
acaaacacac agctctaaat tttttgaaga agatggatcc ttaaaggagg	650
tacataagat aaatgaaatg tatgcttcac tacaagagga attaaagagt	700
atatgcaaaa aagtgaaga cagtgaacaa gcagtagata aactagtaaa	750
ggatgtaaac agattaaaac gagaaattga gaaaaggaga ggagcacaga	800
ttcaggcagc aagagagaag aacatccaaa aagaccctca ggagaacatt	850
tttctttgtc aggcattacg gacctttttt ccaaattctg aattttctca	900
ttcatgtggt atgtctttaa aaaatagaca tgtttctaaa agtagctgta	950
actacaacca ccatctcgat gtagtagaca atctgacctt aatggtagaa	1000
cacactgaca ttcttgaagc tagtccagct agtacaccac aaatcattaa	1050
gcataaagcc ttagacttag atgacagatg gcaattcaag agatctcggg	1100
tggttagatac acaagacaaa cgatctaaag caaatactgg tagtagtaac	1150

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caagataaag catccaaaat gagcagccca gaaacagatg aagaaattga      1200
aaagatgaag ggttttggtg aatattcacg gtctcctaca ttttgatcct      1250
tttaacctta caaggagatt tttttatttg gctgatgggt aaagccaaac      1300
atttctattg tttttactat gttgagctac ttgcagtaag ttcatttggt      1350
tttactatgt tcacctgttt gcagtaatac acagataact cttagtgcac      1400
ttacttcaca aagtactttt tcaaacatca gatgctttta tttccaaacc      1450
tttttttcac ctttcactaa gttgttgagg ggaaggctta cacagacaca      1500
ttctttagaa ttggaaaagt gagaccaggc acagtggctc acacctgtaa      1550
tcccagcact tagggaagac aagtcaggag gattgattga agctaggagt      1600
tagagaccag cctgggcaac gtattgagac catgtctatt aaaaaataaa      1650
atggaaaagc aagaatagcc ttattttcaa aatatggaaa gaaatttata      1700
tgaaaaatta tctgagtcac taaaattctc cttaagtgat acttttttag      1750
aagtacatta tggctagagt tgccagataa aatgctggat atcatgcaat      1800
aaatttgcaa aacatcatct aaaatttaaa aaaaaaaaaa aaaaaaaaaa      1849

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<210> SEQ ID NO 22

<211> LENGTH: 409

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 22

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

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200										205										210									
Glu	Val	His	Lys	Ile	Asn	Glu	Met	Tyr	Ala	Ser	Leu	Gln	Glu	Glu															
				215					220					225															
Leu	Lys	Ser	Ile	Cys	Lys	Lys	Val	Glu	Asp	Ser	Glu	Gln	Ala	Val															
				230					235					240															
Asp	Lys	Leu	Val	Lys	Asp	Val	Asn	Arg	Leu	Lys	Arg	Glu	Ile	Glu															
				245					250					255															
Lys	Arg	Arg	Gly	Ala	Gln	Ile	Gln	Ala	Ala	Arg	Glu	Lys	Asn	Ile															
				260					265					270															
Gln	Lys	Asp	Pro	Gln	Glu	Asn	Ile	Phe	Leu	Cys	Gln	Ala	Leu	Arg															
				275					280					285															
Thr	Phe	Phe	Pro	Asn	Ser	Glu	Phe	Leu	His	Ser	Cys	Val	Met	Ser															
				290					295					300															
Leu	Lys	Asn	Arg	His	Val	Ser	Lys	Ser	Ser	Cys	Asn	Tyr	Asn	His															
				305					310					315															
His	Leu	Asp	Val	Val	Asp	Asn	Leu	Thr	Leu	Met	Val	Glu	His	Thr															
				320					325					330															
Asp	Ile	Pro	Glu	Ala	Ser	Pro	Ala	Ser	Thr	Pro	Gln	Ile	Ile	Lys															
				335					340					345															
His	Lys	Ala	Leu	Asp	Leu	Asp	Asp	Arg	Trp	Gln	Phe	Lys	Arg	Ser															
				350					355					360															
Arg	Leu	Leu	Asp	Thr	Gln	Asp	Lys	Arg	Ser	Lys	Ala	Asn	Thr	Gly															
				365					370					375															
Ser	Ser	Asn	Gln	Asp	Lys	Ala	Ser	Lys	Met	Ser	Ser	Pro	Glu	Thr															
				380					385					390															
Asp	Glu	Glu	Ile	Glu	Lys	Met	Lys	Gly	Phe	Gly	Glu	Tyr	Ser	Arg															
				395					400					405															

Ser Pro Thr Phe

<210> SEQ ID NO 23
<211> LENGTH: 2651
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 23

ggcacagccg cgcgggcgag ggcagagtca gccgagccga gtccagccgg	50
acgagcggac cagcgcgagg cagcccaagc agcgcgagc gaacgcccgc	100
cgccgcccac accctctgcg gtccccgcgg cgcttgccac cttccctcc	150
ttccccgcgt ccccgctcg cgggccagtc agcttgccgg gtctgctgcc	200
ccgcgaaacc ccgaggtcac cagccgcgc ctctgcttcc ctgggcccgc	250
cgccgcctcc acgcccctct tctcccctgg cccggcgctt ggcaccgggg	300
accgttgctt gacgcgaggc ccagctctac ttttcgcccc gcgtctctc	350
cgctgctcg cctcttccac caactccaac tccttctccc tccagctcca	400
ctcgctagtc cccgactccg ccagccctcg gcccgctgcc gtagcgccgc	450
ttcccgctcg gtcccaaagg tgggaacgcg tccgccccgg cccgcaccat	500
ggcacgggtt ggcttgcccg cgcttctctg caccctggca gtgctcagcg	550
ccgcgctgct ggctgccgag ctcaagtoga aaagttgctc ggaagtgcga	600
cgtcttttac tgtccaaagg cttcaacaag aacgatgccc ccctccacga	650

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gatcaacggt gatcatttga agatctgtcc ccagggttct acctgctgct	700
ctcaagagat ggaggagaag tacagcctgc aaagtaaaga tgatttcaaa	750
agtgtggcca gcgaacagtg caatcatttg caagctgtct ttgcttcacg	800
ttacaagaag tttgatgaat tcttcaaaga actacttgaa aatgcagaga	850
aatccctgaa tgatatgttt gtgaagacat atggccatth atacatgcaa	900
aattctgagc tatttaaaga tctcttcgta gagttgaaac gttactacgt	950
ggtgggaaat gtgaacctgg aagaaatgct aaatgacttc tgggctcgcc	1000
tcctggagcg gatgttccgc ctggtgaact ccagtagcca ctttacagat	1050
gagtatctgg aatgtgtgag caagtatacg gagcagctga agcccttcgg	1100
agatgtccct cgcaaattga agctccaggt tactcgtgct tttgtagcag	1150
cccgtacttt cgctcaaggc ttagcggttg cgggagatgt cgtgagcaag	1200
gtctccgtgg taaacccac agccagtggt acccatgccc tgttgaagat	1250
gatctactgc tcccactgcc ggggtctcgt gactgtgaag ccatgttaca	1300
actactgctc aaacatcatg agaggctgtt tggccaacca aggggatctc	1350
gattttgaat ggaacaattt catagatgct atgctgatgg tggcagagag	1400
gctagagggg cctttcaaca ttgaatcggg catggatccc atcgatgtga	1450
agattttcga tgctattatg aacatgcagg ataatagtgt tcaagtgtct	1500
cagaagggtt tccagggatg tggaccccc aagccctcc cagctggacg	1550
aatttctcgt tccatctctg aaagtgcctt cagtgtctgc ttcagaccac	1600
atcacccga ggaacgcca accacagcag ctggcactag tttggaccga	1650
ctggttactg atgtcaagga gaaactgaaa caggccaaga aattctggtc	1700
ctcccttcgg agcaacgttt gcaacgatga gaggatggct gcaggaaacg	1750
gcaatgagga tgactgttgg aatgggaaa gcaaaagcag gtacctgttt	1800
gcagtgcagc gaaatggatt agccaaccag ggcaacaacc cagaggtcca	1850
ggttgacacc agcaaaccag acatactgat ccttcgtcaa atcatggctc	1900
ttcagtgatg gaccagcaag atgaagaatg catacaatgg gaacgacgtg	1950
gacttctttg atatcagtga tgaaagtagt ggagaaggaa gtggaagtgg	2000
ctgtgagtat cagcagtgcc cttcagagtt tgactacaat gccactgacc	2050
atgtcgggaa gagtgcgaat gagaaagcgg acagtgtgg tgtccgtcct	2100
ggggcacagg cctacctcct cactgtcttc tgcacttgt tcctggttat	2150
gcagagagag tggagataat tctcaaactc tgagaaaaag tgttcatcaa	2200
aaagttaaaa ggcaccagtt atcacttttc taccatccta gtgactttgc	2250
tttttaaatg aatggacaac aatgtacagt ttttactatg tggccactgg	2300
tttaagaagt gctgactttg ttttctcatt cagttttggg aggaaaagg	2350
actgtgcatt gagtgggtc ctgctcccc aaaccatgtt aaacgtggct	2400
aacagtgtag gtacagaact atagttagtt gtgcatttgt gattttatca	2450
ctctattatt tgtttgtatg ttttttctc atttcgtttg tgggttttt	2500
tttccaactg tgatctcgcc ttgtttctta caagcaaac agggccctt	2550

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cttggcacgt aacatgtacg tatttctgaa atattaaata gctgtacaga      2600
agcagggtttt atttatcatg ttatcttatt aaaagaaaaa gcccaaaaag      2650
c                                                                2651

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<210> SEQ ID NO 24
<211> LENGTH: 556
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

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<400> SEQUENCE: 24

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

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305					310					315				
Asp	Val	Lys	Ile	Ser	Asp	Ala	Ile	Met	Asn	Met	Gln	Asp	Asn	Ser
320					325					330				
Val	Gln	Val	Ser	Gln	Lys	Val	Phe	Gln	Gly	Cys	Gly	Pro	Pro	Lys
335					340					345				
Pro	Leu	Pro	Ala	Gly	Arg	Ile	Ser	Arg	Ser	Ile	Ser	Glu	Ser	Ala
350					355					360				
Phe	Ser	Ala	Arg	Phe	Arg	Pro	His	His	Pro	Glu	Glu	Arg	Pro	Thr
365					370					375				
Thr	Ala	Ala	Gly	Thr	Ser	Leu	Asp	Arg	Leu	Val	Thr	Asp	Val	Lys
380					385					390				
Glu	Lys	Leu	Lys	Gln	Ala	Lys	Lys	Phe	Trp	Ser	Ser	Leu	Pro	Ser
395					400					405				
Asn	Val	Cys	Asn	Asp	Glu	Arg	Met	Ala	Ala	Gly	Asn	Gly	Asn	Glu
410					415					420				
Asp	Asp	Cys	Trp	Asn	Gly	Lys	Gly	Lys	Ser	Arg	Tyr	Leu	Phe	Ala
425					430					435				
Val	Thr	Gly	Asn	Gly	Leu	Ala	Asn	Gln	Gly	Asn	Asn	Pro	Glu	Val
440					445					450				
Gln	Val	Asp	Thr	Ser	Lys	Pro	Asp	Ile	Leu	Ile	Leu	Arg	Gln	Ile
455					460					465				
Met	Ala	Leu	Arg	Val	Met	Thr	Ser	Lys	Met	Lys	Asn	Ala	Tyr	Asn
470					475					480				
Gly	Asn	Asp	Val	Asp	Phe	Phe	Asp	Ile	Ser	Asp	Glu	Ser	Ser	Gly
485					490					495				
Glu	Gly	Ser	Gly	Ser	Gly	Cys	Glu	Tyr	Gln	Gln	Cys	Pro	Ser	Glu
500					505					510				
Phe	Asp	Tyr	Asn	Ala	Thr	Asp	His	Ala	Gly	Lys	Ser	Ala	Asn	Glu
515					520					525				
Lys	Ala	Asp	Ser	Ala	Gly	Val	Arg	Pro	Gly	Ala	Gln	Ala	Tyr	Leu
530					535					540				
Leu	Thr	Val	Phe	Cys	Ile	Leu	Phe	Leu	Val	Met	Gln	Arg	Glu	Trp
545					550					555				

Arg

<210> SEQ ID NO 25
<211> LENGTH: 870
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 25

ctcgccctca aatgggaacg ctggcctggg actaaagcat agaccaccag	50
gctgagtatc ctgacctgag tcatccccag ggatcaggag cctccagcag	100
ggaaccttcc attatattct tcaagcaact tacagctgca ccgacagtgtg	150
cgatgaaagt tctaattctct tccctcctcc tgttgctgcc actaatgctg	200
atgtccatgg tctctagcag cctgaatcca ggggtcgcca gaggccacag	250
ggaccgagggc caggcttcta ggagatggct ccaggaaggc ggccaagaat	300
gtgagtgcaa agattgggtc ctgagagccc cgagaagaaa attcatgaca	350
tgtgtctgggc tgccaaagaa gcagtgcccc tgtgatcatt tcaagggcaa	400

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tgtgaagaaa acaagacacc aaaggcacca cagaaagcca aacaagcatt	450
ccagagcctg ccagcaattt ctcaacaat gtcagctaag aagctttgct	500
ctgcctttgt aggagctctg agcgccact cttccaatta aacattctca	550
gccaaagaaga cagttagcac acctaccaga cactcttctt ctcccacctc	600
actctccac tgtaccaccc cctaaatcat tccagtgtc tcaaaaagca	650
tgtttttcaa gatcattttg ttgtgtgtc tctctagtgt cttcttctct	700
cgtcagtctt agcctgtgcc ctccccttac ccaggcttag gcttaattac	750
ctgaaagatt ccaggaaact gtagcttcct agctagtgtc atttaacctt	800
aatgcaatc aggaaagtag caaacagaag tcaataaata tttttaaatg	850
tcaaaaaaaaa aaaaaaaaaa	870

<210> SEQ ID NO 26
<211> LENGTH: 119
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 26

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

<210> SEQ ID NO 27
<211> LENGTH: 1371
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 27

ggacgccagc gcctgcagag gctgagcagg gaaaaagcca gtgccccagc	50
ggaagcacag ctgagagctg gtctgccatg gacatcctgg tcccactcct	100
gcagctgctg gtgctgcttc ttaccctgcc cctgcacctc atggctctgc	150
tggtgctgctg gcagcccctg tgcaaaagct acttccccta cctgatggcc	200
gtgctgactc ccaagagcaa ccgcaagatg gagagcaaga aacgggagct	250
cttcagccag ataaaggggc ttacaggagc ctccgggaaa gtggccctac	300
tggaagctggg ctgcggaacc ggagccaact ttcagttcta cccaccgggc	350
tgcagggtca cctgcctaga cccaaatccc cactttgaga agttcctgac	400

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aaagagcatg gctgagaaca ggcacctcca atatgagcgg tttgtggtgg	450
ctcctggaga ggacatgaga cagctggctg atggctccat ggatgtggtg	500
gtctgcactc tgggtgctgtg ctctgtgcag agcccaagga aggtcctgca	550
ggaggtccgg agagtactga gaccgggagg tgtgctcttt ttctgggagc	600
atgtggcaga accatatgga agctgggcct tcatgtggca gcaagttttc	650
gagcccacct ggaaacacat tggggatggc tgctgcctca ccagagagac	700
ctggaaggat cttgagaacg cccagtcttc cgaaatccaa atggaacgac	750
agccccctcc cttgaagtgg ctacctgttg ggccccacat catgggaaag	800
gctgtcaaac aatctttccc aagctccaag gcactcattt gtccttccc	850
cagcctccaa ttagaacaag ccaccacca gcctatctat ctccactga	900
gagggacctc gcagaatgag agaagacatt catgtaccac ctactagtcc	950
ctctctcccc aacctctgcc agggcaatct ctaactcaa tccgccttc	1000
gacagtgaag aagctctact tctacgtga cccagggagg aaacactagg	1050
accctgttgt atcctcaact gcaagtttct ggactagtct cccaacgttt	1100
gcctcccaat gttgtccctt tccttcgttc ccatggtaaa gtcctctctg	1150
ctttcctcct gaggtacac ccatgcgtct ctaggaactg gtcacaaaag	1200
tcatggtgcc tgcacccctg ccaagcccc ctgaccctct ctccccacta	1250
ccaccttctt cctgagctgg gggcaccagg gagaatcaga gatgctgggg	1300
atgccagagc aagactcaaa gaggcagagg ttttgttctc aaatattttt	1350
taataaatag acgaaaccac g	1371

<210> SEQ ID NO 28
<211> LENGTH: 277
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 28

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

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	140		145		150
Gln Ser Pro Arg	Lys Val Leu Gln Glu	Val Arg Arg Val Leu Arg			
	155		160		165
Pro Gly Gly Val	Leu Phe Phe Trp Glu	His Val Ala Glu Pro Tyr			
	170		175		180
Gly Ser Trp Ala	Phe Met Trp Gln Gln	Val Phe Glu Pro Thr Trp			
	185		190		195
Lys His Ile Gly	Asp Gly Cys Cys Leu Thr	Arg Glu Thr Trp Lys			
	200		205		210
Asp Leu Glu Asn	Ala Gln Phe Ser Glu	Ile Gln Met Glu Arg Gln			
	215		220		225
Pro Pro Pro Leu	Lys Trp Leu Pro Val	Gly Pro His Ile Met Gly			
	230		235		240
Lys Ala Val Lys	Gln Ser Phe Pro Ser	Ser Lys Ala Leu Ile Cys			
	245		250		255
Ser Phe Pro Ser	Leu Gln Leu Glu Gln	Ala Thr His Gln Pro Ile			
	260		265		270
Tyr Leu Pro Leu	Arg Gly Thr				
	275				

<210> SEQ ID NO 29
<211> LENGTH: 494
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 29

caatgtttgc ctatccacct cccccaagcc cctttaccta tgetgtgtgt	50
aacgctgtgtg ctgtgtgtgc tgetgtctta aggctcatgc ttggagtggg	100
gactggtcgg tgcccagaaa gtctcttctg ccaactgacgc ccccatcagg	150
gattggggcct tctttccccc ttcctttctg tgtctcctgc ctcatcggcc	200
tgccatgacc tgcagccaag ccagcccg tggggaaggg gagaaagtgg	250
gggatggcta agaaagctgg gagataggga acagaagagg gtagtgggtg	300
ggctaggggg gctgccttat ttaaagtggg tgtttatgat tcttatacta	350
atttatacaa agatattaag gcctgttca ttaagaaatt gttcccttcc	400
cctgtgttca atgtttgtaa agattgttct gtgtaaatat gtctttataa	450
taaacagtta aaagctgaaa aaaaaaaaaa aaaaaaaaaa aaaa	494

<210> SEQ ID NO 30
<211> LENGTH: 73
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 30

Met Leu Leu Leu Thr	Leu Leu Leu Leu Leu Leu Lys Gly	
1	5	10 15
Ser Cys Leu Glu Trp	Gly Leu Val Gly Ala Gln Lys Val Ser Ser	
	20	25 30
Ala Thr Asp Ala Pro	Ile Arg Asp Trp Ala Phe Phe Pro Pro Ser	
	35	40 45
Phe Leu Cys Leu Leu	Pro His Arg Pro Ala Met Thr Cys Ser Gln	
	50	55 60

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Ala Gln Pro Arg Gly Glu Gly Glu Lys Val Gly Asp Gly
65 70

<210> SEQ ID NO 31
<211> LENGTH: 1660
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 31

gtttgaattc cttcaactat acccacagtc caaaagcaga ctcaactgtgt	50
cccaggctac cagttcctcc aagcaagtca tttcccttat ttaaccgatg	100
tgctccctcaa acacctgagt gctactccct atttgcattct gttttgataa	150
atgatgttga caccctccac cgaattctaa gtggaatcat gtcgggaaga	200
gatacaatcc ttggcctgtg tatcctcgca ttagccttgt ctttggccat	250
gatgtttacc ttcagattca tcaccaccct tctgggtcac attttcattt	300
cattgggttat ttggggattg ttgtttgtct gcggtgtttt atggtggctg	350
tattatgact ataccaacga cctcagcata gaattggaca cagaaaggga	400
aaatatgaag tgcgtgctgg ggtttgctat cgtatccaca ggcatcacgg	450
cagtgtctgt cgtcttgatt ttgtttctca gaaagagaat aaaattgaca	500
gttagagcttt tccaaatcac aaataaagcc atcagcagtg ctcccttcct	550
gctgttccag ccactgtgga catttgccat cctcattttc ttctgggtcc	600
tcctgggtggc tgtgtgtgtg agcctgggaa ctgcaggagc tgcccagggt	650
atggaaggcg gccaaagtga atataagccc ctttcgggca ttcggtacat	700
gtggctgtac catttaattg gcctcatctg gactagtga ttcattcctg	750
cgtgccagcaaatgactata gctggggcag tgggtacttg ttatttcaac	800
agaagtaaaa atgatoctcc tgatcatccc atcctttcgt ctctctccat	850
tcctctcttc taccatcaag gaaccgttgt gaaagggcca tttttaatct	900
ctgtgggtgag gattccgaga atcattgtca tgtacatgca aaacgcactg	950
aaagaacagc agcatggtgc attgtccagg tacctgttcc gatgctgcta	1000
ctgctgtttc tgggtgtctg acaaatacct gctccatctc aaccagaatg	1050
catactacta aactgctatt aatgggacag atttctgtac atcagcaaaa	1100
gatgcattca aaatcttgtc caagaactca agtcacttta catctattaa	1150
ctgctttgga gacttcataa tttttctagg aaagggttta gtgggtgtgt	1200
tcactgtttt tggaggactc atggccttta actacaatcg ggcattccag	1250
gtgtgggacg tccctctgtt attggtagct ttttttgctt acttagtagc	1300
ccatagtttt ttatctgtgt ttgaaactgt gctggatgca cttttcctgt	1350
gttttgctgt tgatctggaa acaaatgatg gatcgtcaga aaagccctac	1400
tttatggatc aagaatttct gagtttcgta aaaaggagca acaaatataa	1450
caatgcaagg gcacagcagg acaagcactc attaaggaat gaggagggaa	1500
cagaactcca ggccattgtg agatagatac ccatttaggt atctgtacct	1550
ggaaaacatt tccttctaag agccatttac agaatagaag atgagaccac	1600

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tagagaaaag ttagtgaatt tttttttaaa agacctaata aaccctattc	1650
ttcctcaaaa	1660

<210> SEQ ID NO 32

<211> LENGTH: 445

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 32

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

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320					325					330				
Gly	Lys	Val	Leu	Val	Val	Cys	Phe	Thr	Val	Phe	Gly	Gly	Leu	Met
335					340					345				
Ala	Phe	Asn	Tyr	Asn	Arg	Ala	Phe	Gln	Val	Trp	Ala	Val	Pro	Leu
350					355					360				
Leu	Leu	Val	Ala	Phe	Phe	Ala	Tyr	Leu	Val	Ala	His	Ser	Phe	Leu
365					370					375				
Ser	Val	Phe	Glu	Thr	Val	Leu	Asp	Ala	Leu	Phe	Leu	Cys	Phe	Ala
380					385					390				
Val	Asp	Leu	Glu	Thr	Asn	Asp	Gly	Ser	Ser	Glu	Lys	Pro	Tyr	Phe
395					400					405				
Met	Asp	Gln	Glu	Phe	Leu	Ser	Phe	Val	Lys	Arg	Ser	Asn	Lys	Leu
410					415					420				
Asn	Asn	Ala	Arg	Ala	Gln	Gln	Asp	Lys	His	Ser	Leu	Arg	Asn	Glu
425					430					435				
Glu	Gly	Thr	Glu	Leu	Gln	Ala	Ile	Val	Arg					
440					445									

<210> SEQ ID NO 33
<211> LENGTH: 2773
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 33

gttcgattag ctcctctgag aagaagagaa aaggttcttg gacctctccc	50
tgtttcttcc ttagaataat ttgtatggga tttgtgatgc aggaaagcct	100
aagggaaaaa gaatatcat tctgtgtggt gaaaattttt tgaaaaaaa	150
attgccttct tcaaacaagg gtgtcattct gatatttatg aggactgttg	200
ttctcactat gaaggcatct gttattgaaa tgttccttgt tttgctggtg	250
actggagtac attcaaacia agaaacggca aagaagatta aaaggcccaa	300
gttcactgtg cctcagatca actgcatgtt caaagccgga aagatcatcg	350
atcctgagtt cattgtgaaa tgtccagcag gatgccaaga ccccaaatac	400
catgtttatg gcactgacgt gtatgcatcc tactccagtg tgtgtggcgc	450
tgccgtacac agtgggtgtg ttgataattc aggagggaaa atacttgttc	500
ggaagggttg tggacagtct gggtacaaa ggagttattc caacggtgtc	550
caatcgttat ccctaccacg atggagagaa tcctttatcg tcttagaaag	600
taaacccaaa aagggtgtaa cctaccatc agctcttaca tactcatcat	650
cgaaaagtcc agctgcccaa gcaggtgaga ccacaaaagc ctatcagagg	700
ccacctattc cagggacaac tgcacagccg gtcactctga tgcagttct	750
ggctgtcact gtagctgtgg ccacccccac caccttgcca aggccatccc	800
cttctgtctg ttctaccacc agcatcccca gaccacaatc agtgggccac	850
aggagccagg agatggatct ctggtccact gccacctaca caagcagcca	900
aaacaggccc agagctgata caggtatcca aaggcaagat ccttcaggag	950
ctgccttcca gaaacctgtt ggagcggatg tcagcctggg acttgttcca	1000
aaagaagaat tgagcacaca gtctttggag ccagtatccc tgggagatcc	1050

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aaactgcaaa attgacttgt cgtttttaat tgatgggagc accagcattg	1100
gcaaacggcg attccgaatc cagaagcagc tcctgggtga tgttgcccaa	1150
gctcttgaca ttggccctgc cggtcactg atgggtgttg tccagtatgg	1200
agacaacctt gctactcact ttaacctcaa gacacacagc aattctcgag	1250
atctgaagac agccatagag aaaattactc agagaggagg actttctaata	1300
gtaggtcggg ccatctcctt tgtgaccaag aacttctttt ccaaagccaa	1350
tggaaacaga agcggggctc ccaatgtggt ggtggtgatg gtggatggct	1400
ggccccagga caaagtggag gaggcttcaa gacttgcgag agagtcagga	1450
atcaacattt tcttcatcac cattgaaggt gctgctgaaa atgagaagca	1500
gtatgtggtg gagcccaact ttgcaacaa ggccgtgtgc agaacaaacg	1550
gcttctactc gctccacgtg cagagctggt ttggcctcca caagaccctg	1600
cagcctctgg tgaagcgggt ctgcgacact gaccgcctgg cctgcagcaa	1650
gacctgcttg aactcggtg acattggctt cgtcatcgac ggctccagca	1700
gtgtggggac gggcaacttc cgcaccgtcc tcagtttgt gaccaacctc	1750
accaaaagat ttgagatttc cgacacggac acgcgcctcg gggccgtgca	1800
gtacacctac gaacagcggc tggagtttgg gttcgacaag tacagcagca	1850
agcctgacat cctcaacgcc atcaagaggg tgggctactg gagtggggc	1900
accagcacgg gggctgccat caacttcgcc ctggagcagc tcttcaagaa	1950
gtccaagccc aacaagagga agttaatgat cctcatcacc gacgggaggt	2000
cctacgacga cgtccggatc ccagccatgg ctgccatct gaagggagt	2050
atcacctatg cgataggcgt tgctggggt gcccaagagg agctagaagt	2100
cattgccact caccgccca gagaccactc cttctttgtg gacgagttt	2150
acaacctcca tcagtatgtc ccaggatca tcagaacat ttgtacagag	2200
ttcaactcac agcctcggaa ctgaattcag agcaggcaga gcaccagcaa	2250
gtgctgcttt actaactgac gtgttgacc accccaccgc ttaatggggc	2300
acgcacggtg catcaagtct tgggcagggc atggagaaac aaatgtctg	2350
ttattattct ttgccatcat gctttttcat attccaaaac ttggagtac	2400
aaagatgatc acaaacgtat agaatgagcc aaaaggctac atcatgttg	2450
gggtgctgga gattttacct ttgacaatt gttttcaaaa taaatgttcg	2500
gaatacagtg cagcccttac gacaggctta cgtagagctt ttgtgagatt	2550
tttaagttgt tatttctgat ttgaactctg taaccctcag caagtttcat	2600
ttttgtcatg acaatgtagg aattgctgaa ttaaatgttt agaagatga	2650
aaaaataaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa	2700
aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa	2750
aaaaaaaaaa aaaaaaaaaa aag	2773

<210> SEQ ID NO 34

<211> LENGTH: 678

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

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<400> SEQUENCE: 34

Met	Arg	Thr	Val	Val	Leu	Thr	Met	Lys	Ala	Ser	Val	Ile	Glu	Met	1	5	10	15
Phe	Leu	Val	Leu	Leu	Val	Thr	Gly	Val	His	Ser	Asn	Lys	Glu	Thr	20	25	30	
Ala	Lys	Lys	Ile	Lys	Arg	Pro	Lys	Phe	Thr	Val	Pro	Gln	Ile	Asn	35	40	45	
Cys	Asp	Val	Lys	Ala	Gly	Lys	Ile	Ile	Asp	Pro	Glu	Phe	Ile	Val	50	55	60	
Lys	Cys	Pro	Ala	Gly	Cys	Gln	Asp	Pro	Lys	Tyr	His	Val	Tyr	Gly	65	70	75	
Thr	Asp	Val	Tyr	Ala	Ser	Tyr	Ser	Ser	Val	Cys	Gly	Ala	Ala	Val	80	85	90	
His	Ser	Gly	Val	Leu	Asp	Asn	Ser	Gly	Gly	Lys	Ile	Leu	Val	Arg	95	100	105	
Lys	Val	Ala	Gly	Gln	Ser	Gly	Tyr	Lys	Gly	Ser	Tyr	Ser	Asn	Gly	110	115	120	
Val	Gln	Ser	Leu	Ser	Leu	Pro	Arg	Trp	Arg	Glu	Ser	Phe	Ile	Val	125	130	135	
Leu	Glu	Ser	Lys	Pro	Lys	Lys	Gly	Val	Thr	Tyr	Pro	Ser	Ala	Leu	140	145	150	
Thr	Tyr	Ser	Ser	Ser	Lys	Ser	Pro	Ala	Ala	Gln	Ala	Gly	Glu	Thr	155	160	165	
Thr	Lys	Ala	Tyr	Gln	Arg	Pro	Pro	Ile	Pro	Gly	Thr	Thr	Ala	Gln	170	175	180	
Pro	Val	Thr	Leu	Met	Gln	Leu	Leu	Ala	Val	Thr	Val	Ala	Val	Ala	185	190	195	
Thr	Pro	Thr	Thr	Leu	Pro	Arg	Pro	Ser	Pro	Ser	Ala	Ala	Ser	Thr	200	205	210	
Thr	Ser	Ile	Pro	Arg	Pro	Gln	Ser	Val	Gly	His	Arg	Ser	Gln	Glu	215	220	225	
Met	Asp	Leu	Trp	Ser	Thr	Ala	Thr	Tyr	Thr	Ser	Ser	Gln	Asn	Arg	230	235	240	
Pro	Arg	Ala	Asp	Pro	Gly	Ile	Gln	Arg	Gln	Asp	Pro	Ser	Gly	Ala	245	250	255	
Ala	Phe	Gln	Lys	Pro	Val	Gly	Ala	Asp	Val	Ser	Leu	Gly	Leu	Val	260	265	270	
Pro	Lys	Glu	Glu	Leu	Ser	Thr	Gln	Ser	Leu	Glu	Pro	Val	Ser	Leu	275	280	285	
Gly	Asp	Pro	Asn	Cys	Lys	Ile	Asp	Leu	Ser	Phe	Leu	Ile	Asp	Gly	290	295	300	
Ser	Thr	Ser	Ile	Gly	Lys	Arg	Arg	Phe	Arg	Ile	Gln	Lys	Gln	Leu	305	310	315	
Leu	Ala	Asp	Val	Ala	Gln	Ala	Leu	Asp	Ile	Gly	Pro	Ala	Gly	Pro	320	325	330	
Leu	Met	Gly	Val	Val	Gln	Tyr	Gly	Asp	Asn	Pro	Ala	Thr	His	Phe	335	340	345	
Asn	Leu	Lys	Thr	His	Thr	Asn	Ser	Arg	Asp	Leu	Lys	Thr	Ala	Ile	350	355	360	
Glu	Lys	Ile	Thr	Gln	Arg	Gly	Gly	Leu	Ser	Asn	Val	Gly	Arg	Ala	365	370	375	

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

<210> SEQ ID NO 35
<211> LENGTH: 2095
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 35

c c g a g c a c a g g a g a t t g c c t g c g t t t a g g a g g t g g c t g c g t t g t g g g a a a	50
a g c t a t c a a g g a a g a a a t t g c c a a a c c a t g t c t t t t t t t c t g t t t t c a g a	100
g t a g t t c a c a a c a g a t c t g a g t g t t t t a a t a a g c a t g g a a t a c a g a a a a	150

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caacaaaaa cttaaagcttt aatttcatct ggaattccac agttttctta	200
gctccctgga cccggttgac ctgttggtc tccccgtgg ctgctctatc	250
acgtgggtgct ctccgactac tcaccccgag tgtaaagaac cttcggtcgc	300
cggtgcttctg agctgctgtg gatggcctcg gctctctgga ctgtccttcc	350
gagtaggatg tcaactgagat ccctcaaatg gagcctcctg ctgctgtcac	400
tcctgagttt ctttgtgatg tggtaacctc gccttcccca ctacaatgtg	450
atagaacgcg tgaactggat gtacttctat gagtatgagc cgatttacag	500
acaagacttt cacttcacac ttcgagagca ttcaaactgc tctcatcaaa	550
atccatttct ggtcattctg gtgacctccc acccttcaga tgtgaaagcc	600
aggcaggcca ttagagttac ttggggtgaa aaaaagtctt ggtggggata	650
tgaggttctt acatttttct tattaggcca agaggctgaa aaggaagaca	700
aatgtttggc attgtcctta gaggatgaac accttcttta tggtgacata	750
atccgacaag attttttaga cacatataat aacctgacct tgaaccat	800
tatggcattc aggtgggtaa ctgagttttg cccaatgcc aagtacgtaa	850
tgaagacaga cactgatggt ttcataata ctggcaattt agtgaagtat	900
cttttaaacc taaaccactc agagaagttt ttcacaggtt atcctcta	950
tgataattat tcctatagag gattttacca aaaaacccat atttcttacc	1000
aggagtatcc tttcaagggt ttcctccat actgcagtgg gttgggttat	1050
ataatgtcca gagatttggg gccaaaggatc tatgaaatga tgggtcacgt	1100
aaaacccatc aagtttgaag atgtttatgt cgggatctgt ttgaatttat	1150
taaaagtga cattcatatt ccagaagaca caaatctttt ctttctatat	1200
agaatccatt tggatgtctg tcaactgaga cgtgtgattg cagcccatgg	1250
cttttcttcc aaggagatca tcaacttttg gcaggtcag ctaaggaaca	1300
ccacatgcca ttattaactt cacattctac aaaaagccta gaaggacagg	1350
ataccttggt gaaagtgtta aataaagtag gtactgtgga aaattcatgg	1400
ggaggtcagt gtgctggctt aactgaact gaaactcatg aaaaaccag	1450
actggagact ggagggttac acttgtgatt tattagtcag gcccttcaaa	1500
gatgatatgt ggaggaatta aatataaagg aattggaggt ttttgctaaa	1550
gaaattaata ggaccaaca atttgacat gtcattctgt agactagaat	1600
ttcttaaaag ggtgttactg agttataagc tcaactagct gtaaaaacaa	1650
aacaatgtag agttttatct attgaacaat gtagtcactt gaaggttttg	1700
tgatatatct atgtggatta ccaatttaaa aatatatgta gttctgtgtc	1750
aaaaaacttc ttcactgaag ttatactgaa caaaatttta cctgtttttg	1800
gtcatttata aagtacttca agatgttgca gtatttcaca gttattatta	1850
tttaaaatta cttcaacttt gtgtttttta atgttttgac gatttcaata	1900
caagataaaa aggatagtga atcattcttt acatgcaaac attttccagt	1950
tacttaactg atcagtttat tattgatata tcaactcatt aatgtaaagt	2000
cataggtcat tattgcatat cagtaatctc ttggactttg ttaaatattt	2050

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tactgtggta atatagagaa gaattaaagc aagaaaatct gaaaa

2095

<210> SEQ ID NO 36

<211> LENGTH: 331

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 36

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

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Tyr

<210> SEQ ID NO 37

<211> LENGTH: 2846

<212> TYPE: DNA

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 37

cgctcgggca ccagccgcgg caaggatgga gctgggttgc tggacgcagt	50
tggggctcac ttttcttcag ctccctctca tctcgtcctt gccaaagagag	100
tacacagctca ttaatgaagc ctgccctgga gcagagtgga atatcatgtg	150
tcgggagtgct tgtgaatatg atcagattga gtgcgtctgc cccggaaaga	200
gggaagtctgt gggttatacc atcccttgct gcaggaatga ggagaatgag	250
tgtgactcct gcctgatcca cccaggttgt accatctttg aaaactgcaa	300
gagctgccga aatggctcat ggggggttac cttggatgac ttctatgtga	350
aggggttcta ctgtgcagag tgccgagcag gctggtagcg aggagactgc	400
atgcgatgtg gccaggttct gcgagcccca aagggtcaga tttgtttgga	450
aagctatccc ctaaatgctc actgtgaatg gaccattcat gctaaacctg	500
ggtttgtcat ccaactaaga ttgtcatgt tgagtctgga gtttgactac	550
atgtgccagt atgactatgt tgaggttcgt gatggagaca accgcgatgg	600
ccagatcatc aagcgtgtct gtggcaacga gcggccagct cctatccaga	650
gcataaggatc ctcaactccac gtccctcttc actccgatgg ctccaagaat	700
tttgacggtt tccatgccat ttatgaggag atcacagcat gctcctcatc	750
ccctgttttc catgacggca cgtgcgtcct tgacaaggct ggatcttaca	800
agtggtcctg cttggcaggc tatactgggc agcgtgtga aaatctcctt	850
gaagaaagaa actgctcaga ccctgggggc ccagtcaatg ggtaccagaa	900
aataacaggg ggccctgggc ttatcaacgg acgccatgct aaaattggca	950
ccgtggtgtc tttcttttgt aacaactcct atgttcttag tggcaatgag	1000
aaaagaactt gccagcagaa tggagagtgg tcagggaaac agcccatctg	1050
cataaaagcc tgccgagaac caaagatttc agacctggtg agaaggagag	1100
ttcttccgat gcaggttcag tcaagggaga caccattaca ccagctatac	1150
tcagcggcct tcagcaagca gaaactgcag agtgccccta ccaagaagcc	1200
agcccttccc tttggagatc tgcccatggg ataccaacat ctgcataccc	1250
agctccagta tgagtgcac tcacccttct accgccgcct gggcagcagc	1300
aggaggacat gtctgaggac tgggaagtgg agtgggcggg caccatcctg	1350
catccctatc tgcgggaaaa ttgagaacat cactgctcca aagacccaag	1400
ggttgcgctg gccgtggcag gcagccatct acaggaggac cagcggggtg	1450
catgacggca gcctacacaa gggagcgtgg ttcctagtct gcagcgggtg	1500
cctggtgaat gagcgactg tgggtgtggc tgcccactgt gttactgacc	1550
tggggaaggt caccatgatc aagacagcag acctgaaagt tgttttgggg	1600
aaattctacc gggatgatga cgggatgag aagaccatcc agagcctaca	1650

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gattttctgct atcattctgc atcccaacta tgaccccatc ctgcttgatg 1700
ctgacatcgc catcctgaag ctccctagaca aggcccgat cagcaccga 1750
gtccagccca tctgcctcgc tgccagtcgg gatctcagca ctcccttcca 1800
ggagtcccac atcactgtgg ctggctggaa tgtcctggca gacgtgagga 1850
gccctggctt caagaacgac aactgcgct ctgggggtgg cagtgtggtg 1900
gactcgctgc tgtgtgagga gcagcatgag gacatggca tccagtgag 1950
tgtcactgat aacatgttct gtgccagctg ggaaccact gcccttctg 2000
atatctgcac tgacagaca ggaggcatcg cggtgtgtc ctcccgga 2050
cgagcatctc ctgagccacg ctggcatctg atgggactgg tcagctggag 2100
ctatgataaa acatgcagcc acaggctctc cactgccttc accaaggctc 2150
tgccctttta agactggatt gaaagaaata tgaatgaac catgctcatg 2200
cactccttga gaagtgtttc tgtatatccg tctgtacgtg tgtcattgcg 2250
tgaagcagtg tgggcctgaa gtgtgatttg gcctgtgaac ttggctgtgc 2300
cagggcttct gacttcaggg aaaaactca gtgaagggtg agtagacctc 2350
cattgctggt aggtgatgc cgcgtccact actaggacag ccaattggaa 2400
gatgccaggg cttgcaagaa gtaagtttct tcaagaaga ccatatacaa 2450
aacctctcca ctccactgac ctggtggtct tcccaactt tcagttatac 2500
gaatgccatc agcttgacca gggaagatct gggcttcatg aggcccttt 2550
tgaggctctc aagttctaga gagctgcctg tgggacagcc cagggcagca 2600
gagctgggat gtggtgcatg cttttgtgta catggccaca gtacagtctg 2650
gtccttttcc ttcccatct cttgtacaca ttttaataaa ataagggttg 2700
gcttctgaac tacaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa 2750
aaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa 2800
aaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaa 2846

<210> SEQ ID NO 38

<211> LENGTH: 720

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 38

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

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95					100					105				
Phe	Tyr	Val	Lys	Gly	Phe	Tyr	Cys	Ala	Glu	Cys	Arg	Ala	Gly	Trp
				110					115					120
Tyr	Gly	Gly	Asp	Cys	Met	Arg	Cys	Gly	Gln	Val	Leu	Arg	Ala	Pro
				125					130					135
Lys	Gly	Gln	Ile	Leu	Leu	Glu	Ser	Tyr	Pro	Leu	Asn	Ala	His	Cys
				140					145					150
Glu	Trp	Thr	Ile	His	Ala	Lys	Pro	Gly	Phe	Val	Ile	Gln	Leu	Arg
				155					160					165
Phe	Val	Met	Leu	Ser	Leu	Glu	Phe	Asp	Tyr	Met	Cys	Gln	Tyr	Asp
				170					175					180
Tyr	Val	Glu	Val	Arg	Asp	Gly	Asp	Asn	Arg	Asp	Gly	Gln	Ile	Ile
				185					190					195
Lys	Arg	Val	Cys	Gly	Asn	Glu	Arg	Pro	Ala	Pro	Ile	Gln	Ser	Ile
				200					205					210
Gly	Ser	Ser	Leu	His	Val	Leu	Phe	His	Ser	Asp	Gly	Ser	Lys	Asn
				215					220					225
Phe	Asp	Gly	Phe	His	Ala	Ile	Tyr	Glu	Glu	Ile	Thr	Ala	Cys	Ser
				230					235					240
Ser	Ser	Pro	Cys	Phe	His	Asp	Gly	Thr	Cys	Val	Leu	Asp	Lys	Ala
				245					250					255
Gly	Ser	Tyr	Lys	Cys	Ala	Cys	Leu	Ala	Gly	Tyr	Thr	Gly	Gln	Arg
				260					265					270
Cys	Glu	Asn	Leu	Leu	Glu	Glu	Arg	Asn	Cys	Ser	Asp	Pro	Gly	Gly
				275					280					285
Pro	Val	Asn	Gly	Tyr	Gln	Lys	Ile	Thr	Gly	Gly	Pro	Gly	Leu	Ile
				290					295					300
Asn	Gly	Arg	His	Ala	Lys	Ile	Gly	Thr	Val	Val	Ser	Phe	Phe	Cys
				305					310					315
Asn	Asn	Ser	Tyr	Val	Leu	Ser	Gly	Asn	Glu	Lys	Arg	Thr	Cys	Gln
				320					325					330
Gln	Asn	Gly	Glu	Trp	Ser	Gly	Lys	Gln	Pro	Ile	Cys	Ile	Lys	Ala
				335					340					345
Cys	Arg	Glu	Pro	Lys	Ile	Ser	Asp	Leu	Val	Arg	Arg	Arg	Val	Leu
				350					355					360
Pro	Met	Gln	Val	Gln	Ser	Arg	Glu	Thr	Pro	Leu	His	Gln	Leu	Tyr
				365					370					375
Ser	Ala	Ala	Phe	Ser	Lys	Gln	Lys	Leu	Gln	Ser	Ala	Pro	Thr	Lys
				380					385					390
Lys	Pro	Ala	Leu	Pro	Phe	Gly	Asp	Leu	Pro	Met	Gly	Tyr	Gln	His
				395					400					405
Leu	His	Thr	Gln	Leu	Gln	Tyr	Glu	Cys	Ile	Ser	Pro	Phe	Tyr	Arg
				410					415					420
Arg	Leu	Gly	Ser	Ser	Arg	Arg	Thr	Cys	Leu	Arg	Thr	Gly	Lys	Trp
				425					430					435
Ser	Gly	Arg	Ala	Pro	Ser	Cys	Ile	Pro	Ile	Cys	Gly	Lys	Ile	Glu
				440					445					450
Asn	Ile	Thr	Ala	Pro	Lys	Thr	Gln	Gly	Leu	Arg	Trp	Pro	Trp	Gln
				455					460					465
Ala	Ala	Ile	Tyr	Arg	Arg	Thr	Ser	Gly	Val	His	Asp	Gly	Ser	Leu
				470					475					480

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His	Lys	Gly	Ala	Trp	Phe	Leu	Val	Cys	Ser	Gly	Ala	Leu	Val	Asn
				485					490					495
Glu	Arg	Thr	Val	Val	Val	Ala	Ala	His	Cys	Val	Thr	Asp	Leu	Gly
				500					505					510
Lys	Val	Thr	Met	Ile	Lys	Thr	Ala	Asp	Leu	Lys	Val	Val	Leu	Gly
				515					520					525
Lys	Phe	Tyr	Arg	Asp	Asp	Asp	Arg	Asp	Glu	Lys	Thr	Ile	Gln	Ser
				530					535					540
Leu	Gln	Ile	Ser	Ala	Ile	Ile	Leu	His	Pro	Asn	Tyr	Asp	Pro	Ile
				545					550					555
Leu	Leu	Asp	Ala	Asp	Ile	Ala	Ile	Leu	Lys	Leu	Leu	Asp	Lys	Ala
				560					565					570
Arg	Ile	Ser	Thr	Arg	Val	Gln	Pro	Ile	Cys	Leu	Ala	Ala	Ser	Arg
				575					580					585
Asp	Leu	Ser	Thr	Ser	Phe	Gln	Glu	Ser	His	Ile	Thr	Val	Ala	Gly
				590					595					600
Trp	Asn	Val	Leu	Ala	Asp	Val	Arg	Ser	Pro	Gly	Phe	Lys	Asn	Asp
				605					610					615
Thr	Leu	Arg	Ser	Gly	Val	Val	Ser	Val	Val	Asp	Ser	Leu	Leu	Cys
				620					625					630
Glu	Glu	Gln	His	Glu	Asp	His	Gly	Ile	Pro	Val	Ser	Val	Thr	Asp
				635					640					645
Asn	Met	Phe	Cys	Ala	Ser	Trp	Glu	Pro	Thr	Ala	Pro	Ser	Asp	Ile
				650					655					660
Cys	Thr	Ala	Glu	Thr	Gly	Gly	Ile	Ala	Ala	Val	Ser	Phe	Pro	Gly
				665					670					675
Arg	Ala	Ser	Pro	Glu	Pro	Arg	Trp	His	Leu	Met	Gly	Leu	Val	Ser
				680					685					690
Trp	Ser	Tyr	Asp	Lys	Thr	Cys	Ser	His	Arg	Leu	Ser	Thr	Ala	Phe
				695					700					705
Thr	Lys	Val	Leu	Pro	Phe	Lys	Asp	Trp	Ile	Glu	Arg	Asn	Met	Lys
				710					715					720

<210> SEQ ID NO 39
<211> LENGTH: 2571
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 39

ggttcctaca tcctctcatc tgagaatcag agagcataat cttcttacgg	50
gcccgtgatt tattaacgtg gcttaatctg aaggttctca gtcaaattct	100
ttgtgatcta ctgattgtgg gggcatggca aggtttgctt aaaggagctt	150
ggctgggttg ggcccttgta gctgacagaa ggtggccagg gagaatgcag	200
cacactgtct ggagaatgaa ggcgcttctg ttgctgttct tgccttggtc	250
cagtcctgct aactacattg acaatgtggg caacctgcac ttcctgtatt	300
cagaactctg taaaggtgcc tcccactacg gcctgaccaa agataggaag	350
agggcgtcac aagatggctg tccagacggc tgtgcgagcc tcacagccac	400
ggctccctcc ccagaggttt ctgcagctgc caccatctcc ttaatgacag	450
acgagcctgg cctagacaac cctgcctacg tgtcctcggc agaggacggg	500

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cagccagcaa tcagcccagt ggactctggc cggagcaacc gaactagggc	550
acggcccttt gagagatcca ctattagaag cagatcattt aaaaaataa	600
atcagagcttt gagtgttctt cgaaggacaa agagcgggag tgcagttgcc	650
aaccatgccg accagggcag ggaaaattct gaaaacacca ctgccctga	700
agtctttcca aggttgtacc acctgattcc agatggtgaa attaccagca	750
tcaagatcaa tcgagtagat ccagtgaaa gcctctctat taggctggtg	800
ggaggtagcg aaacccact ggtccatata attatccaac acatttatcg	850
tgatggggtg atcgccagag acggccggct actgccagga gacatcatc	900
taaaggtcaa cgggatggac atcagcaatg tcctcacia ctacgctgtg	950
cgtctctcgc ggagccctg ccaggtgctg tggctgactg tgatgcgtga	1000
acagaagtgc cgcagcagga acaatggaca ggccccgat gcctacagac	1050
cccagatga cagctttcat gtgattctca acaaaagtag ccccgaggag	1100
cagcttgtaa taaaactggt gcgcaagggt gatgagcctg gggttttcat	1150
cttcaatgtg ctggatggcg gtgtggcata tcgacatggt cagcttgagg	1200
agaatgaccg tgtgttagcc atcaatggac atgatcttcg atatggcagc	1250
ccagaaagtg cggctcatct gattcaggcc agtgaaagac gtgttcacct	1300
cgtcgtgtcc cgccagggtc ggcagcggag ccctgacatc ttacaggaag	1350
ccggctggaa cagcaatggc agctggtccc cagggccagg ggagaggagc	1400
aacactccca agcccccca tcctacaatt acttgatcat agaaggtggt	1450
aaatatccaa aaagacccc gtgaatctct cggcatgacc gtcgcagggg	1500
gagcatcaca tagagaatgg gatttgccta tctatgtcat cagtgttgag	1550
cccggaggag tcataagcag agatggaaga ataaaaacag gtgacatttt	1600
gttgaatgtg gatggggtcg aactgacaga ggtcagccg agtgaggcag	1650
tggcattatt gaaaagaaca tcatcctcga tagtactcaa agctttggaa	1700
gtcaaagagt atgagcccca ggaagactgc agcagcccag cagccctgga	1750
ctccaaccac aacatggccc caccagtgga ctggtcccca tcctgggtca	1800
tgtggttgga attaccacgg tgcttgata actgtaaaga tattgtatta	1850
cgaagaaaca cagctggaag tctgggcttc tgcattgtag gaggttatga	1900
agaatacaat ggaacaaac cttttttcat caaatccatt gttgaaggaa	1950
caccagcata caatgatgga agaattagat gtggtgatat tcttcttgct	2000
gtcaatggta gaagtacatc aggaatgata catgcttgct tggcaagact	2050
gctgaaagaa cttaaaggaa gaattactct aactattggt tcttgccctg	2100
gcactttttt atagaatcaa tgatgggtca gaggaaaaca gaaaaatcac	2150
aaataggcta agaagttgaa acactatatt tatcttgta gtttttata	2200
ttaaagaaag aatacattgt aaaaatgtca ggaaaagtat gatcatctaa	2250
tgaaagccag ttacacctca gaaaatatga ttccaaaaaa attaaaacta	2300
ctagtttttt ttcagtgtgg aggatttctc attactctac aacattgttt	2350
atattttttc tattcaataa aaagccctaa aacaactaaa atgattgatt	2400

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tgtatacccc actgaattca agctgattta aatttaaaat ttggtatatg	2450
ctgaagtctg ccaagggtac attatggcca tttttaattt acagctaaaa	2500
tatttttttaa aatgcattgc tgagaaacgt tgctttcatc aaacaagaat	2550
aaatatTTTTT cagaagttaa a	2571

<210> SEQ ID NO 40
 <211> LENGTH: 632
 <212> TYPE: PRT
 <213> ORGANISM: Homo Sapien
 <400> SEQUENCE: 40

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

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

Phe Leu

<210> SEQ ID NO 41

<211> LENGTH: 1964

<212> TYPE: DNA

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 41

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accaggcatt gtatcttcag ttgtcatcaa gttcgcaatc agattggaaa	50
agctcaactt gaagctttct tgctgcagt gaagcagaga gatagatatt	100
attcacgtaa taaaaaacat gggcttcaac ctgactttcc acctttccta	150
caaattccga ttactgttgc tgttgacttt gtgcctgaca gtggttgggt	200
gggccaccag taactacttc gtgggtgccca ttcaagagat tcctaaagca	250
aaggagttca tggctaattt ccataagacc ctcatTTtTg ggaaggga	300
aactctgact aatgaagcat ccacgaaga ggtagaactt gacaactgtc	350
cttctgtgtc tccttacctc agaggccaga gcaagctcat tttcaaacca	400
gatctcactt tggaagaggt acaggcagaa aatcccaag tgtccagagg	450
ccggtatcgc cctcaggaat gtaaagcttt acagagggtc gccatcctcg	500
ttccccacg gaacagagag aaacacctga tgtacctgct ggaacatctg	550
catcccttcc tgcagaggca gcagctggat tatggcatct acgtcatcca	600
ccaggctgaa ggtaaaaagt ttaatcgagc caaactcttg aatgtgggct	650
atctagaagc cctcaaggaa gaaaattggg actgctttat attccacgat	700
gtggacctgg taccgagaa tgactttaac ctttacaagt gtgaggagca	750
tcccagcat ctggtggttg gcaggaaacg cactgggtac aggttacgtt	800
acagtggata ttttgggggt gttactgccc taagcagaga gcagtttttc	850
aagggtaatg gattctctaa caactactgg ggatggggag gcgaagacga	900
tgacctcaga ctcagggttg agctccaaag aatgaaaatt tcccggcccc	950
tgctgaagt gggtaaatat acaatgggtc tccacactag agacaaaggc	1000
aatgaggtga acgcagaacg gatgaagctc ttacaccaag tgtcacgagt	1050
ctggagaaca gatgggttga gtagttgttc ttataaatta gtatctgttg	1100
aacacaatcc tttatatatc aacatcacag tggatttctg gtttggtgca	1150
tgaccttgga tcttttgggt atgtttggaa gaactgattc tttgtttgca	1200
ataattttgg cctagagact tcaaatagta gcacacatta agaacctgtt	1250
acagctcatt gttgagctga atttttcctt tttgtatttt cttagcagag	1300
ctcctgggtga tgtagagtat aaaacagttg taacaagaca gctttcttag	1350
tcattttgat catgagggtt aaatattgta atatggatac ttgaaggact	1400
ttatataaaa ggatgactca aaggataaaa tgaacgctat ttgaggactc	1450
tggttgaagg agattttattt aaatttgaag taatatatta tgggataaaa	1500
ggccacagga aataagactg ctgaatgtct gagagaacca gagttgttct	1550
cgtccaaggt agaaaggtag gaagatacaa tactgttatt catTTtatcct	1600
gtacaatcat ctgtgaagtg gtggtgtcag gtgagaaggc gtccacaaaa	1650
gaggggagaa aaggcgacga atcaggacac agtgaacttg ggaatgaaga	1700
ggtagcagga ggggtggagt tcggctgcaa aggcagcagt agctgagctg	1750
gttgacagtg ctgatagcct tcaggggagg acctgccag gtatgccttc	1800
cagtgatgcc caccagagaa tacattctct attagttttt aaagagtttt	1850
tgtaaaatga ttttgtacaa gtaggatag aattagcagt ttacaagttt	1900

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acatatataac taataataaaa tatgtctatc aaatacctct gtagtaaaat 1950
gtgaaaaaagc aaaa 1964

<210> SEQ ID NO 42
<211> LENGTH: 344
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 42

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

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Gly Leu Ser Ser Cys Ser Tyr Lys Leu Val Ser Val Glu His Asn
320 325 330

Pro Leu Tyr Ile Asn Ile Thr Val Asp Phe Trp Phe Gly Ala
335 340

<210> SEQ ID NO 43
<211> LENGTH: 485
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 43
gctcaagacc cagcagtggg acagccagac agacggcacg atggcactga 50
gctcccatgat ctgggccgct tgcctcctgc tcctcctcct cctcgccagc 100
ctgaccagtgt gctctgtttt cccacaacag acgggacaac ttgcagagct 150
gcaaccccag gacagagctg gagccagggc cagctggatg cccatgttcc 200
agagcggaag gaggcgagac acccacttcc ccatctgcat tttctgctgc 250
ggctgctgtc atcgatcaaa gtgtgggatg tgctgcaaga cgtagaacct 300
acctgccctg cccccgtccc ctccccttct tatttattcc tgctgcccc 350
gaacataggt cttggaataa aatggctggt tcttttgttt tccaaaaaaa 400
aaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa 450
aaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaa 485

<210> SEQ ID NO 44
<211> LENGTH: 84
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 44
Met Ala Leu Ser Ser Gln Ile Trp Ala Ala Cys Leu Leu Leu Leu
1 5 10 15
Leu Leu Leu Ala Ser Leu Thr Ser Gly Ser Val Phe Pro Gln Gln
20 25 30
Thr Gly Gln Leu Ala Glu Leu Gln Pro Gln Asp Arg Ala Gly Ala
35 40 45
Arg Ala Ser Trp Met Pro Met Phe Gln Arg Arg Arg Arg Arg Asp
50 55 60
Thr His Phe Pro Ile Cys Ile Phe Cys Cys Gly Cys Cys His Arg
65 70 75
Ser Lys Cys Gly Met Cys Cys Lys Thr
80

<210> SEQ ID NO 45
<211> LENGTH: 1076
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 45
gtggcttcat ttcagtggct gacttcaga gagcaatatg gctggttccc 50
caacatgcct caccctcatc tatatccttt ggcagctcac agggtcagca 100
gcctctggac ccgtgaaaga gctggtcggt tccgttggtg gggccgtgac 150
tttccccctg aagtccaaag taaagcaagt tgactctatt gtctggacct 200

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<210> SEQ ID NO 46
<211> LENGTH: 335
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 46
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Met	Ala	Gly	Ser	Pro	Thr	Cys	Leu	Thr	Leu	Ile	Tyr	Ile	Leu	Trp
1				5					10					15
Gln	Leu	Thr	Gly	Ser	Ala	Ala	Ser	Gly	Pro	Val	Lys	Glu	Leu	Val
				20					25					30
Gly	Ser	Val	Gly	Gly	Ala	Val	Thr	Phe	Pro	Leu	Lys	Ser	Lys	Val
				35					40					45
Lys	Gln	Val	Asp	Ser	Ile	Val	Trp	Thr	Phe	Asn	Thr	Thr	Pro	Leu
				50					55					60
Val	Thr	Ile	Gln	Pro	Glu	Gly	Gly	Thr	Ile	Ile	Val	Thr	Gln	Asn
				65					70					75
Arg	Asn	Arg	Glu	Arg	Val	Asp	Phe	Pro	Asp	Gly	Gly	Tyr	Ser	Leu
				80					85					90
Lys	Leu	Ser	Lys	Leu	Lys	Lys	Asn	Asp	Ser	Gly	Ile	Tyr	Tyr	Val
				95					100					105
Gly	Ile	Tyr	Ser	Ser	Ser	Leu	Gln	Gln	Pro	Ser	Thr	Gln	Glu	Tyr
				110					115					120
Val	Leu	His	Val	Tyr	Glu	His	Leu	Ser	Lys	Pro	Lys	Val	Thr	Met
				125					130					135
Gly	Leu	Gln	Ser	Asn	Lys	Asn	Gly	Thr	Cys	Val	Thr	Asn	Leu	Thr
				140					145					150
Cys	Cys	Met	Glu	His	Gly	Glu	Glu	Asp	Val	Ile	Tyr	Thr	Trp	Lys
				155					160					165

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Ala	Leu	Gly	Gln	Ala	Ala	Asn	Glu	Ser	His	Asn	Gly	Ser	Ile	Leu
	170								175					180
Pro	Ile	Ser	Trp	Arg	Trp	Gly	Glu	Ser	Asp	Met	Thr	Phe	Ile	Cys
	185								190					195
Val	Ala	Arg	Asn	Pro	Val	Ser	Arg	Asn	Phe	Ser	Ser	Pro	Ile	Leu
	200								205					210
Ala	Arg	Lys	Leu	Cys	Glu	Gly	Ala	Ala	Asp	Asp	Pro	Asp	Ser	Ser
	215								220					225
Met	Val	Leu	Leu	Cys	Leu	Leu	Leu	Val	Pro	Leu	Leu	Leu	Ser	Leu
	230								235					240
Phe	Val	Leu	Gly	Leu	Phe	Leu	Trp	Phe	Leu	Lys	Arg	Glu	Arg	Gln
	245								250					255
Glu	Glu	Tyr	Ile	Glu	Glu	Lys	Lys	Arg	Val	Asp	Ile	Cys	Arg	Glu
	260								265					270
Thr	Pro	Asn	Ile	Cys	Pro	His	Ser	Gly	Glu	Asn	Thr	Glu	Tyr	Asp
	275								280					285
Thr	Ile	Pro	His	Thr	Asn	Arg	Thr	Ile	Leu	Lys	Glu	Asp	Pro	Ala
	290								295					300
Asn	Thr	Val	Tyr	Ser	Thr	Val	Glu	Ile	Pro	Lys	Lys	Met	Glu	Asn
	305								310					315
Pro	His	Ser	Leu	Leu	Thr	Met	Pro	Asp	Thr	Pro	Arg	Leu	Phe	Ala
	320								325					330
Tyr	Glu	Asn	Val	Ile										
	335													

<210> SEQ ID NO 47
<211> LENGTH: 766
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 47

ggctcgagcg tttctgagcc aggggtgacc atgacctgct gcgaaggatg	50
gacatcctgc aatggattca gcctgctggt tctactgctg ttaggagtag	100
ttctcaatgc gatacctcta attgtcagct tagttgagga agaccaattt	150
tctcaaaacc ccatctcttg ctttgagtgg tgggtcccag gaattatag	200
agcagggtctg atggccattc cagcaacaac aatgtccttg acagcaagaa	250
aaagagcgtg ctgcaacaac agaactggaa tgtttctttc atcatttttc	300
agtgatgatca cagtcattgg tgctctgtat tgcagtctga tatccatcca	350
ggctctctta aaaggtcctc tcatgtgtaa ttctccaagc aacagtaatg	400
ccaattgtga attttcattg aaaaacatca gtgacattca tccagaatcc	450
ttcaacttgc agtgggtttt caatgactct tgtgcacctc ctactggttt	500
caataaacc accagtaacg acaccatggc gagtggctgg agagcatcta	550
gtttccactt cgattctgaa gaaaacaaac ataggcttat ccacttctca	600
gtatttttag gtctattgct tgttggaatt ctggaggtcc tgtttgggct	650
cagtcagata gtcatcgggt tccttggtg tctgtgtgga gtctctaagc	700
gaagaagtca aattgtgtag tttaatggga ataaaatgta agtatcagta	750
gtttgaaaaa aaaaaa	766

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<210> SEQ ID NO 48
<211> LENGTH: 229
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 48

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

<210> SEQ ID NO 49
<211> LENGTH: 636
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 49

atccgtttctc tgcgctgccca gctcaggtga gccctcgcca aggtgacctc 50
gcagagacct ggtgaaggag cagtgaaggaa cctgcagagt cacacagttg 100
ctgaccaatt gagctgtgag cctggagcag atccgtgggc tgcagacccc 150
cgccccagtg cctctcccc tgcagccctg cccctcgaac tgtgacatgg 200
agagagtgc cctggccctt ctectactgg caggcctgac tgccttgga 250
gccaatgacc catttgccaa taaagacgat cccttctact atgactggaa 300

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aaacctgcag ctgagcggac tgatctgcgg agggctcctg gccattgctg	350
ggatcgcggc agttctgagt ggcaaatgca aatacaagag cagccagaag	400
cagcacagtc ctgtacctga gaaggccatc ccactcatca ctccaggctc	450
tgccactact tgctgagcac aggactggcc tccagggatg gcctgaagcc	500
taacactggc cccagcacc tcctcccctg ggaggcctta tcctcaagga	550
aggacttctc tccaagggca ggctgttagg cccctttctg atcaggaggc	600
ttctttatga attaaactcg cccaccacc ccctca	636

<210> SEQ ID NO 50
<211> LENGTH: 89
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 50

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

<210> SEQ ID NO 51
<211> LENGTH: 1734
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 51

gtggactctg agaagcccag gcagttgagg acaggagaga gaaggctgca	50
gaccagaggg gagggaggac agggagtctg aaggaggagg acagaggagg	100
gcacagagac gcagagcaag gccggcaagg aggagaccct ggtgggagga	150
agacactctg gagagagagg gggctgggca gagatgaagt tccaggggcc	200
cctggcctcg ctctgctgg ccctctgcct gggcagtggt gaggtggcc	250
ccctgcagag cggagaggaa agcactggga caaatattgg ggaggccctt	300
ggacatggcc tgggagacgc cctgagcgaa ggggtgggaa aggccattgg	350
caaagaggcc ggaggggcag ctggctctaa agtcagttag gcccttggcc	400
aagggaccag agaagcagtt ggcactggag tcaggcaggt tccaggcttt	450
ggcgacagc atgctttggg caacagggtc ggggaagcag cccatgctct	500
gggaaacact gggcacgaga ttggcagaca ggcagaagat gtcattcgac	550
acggagcaga tgctgtccgc ggctcctggc aggggggtgcc tggccacagt	600
ggtgcttggg aaacttctgg aggccatggc atctttggct ctcaaggtgg	650
ccttgagggc cagggccagg gcaatcctgg aggtctgggt actccgtggg	700

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tccacggata ccccggaac tcagcaggca gctttggaat gaatcctcag	750
ggagctccct ggggtcaagg aggcaatgga gggccaccaa actttgggac	800
caacactcag ggagctgtgg ccagcctgg ctatggttca gtgagagcca	850
gcaaccagaa tgaaggggtgc acgaatcccc caccatctgg ctgaggtgga	900
ggctccagca actctggggg aggcagcggc tcacagtcgg gcagcagtg	950
cagtggcagc aatggtgaca acaacaatgg cagcagcagt ggtggcagca	1000
gcagtggcag cagcagtggc agcagcagtg gcggcagcag tggcggcagc	1050
agtgtgtgca gcagtggcaa cagtgtgtgc agcagaggtg acagcggcag	1100
tgagtcctcc tggggatcca gcaccggctc ctctccggc aaccacggtg	1150
ggagcggcgg aggaaatgga cataaaccgg ggtgtgaaaa gccagggaat	1200
gaagcccgcg ggagcgggga atctgggatt cagggtctca gaggacagg	1250
agtttccagc aacatgaggg aaataagcaa agagggcaat cgcttccttg	1300
gaggctctgg agacaattat cgggggcaag ggtcgagctg ggcagtgga	1350
ggaggtgacg ctgttggtgg agtcaatact gtgaactctg agacgtctcc	1400
tgggatgttt aactttgaca ctttctggaa gaattttaaa tccaagctgg	1450
gtttcatcaa ctgggatgcc ataaacaagg accagagaag ctctcgcatc	1500
ccgtgacctc cagacaagga gccaccagat tggatgggag cccccacact	1550
ccctccttaa aacaccaccc tctcatcact aatctcagcc cttgcccttg	1600
aaataaacct tagctgcccc acaaaaaaaaa aaaaaaaaaa aaaaaaaaaa	1650
aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa	1700
aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaa	1734

<210> SEQ ID NO 52
<211> LENGTH: 440
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 52

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

-continued

Pro Gly His Ser Gly Ala Trp Glu Thr Ser Gly Gly His Gly Ile
140 145 150

Phe Gly Ser Gln Gly Gly Leu Gly Gly Gln Gly Gln Gly Asn Pro
155 160 165

Gly Gly Leu Gly Thr Pro Trp Val His Gly Tyr Pro Gly Asn Ser
170 175 180

Ala Gly Ser Phe Gly Met Asn Pro Gln Gly Ala Pro Trp Gly Gln
185 190 195

Gly Gly Asn Gly Gly Pro Pro Asn Phe Gly Thr Asn Thr Gln Gly
200 205 210

Ala Val Ala Gln Pro Gly Tyr Gly Ser Val Arg Ala Ser Asn Gln
215 220 225

Asn Glu Gly Cys Thr Asn Pro Pro Pro Ser Gly Ser Gly Gly Gly
230 235 240

Ser Ser Asn Ser Gly Gly Gly Ser Gly Ser Gln Ser Gly Ser Ser
245 250 255

Gly Ser Gly Ser Asn Gly Asp Asn Asn Asn Gly Ser Ser Ser Gly
260 265 270

Gly Ser Ser Ser Gly Ser Ser Ser Gly Ser Ser Ser Gly Gly Ser
275 280 285

Ser Gly Gly Ser Ser Gly Gly Ser Ser Gly Asn Ser Gly Gly Ser
290 295 300

Arg Gly Asp Ser Gly Ser Glu Ser Ser Trp Gly Ser Ser Thr Gly
305 310 315

Ser Ser Ser Gly Asn His Gly Gly Ser Gly Gly Gly Asn Gly His
320 325 330

Lys Pro Gly Cys Glu Lys Pro Gly Asn Glu Ala Arg Gly Ser Gly
335 340 345

Glu Ser Gly Ile Gln Gly Phe Arg Gly Gln Gly Val Ser Ser Asn
350 355 360

Met Arg Glu Ile Ser Lys Glu Gly Asn Arg Leu Leu Gly Gly Ser
365 370 375

Gly Asp Asn Tyr Arg Gly Gln Gly Ser Ser Trp Gly Ser Gly Gly
380 385 390

Gly Asp Ala Val Gly Gly Val Asn Thr Val Asn Ser Glu Thr Ser
395 400 405

Pro Gly Met Phe Asn Phe Asp Thr Phe Trp Lys Asn Phe Lys Ser
410 415 420

Lys Leu Gly Phe Ile Asn Trp Asp Ala Ile Asn Lys Asp Gln Arg
425 430 435

Ser Ser Arg Ile Pro
440

<210> SEQ ID NO 53
<211> LENGTH: 1676
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 53

ggagaagagg ttgtgtggga caagctgctc cgcacagaag gatgtcgctg 50

ctgagcctgc cctggctggg cctcagaccg gtggcaatgt ccccatggct 100

-continued

actcctgctg ctggttggtg gctcctggct actcgccgc atcctggctt	150
ggacctatgc cttctataac aactgccgc ggctccagt tttccacag	200
ccccaaaaa ggaactggtt ttggggtcac ctgggcctga tcaactctac	250
agaggagggc ttgaaggact cgaccagat gtcggccacc tattcccagg	300
gctttacggt atggctgggt cccatcatcc ccttcacgt tttatgccac	350
cctgacacca tccggtctat caccaatgcc tcagctgcca ttgcacccaa	400
ggataatctc ttcacaggt tcctgaagcc ctggctggga gaagggatac	450
tgctgagtgg cggtgacaag tggagccgc accgtcggat gctgacgccc	500
gccttcatt tcaacatcct gaagtcctat ataacgatct tcaacaagag	550
tgcaaacatc atgcttgaca agtggcagca cctggcctca gagggcagca	600
gtcgtctgga catgtttgag cacatcagcc tcatgacctt ggacagtcta	650
cagaaatgca tcttcagctt tgacagccat tgtcaggaga ggcccagtga	700
atatattgcc accatcttgg agctcagtgc ccttgtagag aaaagaagcc	750
agcatatcct ccagcacatg gactttctgt attacctc ccatgacggg	800
cggcgcttcc acagggcctg ccgcctgtg catgacttca cagacgctgt	850
catccgggag cggcgctcga ccctccccc tcagggtatt gatgattttt	900
tcaaagacaa agccaagtcc aagactttgg atttcattga tgtgcttctg	950
ctgagcaagg atgaagatgg gaaggcattg tcagatgagg atataagagc	1000
agaggctgac accttcattg ttggaggcca tgacaccacg gccagtggcc	1050
tctcctgggt cctgtacaac cttgcgagc acccagaata ccaggagcgc	1100
tgccgacagg aggtgcaaga gcttctgaag gaccgcgatc ctaaagagat	1150
tgaatgggac gacctggccc agctgccctt cctgacctg tgcgtgaagg	1200
agagcctgag gttacatccc ccagctccct tcactctccg atgctgcacc	1250
caggacattg ttctcccaga tggccgagtc atcccaaag gcattacctg	1300
cctcatcgat attatagggg tccatcacia ccaactgtg tggccggatc	1350
ctgagggtcta cgacccttc cgctttgacc cagagaacag caaggggagg	1400
tcacctctgg cttttattcc ttctccgca gggcccagga actgcatcgg	1450
gcaggcgctc gccatggcgg agatgaaagt ggtcctggcg ttgatgctgc	1500
tgcacttccg gttcctgcca gaccacactg agccccgag gaagctggaa	1550
ttgatcatgc gcgccgagg cggtctttg ctgcgggtgg agccctgaa	1600
tgtaggcttg cagtgacttt ctgacccatc cacctgtttt tttgcagatt	1650
gtcatgaata aaacggtgct gtcaaa	1676

<210> SEQ ID NO 54

<211> LENGTH: 524

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 54

Met	Ser	Leu	Leu	Ser	Leu	Pro	Trp	Leu	Gly	Leu	Arg	Pro	Val	Ala
1				5					10				15	

Met Ser Pro Trp Leu Leu Leu Leu Val Val Gly Ser Trp Leu

-continued

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

-continued

Ile Val Leu Pro Asp Gly Arg Val Ile Pro Lys Gly Ile Thr Cys
410 415 420
Leu Ile Asp Ile Ile Gly Val His His Asn Pro Thr Val Trp Pro
425 430 435
Asp Pro Glu Val Tyr Asp Pro Phe Arg Phe Asp Pro Glu Asn Ser
440 445 450
Lys Gly Arg Ser Pro Leu Ala Phe Ile Pro Phe Ser Ala Gly Pro
455 460 465
Arg Asn Cys Ile Gly Gln Ala Phe Ala Met Ala Glu Met Lys Val
470 475 480
Val Leu Ala Leu Met Leu Leu His Phe Arg Phe Leu Pro Asp His
485 490 495
Thr Glu Pro Arg Arg Lys Leu Glu Leu Ile Met Arg Ala Glu Gly
500 505 510
Gly Leu Trp Leu Arg Val Glu Pro Leu Asn Val Gly Leu Gln
515 520

<210> SEQ ID NO 55
<211> LENGTH: 644
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 55

atcgcatcaa ttgggagtac catcttcctc atgggaccag tgaaacagct 50
gaagcgaatg tttgagccta ctcgtttgat tgcaactatc atggtgctgt 100
tgtgttttgc acttaccctg tgttctgcct tttggtggca taacaaggga 150
cttgacactta tcttctgcat tttgcagtct ttggcattga cgtggtacag 200
cctttccttc ataccatttg caagggatgc tgtgaagaag tgttttgccg 250
tgtgtcttgc ataattcatg gccagtttta tgaagctttg gaaggcacta 300
tggaacagaag ctggtggaca gttttgtaac tatcttcgaa acctctgtct 350
tacagacatg tgccttttat cttgcagcaa tgtgttgctt gtgattcgaa 400
catttgaggg ttacttttgg aagcaacaat acattctcga acctgaatgt 450
cagtagcaca ggatgagaag tgggttctgt atcttgtgga gtggaatctt 500
cctcatgtac ctgtttcctc tctggatgtt gtccactga attcccatga 550
atacaaacct attcagcaac agcaaaaaaa aaaaaaaaaa aaaaaaaaaa 600
aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaa 644

<210> SEQ ID NO 56
<211> LENGTH: 77
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 56

Met Gly Pro Val Lys Gln Leu Lys Arg Met Phe Glu Pro Thr Arg
1 5 10 15
Leu Ile Ala Thr Ile Met Val Leu Leu Cys Phe Ala Leu Thr Leu
20 25 30
Cys Ser Ala Phe Trp Trp His Asn Lys Gly Leu Ala Leu Ile Phe
35 40 45

-continued

Cys	Ile	Leu	Gln	Ser	Leu	Ala	Leu	Thr	Trp	Tyr	Ser	Leu	Ser	Phe
			50						55					60
Ile	Pro	Phe	Ala	Arg	Asp	Ala	Val	Lys	Lys	Cys	Phe	Ala	Val	Cys
			65						70					75

Leu Ala

<210> SEQ ID NO 57
<211> LENGTH: 3334
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 57

cggtctcgagc	tcgagccgaa	tcggctcgag	gggcagtgga	gcacccagca	50
ggccgccaac	atgctctgtc	tgtgcctgta	cgtgccggtc	atcggggaag	100
cccagaccga	gttccagtac	tttagtcga	aggggctccc	tgccgagctg	150
aagtccattt	tcaagctcag	tgtcttcac	ccctcccagg	aattctccac	200
ctaccgccag	tggaagcaga	aaattgtaca	agctggagat	aaggaccttg	250
atgggcagct	agactttgaa	gaatttgtcc	attatctcca	agatcatgag	300
aagaagctga	ggctggtgtt	taagattttg	gacaaaaaga	atgatggacg	350
cattgacgog	caggagatca	tcagtcacct	gcgggacttg	ggagtcaaga	400
tatctgaaca	gcaggcagaa	aaaattctca	agagcatgga	taaaaacggc	450
acgatgacca	tcgactggaa	cgagtggaga	gactaccacc	tcctccaccc	500
cgtggaaaac	atccccgaga	tcctctctca	ctggaagcat	tccacgatct	550
ttgatgtggg	tgagaatcta	acggtcccgg	atgagttcac	agtggaggag	600
aggcagacgg	ggatgtggtg	gagacacctg	gtggcaggag	gtggggcagg	650
ggccgtatcc	agaacctgca	cggccccctc	ggacaggctc	aagggtctca	700
tgccaggtcca	tgctctccgc	agcaacaaca	tgggcatcgt	tggtggcttc	750
actcagatga	ttcgagaagg	aggggccagg	tcactctggc	ggggcaatgg	800
catcaacgtc	ctcaaaattg	ccccgaatc	agccatcaaa	ttcatggcct	850
atgagcagat	caagcgccct	gttggtatgt	accaggagac	tctgaggatt	900
cacgagaggg	ttgtggcagg	gtccttgga	ggggccatcg	cccagagcag	950
catctaccca	atggaggtcc	tgaagacctg	gatggcgctg	cggaagacag	1000
gccagtactc	aggaatgctg	gactgcgcga	ggaggatcct	ggccagagag	1050
ggggtggccg	ccttctacaa	aggtatgtc	cccaacatgc	tgggcatcat	1100
cccctatgcc	ggcatcgacc	ttgcagtcta	cgagacgctc	aagaatgcct	1150
ggctgcagca	ctatgcagtg	aacagcgcg	accccgcggt	gtttgtgctc	1200
ctggcctgtg	gcaccatgtc	cagtacctgt	ggccagctgg	ccagctaccc	1250
cctggcccta	gtcaggaccc	ggatgcaggc	gcaagcctct	attgagggcg	1300
ctccggagggt	gaccatgagc	agcctcttca	aacatatact	gcggaccgag	1350
ggggccttcg	ggctgtacag	ggggctggcc	cccaacttca	tgaaggtcat	1400
cccagctgtg	agcatcagct	acgtggtcta	cgagaacctg	aagatcaccc	1450
tgggcgtgca	gtcgcggtga	cggggggagg	gccgcccggc	agtggactcg	1500

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ctgatcctgg gccgcagcct ggggtgtgca gccatctcat tctgtgaatg	1550
tgccaacact aagctgtctc gagccaagct gtgaaaacc tagacgcacc	1600
cgcagggagg gtggggagag ctggcaggcc cagggcttgt cctgctgacc	1650
ccagcagacc ctctgttgg ttccagcgaa gaccacaggc attccttagg	1700
gtccagggtc agcaggctcc gggctcacat gtgtaaggac aggacatttt	1750
ctgcagtgcc tgccaatagt gagcttgag cctggaggcc ggcttagttc	1800
ttccatttca cccttgacg cagctgttgg ccacggcccc tgccctctgg	1850
tctgccgtgc atctccctgt gccctcttgc tgccctgcctg tctgctgagg	1900
taaggtggga ggagggtac agcccacatc ccacccctc gtccaatccc	1950
ataatccatg atgaaagtg aggtcacgtg gcctccagg cctgaattcc	2000
caacctacag cattgacgcc aacttggtg tgaaggaga ggaaggatc	2050
tggccttgtg gtcaactggca tctgagccct gctgatggct ggggctctcg	2100
ggcatgcttg ggagtgcagg gggctcgggc tgccctggcct ggctgcacag	2150
aaggcaagtg ctggggctca tgggtgctcg agctggcctg gacctgtca	2200
ggatggggcc caccctagaa ccaaactcac tgtcccact gtggcatgag	2250
ggcagtggag caccatgttt gagggcgaa ggacagagcgt ttgtgtgttc	2300
tggggaggga aggaaaaggt gttggaggcc ttaattatgg actgttggga	2350
aaagggtttt gtccagaagg acaagccgga caaatgagcg acttctgtgc	2400
ttccagagga agacgaggga gcaggagctt ggctgactgc tcagagtctg	2450
ttctgacgcc ctgggggttc ctgtccaacc ccagcagggg cgcagcggga	2500
ccagcccac attccacttg tgtcactgct tggaacctat ttattttgta	2550
tttatttgaa cagagttagt tcctaactat ttttatagat ttgtttaatt	2600
aatagcttgt cattttcaag ttcatTTTT attcataatt atgttcatgg	2650
ttgattgtac cttcccaagc ccgcccagtg ggatgggagg aggaggagaa	2700
ggggggcctt gggccgctgc agtcacatct gtccagagaa attccttttg	2750
ggactggagg cagaaaagcg gccagaagcg agcagccctg gtccttttcc	2800
tttggcagggt tggggaaggg cttgccccca gccttaggat ttcagggttt	2850
gactgggggg gtggagagag agggagggaac ctcaataacc ttgaaggagg	2900
aatccagtta tttcctgcgc tgcgagggtt tctttatttc actcttttct	2950
gaatgtcaag gcagtgagggt gcctctcact gtgaatttgt ggtgggcggg	3000
ggctggagga gaggggggg ggctggctcc gtccctccca gccttctgct	3050
gcccttgctt aacaatgccg gccaaactggc gacctcacgg ttgcacttcc	3100
attccaccag aatgacctga tgaggaaatc ttcaatagga tgcaaagatc	3150
aatgcaaaaa ttgttatata tgaacatata actggagtcg tcaaaaagca	3200
aattaagaaa gaattggacg ttagaagttg tcatttaag cagccttcta	3250
ataaagtgtt ttcaaagctg aaaaaaaaa aaaaaaaaa aaaaaaaaa	3300
aaaaaaaaa aaaaaaaaa aaaaaaaaa aaaa	3334

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<211> LENGTH: 469

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 58

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

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	350		355		360
Lys Asn Ala Trp	Leu Gln His Tyr Ala	Val Asn Ser Ala Asp	Pro		
	365	370	375		
Gly Val Phe Val	Leu Leu Ala Cys Gly	Thr Met Ser Ser Thr	Cys		
	380	385	390		
Gly Gln Leu Ala	Ser Tyr Pro Leu Ala	Leu Val Arg Thr Arg	Met		
	395	400	405		
Gln Ala Gln Ala	Ser Ile Glu Gly Ala	Pro Glu Val Thr Met	Ser		
	410	415	420		
Ser Leu Phe Lys	His Ile Leu Arg Thr	Glu Gly Ala Phe Gly	Leu		
	425	430	435		
Tyr Arg Gly Leu	Ala Pro Asn Phe Met	Lys Val Ile Pro Ala	Val		
	440	445	450		
Ser Ile Ser Tyr	Val Val Tyr Glu Asn	Leu Lys Ile Thr Leu	Gly		
	455	460	465		
Val Gln Ser Arg					

<210> SEQ ID NO 59
<211> LENGTH: 1658
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 59

ggaaggcagc ggcagctcca ctcagccagt acccagatac gctgggaacc	50
ttccccagcc atggcttccc tggggcagat cctcttctgg agcataatta	100
gcatcatcat tattctggct ggagcaattg cactcatcat tggctttggt	150
atttcagggg gacactccat cacagtcaact actgtcgctt cagctgggaa	200
cattggggag gatggaatcc tgagctgcac ttttgaacct gacatcaaac	250
ttttctgatat cgtgatacaa tggctgaagg aaggtgtttt aggcttggtc	300
catgagtcca aagaaggcaa agatgagctg tcggagcagg atgaaatgtt	350
cagaggcccg acagcagtgt ttgctgatca agtgatagtt ggcaatgcct	400
ctttgcggct gaaaaacgtg caactcacag atgctggcac ctacaaatgt	450
tatatcatca cttctaaagg caagggaat gctaaccttg agtataaaac	500
tggagccttc agcatgccgg aagtgaatgt ggactataat gccagctcag	550
agaccttgcg gtgtgaggct ccccgatggt tccccagcc cacagtggtc	600
tgggcatccc aagttgacca gggagccaac ttctcggaag tctccaatac	650
cagcttttag ctgaactctg agaatgtgac catgaaggtt gtgtctgtgc	700
tctacaatgt tacgatcaac aacacatact cctgtatgat tgaaaatgac	750
attgccaaag caacagggga tatcaaagt acagaatcgg agatcaaaag	800
gcggagtca ctcagctgc taaactcaaa ggcttctctg tgtgtctctt	850
ctttctttgc catcagctgg gcacttctgc ctctcagccc ttacctgatg	900
ctaaaataat gtgccttggc cacaaaaaag catgcaaagt cattgttaca	950
acagggatct acagaactat ttcaccacca gatatgacct agttttatat	1000
ttctgggagg aatgaattc atatctagaa gtctggagtg agcaacaag	1050
agcaagaaac aaaaagaagc caaaagcaga aggctccaat atgaacaaga	1100

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taaattctatc ttcaaagaca tattagaagt tgggaaaata attcatgtga	1150
actagacaag tgtgttaaga gtgataagta aaatgcacgt ggagacaagt	1200
gcatccccag atctcagga cctccccctg cctgtcacct ggggagtga	1250
aggacaggat agtgcattgt ctttgtctct gaatttttag ttatatgtgc	1300
tgtaatgttg ctctgaggaa gcccctggaa agtctatccc aacatatcca	1350
catcttatat tccacaaatt aagctgtagt atgtacccta agacgtgct	1400
aattgactgc cacttcgcaa ctacagggcg gctgcatttt agtaatgggt	1450
caaagtattc actttttatg atgtctccaa aggtgccttg gcttctcttc	1500
ccaactgaca aatgccaaag ttgagaaaaa tgatcataat tttagcataa	1550
acagagcagt cggggacacc gatattataa ataaactgag caccttcttt	1600
ttaaacaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa	1650
aaaaaaaaa	1658

<210> SEQ ID NO 60

<211> LENGTH: 282

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 60

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

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215	220	225
Met Ile Glu Asn Asp Ile Ala Lys Ala Thr Gly Asp Ile Lys Val		
230	235	240
Thr Glu Ser Glu Ile Lys Arg Arg Ser His Leu Gln Leu Leu Asn		
245	250	255
Ser Lys Ala Ser Leu Cys Val Ser Ser Phe Phe Ala Ile Ser Trp		
260	265	270
Ala Leu Leu Pro Leu Ser Pro Tyr Leu Met Leu Lys		
275	280	

<210> SEQ ID NO 61
<211> LENGTH: 1617
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 61

tgacgtcaga atcaccatgg ccagctatcc ttaccggcag ggtgcccag	50
gagctgcagg acaagcacca ggagcccctc cgggtagcta ctaccctgga	100
cccccaata gtggagggca gtatggtagt gggctacccc ctggtggtgg	150
ttatgggggt cctgcccctg gagggcctta tggaccacca gctggtggag	200
ggccctatgg acacccaat cctgggatgt tcccctctgg aactccagga	250
ggaccatatg gcggtgcagc tcccgggggc ccctatggtc agccacctcc	300
aagttcctac ggtgcccagc agcctgggct ttatggacag ggtggcgccc	350
ctcccaatgt ggatcctgag gcctactcct ggttcagtc ggtggactca	400
gatcacagtg gctatatctc catgaaggag ctaaagcagg ccctgggtcaa	450
ctgcaattgg tcttcattca atgatgagac ctgcctcatg atgataaaca	500
tgtttgacaa gaccaagtca ggccgcctcg atgtctacgg cttctcagcc	550
ctgtggaaat tcatccagca gtggaagaac ctcttcagc agtatgaccg	600
ggaccgctcg ggctccatta gctacacaga gctgcagcaa gctctgtccc	650
aaatgggcta caacctgagc cccagttca cccagcttct ggtctcccgc	700
tactgccac gctctgccaa tcctgccatg cagcttgacc gcttcatcca	750
ggtgtgcacc cagctgcagg tgctgacaga ggccttccgg gagaaggaca	800
cagctgtaca aggcaacatc cggctcagct tcgaggactt cgtcaccatg	850
acagcttctc ggatgctatg acccaacat ctgtggagag tggagtgcac	900
cagggacctt tcctggcttc ttagagtgag agaagtatgt ggacatctct	950
tcttttctcg tccctctaga agaacattct cccttgcttg atgcaacact	1000
gttcacaaaag aggttgaga gtccctgcac atagccacca aatagtgagg	1050
accggggctg aggccacaca gataggggcc tgatggagga gaggatagaa	1100
gttgaatgct ctgatggcca tgagcagttg agtggcacag cctggcacca	1150
ggagcaggtc cttgtaatgg agttagtgtc cagtcagctg agctccaccc	1200
tgatgccagt ggtgagtgtt catcggcctg ttaccgttag tacctgtgtt	1250
ccctcaccag gccatcctgt caaacagacc catcttctcc aaagtggaat	1300
ctgaccaagc atgagagaga tctgtctatg ggaccagtgg cttggattct	1350

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gccacaccca taaatccttg tgtgttaact tctagctgcc tggggctggc      1400
cctgctcaga caaatctgct ccctgggcat ctttggccag gcttctgccc      1450
cctgcagctg ggacccctca ctgcctgcc atgctctgct cggttcagt      1500
ctccaggaga cagtggctac ctctccctgc caatactttt ttttaatttgc      1550
attttttttc atttggggcc aaaagtcag tgaaattgta agcttcaata      1600
aaaggatgaa actctga      1617

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<210> SEQ ID NO 62

<211> LENGTH: 284

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 62

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

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<210> SEQ ID NO 63
<211> LENGTH: 1234
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 63

caggatgcag ggcgcgtgg caggagctg cgctcctctg ggcctgctcc 50
tggtctgtct tcatctccca ggctctttg cccggagcat cgggtgtgtg 100
gaggagaaa tttcccaaaa cttcgggacc aacttgctc agctcggaca 150
accttctctc actggccct ctaactctga acatccgcag cccgctctgg 200
accctaggtc taatgacttg gcaagggttc ctctgaagct cagcgtgcct 250
ccatcagatg gcttcccacc tgcaggaggt tctgcagtgc agaggtggcc 300
tccatcgtgg gggctgcctg ccattggattc ctggcccctt gaggatcctt 350
ggcagatgat ggctgctgcg gctgaggacc gcctggggga agcgtgcct 400
gaagaactct cttacctctc cagtgtctcg gccctcgtc cgggcagtgg 450
ccctttgcct ggggagtctt ctcccgatgc cacaggcctc tcacctgagg 500
cttactcct ccaccaggac tcggagtcca gacgactgcc ccgttcta 550
tactgggag ccgggggaaa aatcctttcc caacgccctc cctggtctct 600
catccacagg gttctgcctg atcacccttg gggtaacctg aatccagt 650
tgtctctggg aggtggaggc cctgggactg gttggggaac gaggcccatg 700
ccacacctg agggaatctg gggatatcaat aatcaacccc cagggtaccag 750
ctggggaaa attaatcgg atccaggagg cagctgggga aatattaatc 800
ggtatccagg aggcagctgg gggaatatta atcggtatcc aggaggcagc 850
tgggggaata ttcatctata ccaggtatc aataacccat ttctcctctg 900
agttctccg cctcctggct cttcttgaa catccagct ggcttcctta 950
atcctccaag ccctagggtg cagtggggct agagcacgat agagggaaac 1000
ccaacattgg gagttagagt cctgctcccg ccccttgctg tgtgggctca 1050
atccaggccc tgtaaacatg tttccagcac tatccctact ttccagtgc 1100
tccctgctc atctccaata aaataaaagc acttatgaaa aaaaaaaaaa 1150
aaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa 1200
aaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaa 1234

<210> SEQ ID NO 64
<211> LENGTH: 325
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 64

Met Gln Gly Arg Val Ala Gly Ser Cys Ala Pro Leu Gly Leu Leu 15
1 5 10

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

Val Val Glu Glu Lys Val Ser Gln Asn Phe Gly Thr Asn Leu Pro 45
35 40 45

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Gln	Leu	Gly	Gln	Pro	Ser	Ser	Thr	Gly	Pro	Ser	Asn	Ser	Glu	His	
				50					55					60	
Pro	Gln	Pro	Ala	Leu	Asp	Pro	Arg	Ser	Asn	Asp	Leu	Ala	Arg	Val	
				65					70					75	
Pro	Leu	Lys	Leu	Ser	Val	Pro	Pro	Ser	Asp	Gly	Phe	Pro	Pro	Ala	
				80					85					90	
Gly	Gly	Ser	Ala	Val	Gln	Arg	Trp	Pro	Pro	Ser	Trp	Gly	Leu	Pro	
				95					100					105	
Ala	Met	Asp	Ser	Trp	Pro	Pro	Glu	Asp	Pro	Trp	Gln	Met	Met	Ala	
				110					115					120	
Ala	Ala	Ala	Glu	Asp	Arg	Leu	Gly	Glu	Ala	Leu	Pro	Glu	Glu	Leu	
				125					130					135	
Ser	Tyr	Leu	Ser	Ser	Ala	Ala	Ala	Leu	Ala	Pro	Gly	Ser	Gly	Pro	
				140					145					150	
Leu	Pro	Gly	Glu	Ser	Ser	Pro	Asp	Ala	Thr	Gly	Leu	Ser	Pro	Glu	
				155					160					165	
Ala	Ser	Leu	Leu	His	Gln	Asp	Ser	Glu	Ser	Arg	Arg	Leu	Pro	Arg	
				170					175					180	
Ser	Asn	Ser	Leu	Gly	Ala	Gly	Gly	Lys	Ile	Leu	Ser	Gln	Arg	Pro	
				185					190					195	
Pro	Trp	Ser	Leu	Ile	His	Arg	Val	Leu	Pro	Asp	His	Pro	Trp	Gly	
				200					205					210	
Thr	Leu	Asn	Pro	Ser	Val	Ser	Trp	Gly	Gly	Gly	Gly	Pro	Gly	Thr	
				215					220					225	
Gly	Trp	Gly	Thr	Arg	Pro	Met	Pro	His	Pro	Glu	Gly	Ile	Trp	Gly	
				230					235					240	
Ile	Asn	Asn	Gln	Pro	Pro	Gly	Thr	Ser	Trp	Gly	Asn	Ile	Asn	Arg	
				245					250					255	
Tyr	Pro	Gly	Gly	Ser	Trp	Gly	Asn	Ile	Asn	Arg	Tyr	Pro	Gly	Gly	
				260					265					270	
Ser	Trp	Gly	Asn	Ile	Asn	Arg	Tyr	Pro	Gly	Gly	Ser	Trp	Gly	Asn	
				275					280					285	
Ile	His	Leu	Tyr	Pro	Gly	Ile	Asn	Asn	Pro	Phe	Pro	Pro	Gly	Val	
				290					295					300	
Leu	Arg	Pro	Pro	Gly	Ser	Ser	Trp	Asn	Ile	Pro	Ala	Gly	Phe	Pro	
				305					310					315	
Asn	Pro	Pro	Ser	Pro	Arg	Leu	Gln	Trp	Gly						
				320					325						

<210> SEQ ID NO 65

<211> LENGTH: 422

<212> TYPE: DNA

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 65

aaggagagggc caccgggact tcagtgtctc ctccatccca ggagcgcagt	50
ggccactatg gggctctgggc tgccccttgt cctcctcttg accctccttg	100
gcagctcaca tggaacaggg cggggtatga ctttgcaact gaagctgaag	150
gagtccttttc tgacaaattc ctctctatgag tccagcttcc tggaattgct	200
tgaaaagctc tgcctctctc tccatctccc ttcagggacc agcgtcaccc	250
tccaccatgc aagatctcaa caccatgttg totgcaacac atgacagcca	300

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ttgaagcctg tgtccttctt ggcccgggct tttgggcccg ggatgcagga	350
ggcaggcccc gaccctgtct ttcagcaggc cccaccctc ctgagtggca	400
ataaataaaa ttcggtatgc tg	422

<210> SEQ ID NO 66
<211> LENGTH: 78
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 66

Met Gly Ser Gly Leu Pro Leu Val Leu Leu Leu Thr Leu Leu Gly	
1 5 10 15	
Ser Ser His Gly Thr Gly Pro Gly Met Thr Leu Gln Leu Lys Leu	
20 25 30	
Lys Glu Ser Phe Leu Thr Asn Ser Ser Tyr Glu Ser Ser Phe Leu	
35 40 45	
Glu Leu Leu Glu Lys Leu Cys Leu Leu Leu His Leu Pro Ser Gly	
50 55 60	
Thr Ser Val Thr Leu His His Ala Arg Ser Gln His His Val Val	
65 70 75	
Cys Asn Thr	

<210> SEQ ID NO 67
<211> LENGTH: 744
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 67

acggaccgag ggttcgaggg agggacacgg accaggaacc tgagctaggt	50
caaagacgcc cgggccaggt gccccgtcgc aggtgccctt ggcggagat	100
gcggtaggag gggcgagcgc gagaagcccc ttcctcggcg ctgccaaacc	150
gccaccacgc ccatggcgaa ccccgggctg gggctgcttc tggcgctggg	200
cctgccgttc ctgctggccc gctggggccg agcctggggg caaatacaga	250
ccacttctgc aaatgagaat agcactgttt tgccttcata caccagctcc	300
agctccgatg gcaacctgcg tccggaagcc atcactgcta tcatcgtggt	350
cttctccctc ttggctgcct tgctcctggc tgtggggctg gcaactgttg	400
tgcggaagct tcgggagaag cggcagacgg agggcaccta ccggcccagt	450
agcgaggagc agttctccca tgcagccgag gcccgggccc ctgaggactc	500
caaggagacg gtgcagggct gcctgccc ataggtcccc tctcctgcat	550
ctgtctccct tcattgtgtg gtgaccttgg ggaaaggcag tgccctctct	600
gggcagtcat atccaccacg tgcttaatat cagggaagaa ggtacttcaa	650
agactctgcc cctgaggtca agagaggatg gggctattca cttttatata	700
tttatataaa attagtagtg agatgtaaaa aaaaaaaaaa aaaa	744

<210> SEQ ID NO 68
<211> LENGTH: 123
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

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<400> SEQUENCE: 68

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

Leu Pro Ile

<210> SEQ ID NO 69
<211> LENGTH: 3265
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 69

gccaggaata actagagagg aacaatgggg ttattcagag gttttgtttt	50
cctcttagtt ctgtgcctgc tgcaccagtc aaatacttcc ttcattaagc	100
tgaataataa tggctttgaa gatattgtca ttgttataga tcctagtgtg	150
ccagaagatg aaaaaataat tgaacaaata gaggatatgg tgactacagc	200
ttctacgtac ctgtttgaag ccacagaaaa aagatTTTTT ttcaaaaatg	250
tatctatatt aattcctgag aattggaagg aaaatcctca gtacaaaagg	300
ccaaaacatg aaaaccataa acatgctgat gttatagttg caccacctac	350
actcccaggt agagatgaac catacaccaa gcagttcaca gaatgtggag	400
agaaaggcga atacattcac ttcacccctg accttctact tggaaaaaaa	450
caaaatgaat atggaccacc aggcaaaactg tttgtccatg agtgggctca	500
cctccggtgg ggagtgtttg atgagtacaa tgaagatcag cctttctacc	550
gtgctaagtc aaaaaaaatc gaagcaacaa ggtgttccgc aggtatctct	600
ggtagaaata gagtttataa gtgtcaagga ggcagctgtc ttagtagagc	650
atgcagaatt gattctacaa caaaactgta tggaaaagat tgtcaattct	700
ttcctgataa agtacaaaca gaaaaagcat ccataatggt tatgcaaagt	750
attgattctg ttgttgaatt ttgtaacgaa aaaaccata atcaagaagc	800
tccaagccta caaacataa agtgcaattt tagaagtaca tgggaggtga	850
ttagcaattc tgaggatttt aaaaacacca taccatggt gacaccacct	900
cctccacctg tcttctcatt gctgaagatc agtcaaagaa ttgtgtgctt	950
agttcttgat aagtctggaa gcatgggggg taaggaccgc ctaaatacgaa	1000

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tgaatcaagc agcaaaacat ttcctgctgc agactgttga aaatggatcc	1050
tgggtgggga tggttcactt tgatagtact gccactattg taaataagct	1100
aatccaaata aaaagcagtg atgaaagaaa cacactcatg gcaggattac	1150
ctacatatcc tctgggagga acttccatct gctctggaat taaatatgca	1200
tttcagggtga ttggagagct acattcccaa ctcgatggat ccgaagtact	1250
gctgctgact gatggggagg ataacactgc aagttcttgt attgatgaag	1300
tgaacaaaag tggggccatt gttcatttta ttgctttggg aagagctgct	1350
gatgaagcag taatagagat gagcaagata acaggaggaa gtcattttta	1400
tgtttcagat gaagctcaga acaatggcct cattgatgct tttggggctc	1450
ttacatcagg aaatactgat ctctcccaga agtcccttca gctcgaaagt	1500
aagggattaa cactgaatag taatgcctgg atgaacgaca ctgtcataat	1550
tgatagtaca gtgggaaagg acacgttctt tctcatcaca tggaacagtc	1600
tgccctccag tatttctctc tgggatccca gtggaacaat aatggaaaat	1650
ttcacagtgg atgcaacttc caaaatggcc tatctcagta ttccaggaac	1700
tgcaaaagtg ggcacttggg catacaatct tcaagccaaa gcgaaccag	1750
aaacattaac tattacagta acttctcgag cagcaaatc tctgtgcct	1800
ccaatcacag tgaatgctaa aatgaataag gacgtaaaca gtttcccag	1850
cccaatgatt gtttacgag aaattctaca aggatatgta cctgttcttg	1900
gagccaatgt gactgctttc attgaatcac agaatggaca tacagaagtt	1950
ttggaacttt tggataatgg tgcaggcgct gattctttca agaatgatgg	2000
agttactacc aggtatttta cagcatatac agaaaatggc agatatagct	2050
taaaagttog ggctcatgga ggagcaaaca ctgccaggct aaaattacgg	2100
cctccactga atagagccgc gtacatacca ggctgggtag tgaacgggga	2150
aattgaagca aaccgcgcaa gacctgaaat tgatgaggat actcagacca	2200
ccttgaggga tttcagccga acagcatccg gaggtgcatt tgtggtatca	2250
caagtcccaa gccttcctt gcctgaccaa taccaccaa gtcaaatcac	2300
agaccttgat gccacagttc atgaggataa gattattctt acatggacag	2350
caccaggaga taattttgat gttgaaaag ttcaacgtta tatcataaga	2400
ataagtgcaa gtattcttga tctaagagac agttttgatg atgctcttca	2450
agtaataact actgatctgt caccaaagga ggccaactcc aaggaaagct	2500
ttgcatttaa accagaaaat atctcagaag aaaatgcaac ccacatattt	2550
attgccatta aaagtataga taaaagcaat ttgacatcaa aagtatccaa	2600
cattgcacaa gtaactttgt ttatccctca agcaaatcct gatgacattg	2650
atcctacacc tactcctact cctactccta ctctgataa aagtcataat	2700
tctggagtta atattttctac gctggtattg tctgtgattg ggtctgtgt	2750
aattgttaac tttattttta gtaccacat ttgaacctta acgaagaaaa	2800
aaatcttcaa gtagacctag aagagagttt taaaaacaa aacaatgtaa	2850
gtaaaggata tttctgaatc ttaaaattca tccatgtgt gatcataaac	2900

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tcataaaaaat aattttaaga tgtcggaaaa ggatactttg attaaataaa	2950
aacactcatg gatatgtaaa aactgtcaag attaaaattt aatagtttca	3000
ttttattgtt attttatttg taagaaatag tgatgaacaa agatcctttt	3050
tcatactgat acctggttgt atattatttg atgcaacagt tttctgaaat	3100
gatatttcaa attgcatcaa gaaattaaaa tcattctatct gagtagtcaa	3150
aatacaagta aaggagagca aataacaac atttgaaaa aaaaaaaaaa	3200
aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa	3250
aaaaaaaaaa aaaaaa	3265

<210> SEQ ID NO 70

<211> LENGTH: 919

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 70

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

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260					265					270				
Ser	Thr	Trp	Glu	Val	Ile	Ser	Asn	Ser	Glu	Asp	Phe	Lys	Asn	Thr
				275						280				285
Ile	Pro	Met	Val	Thr	Pro	Pro	Pro	Pro	Pro	Val	Phe	Ser	Leu	Leu
				290						295				300
Lys	Ile	Ser	Gln	Arg	Ile	Val	Cys	Leu	Val	Leu	Asp	Lys	Ser	Gly
				305						310				315
Ser	Met	Gly	Gly	Lys	Asp	Arg	Leu	Asn	Arg	Met	Asn	Gln	Ala	Ala
				320						325				330
Lys	His	Phe	Leu	Leu	Gln	Thr	Val	Glu	Asn	Gly	Ser	Trp	Val	Gly
				335						340				345
Met	Val	His	Phe	Asp	Ser	Thr	Ala	Thr	Ile	Val	Asn	Lys	Leu	Ile
				350						355				360
Gln	Ile	Lys	Ser	Ser	Asp	Glu	Arg	Asn	Thr	Leu	Met	Ala	Gly	Leu
				365						370				375
Pro	Thr	Tyr	Pro	Leu	Gly	Gly	Thr	Ser	Ile	Cys	Ser	Gly	Ile	Lys
				380						385				390
Tyr	Ala	Phe	Gln	Val	Ile	Gly	Glu	Leu	His	Ser	Gln	Leu	Asp	Gly
				395						400				405
Ser	Glu	Val	Leu	Leu	Leu	Thr	Asp	Gly	Glu	Asp	Asn	Thr	Ala	Ser
				410						415				420
Ser	Cys	Ile	Asp	Glu	Val	Lys	Gln	Ser	Gly	Ala	Ile	Val	His	Phe
				425						430				435
Ile	Ala	Leu	Gly	Arg	Ala	Ala	Asp	Glu	Ala	Val	Ile	Glu	Met	Ser
				440						445				450
Lys	Ile	Thr	Gly	Gly	Ser	His	Phe	Tyr	Val	Ser	Asp	Glu	Ala	Gln
				455						460				465
Asn	Asn	Gly	Leu	Ile	Asp	Ala	Phe	Gly	Ala	Leu	Thr	Ser	Gly	Asn
				470						475				480
Thr	Asp	Leu	Ser	Gln	Lys	Ser	Leu	Gln	Leu	Glu	Ser	Lys	Gly	Leu
				485						490				495
Thr	Leu	Asn	Ser	Asn	Ala	Trp	Met	Asn	Asp	Thr	Val	Ile	Ile	Asp
				500						505				510
Ser	Thr	Val	Gly	Lys	Asp	Thr	Phe	Phe	Leu	Ile	Thr	Trp	Asn	Ser
				515						520				525
Leu	Pro	Pro	Ser	Ile	Ser	Leu	Trp	Asp	Pro	Ser	Gly	Thr	Ile	Met
				530						535				540
Glu	Asn	Phe	Thr	Val	Asp	Ala	Thr	Ser	Lys	Met	Ala	Tyr	Leu	Ser
				545						550				555
Ile	Pro	Gly	Thr	Ala	Lys	Val	Gly	Thr	Trp	Ala	Tyr	Asn	Leu	Gln
				560						565				570
Ala	Lys	Ala	Asn	Pro	Glu	Thr	Leu	Thr	Ile	Thr	Val	Thr	Ser	Arg
				575						580				585
Ala	Ala	Asn	Ser	Ser	Val	Pro	Pro	Ile	Thr	Val	Asn	Ala	Lys	Met
				590						595				600
Asn	Lys	Asp	Val	Asn	Ser	Phe	Pro	Ser	Pro	Met	Ile	Val	Tyr	Ala
				605						610				615
Glu	Ile	Leu	Gln	Gly	Tyr	Val	Pro	Val	Leu	Gly	Ala	Asn	Val	Thr
				620						625				630
Ala	Phe	Ile	Glu	Ser	Gln	Asn	Gly	His	Thr	Glu	Val	Leu	Glu	Leu
				635						640				645

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Leu Asp Asn Gly Ala Gly Ala Asp Ser Phe Lys Asn Asp Gly Val
650 655 660

Tyr Ser Arg Tyr Phe Thr Ala Tyr Thr Glu Asn Gly Arg Tyr Ser
665 670 675

Leu Lys Val Arg Ala His Gly Gly Ala Asn Thr Ala Arg Leu Lys
680 685 690

Leu Arg Pro Pro Leu Asn Arg Ala Ala Tyr Ile Pro Gly Trp Val
695 700 705

Val Asn Gly Glu Ile Glu Ala Asn Pro Pro Arg Pro Glu Ile Asp
710 715 720

Glu Asp Thr Gln Thr Thr Leu Glu Asp Phe Ser Arg Thr Ala Ser
725 730 735

Gly Gly Ala Phe Val Val Ser Gln Val Pro Ser Leu Pro Leu Pro
740 745 750

Asp Gln Tyr Pro Pro Ser Gln Ile Thr Asp Leu Asp Ala Thr Val
755 760 765

His Glu Asp Lys Ile Ile Leu Thr Trp Thr Ala Pro Gly Asp Asn
770 775 780

Phe Asp Val Gly Lys Val Gln Arg Tyr Ile Ile Arg Ile Ser Ala
785 790 795

Ser Ile Leu Asp Leu Arg Asp Ser Phe Asp Asp Ala Leu Gln Val
800 805 810

Asn Thr Thr Asp Leu Ser Pro Lys Glu Ala Asn Ser Lys Glu Ser
815 820 825

Phe Ala Phe Lys Pro Glu Asn Ile Ser Glu Glu Asn Ala Thr His
830 835 840

Ile Phe Ile Ala Ile Lys Ser Ile Asp Lys Ser Asn Leu Thr Ser
845 850 855

Lys Val Ser Asn Ile Ala Gln Val Thr Leu Phe Ile Pro Gln Ala
860 865 870

Asn Pro Asp Asp Ile Asp Pro Thr Pro Thr Pro Thr Pro Thr Pro
875 880 885

Thr Pro Asp Lys Ser His Asn Ser Gly Val Asn Ile Ser Thr Leu
890 895 900

Val Leu Ser Val Ile Gly Ser Val Val Ile Val Asn Phe Ile Leu
905 910 915

Ser Thr Thr Ile

<210> SEQ ID NO 71
<211> LENGTH: 3877
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 71

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aagaccatac gtccccgggc aggggtgaca acaggtgtca tctttttgat	100
ctcgtgtgtg gctgccttcc tatttcaagg aaagacgcca aggtaatttt	150
gaccagagg agcaatgatg tagccacctc ctaaccttcc cttcttgaac	200
ccccagttat gccaggattt actagagagt gtcaactcaa ccagcaagcg	250
gtcctctcgg cttaacttgt ggttgaggga gagaaccttt gtggggctgc	300

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gttctcttag cagtgtctag aagtgacttg cctgagggtg gaccagaaga	350
aaggaaaggt cccctcttgc tgttggtctg acatcaggaa ggctgtgatg	400
ggaatgaagg tgaaaacttg gagatttcac ttcagtcatt gcttctgcct	450
gcaagatcat cctttaaaag tagagaagct gctctgtgtg gtggttaact	500
ccaagaggca gaactcgttc tagaaggaaa tggatgcaag cagctccggg	550
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ggggcccggt ctttgaggga tgccaccggt tctggacgca tggctgattc	650
ctgaatgatg atggttcgcc gggggctgct tgcgtggatt tcccgggtgg	700
tggttttgct ggtgctcctc tgctgtgcta tctctgtcct gtacatgttg	750
gcctgcaccc caaaagtga caggagcag ctggcactgc ccagggccaa	800
cagccccag gggaaggagg ggtaccagc cgtccttcag gagtgggagg	850
agcagcaccg caactacgtg agcagcctga agcggcagat cgcacagctc	900
aaggaggagc tgcaggagag gagttagcag ctcaggaatg ggcagtacca	950
agccagcgat gctgctggcc tgggtctgga caggagcccc ccagagaaaa	1000
cccaggccga cctcctggcc ttcctgcact cgcagggtga caaggcagag	1050
gtgaatgctg gcgtcaagct ggccacagag tatgcagcag tgcctttcga	1100
tagctttact ctacagaagg tgtaccagct ggagactggc cttaccggcc	1150
accccagga gaagcctgtg aggaaggaca agcgggatga gttggtgaa	1200
gccattgaat cagccttgga gaccctgaac aatcctgcag agaacagccc	1250
caatcacggt ccttacacgg cctctgattt catagaaggg atctaccgaa	1300
cagaaaggga caaagggaca ttgtatgagc tcaccttcaa aggggaccac	1350
aaacacgaat tcaaacggct catcttattt cgaccattca gccccatcat	1400
gaaagtgaat aatgaaaagc tcaacatggc caacacgctt atcaatgtta	1450
tcgtgcctct agcaaaaagg gtggacaagt tccggcagtt catgcagaat	1500
ttcagggaga tgtgcattga gcaggatggg agagtccatc tctactgtgt	1550
ttactttggg aaagaagaaa taaatgaagt caaaggaata cttgaaaaa	1600
cttccaaagc tgccaacttc aggaacttta ccttcattca gctgaatgga	1650
gaattttctc ggggaaaggg acttgatgtt ggagcccgtc tctggaaggg	1700
aagcaacgtc cttctctttt tctgtgatgt ggacatctac ttcacatctg	1750
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tatccagttc ttttcagtca gtacaatcct ggcataatat acggccacca	1850
tgatgcagtc cctcccttg aacagcagct ggtcataaag aaggaaactg	1900
gattttggag agactttgga tttgggatga cgtgtcagta tcggtcagac	1950
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ggacgcctgt gcgaggactc ttccacctct ggcatgagaa gcgctgcatg	2100
gacgagctga ccccgagca gtacaagatg tgcatgcagt ccaaggccat	2150
gaacgaggca tcccacggcc agctgggcat gctggtgttc aggcacgaga	2200

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tagaggctca ccttcgcaaa cagaaacaga agacaagtag caaaaaaaca	2250
tgaactccca gagaaggatt gtgggagaca ctttttcttt ccttttgcaa	2300
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agaattggac tgatgggtca gagatgagaa agcctccgat ttctctctgt	2400
tgggcttttt acaacagaaa tcaaaatctc cgctttgcct gcaaaagtaa	2450
cccagttgca ccctgtgaag tgtctgacaa aggcagaatg cttgtgagat	2500
tataagccta atggtgtgga ggttttgatg gtgtttacaa tacactgaga	2550
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gtaaaaaatt cattagcatg aaaggcaagc atatttctcc tcatatgaat	2650
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taaaatggac cagaaaagaa aagaaacat aaatatcgtg tcatattttc	2800
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acaagttggc ctacattttt atatttttta agaagatact ttgagatgca	3000
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gggaaggaat ggtttgtact aatacagacg tacagatact ttctctgaag	3150
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cttacctggt taaataaacc aaagtatacc gtgtgaacca aacaatctct	3350
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gagaaaaata tatatatata tatatatatt gtgaaagatc aatccatctg	3450
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cagggtgaag taactgaatt attttttaaa ttaagcagtt ctactcaatc	3550
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gctgttggtg tgtaaaaaa tgcattgtat tgatttgtac tggtagttta	3800
tgaaatttaa ttaaaacaca ggccatgaat ggaagggtgg attgcacagc	3850
taataaaaata tgatttggtg atatgaa	3877

<210> SEQ ID NO 72

<211> LENGTH: 532

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 72

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Met	Met	Met	Val	Arg	Arg	Gly	Leu	Leu	Ala	Trp	Ile	Ser	Arg	Val	1	5	10	15
Val	Val	Leu	Leu	Val	Leu	Leu	Cys	Cys	Ala	Ile	Ser	Val	Leu	Tyr	20	25	30	
Met	Leu	Ala	Cys	Thr	Pro	Lys	Gly	Asp	Glu	Glu	Gln	Leu	Ala	Leu	35	40	45	
Pro	Arg	Ala	Asn	Ser	Pro	Thr	Gly	Lys	Glu	Gly	Tyr	Gln	Ala	Val	50	55	60	
Leu	Gln	Glu	Trp	Glu	Glu	Gln	His	Arg	Asn	Tyr	Val	Ser	Ser	Leu	65	70	75	
Lys	Arg	Gln	Ile	Ala	Gln	Leu	Lys	Glu	Glu	Leu	Gln	Glu	Arg	Ser	80	85	90	
Glu	Gln	Leu	Arg	Asn	Gly	Gln	Tyr	Gln	Ala	Ser	Asp	Ala	Ala	Gly	95	100	105	
Leu	Gly	Leu	Asp	Arg	Ser	Pro	Pro	Glu	Lys	Thr	Gln	Ala	Asp	Leu	110	115	120	
Leu	Ala	Phe	Leu	His	Ser	Gln	Val	Asp	Lys	Ala	Glu	Val	Asn	Ala	125	130	135	
Gly	Val	Lys	Leu	Ala	Thr	Glu	Tyr	Ala	Ala	Val	Pro	Phe	Asp	Ser	140	145	150	
Phe	Thr	Leu	Gln	Lys	Val	Tyr	Gln	Leu	Glu	Thr	Gly	Leu	Thr	Arg	155	160	165	
His	Pro	Glu	Glu	Lys	Pro	Val	Arg	Lys	Asp	Lys	Arg	Asp	Glu	Leu	170	175	180	
Val	Glu	Ala	Ile	Glu	Ser	Ala	Leu	Glu	Thr	Leu	Asn	Asn	Pro	Ala	185	190	195	
Glu	Asn	Ser	Pro	Asn	His	Arg	Pro	Tyr	Thr	Ala	Ser	Asp	Phe	Ile	200	205	210	
Glu	Gly	Ile	Tyr	Arg	Thr	Glu	Arg	Asp	Lys	Gly	Thr	Leu	Tyr	Glu	215	220	225	
Leu	Thr	Phe	Lys	Gly	Asp	His	Lys	His	Glu	Phe	Lys	Arg	Leu	Ile	230	235	240	
Leu	Phe	Arg	Pro	Phe	Ser	Pro	Ile	Met	Lys	Val	Lys	Asn	Glu	Lys	245	250	255	
Leu	Asn	Met	Ala	Asn	Thr	Leu	Ile	Asn	Val	Ile	Val	Pro	Leu	Ala	260	265	270	
Lys	Arg	Val	Asp	Lys	Phe	Arg	Gln	Phe	Met	Gln	Asn	Phe	Arg	Glu	275	280	285	
Met	Cys	Ile	Glu	Gln	Asp	Gly	Arg	Val	His	Leu	Thr	Val	Val	Tyr	290	295	300	
Phe	Gly	Lys	Glu	Glu	Ile	Asn	Glu	Val	Lys	Gly	Ile	Leu	Glu	Asn	305	310	315	
Thr	Ser	Lys	Ala	Ala	Asn	Phe	Arg	Asn	Phe	Thr	Phe	Ile	Gln	Leu	320	325	330	
Asn	Gly	Glu	Phe	Ser	Arg	Gly	Lys	Gly	Leu	Asp	Val	Gly	Ala	Arg	335	340	345	
Phe	Trp	Lys	Gly	Ser	Asn	Val	Leu	Leu	Phe	Phe	Cys	Asp	Val	Asp	350	355	360	
Ile	Tyr	Phe	Thr	Ser	Glu	Phe	Leu	Asn	Thr	Cys	Arg	Leu	Asn	Thr	365	370	375	

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Gln	Pro	Gly	Lys	Lys	Val	Phe	Tyr	Pro	Val	Leu	Phe	Ser	Gln	Tyr
			380						385				390	
Asn	Pro	Gly	Ile	Ile	Tyr	Gly	His	His	Asp	Ala	Val	Pro	Pro	Leu
			395						400				405	
Glu	Gln	Gln	Leu	Val	Ile	Lys	Lys	Glu	Thr	Gly	Phe	Trp	Arg	Asp
			410						415				420	
Phe	Gly	Phe	Gly	Met	Thr	Cys	Gln	Tyr	Arg	Ser	Asp	Phe	Ile	Asn
			425						430				435	
Ile	Gly	Gly	Phe	Asp	Leu	Asp	Ile	Lys	Gly	Trp	Gly	Gly	Glu	Asp
			440						445				450	
Val	His	Leu	Tyr	Arg	Lys	Tyr	Leu	His	Ser	Asn	Leu	Ile	Val	Val
			455						460				465	
Arg	Thr	Pro	Val	Arg	Gly	Leu	Phe	His	Leu	Trp	His	Glu	Lys	Arg
			470						475				480	
Cys	Met	Asp	Glu	Leu	Thr	Pro	Glu	Gln	Tyr	Lys	Met	Cys	Met	Gln
			485						490				495	
Ser	Lys	Ala	Met	Asn	Glu	Ala	Ser	His	Gly	Gln	Leu	Gly	Met	Leu
			500						505				510	
Val	Phe	Arg	His	Glu	Ile	Glu	Ala	His	Leu	Arg	Lys	Gln	Lys	Gln
			515						520				525	
Lys	Thr	Ser	Ser	Lys	Lys	Thr								
			530											

<210> SEQ ID NO 73
<211> LENGTH: 1701
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien
<220> FEATURE:
<221> NAME/KEY: unsure
<222> LOCATION: 1528
<223> OTHER INFORMATION: unknown base

<400> SEQUENCE: 73	
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tggaagccca cagagacaga gacagcaaga gaagcagaga taaatacact	150
cacgccagga gctcgctcgc tctctctctc tctctctcac tcctccctcc	200
ctctctctct gcctgtccta gtccctctagt cctcaaattc ccagtcccct	250
gcaccccttc ctgggacact atgttgttct ccgccctcct gctggagggtg	300
atttggatcc tggctgcaga tgggggtcaa cactggacgt atgaggggcc	350
acatggtcag gaccattggc cagcctctta ccctgagtgt ggaacaatg	400
cccagtcgcc catcgatatt cagacagaca gtgtgacatt tgaccctgat	450
ttgcctgctc tgcagcccca cggatatgac cagcctggca ccgagccttt	500
ggacctgcac aacaatggcc acacagtgca actctctctg ccctctaccc	550
tgtatctggg tggacttccc cgaaaatatg tagctgccca gctccacctg	600
cactgggggc agaaaggatc ccagggggg tcagaacacc agatcaacag	650
tgaagccaca tttgcagagc tccacattgt acattatgac tctgattcct	700
atgacagctt gagtgaggct gctgagaggc ctcaaggcct ggctgtcctg	750
ggcatcctaa ttgaggtggg tgagactaag aatatagctt atgaacacat	800

-continued

tctgagtcac ttgcatgaag tcaggcataa agatcagaag acctcagtgc 850
ctcccttcaa cctaagagag ctgctcccca aacagctggg gcagtacttc 900
cgctacaatg gctcgctcac aactccccct tgctaccaga gtgtgctctg 950
gacagttttt tatagaaggt cccagatttc aatggaacag ctggaaaagc 1000
ttcaggggac attgttctcc acagaagagg agccctctaa gcttctggta 1050
cagaactacc gagcccttca gcctctcaat cagcgcatgg tctttgcttc 1100
tttcatccaa gcaggatcct cgtataccac aggtgaaatg ctgagtctag 1150
gtgtaggaat cttggttggc tgtctctgcc ttctcctggc tgtttatttc 1200
attgctagaa agattcggaa gaagaggctg gaaaaccgaa agagtgtggt 1250
cttcacctca gcacaagcca cgactgaggc ataaattcct tctcagatac 1300
catggatgtg gatgacttcc cttcatgcct atcaggaagc ctctaaaatg 1350
gggtgtagga tctggccaga aacactgtag gagtagtaag cagatgtcct 1400
ccttccccctg gacatctctt agagaggaat ggaccaggc tgtcattcca 1450
ggaagaactg cagagccttc agcctctcca aacatgtagg aggaaatgag 1500
gaaatcgctg tgttgtaaat gcagaganca aactctgttt agttgcaggg 1550
gaagtttggg atatacccca aagtcctcta cccctcact tttatggccc 1600
tttccttaga tatactgcgg gatctctcct taggataaag agttgctgtt 1650
gaagttgtat atttttgatc aatatatttg gaaattaaag tttctgactt 1700
t 1701

<210> SEQ ID NO 74

<211> LENGTH: 337

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 74

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

-continued

Ala Glu Arg Pro Gln Gly Leu Ala Val Leu Gly Ile Leu Ile Glu
155 160 165
Val Gly Glu Thr Lys Asn Ile Ala Tyr Glu His Ile Leu Ser His
170 175 180
Leu His Glu Val Arg His Lys Asp Gln Lys Thr Ser Val Pro Pro
185 190 195
Phe Asn Leu Arg Glu Leu Leu Pro Lys Gln Leu Gly Gln Tyr Phe
200 205 210
Arg Tyr Asn Gly Ser Leu Thr Thr Pro Pro Cys Tyr Gln Ser Val
215 220 225
Leu Trp Thr Val Phe Tyr Arg Arg Ser Gln Ile Ser Met Glu Gln
230 235 240
Leu Glu Lys Leu Gln Gly Thr Leu Phe Ser Thr Glu Glu Glu Pro
245 250 255
Ser Lys Leu Leu Val Gln Asn Tyr Arg Ala Leu Gln Pro Leu Asn
260 265 270
Gln Arg Met Val Phe Ala Ser Phe Ile Gln Ala Gly Ser Ser Tyr
275 280 285
Thr Thr Gly Glu Met Leu Ser Leu Gly Val Gly Ile Leu Val Gly
290 295 300
Cys Leu Cys Leu Leu Leu Ala Val Tyr Phe Ile Ala Arg Lys Ile
305 310 315
Arg Lys Lys Arg Leu Glu Asn Arg Lys Ser Val Val Phe Thr Ser
320 325 330
Ala Gln Ala Thr Thr Glu Ala
335

<210> SEQ ID NO 75
<211> LENGTH: 1743
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 75

tgccgctgcc gccgctgctg ctgttgctcc tggcgggcgcc ttggggacgg 50
gcagttccct gtgtctctgg tggtttgctt aaacctgcaa acatcacctt 100
cttatccatc aacatgaaga atgtcctaca atggactcca ccagagggtc 150
ttcaaaggagt taaagttact tacactgtgc agtatttcat cacaaattgg 200
cccaccagag gtggcactga ctacagatga gaagtccatt tctgttgtcc 250
tgacagctcc agagaagtgg aagagaaatc cagaagacct tcctgtttcc 300
atgcaacaaa tatactcaa tctgaagtat aacgtgtctg tgttgaatac 350
taaatacaaa agaacgtggt cccagtgtgt gaccaaccac acgctgggtg 400
tcacctggct ggagccgaac actctttact gcgtacacgt ggagtccttc 450
gtcccagggg ccctcgcgg tgctcagcct tctgagaagc agtgtgccag 500
gactttgaaa gatcaatcat cagagttcaa ggctaaaatc atcttctggt 550
atgttttgcc catatctatt accgtgtttc ttttttctgt gatgggctat 600
tccatctacc gatatatcca cgttggcata gagaaacacc cagcaaattt 650
gattttgatt tatggaaatg aatttgacaa aagattcttt gtgcctgctg 700

-continued

aaaaaatcgt gattaacttt atcaccctca atatctcgga tgattctaaa	750
atttctcatc aggatatgag ttacttgga aaaagcagtg atgtatccag	800
ccttaatgat cctcagccca gcgggaacct gagccccct caggaggaag	850
aggaggtgaa acatttaggg tatgcttcgc atttgatgga aattttttgt	900
gactctgaag aaaacacgga aggtacttct ctcaccagc aagagtcct	950
cagcagaaca atacccccg ataaaacagt cattgaatat gaatatgat	1000
tcagaaccac tgacatttgt gcggggcctg aagagcagga gctcagttg	1050
caggaggagg tgtccacaca aggaacatta ttggagtcgc aggcagcgtt	1100
ggcagtcctt ggcccgcaa cgttacagta ctcatacacc cctcagctcc	1150
aagacttaga cccctggcg caggagcaca cagactcgga ggaggggccc	1200
gaggaagagc catcgacgac cctggtcgac tgggatcccc aaactggcag	1250
gctgtgtatt ccttcgctgt ccagcttcga ccaggattca gagggctgcg	1300
agccttctga gggggatggg ctgggagagg agggcttct atctagactc	1350
tatgaggagc cggtccaga caggccacca ggagaaaatg aaacctatct	1400
catgcaattc atggaggaat gggggttata tgtgcagatg gaaaactgat	1450
gccaacactt ccttttgctt ttgttttcct gtgcaacaa gtgagtcacc	1500
cctttgatcc cagccataaa gtacctggga tgaaagaagt tttttccagt	1550
ttgtcagtg ctgtgagaat tacttatttc ttttctctat tctcatagca	1600
cgtgtgtgat tggttcatgc atgtaggctt ctaacaatg atgggtggcc	1650
tctggagtcc aggggctggc cggttgttct atgcagagaa agcagtcaat	1700
aaatgtttgc cagactgggt gcagaattta ttcaggtggg tgt	1743

<210> SEQ ID NO 76

<211> LENGTH: 442

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 76

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

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Phe Lys Ala Lys Ile Ile Phe Trp Tyr Val Leu Pro Ile Ser Ile
140 145 150
Thr Val Phe Leu Phe Ser Val Met Gly Tyr Ser Ile Tyr Arg Tyr
155 160 165
Ile His Val Gly Lys Glu Lys His Pro Ala Asn Leu Ile Leu Ile
170 175 180
Tyr Gly Asn Glu Phe Asp Lys Arg Phe Phe Val Pro Ala Glu Lys
185 190 195
Ile Val Ile Asn Phe Ile Thr Leu Asn Ile Ser Asp Asp Ser Lys
200 205 210
Ile Ser His Gln Asp Met Ser Leu Leu Gly Lys Ser Ser Asp Val
215 220 225
Ser Ser Leu Asn Asp Pro Gln Pro Ser Gly Asn Leu Arg Pro Pro
230 235 240
Gln Glu Glu Glu Glu Val Lys His Leu Gly Tyr Ala Ser His Leu
245 250 255
Met Glu Ile Phe Cys Asp Ser Glu Glu Asn Thr Glu Gly Thr Ser
260 265 270
Leu Thr Gln Gln Glu Ser Leu Ser Arg Thr Ile Pro Pro Asp Lys
275 280 285
Thr Val Ile Glu Tyr Glu Tyr Asp Val Arg Thr Thr Asp Ile Cys
290 295 300
Ala Gly Pro Glu Glu Gln Glu Leu Ser Leu Gln Glu Glu Val Ser
305 310 315
Thr Gln Gly Thr Leu Leu Glu Ser Gln Ala Ala Leu Ala Val Leu
320 325 330
Gly Pro Gln Thr Leu Gln Tyr Ser Tyr Thr Pro Gln Leu Gln Asp
335 340 345
Leu Asp Pro Leu Ala Gln Glu His Thr Asp Ser Glu Glu Gly Pro
350 355 360
Glu Glu Glu Pro Ser Thr Thr Leu Val Asp Trp Asp Pro Gln Thr
365 370 375
Gly Arg Leu Cys Ile Pro Ser Leu Ser Ser Phe Asp Gln Asp Ser
380 385 390
Glu Gly Cys Glu Pro Ser Glu Gly Asp Gly Leu Gly Glu Glu Gly
395 400 405
Leu Leu Ser Arg Leu Tyr Glu Glu Pro Ala Pro Asp Arg Pro Pro
410 415 420
Gly Glu Asn Glu Thr Tyr Leu Met Gln Phe Met Glu Glu Trp Gly
425 430 435
Leu Tyr Val Gln Met Glu Asn
440

<210> SEQ ID NO 77
<211> LENGTH: 1636
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 77

gaggagcggg ccgaggactc cagcgtgccc aggtctggca tcctgcactt 50
gctgccctct gacacctggg aagatggcgg gcccgtagac cttcacccctt 100

-continued

ctctgtggtt tgctggcagc caccttgatc caagccaccc tcagtcccac	150
tgcagtttctc atcctcggcc caaaagtcac caaagaaaag ctgacacagg	200
agctgaagga ccacaacgcc accagcatcc tgcagcagct gccgctgctc	250
agtgccatgc gggaaaagcc agccggaggc atccctgtgc tgggcagcct	300
ggtgaacacc gtcctgaagc acatcatctg gctgaaggtc atcacagcta	350
acatcctcca gctgcagggtg aagccctcgg ccaatgacca ggagctgcta	400
gtcaagatcc ccctggacat ggtggctgga ttcaacacgc ccctggtcaa	450
gaccatcgtg gagtccaca tgacgactga ggcccaagcc accatccgca	500
tggacaccag tgcaagtggc cccacccgcc tggtcctcag tgactgtgcc	550
accagccatg ggagcctgcg catccaactg ctgtataagc tctccttcct	600
ggtgaacgcc ttagctaagc aggtcatgaa cctcctagtg ccatccctgc	650
ccaatctagt gaaaaaccag ctgtgtcccg tgatcgaggc ttccttcaat	700
ggcatgtatg cagacctcct gcagctgggtg aaggtgccca ttccctcag	750
cattgaccgt ctggagtttg accttctgta tcctgccatc aagggtgaca	800
ccattcagct ctacctgggg gccaaattgt tggactcaca gggaaagggtg	850
accaagtggg tcaataactc tgcagcttcc ctgacaatgc ccaccctgga	900
caacatcccg ttcagcctca tcgtgagtca ggacgtgggtg aaagctgcag	950
tggctgctgt gctctctcca gaagaattca tggtcctggt ggactctgtg	1000
cttcctgaga gtgcccatcg gctgaagtca agcatcgggc tgatcaatga	1050
aaaggctgca gataagctgg gatctacca gatcgtgaag atcctaactc	1100
aggacactcc cgagtttttt atagaccaag gccatgcaa ggtggcccaa	1150
ctgatcgtgc tggaagtgtt tcctccaggt gaagccctcc gccotttgtt	1200
caccttgggc atcgaagcca gctcggaagc tcagttttac accaaagggtg	1250
accaacttat actcaacttg aataacatca gctctgatcg gatccagctg	1300
atgaactctg ggattggctg gttccaacct gatgttctga aaaacatcat	1350
cactgagatc atccactcca tcctgctgcc gaaccagaat ggcaaattaa	1400
gatctggggg cccagtgtca ttggtgaagg ccttgggatt cgaggcagct	1450
gagtcctcac tgaccaagga tgccttgtg cttactccag cctccttgtg	1500
gaaaccacgc tctcctgtct cccagtgaag acttggatgg cagccatcag	1550
ggaaggctgg gtcccagctg ggagtatggg tgtgagctct atagaccatc	1600
cctctctgca atcaataaac acttgccctgt gaaaaa	1636

<210> SEQ ID NO 78
<211> LENGTH: 484
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 78

Met Ala Gly Pro Trp Thr Phe Thr Leu Leu Cys Gly Leu Leu Ala
1 5 10 15

Ala Thr Leu Ile Gln Ala Thr Leu Ser Pro Thr Ala Val Leu Ile
20 25 30

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

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410					415					420				
Val	Leu	Lys	Asn	Ile	Ile	Thr	Glu	Ile	Ile	His	Ser	Ile	Leu	Leu
425					430					435				
Pro	Asn	Gln	Asn	Gly	Lys	Leu	Arg	Ser	Gly	Val	Pro	Val	Ser	Leu
440					445					450				
Val	Lys	Ala	Leu	Gly	Phe	Glu	Ala	Ala	Glu	Ser	Ser	Leu	Thr	Lys
455					460					465				
Asp	Ala	Leu	Val	Leu	Thr	Pro	Ala	Ser	Leu	Trp	Lys	Pro	Ser	Ser
470					475					480				
Pro Val Ser Gln														

<210> SEQ ID NO 79
<211> LENGTH: 1475
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 79

gagagaagtc agcctggcag agagactctg aaatgaggga ttagaggtgt	50
tcaaggagca agagcttcag cctgaagaca agggagcagt ccctgaagac	100
gcttctactg agaggtctgc catggcctct cttggcctcc aacttgtggg	150
ctacatccta ggccttcttg ggcttttggg cacactgggt gccatgctgc	200
tccccagctg gaaaacaagt tcttatgtcg gtgccagcat tgtgacagca	250
gttggtctct ccaagggcct ctggatggaa tgtgccacac acagcacagg	300
catcaccagc tgtgacatct atagcaccct tctgggcctg cccgctgaca	350
tccaggctgc ccaggccatg atggtgacat ccagtgcaat ctccctccctg	400
gcctgcatta tctctgtggt gggcatgaga tgcacagtct tctgccagga	450
atcccagagc aaagacagag tggcggtagc aggtggagtc tttttcatcc	500
ttggaggcct cctgggattc attcctgttg cctggaatct tcatgggatc	550
ctacgggact tctactcacc actggcgcct gacagcatga aatttgagat	600
tggagaggct ctttacttgg gcattatttc ttccctgttc tccctgatag	650
ctggaatcat cctctgcttt tcctgctcat ccagagaaa tcgctccaac	700
tactacgatg cctaccaagc ccaacctctt gccacaagga gctctccaag	750
gcctggtcaa cctcccaaag tcaagagtga gttcaattcc tacagcctga	800
cagggtatgt gtgaagaacc aggggccaga gctggggggt ggctgggtct	850
gtgaaaaaca gtggacagca ccccgagggc cacaggtgag ggacactacc	900
actggatcgt gtcagaaggt gctgctgagg atagactgac tttggccatt	950
ggattgagca aaggcagaaa tgggggctag tgtaacagca tgcaggttga	1000
attgccaaag atgctcgcca tgccagcctt tctgttttcc tcaccttgct	1050
gtccccctgc cctaagtccc caacctcaa cttgaaaccc cattccctta	1100
agccaggact cagaggatcc ctttgccctc tggtttacct gggactccat	1150
ccccaaaccc actaatcaca tcccactgac tgaccctctg tgatcaaaga	1200
ccctctctct ggctgaggtt ggctcttagc tcattgctgg ggatgggaag	1250
gagaagcagt ggcttttgtg ggcattgctc taacctactt ctcaagcttc	1300

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cctccaaaga aactgattgg ccctggaacc tccatccac tctgttatg	1350
actccacagt gtccagacta atttgtgcat gaactgaaat aaaaccatcc	1400
tacggtatcc agggaacaga aagcaggatg caggatggga ggacaggaag	1450
gcagcctggg acatttaaaa aaata	1475

<210> SEQ ID NO 80
 <211> LENGTH: 230
 <212> TYPE: PRT
 <213> ORGANISM: Homo Sapien

<400> SEQUENCE: 80

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

<210> SEQ ID NO 81
 <211> LENGTH: 1732
 <212> TYPE: DNA
 <213> ORGANISM: Homo Sapien

<400> SEQUENCE: 81

cccacgcgtc cgccctctc ccttctgctg gaccttcctt cgtctctcca	50
ttctctccctc ctttcccgc gttctctttc cacctttctc ttcttccac	100

-continued	
cttagacctc ccttcctgcc ctcccttccct gccaccgct gcttcctggc	150
ccttctccga ccccgctcta gcagcagacc tcctggggtc tgtgggttga	200
tctgtggccc ctgtgcctcc gtgtcccttt cgtctccctt cctcccgact	250
ccgctcccggt accagcggcc tgaccctggg gaaaggatgg ttcccagggt	300
gagggtcctc tcctccttgc tgggactcgc gctgctctgg ttccccctgg	350
actcccacgc tcgagccgc ccagacatgt tctgcctttt ccatgggaag	400
agatactccc ccggcgagag ctggcaccct tacttgagc cacaaggcct	450
gatgtactgc ctgcgctgta cctgctcaga gggcgcccat gtgagtgtt	500
accgcctcca ctgtccgct gtccactgcc ccagcctgt gacggagcca	550
cagcaatgct gtcccaagtg tgtggaacct cacactccct ctggactccg	600
ggccccacca aagtcctgcc agcacaacgg gaccatgtac caacacggag	650
agatcttcag tgcccatgag ctgttcccct ccgcctgcc caaccagtgt	700
gtcctctgca gctgcacaga gggccagatc tactgcggcc tcacaacctg	750
ccccgaacca ggctgcccag cccccctccc actgccagac tcctgctgcc	800
aagcctgcaa agatgaggca agtgagcaat cggatgaaga ggacagtgtg	850
cagtcgctcc atggggtgag acatcctcag gatccatgtt ccagtgatgc	900
tggggagaaag agaggcccg gacccccagc cccactggc ctcagcgccc	950
ctctgagctt catccctcgc cacttcagac ccaagggagc aggcagcaca	1000
actgtcaaga tcgtcctgaa ggagaacat aagaaagcct gtgtgcatgg	1050
cgggaagacg tactcccacg gggagggtgt gcacccggcc ttccgtgcct	1100
tcggcccctt gccctgcac ctatgcacct gtgaggatgg ccgccaggac	1150
tgccagcgtg tgacctgtcc caccgagtac cctgcccgtc accccgagaa	1200
agtggtctgg aagtgtgca agatttgccc agaggacaaa gcagaccctg	1250
gccacagtga gatcagttct accaggtgtc ccaaggcacc gggccgggtc	1300
ctcgtccaca catcggtatc cccaagccca gacaacctgc gtcgctttgc	1350
cctggaacac gaggcctcg acttggtgga gatctacctc tggaagctgg	1400
taaaagatga ggaaactgag gctcagagag gtgaagtacc tggcccaagg	1450
ccacacagcc agaattctcc acttgactca gatcaagaaa gtcaggaagc	1500
aagacttcca gaaagaggca cagcacttcc gactgctcgc tggccccac	1550
gaaggtcact ggaacgtctt cctagcccag accctggagc tgaaggtcac	1600
ggccagtcca gacaaagtga ccaagacata acaaagacct aacagttgca	1650
gatatgagct gtataattgt tgttattata tattaataaa taagaagttg	1700
cattaccctc aaaaaaaaaa aaaaaaaaaa aa	1732

<210> SEQ ID NO 82
<211> LENGTH: 451
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 82

Met Val Pro Glu Val Arg Val Leu Ser Ser Leu Leu Gly Leu Ala
1 5 10 15

-continued

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

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Ser	Gln	Asn	Leu	Pro	Leu	Asp	Ser	Asp	Gln	Glu	Ser	Gln	Glu	Ala
			395						400					405
Arg	Leu	Pro	Glu	Arg	Gly	Thr	Ala	Leu	Pro	Thr	Ala	Arg	Trp	Pro
			410						415					420
Pro	Arg	Arg	Ser	Leu	Glu	Arg	Leu	Pro	Ser	Pro	Asp	Pro	Gly	Ala
			425						430					435
Glu	Gly	His	Gly	Gln	Ser	Arg	Gln	Ser	Asp	Gln	Asp	Ile	Thr	Lys
			440						445					450

Thr

<210> SEQ ID NO 83
<211> LENGTH: 2052
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 83

gacagctgtg tctcgatgga gtagactctc agaacagcgc agtttgcct	50
ccgctcacgc agagcctctc cgtggcttcc gcaccttgag cattaggcca	100
gttctcctct tctctctaata ccatccgtca cctctcctgt catccgtttc	150
catgccgtga ggtccattca cagaacacat ccatggctct catgctcagt	200
ttggttctga gtctcctcaa gctgggatca gggcagtggc aggtgtttgg	250
gccagacaag cctgtccagg ccttggtggg ggaggacgca gcattctcct	300
gtttcctgtc tcctaagacc aatgcagagg ccatggaagt gcggttcttc	350
aggggccagt tctctagcgt ggtccacctc tacagggacg ggaaggacca	400
gccatttatg cagatgccac agtatcaagg caggacaaaa ctggtgaagg	450
attctattgc ggaggggcgc atctctctga ggctggaaaa cattactgtg	500
ttggatgctg gcctctatgg gtgcaggatt agttcccagt cttactacca	550
gaaggccatc tgggagctac aggtgtcagc actgggctca gttcctctca	600
tttccatcac gggatatgtt gatagagaca tccagctact ctgtcagtcc	650
tcgggctggt tccccggcc cacagcgaag tggaaaggtc cacaaggaca	700
ggatttgtcc acagactcca ggacaacag agacatgcat ggcctgtttg	750
atgtggagat ctctctgacc gtccaagaga acgccgggag catatcctgt	800
tccatgcggc atgctcatct gagccgagag gtggaatcca ggttacagat	850
aggagatacc tttttcgagc ctatatcgtg gcacctggct accaaagtac	900
tgggaatact ctgctgtggc ctattttttg gcattgtttg actgaagatt	950
ttctttccca aattccagtg gaaaatccag gcggaactgg actggagaag	1000
aaagcacgga caggcagaat tgagagacgc ccggaacac gcagtggagg	1050
tgactctgga tccagagacg gctcaccoga agctctgcgt ttctgatctg	1100
aaaactgtaa cccatagaaa agctccccag gaggtgcctc actctgagaa	1150
gagatttaca aggaagagtg tggtgcttc tcagagtctc caagcaggga	1200
aacattactg ggaggtggac ggaggacaca ataaaagggt gcgcgtggga	1250
gtgtgccggg atgatgtgga caggaggaag gactacgtga ctttgtctcc	1300
cgatcatggg tactgggtcc tcagactgaa tggagaacat ttgtatttca	1350

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cattaaatcc ccgttttata gcgtctttcc ccaggacccc acctacaaaa	1400
ataggggtct tcctggacta tgagtgtggg accatctcct tcttcaacat	1450
aatgaccag tcccttattt ataccctgac atgtcggttt gaaggcttat	1500
tgaggcccta cattgagtat ccgtcctata atgagcaaaa tggaactccc	1550
atagtcatct gcccagtcac ccaggaatca gagaaagagg cctcttggca	1600
aagggcctct gcaatcccag agacaagcaa cagtgagtcc tcctcacag	1650
caaccacgcc cttcctcccc aggggtgaaa tgtaggatga atcacatccc	1700
acattcttct ttagggatat taaggtctct ctcccagatc caaagtcccg	1750
cagcagccgg ccaaggtggc ttccagatga agggggactg gcctgtccac	1800
atgggagtcga ggtgtcatgg ctgccctgag ctgggaggga agaaggctga	1850
cattacattt agtttgctct cactccatct ggctaagtga tcttgaata	1900
ccacctctca ggtgaagaac cgtcaggaat tcccatctca caggctgtgg	1950
tgtagattaa gtagacaagg aatgtgaata atgcttagat cttattgatg	2000
acagagtgtg tcctaattgt ttgttcatta tattacactt tcagtaaaaa	2050
aa	2052

<210> SEQ ID NO 84

<211> LENGTH: 500

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 84

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

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Thr	Val	Gln	Glu	Asn	Ala	Gly	Ser	Ile	Ser	Cys	Ser	Met	Arg	His
				200					205					210
Ala	His	Leu	Ser	Arg	Glu	Val	Glu	Ser	Arg	Val	Gln	Ile	Gly	Asp
				215					220					225
Thr	Phe	Phe	Glu	Pro	Ile	Ser	Trp	His	Leu	Ala	Thr	Lys	Val	Leu
				230					235					240
Gly	Ile	Leu	Cys	Cys	Gly	Leu	Phe	Phe	Gly	Ile	Val	Gly	Leu	Lys
				245					250					255
Ile	Phe	Phe	Ser	Lys	Phe	Gln	Trp	Lys	Ile	Gln	Ala	Glu	Leu	Asp
				260					265					270
Trp	Arg	Arg	Lys	His	Gly	Gln	Ala	Glu	Leu	Arg	Asp	Ala	Arg	Lys
				275					280					285
His	Ala	Val	Glu	Val	Thr	Leu	Asp	Pro	Glu	Thr	Ala	His	Pro	Lys
				290					295					300
Leu	Cys	Val	Ser	Asp	Leu	Lys	Thr	Val	Thr	His	Arg	Lys	Ala	Pro
				305					310					315
Gln	Glu	Val	Pro	His	Ser	Glu	Lys	Arg	Phe	Thr	Arg	Lys	Ser	Val
				320					325					330
Val	Ala	Ser	Gln	Ser	Phe	Gln	Ala	Gly	Lys	His	Tyr	Trp	Glu	Val
				335					340					345
Asp	Gly	Gly	His	Asn	Lys	Arg	Trp	Arg	Val	Gly	Val	Cys	Arg	Asp
				350					355					360
Asp	Val	Asp	Arg	Arg	Lys	Glu	Tyr	Val	Thr	Leu	Ser	Pro	Asp	His
				365					370					375
Gly	Tyr	Trp	Val	Leu	Arg	Leu	Asn	Gly	Glu	His	Leu	Tyr	Phe	Thr
				380					385					390
Leu	Asn	Pro	Arg	Phe	Ile	Ser	Val	Phe	Pro	Arg	Thr	Pro	Pro	Thr
				395					400					405
Lys	Ile	Gly	Val	Phe	Leu	Asp	Tyr	Glu	Cys	Gly	Thr	Ile	Ser	Phe
				410					415					420
Phe	Asn	Ile	Asn	Asp	Gln	Ser	Leu	Ile	Tyr	Thr	Leu	Thr	Cys	Arg
				425					430					435
Phe	Glu	Gly	Leu	Leu	Arg	Pro	Tyr	Ile	Glu	Tyr	Pro	Ser	Tyr	Asn
				440					445					450
Glu	Gln	Asn	Gly	Thr	Pro	Ile	Val	Ile	Cys	Pro	Val	Thr	Gln	Glu
				455					460					465
Ser	Glu	Lys	Glu	Ala	Ser	Trp	Gln	Arg	Ala	Ser	Ala	Ile	Pro	Glu
				470					475					480
Thr	Ser	Asn	Ser	Glu	Ser	Ser	Ser	Gln	Ala	Thr	Thr	Pro	Phe	Leu
				485					490					495
Pro	Arg	Gly	Glu	Met										
				500										

<210> SEQ ID NO 85
<211> LENGTH: 1665
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 85

aacagacggtt ccctcgcggc cctggcacct ctaaccccag acatgctgct	50
gctgctgctg ccctgctctt gggggaggga gagggcgga ggacagacaa	100

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gtaaactgct gacgatgcag agttccgtga cgggtgcagga aggcctgtgt	150
gtccatgtgc cctgctcctt ctctacccc tcgcatggct ggatttacc	200
tggcccaagta gttcatggct actggttccg ggaaggggcc aatacagacc	250
aggatgctcc agtggccaca aacaaccag ctcgggcagt gtgggaggag	300
actcgggacc gattccacct ccttggggac ccacatacca agaattgcac	350
cctgagcatc agagatgcca gaagaagtga tgcggggaga tacttctttc	400
gtatggagaa aggaagtata aaatggaatt ataaacatca cgggtctctt	450
gtgaatgtga cagccttgac ccacaggccc aacatcctca tcccaggcac	500
cctggagtcc ggctgcccc agaattctgac ctgctctgtg ccttgggcct	550
gtgagcaggg gacccccct atgatctcct ggatagggac ctccgtgtcc	600
cccctggacc cctccaccac ccgctcctcg gtgctcacc tcacccaca	650
gcccaggac catggcacca gcctcacctg tcaggtgacc ttccctggg	700
ccagcgtgac cacgaacaag accgtccatc tcaacgtgtc ctaccgcct	750
cagaacttga ccatgactgt cttccaagga gacggcacag tatccacagt	800
cttgggaaat ggctcatctc tgctactccc agagggccag tctctgcgc	850
tggtctgtgc agttgatgca gttgacagca atccccctgc caggctgagc	900
ctgagctgga gaggcctgac cctgtgcccc tcacagccct caaaccggg	950
gggtgtggag ctgccttggg tgcacctgag ggatgcagct gaattcacct	1000
gcagagctca gaacctctc ggctctcagc aggtctacct gaacgtctcc	1050
ctgcagagca aagccacatc aggagtgact cagggggtgg tcgggggagc	1100
tggagccaca gccctggtct tcctgtcctt ctgcgtcatc ttogttgtag	1150
tgaggctctg caggaagaaa tcggcaaggc cagcagcggg cgtgggagat	1200
acgggcatag aggatgcaaa cgctgtcagg ggttcagcct ctacggggcc	1250
cctgactgaa ccttgggcag aagacagtcc ccagaccag cctccccag	1300
cttctgcccg ctctcagtg ggggaaggag agctccagta tgcattccctc	1350
agcttccaga tggatgaagc ttgggactcg cggggacagg aggccactga	1400
caccgagtac tcggagatca agatccacag atgagaaact gcagagactc	1450
accctgattg agggatcaca gcccctccag gcaagggaga agtcagaggc	1500
tgattcttgt agaattaaca gccctcaacg tgatgagcta tgataacact	1550
atgaattatg tgcagagtga aaagcacaca ggcttttagag tcaaagtatc	1600
tcaaacctga atccacactg tgccctccct tttatTTTTT taactaaaag	1650
acagacaaat tccta	1665

<210> SEQ ID NO 86

<211> LENGTH: 463

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 86

Met	Leu	Leu	Leu	Leu	Leu	Pro	Leu	Leu	Trp	Gly	Arg	Glu	Arg	Ala
1				5					10					15

Glu	Gly	Gln	Thr	Ser	Lys	Leu	Leu	Thr	Met	Gln	Ser	Ser	Val	Thr
-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----

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Glu	Pro	Trp	Ala	Glu	Asp	Ser	Pro	Pro	Asp	Gln	Pro	Pro	Pro	Ala
			410						415					420
Ser	Ala	Arg	Ser	Ser	Val	Gly	Glu	Gly	Glu	Leu	Gln	Tyr	Ala	Ser
			425						430					435
Leu	Ser	Phe	Gln	Met	Val	Lys	Pro	Trp	Asp	Ser	Arg	Gly	Gln	Glu
			440						445					450
Ala	Thr	Asp	Thr	Glu	Tyr	Ser	Glu	Ile	Lys	Ile	His	Arg		
			455						460					

<210> SEQ ID NO 87
<211> LENGTH: 1176
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 87

agaaagctgc actctgttga gctccagggc gcagtggagg gagggagtga	50
aggagctctc tgtacccaag gaaagtgcag ctgagactca gacaagatta	100
caatgaacca actcagcttc ctgctgtttc tcatagcgac caccagagga	150
tggagtacag atgaggctaa tacttacttc aaggaatgga cctgttcttc	200
gtctccatct ctgcccagaa gctgcaagga aatcaaagac gaatgtccta	250
gtgcatttga tggcctgtat tttctccgca ctgagaatgg tgttatctac	300
cagaccttct gtgacatgac ctctgggggt ggcggctgga ccctggtggc	350
cagcgtgcat gagaatgaca tgcgtgggaa gtgcacggtg ggcgacgct	400
ggtccagtca gcagggcagc aaagcagact acccagaggg ggacggcaac	450
tgggccaaact acaacacctt tggatctgca gaggcggcca cgagcgatga	500
ctacaagaac cctggctact acgacatcca ggccaaggac ctgggcatct	550
ggcacgtgcc caataagtcc cccatgcagc actggagaaa cagctccctg	600
ctgaggtacc gcacggacac tggcttcctc cagacactgg gacataatct	650
gtttggcatc taccagaaat atccagtga atatggagaa ggaaagtgtt	700
ggactgacaa cggcccgggtg atccctgtgg tctatgattt tggcgacgcc	750
cagaaaacag catcttatta ctacacctat ggccagcggg aattcaactgc	800
gggatttggt cagttcaggg tatttaataa cgagagagca gccaacgcct	850
tgtgtgctgg aatgagggtc accggatgta aactgagca tcaactgcatt	900
ggtggaggag gatactttcc agaggccagt cccagcagt gtggagattt	950
ttctggtttt gattggagtg gatatggaac tcatgttggt tacagcagca	1000
gccgtgagat aactgaggca gctgtgcttc tattctatcg ttgagagttt	1050
tgtgggaggg aaccagacc tctcctccca accatgagat cccaaggatg	1100
gagaacaact taccagtag ctagaatgtt aatggcagaa gagaaaacaa	1150
taaatcatat tgactcaaga aaaaaa	1176

<210> SEQ ID NO 88
<211> LENGTH: 313
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 88

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

<210> SEQ ID NO 89
<211> LENGTH: 759
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 89

ctagatttgt cggcttgcg ggagacttca ggagtcgctg tctctgaact 50
tccagcctca gagaccgcc ccctgtccc cgagggccat gggccgggtc 100

-continued

tcagggcttg tgccctctcg cttcctgacg ctcctggcgc atctggtggt	150
cgctcatcacc ttattctggt cccgggacag caacatacag gcctgcctgc	200
ctctcacgtt cacccccag gagtatgaca agcaggacat tcagctggtg	250
gccgcgctct ctgtcacctt gggcctcttt gcagtggagc tggccggttt	300
cctctcagga gtctccatgt tcaacagcac ccagagcctc atctccattg	350
gggctcactg tagtgcattc gtggccctgt ccttcttcat attcgagcgt	400
tgggagtgcg ctacgtattg gtacattttt gtcttctgca gtgcccttcc	450
agctgtcact gaaatggctt tattcgtcac cgtctttggg ctgaaaaaga	500
aacccttctg attaccttca tgacgggaac ctaaggacga agcctacagg	550
ggcaagggcc gcttcgtatt cctggaagaa ggaaggcata ggcttcggtt	600
ttccctcgg aaactgcttc tgctggagga tatgtgttg aataattacg	650
tcttgagtct gggattatcc gcattgtatt tagtgctttg taataaaata	700
tgttttgtag taacattaag acttatatac agttttaggg gacaattaa	750
aaaaaaaa	759

<210> SEQ ID NO 90
<211> LENGTH: 140
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 90

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

<210> SEQ ID NO 91
<211> LENGTH: 1871
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 91

ctgggacccc gaaaagagaa ggggagagcg aggggacgag agcggaggag	50
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gaagatgcaa ctgactcgct gctgcttcgt gttcctgggtg cagggtagcc	100
tctatctggt catctgtggc caggatgatg gtcctcccg ctcagaggac	150
cctgagcgtg atgaccacga gggccagccc cggccccggg tgcctcgga	200
gcggggccac atctcaccta agtcccgccc catggccaat tccactctcc	250
tagggctgct gggcccgctt ggggaggctt ggggcattct tgggcagccc	300
cccaaccgcc cgaaccacag cccccaccc tcagccaagg tgaagaaaat	350
ctttggctgg ggcgacttct actccaacat caagacgggtg gccctgaacc	400
tgctcgtcac agggaagatt gtggaccatg gcaatgggac cttcagcgtc	450
cacttccaac acaatgccac aggccaggga aacatctcca tcagcctcgt	500
gccccccagt aaagctgtag agttccacca ggaacagcag atcttcacgt	550
aagccaaggc ctccaaaatc ttcaactgcc ggatggagtg ggagaaggta	600
gaacggggcc gccggacctc gctttgcacc cagcaccag ccaagatctg	650
ctcccgagac cagctcaga gctcagccac ctggagctgc tcccagccct	700
tcaaagtcgt ctgtgtctac atcgcttctt acagcacgga ctatcggtg	750
gtccagaagg tgtgccaga ttacaactac catagtata cccctacta	800
cccctctggg tgacccgggg caggccacag aggccaggcc agggctggaa	850
ggacaggcct gcccatgcag gagaccatct ggacaccggg cagggaaggg	900
gttgggcctc aggcaggagg gggggtggag acgaggagat gccaaagggt	950
gccaggggcca agtctcaagt ggcagagaaa ggggcccaag tgctggtccc	1000
aacctgaagc tgtggagtga ctagatcaca ggagcactgg aggaggagt	1050
ggctctctgt gcagcctcac agggctttgc caggagcca cagagagatg	1100
ctgggtcccc gaggcctgtg ggcaggccga tcagtgtggc cccagatcaa	1150
gtcatgggag gaagctaagc ccttggttct tgccatcctg aggaaagata	1200
gcaacaggga gggggagatt tcatcagtgt ggacagcctg tcaacttagg	1250
atggatggct gagagggtt cctaggagcc agtcagcagg gtgggtggg	1300
gccagaggag ctctccagcc ctgcctagtg ggcgccctga gcccttgtc	1350
gtgtgctgag catggcatga ggctgaagtg gcaaccctgg ggtctttgat	1400
gtcttgacag attgaccatc tgtctccagc caggccaccc ctttccaaa	1450
ttccctcttc tgccagtact cccctgtac caccattgc tgatggcaca	1500
cccctcctta agctaagaca ggacgattgt ggtcctccca cactaaggcc	1550
acagcccatc cgcgtgctgt gtgtccctct tccaccccaa cccctgctgg	1600
ctcctctggg agcatccatg tcccggagag gggccctca acagtcagcc	1650
tcacctgtca gaccgggggt ctcccgatc tggatggcgc cgcctctca	1700
gcagcgggca cgggtggggc ggggcggg cgcagagcat gtgctggatc	1750
tgttctgtgt gtctgtctgt ggggtggggg aggggagga agtcttgtga	1800
aaccgctgat tgctgacttt tgtgtgaaga atcgtgttct tggagcagga	1850
aataaagctt gccccggggc a	1871

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<211> LENGTH: 252
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 92
Met Gln Leu Thr Arg Cys Cys Phe Val Phe Leu Val Gln Gly Ser
1 5 10 15
Leu Tyr Leu Val Ile Cys Gly Gln Asp Asp Gly Pro Pro Gly Ser
20 25 30
Glu Asp Pro Glu Arg Asp Asp His Glu Gly Gln Pro Arg Pro Arg
35 40 45
Val Pro Arg Lys Arg Gly His Ile Ser Pro Lys Ser Arg Pro Met
50 55 60
Ala Asn Ser Thr Leu Leu Gly Leu Leu Ala Pro Pro Gly Glu Ala
65 70 75
Trp Gly Ile Leu Gly Gln Pro Pro Asn Arg Pro Asn His Ser Pro
80 85 90
Pro Pro Ser Ala Lys Val Lys Lys Ile Phe Gly Trp Gly Asp Phe
95 100 105
Tyr Ser Asn Ile Lys Thr Val Ala Leu Asn Leu Leu Val Thr Gly
110 115 120
Lys Ile Val Asp His Gly Asn Gly Thr Phe Ser Val His Phe Gln
125 130 135
His Asn Ala Thr Gly Gln Gly Asn Ile Ser Ile Ser Leu Val Pro
140 145 150
Pro Ser Lys Ala Val Glu Phe His Gln Glu Gln Gln Ile Phe Ile
155 160 165
Glu Ala Lys Ala Ser Lys Ile Phe Asn Cys Arg Met Glu Trp Glu
170 175 180
Lys Val Glu Arg Gly Arg Arg Thr Ser Leu Cys Thr His Asp Pro
185 190 195
Ala Lys Ile Cys Ser Arg Asp His Ala Gln Ser Ser Ala Thr Trp
200 205 210
Ser Cys Ser Gln Pro Phe Lys Val Val Cys Val Tyr Ile Ala Phe
215 220 225
Tyr Ser Thr Asp Tyr Arg Leu Val Gln Lys Val Cys Pro Asp Tyr
230 235 240
Asn Tyr His Ser Asp Thr Pro Tyr Tyr Pro Ser Gly
245 250

<210> SEQ ID NO 93
<211> LENGTH: 902
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 93

cggtggccat gactgcggcc gtgttcttcg gctgcgcctt cattgccttc	50
gggcctgcgc tcgcccttta tgtcttcacc atcgccatcg agccgttgcg	100
tatcatcttc ctcatcgccg gagctttctt ctggttggtg tctctactga	150
tttcgtccct tgtttggttc atggcaagag tcattattga caacaaagat	200
ggaccaacac agaaatatct gctgatcttt ggagcgtttg tctctgtcta	250
tatccaagaa atgttccgat ttgcatatta taaactctta aaaaaagcca	300

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gtgaagggtt gaagagtata aaccaggtg agacagcacc ctctatgcga      350
ctgctggcct atgtttcttg cttgggcttt ggaatcatga gtggagtatt      400
ttcctttgtg aataccctat ctgactcctt ggggccaggc acagtgggca      450
ttcatggaga ttctcctcaa ttcttccttt attcagcttt catgacgctg      500
gtcattatct tgctgcatgt attctggggc attgtatttt ttgatggctg      550
tgagaagaaa aagtggggca tcctccttat cgttctcctg acccacctgc      600
tggtgtcagc ccagaccttc ataagttctt attatggaat aaacctggcg      650
tcagcattta taatcctggt gctcatgggc acctgggcat tcttagctgc      700
gggaggcagc tgccgaagcc tgaactctg cctgctctgc caagacaaga      750
actttctctt ttacaaccag cgctccagat aacctcaggg aaccagcact      800
tcccaaaccg cagactacat cttagagga agcacaactg tgcctttttc      850
tgaaaatccc tttttctggt ggaattgaga aagaaataaa actatgcaga      900
ta                                                                902
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<210> SEQ ID NO 94

<211> LENGTH: 257

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 94

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

<210> SEQ ID NO 95
<211> LENGTH: 1073
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 95
aatttttcac cagagtaaacc ttgagaaacc aactggacct tgagtattgt 50
acatttttgcc tcgtggaccc aaaggtagca atctgaaaca tgaggagtac 100
gattctactg ttttgtcttc taggatcaac tcggtcatta ccacagctca 150
aacctgcttt gggactccct cccacaaaac tggctccgga tcaggggaaca 200
ctaccaaacc aacagcagtc aaatcaggtc tttccttctt taagtctgat 250
accattaaca cagatgctca cactggggcc agatctgcat ctgttaaata 300
ctgctgcagg aatgacacct ggtaccacaga cccaccatt gacctggga 350
gggttgaatg tacaacagca actgcaccca catgtgttac caatttttgt 400
cacacaactt ggagcccagg gcactatcct aagctcagag gaattgccac 450
aaatcttcac gagcctcatc atccattcct tgttcccggg aggcatactg 500
cccaccagtc aggcaggggc taatccagat gtccaggatg gaagccttcc 550
agcaggagga gcaggtgtaa atcctgccac ccagggaacc ccagcaggcc 600
gcctcccaac tcccagtggc acagatgacg actttgcagt gaccaccct 650
gcaggcatcc aaaggagcac acatgccatc gaggaagcca ccacagaatc 700
agcaaatgga attcagtaag ctgtttcaaa ttttttcaac taagctgcct 750
cgaatttggg gatacatgtg aatctttatc attgattata ttatggaata 800
gattgagaca cattggatag tcttagaaga aattaattct taatttacct 850
gaaaatatc ttgaaatttc agaaaatatg ttctatgtag agaatcccaa 900
cttttaaaaa caataattca atggataaat ctgtctttga aatataacat 950
tatgtctgcct ggatgatatg catattaaaa catatttgga aaactggaaa 1000
aaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa 1050
aaaaaaaaa aaaaaaaaaa aaa 1073

<210> SEQ ID NO 96
<211> LENGTH: 209
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 96
Met Arg Ser Thr Ile Leu Leu Phe Cys Leu Leu Gly Ser Thr Arg
1 5 10 15
Ser Leu Pro Gln Leu Lys Pro Ala Leu Gly Leu Pro Pro Thr Lys

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

<210> SEQ ID NO 97
<211> LENGTH: 2848
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 97

gctcaagtgc cctgccttgc cccacccagc ccagcctggc cagagccccc	50
tgagagaagg gctctcttct tgcttggcag ctggaccaag ggagccagtc	100
ttgggcgctg gagggcctgt cctgacctg gtccctgcct ggctgtggct	150
gctttgtgtc tccgtccccc aggtctctcc caaggcccag cctgcagagc	200
tgtctgtgga agttccagaa aactatggtg gaaatttccc ttataacctg	250
accaagttgc cgctgccccg tgagggggct gaaggccaga tcgtgctgtc	300
aggggactca ggcaaggcaa ctgagggccc atttgctatg gatccagatt	350
ctggcttctc gctggtgacc agggccctgg accgagagga gcaggcagag	400
taccagctac aggtcaccct ggagatgcag gatggacatg tcttgtgggg	450
tccacagcct gtgcttgtgc acgtgaagga tgagaatgac caggtgcccc	500
atttctctca agccatctac agagctcggc tgagccgggg taccaggcct	550
ggcatccctc tcctcttctc tgaggcttca gaccgggatg agccaggcac	600
agccaactcg gatcttcgat tccacatcct gagccaggct ccagcccagc	650
cttccccaga catgttccag ctggagcctc ggctgggggc tctggccctc	700
agccccaagg ggagcaccag ccttgaccac gccctggaga ggacctacca	750

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gctgttgta caggtcaagg acatgggtga ccaggcctca ggccaccagg	800
ccactgccac cgtggaagtc tccatcatag agagcacctg ggtgtcccta	850
gagcctatcc acctggcaga gaatctcaaa gtcctatacc cgcaccacat	900
ggcccaghta cactggagtg ggggtgatgt gcactatcac ctggagagcc	950
atcccccggt accctttgaa gtgaatgcag agggaaacct ctacgtgacc	1000
agagagctgg acagagaagc ccaggctgag tacctgctcc aggtgcgggc	1050
tcagaattcc catggcgagg actatgcggc ccctctggag ctgcacgtgc	1100
tgggtgatga tgagaatgac aacgtgccta tctgccctcc ccgtgacccc	1150
acagtacga tccctgagct cagtccacca ggtactgaag tgactagact	1200
gtcagcagag gatgcagatg cccccggctc cccaattcc caggttgtgt	1250
atcagctcct gagccctgag cctgaggatg gggtagaggg gagagccttc	1300
caggtggacc ccacttcagg cagtgtgacg ctgggggtgc tccactccg	1350
agcaggccag aacatcctgc ttctggtgct ggccatggac ctggcaggcg	1400
cagaggggtg cttcagcagc acgtgtgaag tcgaagtcgc agtcacagat	1450
atcaatgatc acgcccctga gttcatcact tcccagattg ggcctataag	1500
cctccctgag gatgtggagc cgggactct ggtggccatg ctaacagcca	1550
ttgatgctga cctcgagccc gccttcggc tcattggattt tgccattgag	1600
aggggagaca cagaaggagc ttttggcctg gattgggagc cagactctgg	1650
gcatgttaga ctcagactct gcaagaacct cagttatgag gcagctccaa	1700
gtcatgaggt ggtgggtggt gtgcagagtg tggcgaagct ggtggggcca	1750
ggcccaggcc ctggagccac cggccagggt actgtgctag tggagagagt	1800
gatgccaccc cccaagtgg accaggagag ctacgaggcc agtgtccca	1850
tcagtgcctc agccggctct ttctctgctga ccatccagcc ctccgacccc	1900
atcagccgaa ccctcagggt ctccctagtc aatgactcag agggctggct	1950
ctgcattgag aaattctccg gggaggtgca caccgcccag tccctgcagg	2000
gcgcccagcc tggggacacc tacacggtgc ttgtggaggc ccaggataca	2050
gccctgactc ttgcccctgt gccctcccaa tacctctgca cccccgcca	2100
agaccatggc ttgatcgtga gtggacccag caaggacccc gatctggcca	2150
gtgggcacgg tccctacagc ttcaccttg gtcccaaccc caggtgcaa	2200
cgggattggc gcctccagac tctcaatggt tcccatgcct acctcacctt	2250
ggccctgcat tgggtggagc cactgaaca cataatcccc gtggtggtca	2300
gccacaatgc ccagatgtg cagctcctgg ttcgagtgat cgtgtgtcgc	2350
tgaacgtgg aggggcagtg catgcgcaag gtgggcccga tgaaggcat	2400
gccacgaag ctgtcggcag tgggcatcct tgtaggcacc ctggtagcaa	2450
taggaatctt cctcatcctc attttcaccc actggacat gtcaaggag	2500
aaggacccgg atcaaccagc agacagcgtg cccctgaagg cgactgtctg	2550
aatggcccag gcagctctag ctgggagctt ggcctctggc tccatctgag	2600
tccctggga gagagcccag caccgaagat ccagcagggg acaggacaga	2650

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gtagaagccc ctccatctgc cctgggggtgg aggcaccatc accatcacca	2700
ggcatgtctg cagagcctgg acaccaactt tatggactgc ccatgggagt	2750
gtcccaaatg tcaggggtgt tgcccaataa taaagcccca gagaactggg	2800
ctggggcccta tgggaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaag	2848

<210> SEQ ID NO 98

<211> LENGTH: 807

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 98

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

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

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680					685					690				
Gln	Arg	Asp	Trp	Arg	Leu	Gln	Thr	Leu	Asn	Gly	Ser	His	Ala	Tyr
695					700					705				
Leu	Thr	Leu	Ala	Leu	His	Trp	Val	Glu	Pro	Arg	Glu	His	Ile	Ile
710					715					720				
Pro	Val	Val	Val	Ser	His	Asn	Ala	Gln	Met	Trp	Gln	Leu	Leu	Val
725					730					735				
Arg	Val	Ile	Val	Cys	Arg	Cys	Asn	Val	Glu	Gly	Gln	Cys	Met	Arg
740					745					750				
Lys	Val	Gly	Arg	Met	Lys	Gly	Met	Pro	Thr	Lys	Leu	Ser	Ala	Val
755					760					765				
Gly	Ile	Leu	Val	Gly	Thr	Leu	Val	Ala	Ile	Gly	Ile	Phe	Leu	Ile
770					775					780				
Leu	Ile	Phe	Thr	His	Trp	Thr	Met	Ser	Arg	Lys	Lys	Asp	Pro	Asp
785					790					795				
Gln	Pro	Ala	Asp	Ser	Val	Pro	Leu	Lys	Ala	Thr	Val			
800					805									

<210> SEQ ID NO 99
<211> LENGTH: 2436
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 99

ggctgaccgt gctacattgc ctggaggaag cctaaggaac ccaggcatcc	50
agctgcccac gcctgagtcc aagattcttc ccaggaacac aaacgtagga	100
gacccacgct cctggaagca ccagccttta tctcttcacc ttcaagtccc	150
ctttctcaag aatcctctgt tctttgccct ctaaagtctt ggtacatcta	200
ggacccaggc atcttgcttt ccagccacaa agagacagat gaagatgcag	250
aaaggaaatg ttctccttat gtttggtcta ctattgcatt tagaagctgc	300
aacaaattcc aatgagacta gcacctctgc caacactgga tccagtgtga	350
tctccagtgg agccagcaca gccaccaact ctgggtccag tgtgacctcc	400
agtggggcca gcacagccac catctcaggg tccagcgtga cctccaatgg	450
ggtcagcata gtcaccaact ctgagtcca tacaacctcc agtgggatca	500
gcacagccac caactctgag ttcagcacag cgtccagtgg gatcagcata	550
gccaccaact ctgagtccag cacaacctcc agtggggcca gcacagccac	600
caactctgag tccagcacac cctccagtgg ggccagcaca gtcaccaact	650
ctgggtccag tgtgacctcc agtggagcca gcaactgccac caactctgag	700
tccagcacag tgtccagtag ggccagcact gccaccaact ctgagtctag	750
cacactctcc agtggggcca gcacagccac caactctgac tccagcacia	800
cctccagtgg ggctagcaca gccaccaact ctgagtccag cacaacctcc	850
agtggggcca gcacagccac caactctgag tccagcacag tgtccagtag	900
ggccagcact gccaccaact ctgagtccag cacaacctcc agtggggcca	950
gcacagccac caactctgag tccagaacga cctccaatgg ggtgggcaca	1000
gccaccaact ctgagtccag cagacctcc agtggggcca gcacagccac	1050

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caactctgac tccagcacag tgtccagtgg ggccagcact gccaccaact	1100
ctgagtccag cacgacctcc agtggggcca gcacagccac caactctgag	1150
tccagcacga cctccagtgg ggctagcaca gccaccaact ctgactccag	1200
cacaacctcc agtggggccg gcacagccac caactctgag tccagcacag	1250
tgtccagtgg gatcagcaca gtcaccaatt ctgagtccag cacacctcc	1300
agtggggcca acacagccac caactctgag tccagtacga cctccagtgg	1350
ggccaacaca gccaccaact ctgagtccag cacagtgtcc agtggggcca	1400
gcaactgccac caactctgag tccagcaca cctccagtgg ggtcagcaca	1450
gccaccaact ctgagtccag cacaacctcc agtggggcta gcacagccac	1500
caactctgac tccagcaca cctccagtga ggccagcaca gccaccaact	1550
ctgagtctag cacagtgtcc agtgggatca gcacagtcac caattctgag	1600
tccagcaca cctccagtgg ggccaacaca gccaccaact ctgggtccag	1650
tgtgacctct gcaggctctg gaacagcagc tctgactgga atgcacacaa	1700
cttcccatag tgcattctact gcagtgagtg aggcaaagcc tgggtgggtcc	1750
ctggtgccgt gggaaatctt cctcatcacc ctggtctcgg ttgtggcggc	1800
cgtggggctc tttgctgggc tcttcttctg tgtgagaaac agcctgtccc	1850
tgagaaacac cttaacaca gctgtctacc accctcatgg cctcaacctat	1900
ggccttggtc caggccctgg agggaaatcat ggagccccc acaggcccag	1950
gtggagtcc aactggttct ggaggagacc agtatcatcg atagccatgg	2000
agatgagcgg gaggaacagc ggccctgag cagccccgga agcaagtgc	2050
gcattcttca ggaaggaaga gacctgggca cccaagacct ggtttccttt	2100
cattcatccc aggagacccc tcccagcttt gtttgagatc ctgaaaatct	2150
tgaagaaggt attcctcacc tttcttgctt ttaccagaca ctggaaagag	2200
aatactatat tgctcattta gctaagaaat aaatacatct catctaacac	2250
acacgacaaa gagaagctgt gcttgccccg ggggtgggtat ctgctctga	2300
gatgaactca gttataggag aaaacctcca tgctggactc catctggcat	2350
tcaaaatctc cacagtaaaa tccaaagacc tcaaaaaaaaa aaaaaaaaaa	2400
aaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaa	2436

<210> SEQ ID NO 100

<211> LENGTH: 596

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 100

Met	Lys	Met	Gln	Lys	Gly	Asn	Val	Leu	Leu	Met	Phe	Gly	Leu	Leu
1				5					10					15
Leu	His	Leu	Glu	Ala	Ala	Thr	Asn	Ser	Asn	Glu	Thr	Ser	Thr	Ser
				20					25					30
Ala	Asn	Thr	Gly	Ser	Ser	Val	Ile	Ser	Ser	Gly	Ala	Ser	Thr	Ala
				35					40					45
Thr	Asn	Ser	Gly	Ser	Ser	Val	Thr	Ser	Ser	Gly	Val	Ser	Thr	Ala
				50					55					60

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Thr	Ile	Ser	Gly	Ser	Ser	Val	Thr	Ser	Asn	Gly	Val	Ser	Ile	Val
			65						70					75
Thr	Asn	Ser	Glu	Phe	His	Thr	Thr	Ser	Ser	Gly	Ile	Ser	Thr	Ala
			80						85					90
Thr	Asn	Ser	Glu	Phe	Ser	Thr	Ala	Ser	Ser	Gly	Ile	Ser	Ile	Ala
			95						100					105
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Thr	Ser	Ser	Gly	Ala	Ser	Thr	Ala
			110						115					120
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Pro	Ser	Ser	Gly	Ala	Ser	Thr	Val
			125						130					135
Thr	Asn	Ser	Gly	Ser	Ser	Val	Thr	Ser	Ser	Gly	Ala	Ser	Thr	Ala
			140						145					150
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Val	Ser	Ser	Arg	Ala	Ser	Thr	Ala
			155						160					165
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Leu	Ser	Ser	Gly	Ala	Ser	Thr	Ala
			170						175					180
Thr	Asn	Ser	Asp	Ser	Ser	Thr	Thr	Ser	Ser	Gly	Ala	Ser	Thr	Ala
			185						190					195
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Thr	Ser	Ser	Gly	Ala	Ser	Thr	Ala
			200						205					210
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Val	Ser	Ser	Arg	Ala	Ser	Thr	Ala
			215						220					225
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Thr	Ser	Ser	Gly	Ala	Ser	Thr	Ala
			230						235					240
Thr	Asn	Ser	Glu	Ser	Arg	Thr	Thr	Ser	Asn	Gly	Ala	Gly	Thr	Ala
			245						250					255
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Thr	Ser	Ser	Gly	Ala	Ser	Thr	Ala
			260						265					270
Thr	Asn	Ser	Asp	Ser	Ser	Thr	Val	Ser	Ser	Gly	Ala	Ser	Thr	Ala
			275						280					285
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Thr	Ser	Ser	Gly	Ala	Ser	Thr	Ala
			290						295					300
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Thr	Ser	Ser	Gly	Ala	Ser	Thr	Ala
			305						310					315
Thr	Asn	Ser	Asp	Ser	Ser	Thr	Thr	Ser	Ser	Gly	Ala	Gly	Thr	Ala
			320						325					330
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Val	Ser	Ser	Gly	Ile	Ser	Thr	Val
			335						340					345
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Pro	Ser	Ser	Gly	Ala	Asn	Thr	Ala
			350						355					360
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Thr	Ser	Ser	Gly	Ala	Asn	Thr	Ala
			365						370					375
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Val	Ser	Ser	Gly	Ala	Ser	Thr	Ala
			380						385					390
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Thr	Ser	Ser	Gly	Val	Ser	Thr	Ala
			395						400					405
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Thr	Ser	Ser	Gly	Ala	Ser	Thr	Ala
			410						415					420
Thr	Asn	Ser	Asp	Ser	Ser	Thr	Thr	Ser	Ser	Glu	Ala	Ser	Thr	Ala
			425						430					435
Thr	Asn	Ser	Glu	Ser	Ser	Thr	Val	Ser	Ser	Gly	Ile	Ser	Thr	Val

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440	445	450
Thr Asn Ser Glu Ser Ser Thr Thr Ser	Ser Gly Ala Asn Thr Ala	
455	460	465
Thr Asn Ser Gly Ser Ser Val Thr Ser	Ala Gly Ser Gly Thr Ala	
470	475	480
Ala Leu Thr Gly Met His Thr Thr Ser	His Ser Ala Ser Thr Ala	
485	490	495
Val Ser Glu Ala Lys Pro Gly Gly Ser	Leu Val Pro Trp Glu Ile	
500	505	510
Phe Leu Ile Thr Leu Val Ser Val Val	Ala Ala Val Gly Leu Phe	
515	520	525
Ala Gly Leu Phe Phe Cys Val Arg Asn	Ser Leu Ser Leu Arg Asn	
530	535	540
Thr Phe Asn Thr Ala Val Tyr His Pro	His Gly Leu Asn His Gly	
545	550	555
Leu Gly Pro Gly Pro Gly Gly Asn His	Gly Ala Pro His Arg Pro	
560	565	570
Arg Trp Ser Pro Asn Trp Phe Trp Arg	Arg Pro Val Ser Ser Ile	
575	580	585
Ala Met Glu Met Ser Gly Arg Asn Ser	Gly Pro	
590	595	

<210> SEQ ID NO 101
<211> LENGTH: 1728
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 101

ggccggagcg ctcgcgtta cgggatgaat taacggcggg ttccgcacgg	50
aggttgtgac ccctacggag cccacgcttg cccacgcacc ccactcggcg	100
tcgcgcggcg tgccctgctt gtcacaggtg ggaggctgga actatcaggc	150
tgaaaaacag agtgggtact ctcttctggg aagctggcaa caaatggatg	200
atgtgatata tgcattccag gggaaggaa attgtggtgc ttctgaaccc	250
atggtcaatt aacgaggcag tttctagcta ctgcacgtac ttcataaagc	300
aggactctaa aagctttgga atcatggtgt catggaaagg gatttacttt	350
atactgactc tgttttgggg aagctttttt ggaagcattt tcatgctgag	400
tcccttttta cctttgatgt ttgtaaaccc atcttggtat cgctggatca	450
acaaccgcct tgtggcaaca tggtcacccc tacctgtggc attattggag	500
accatgtttg gtgtaaaagt gattataact ggggatgcat ttgttcctgg	550
agaaagaagt gtcattatca tgaaccatcg gacaagaatg gactggatgt	600
tcctgtggaa ttgcctgatg cgatatagct acctcagatt ggagaaaatt	650
tgccctcaaag cgagtctcaa aggtgttcct ggatttggtt gggccatgca	700
ggctgctgcc tatatcttca ttcataggaa atggaaggat gacaagagcc	750
atctcgaaga catgattgat tacttttgty atattcacga accacttcaa	800
ctcctcatat tcccagaagg gactgatctc acagaaaaca gcaagtctcg	850
aagtaatgca tttgctgaaa aaaatggact tcagaaatat gaatatgttt	900

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tacatccaag aactacaggc ttacttttg tggtagaccg tctaagagaa	950
ggtaagaacc ttgatgctgt ccatgatatc actgtggcgt atcctcacia	1000
cattcctcaa tcagagaagc acctcctcca aggagacttt cccagggaaa	1050
tccactttca cgtccaccgg tatccaatag acaccctccc cacatccaag	1100
gaggaccttc aactctggtg ccacaaacgg tgggaagaga aagaagagag	1150
gctgcgttcc ttctatcaag gggagaagaa tttttatttt accggacaga	1200
gtgtcattcc accttgcaag tctgaactca gggtccttgt ggtcaaattg	1250
ctctctatac tgtattggac cctgttcagc cctgcaatgt gcctactcat	1300
atatttgtac agtcttgtaa agtggtattt tataatcacc attgtaatct	1350
ttgtgctgca agagagaata ttgtgtggac tggagatcat agaacttgca	1400
tgttaccgac ttttacacia acagccacat ttaaattcaa agaaaaatga	1450
gtaagattat aaggtttgcc atgtgaaaac ctagagcata ttttggaat	1500
gttctaaacc tttctaagct cagatgcatt ttgcatgac tatgtcgaat	1550
atttcttact gccatcatta ttgttaaag atattttgca cttaattttg	1600
tgggaaaaat attgctacaa ttttttttaa tctctgaatg taatttcgat	1650
actgtgtaca tagcagggag tgatcggggt gaaataactt gggccagaat	1700
attattaaac aatcatcagg cttttaaa	1728

<210> SEQ ID NO 102
<211> LENGTH: 414
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 102

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

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	170		175		180
His Phe Glu Asp	Met Ile Asp Tyr Phe	Cys Asp Ile His Glu	Pro		
	185		190		195
Leu Gln Leu Leu	Ile Phe Pro Glu Gly	Thr Asp Leu Thr Glu	Asn		
	200		205		210
Ser Lys Ser Arg	Ser Asn Ala Phe Ala	Glu Lys Asn Gly Leu	Gln		
	215		220		225
Lys Tyr Glu Tyr	Val Leu His Pro Arg	Thr Thr Gly Phe Thr	Phe		
	230		235		240
Val Val Asp Arg	Leu Arg Glu Gly Lys	Asn Leu Asp Ala Val	His		
	245		250		255
Asp Ile Thr Val	Ala Tyr Pro His Asn	Ile Pro Gln Ser Glu	Lys		
	260		265		270
His Leu Leu Gln	Gly Asp Phe Pro Arg	Glu Ile His Phe His	Val		
	275		280		285
His Arg Tyr Pro	Ile Asp Thr Leu Pro	Thr Ser Lys Glu Asp	Leu		
	290		295		300
Gln Leu Trp Cys	His Lys Arg Trp Glu	Glu Lys Glu Glu Arg	Leu		
	305		310		315
Arg Ser Phe Tyr	Gln Gly Glu Lys Asn	Phe Tyr Phe Thr Gly	Gln		
	320		325		330
Ser Val Ile Pro	Pro Cys Lys Ser Glu	Leu Arg Val Leu Val	Val		
	335		340		345
Lys Leu Leu Ser	Ile Leu Tyr Trp Thr	Leu Phe Ser Pro Ala	Met		
	350		355		360
Cys Leu Leu Ile	Tyr Leu Tyr Ser Leu	Val Lys Trp Tyr Phe	Ile		
	365		370		375
Ile Thr Ile Val	Ile Phe Val Leu Gln	Glu Arg Ile Phe Gly	Gly		
	380		385		390
Leu Glu Ile Ile	Glu Leu Ala Cys Tyr	Arg Leu Leu His Lys	Gln		
	395		400		405
Pro His Leu Asn	Ser Lys Lys Asn Glu				
	410				

<210> SEQ ID NO 103
<211> LENGTH: 2403
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 103

cggtctcgagc ggctcgagtg aagagcctct ccacggctcc tgcgcctgag	50
acagctggcc tgacctccaa atcatccatc caccctgct gtcattctgtt	100
ttcatagtgt gagatcaacc cacaggaata tccatggctt ttgtgctcat	150
tttggttctc agttttctac agctgggtgc aggacagtgg caagtcactg	200
gaccgggcaa gtttgtccag gccttggtgg gggaggacgc cgtgtttctcc	250
tgctccctct ttctgagac cagtgcagag gctatggaag tgcggttctt	300
caggaatcag ttccatgctg tgggtccacct ctacagagat ggggaagact	350
gggaatctaa gcagatgcca cagtatcgag ggagaactga gtttgtgaag	400
gactccattg caggggggcg tgtctctcta aggctaaaaa acatcactcc	450

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ctcggacatc ggcctgtatg ggtgctggtt cagttcccag atttacgatg	500
aggaggccac ctgggagctg cgggtggcag cactgggctc acttcctctc	550
atttccatcg tgggatatgt tgacggaggt atccagttac tctgcctgtc	600
ctcaggctgg ttcccccagc ccacagccaa gtggaaggt ccacaaggac	650
aggatttgtc ttcagactcc agagcaaatg cagatgggta cagcctgtat	700
gatgtggaga tctccattat agtccaggaa aatgctggga gcatattgtg	750
ttccatccac cttgctgagc agagtcatga ggtggaatcc aaggatttga	800
taggagagac gtttttccag ccctcacctt ggcgcctggc ttctatttta	850
ctcgggttac tctgtgtgct cctgtgtggt gttgtcatgg ggaatgataat	900
tgttttcttc aaatccaaag ggaaaatcca ggcggaactg gactggagaa	950
gaaagcacgg acaggcagaa ttgagagacg cccggaaaca cgagtgagg	1000
gtgactctgg atccagagac ggctcaccgg aagctctgctg tttctgatct	1050
gaaaactgta acccatagaa aagctcccca ggaggctcct cactctgaga	1100
agagatttac aaggaagagt gtggtggctt ctcagggttt ccaagcagg	1150
agacattact gggaggtgga cgtgggacaa aatgtagggt ggtatgtggg	1200
agtgtgtcgg gatgacgtag acaggggaa gaacaatgtg actttgtctc	1250
ccaacaatgg gtattgggtc ctcagactga caacagaaca ttgtatttc	1300
acattcaatc cccattttat cagcctcccc cccagcacc ctcctacacg	1350
agtagggttc ttcttgact atgagggtgg gaccatctcc ttcttcaata	1400
caaatgacca gtcccttatt tatacctgc tgacatgtca gtttgaaggc	1450
ttgttgagac cctatatcca gcatgcgatg tatgacgagg aaaaggggac	1500
tcccatattc atatgtccag tgcctgggg atgagacaga gaagacctg	1550
cttaaaggcg cccacaccac agaccagac acagccaagg gagagtgtc	1600
ccgacaggtg gcccagctt cctctccgga gctgcgac agagagtcac	1650
gccccact ctcctttagg gagctgaggt tctctgtccc tgagccctgc	1700
agcagcggca gtcacagctt ccagatgagg ggggattggc ctgaccctgt	1750
gggagtcaga agccatggct gccctgaagt ggggacggaa tagactcaca	1800
ttaggtttag tttgtgaaa ctcacccag ctaagcgatc ttgaacaagt	1850
cacaacctcc caggctcctc atttgctagt cacggacagt gattcctgcc	1900
tcacaggtga agattaaaga gacaacgaat gtgaatcatg cttgcaggtt	1950
tgagggcaca gtgtttgcta atgatgtgtt tttatattat acattttccc	2000
accataaact ctgtttgctt attccacatt aatttacttt tctctatacc	2050
aaatcaccca tggaatagtt attgaacacc tgctttgtga ggctcaaaga	2100
ataaagagga ggtaggattt ttcactgatt ctataagccc agcattacct	2150
gataccaaaa ccaggcaag aaaacagaag aagaggaagg aaaactacag	2200
gtccatatcc ctcattaaca cagacacaaa aattctaat aaaattttaa	2250
caaattaaac taaacaatat atttaaagat gatataaac tactcagtgt	2300
ggtttgtccc acaaatgcag agttggttta atatttaaat atcaaccagt	2350

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gtaattcagc acattaataa agtaaaaaag aaaaccataa aaaaaaaaaa	2400
aaa	2403

<210> SEQ ID NO 104

<211> LENGTH: 466

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 104

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

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				320					325					330
Val	Ala	Ser	Gln	Gly	Phe	Gln	Ala	Gly	Arg	His	Tyr	Trp	Glu	Val
				335					340					345
Asp	Val	Gly	Gln	Asn	Val	Gly	Trp	Tyr	Val	Gly	Val	Cys	Arg	Asp
				350					355					360
Asp	Val	Asp	Arg	Gly	Lys	Asn	Asn	Val	Thr	Leu	Ser	Pro	Asn	Asn
				365					370					375
Gly	Tyr	Trp	Val	Leu	Arg	Leu	Thr	Thr	Glu	His	Leu	Tyr	Phe	Thr
				380					385					390
Phe	Asn	Pro	His	Phe	Ile	Ser	Leu	Pro	Pro	Ser	Thr	Pro	Pro	Thr
				395					400					405
Arg	Val	Gly	Val	Phe	Leu	Asp	Tyr	Glu	Gly	Gly	Thr	Ile	Ser	Phe
				410					415					420
Phe	Asn	Thr	Asn	Asp	Gln	Ser	Leu	Ile	Tyr	Thr	Leu	Leu	Thr	Cys
				425					430					435
Gln	Phe	Glu	Gly	Leu	Leu	Arg	Pro	Tyr	Ile	Gln	His	Ala	Met	Tyr
				440					445					450
Asp	Glu	Glu	Lys	Gly	Thr	Pro	Ile	Phe	Ile	Cys	Pro	Val	Ser	Trp
				455					460					465

Gly

<210> SEQ ID NO 105
<211> LENGTH: 2103
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 105

ccttcacagg	actcttcatt	gctggttggc	aatgatgtat	cggccagatg	50
tggtgagggc	taggaaaaga	gtttgttggg	aaccctgggt	tatcggcctc	100
gtcatcttca	tatccctgat	tgtcctggca	gtgtgcattg	gactcactgt	150
tcattatgtg	agatataatc	aaaagaagac	ctacaattac	tatagcacat	200
tgtcatttac	aactgacaaa	ctatatgctg	agtttggcag	agaggcttct	250
aacaatttta	cagaaatgag	ccagagactt	gaatcaatgg	tgaaaaatgc	300
attttataaa	tctccattaa	gggaagaatt	tgtcaagtct	cagggttatca	350
agttcagtc	acagaagcat	ggagtgttgg	ctcatatgct	gttgatttgt	400
agatttcact	ctactgagga	tcctgaaact	gtagataaaa	ttgttcaact	450
tgttttacat	gaaaagctgc	aagatgctgt	aggaccccct	aaagtagatc	500
ctcactcagt	taaaattaaa	aaaatcaaca	agacagaaac	agacagctat	550
ctaaaccatt	gctgcggaac	acgaagaagt	aaaactctag	gtcagagtct	600
caggatcggt	gggtggacag	aagtagaaga	gggtgaatgg	ccctggcagg	650
ctagcctgca	gtgggatggg	agtcatcgct	gtggagcaac	cttaattaat	700
gccacatggc	ttgtgagtgc	tgctcactgt	tttacaacat	ataagaaccc	750
tgccagatgg	actgcttcct	ttggagtaac	aataaaacct	tcgaaaatga	800
aacggggtct	ccggagaata	attgtccatg	aaaaatacaa	acacccatca	850
catgactatg	atatttctct	tgacagcctt	tctagccctg	ttccctacac	900
aatgcagta	catagagttt	gtctccctga	tgcatcctat	gagtttcaac	950

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caggatgatgt gatgtttgtg acaggatttg gagcactgaa aaatgatggt      1000
tacagtcaaa atcatcttcg acaagcacag gtgactctca tagacgctac      1050
aacttgcaat gaacctcaag cttaaatga cgccataact cctagaatgt      1100
tatgtgctgg ctcccttagaa ggaaaaacag atgcatgccca gggtgactct      1150
ggaggaccac tggtagttc agatgctaga gatatctggt accttgctgg      1200
aatagtgagc tggggagatg aatgtgcgaa acccaacaag cctgggtgtt      1250
atactagagt tacggccttg cgggactgga ttacttcaaa aactgggtatc      1300
taagagacaa aagcctcatg gaacagataa cttttttttt tgttttttgg      1350
gtgtggaggc cttttttaga gatacagaat tggagaagac ttgcaaaaca      1400
gctagatttg actgatctca ataaactggt tgcttgatgc atgtattttc      1450
ttcccagctc tgttccgcac gtaagcatcc tgcttctgcc agatcaactc      1500
tgtcatctgt gagcaatagt tgaaacttta tgtacataga gaaatagata      1550
atacaatatt acattacagc ctgtattcat ttgttctcta gaagttttgt      1600
cagaattttg acttgttgac ataaatttgt aatgcatata tacaatttga      1650
agcactcctt ttcttcagtt cctcagctcc tctcatttca gcaaatatcc      1700
attttcaagg tgcagaacaa ggagtgaaag aaaatataag aagaaaaaaaa      1750
tcccctacat tttattggca cagaaaagta ttagggtgtt ttcttagtgg      1800
aatattagaa atgatcatat tcattatgaa aggtcaagca aagacagcag      1850
aataccaatc acttcatcat ttaggaagta tgggaactaa gttaagggaag      1900
tccagaaaga agccaagata tacccttatt ttcatttcca aacaactact      1950
atgataaatg tgaagaagat tctgtttttt tgtgacctat aataattata      2000
caaacttcat gcaatgtact tgttctaagc aaattaaagc aaatatttat      2050
ttaacattgt tactgaggat gtcaacatat aacaataaaa tataaatcac      2100
cca                                                                2103
```

<210> SEQ ID NO 106
<211> LENGTH: 423
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 106

```
Met Met Tyr Arg Pro Asp Val Val Arg Ala Arg Lys Arg Val Cys
 1          5          10          15
Trp Glu Pro Trp Val Ile Gly Leu Val Ile Phe Ile Ser Leu Ile
 20         25         30
Val Leu Ala Val Cys Ile Gly Leu Thr Val His Tyr Val Arg Tyr
 35         40         45
Asn Gln Lys Lys Thr Tyr Asn Tyr Tyr Ser Thr Leu Ser Phe Thr
 50         55         60
Thr Asp Lys Leu Tyr Ala Glu Phe Gly Arg Glu Ala Ser Asn Asn
 65         70         75
Phe Thr Glu Met Ser Gln Arg Leu Glu Ser Met Val Lys Asn Ala
 80         85         90
Phe Tyr Lys Ser Pro Leu Arg Glu Glu Phe Val Lys Ser Gln Val
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95										100										105									
Ile	Lys	Phe	Ser	Gln	Gln	Lys	His	Gly	Val	Leu	Ala	His	Met	Leu															
				110					115					120															
Leu	Ile	Cys	Arg	Phe	His	Ser	Thr	Glu	Asp	Pro	Glu	Thr	Val	Asp															
				125					130					135															
Lys	Ile	Val	Gln	Leu	Val	Leu	His	Glu	Lys	Leu	Gln	Asp	Ala	Val															
				140					145					150															
Gly	Pro	Pro	Lys	Val	Asp	Pro	His	Ser	Val	Lys	Ile	Lys	Lys	Ile															
				155					160					165															
Asn	Lys	Thr	Glu	Thr	Asp	Ser	Tyr	Leu	Asn	His	Cys	Cys	Gly	Thr															
				170					175					180															
Arg	Arg	Ser	Lys	Thr	Leu	Gly	Gln	Ser	Leu	Arg	Ile	Val	Gly	Gly															
				185					190					195															
Thr	Glu	Val	Glu	Glu	Gly	Glu	Trp	Pro	Trp	Gln	Ala	Ser	Leu	Gln															
				200					205					210															
Trp	Asp	Gly	Ser	His	Arg	Cys	Gly	Ala	Thr	Leu	Ile	Asn	Ala	Thr															
				215					220					225															
Trp	Leu	Val	Ser	Ala	Ala	His	Cys	Phe	Thr	Thr	Tyr	Lys	Asn	Pro															
				230					235					240															
Ala	Arg	Trp	Thr	Ala	Ser	Phe	Gly	Val	Thr	Ile	Lys	Pro	Ser	Lys															
				245					250					255															
Met	Lys	Arg	Gly	Leu	Arg	Arg	Ile	Ile	Val	His	Glu	Lys	Tyr	Lys															
				260					265					270															
His	Pro	Ser	His	Asp	Tyr	Asp	Ile	Ser	Leu	Ala	Glu	Leu	Ser	Ser															
				275					280					285															
Pro	Val	Pro	Tyr	Thr	Asn	Ala	Val	His	Arg	Val	Cys	Leu	Pro	Asp															
				290					295					300															
Ala	Ser	Tyr	Glu	Phe	Gln	Pro	Gly	Asp	Val	Met	Phe	Val	Thr	Gly															
				305					310					315															
Phe	Gly	Ala	Leu	Lys	Asn	Asp	Gly	Tyr	Ser	Gln	Asn	His	Leu	Arg															
				320					325					330															
Gln	Ala	Gln	Val	Thr	Leu	Ile	Asp	Ala	Thr	Thr	Cys	Asn	Glu	Pro															
				335					340					345															
Gln	Ala	Tyr	Asn	Asp	Ala	Ile	Thr	Pro	Arg	Met	Leu	Cys	Ala	Gly															
				350					355					360															
Ser	Leu	Glu	Gly	Lys	Thr	Asp	Ala	Cys	Gln	Gly	Asp	Ser	Gly	Gly															
				365					370					375															
Pro	Leu	Val	Ser	Ser	Asp	Ala	Arg	Asp	Ile	Trp	Tyr	Leu	Ala	Gly															
				380					385					390															
Ile	Val	Ser	Trp	Gly	Asp	Glu	Cys	Ala	Lys	Pro	Asn	Lys	Pro	Gly															
				395					400					405															
Val	Tyr	Thr	Arg	Val	Thr	Ala	Leu	Arg	Asp	Trp	Ile	Thr	Ser	Lys															
				410					415					420															
Thr	Gly	Ile																											

<210> SEQ ID NO 107
<211> LENGTH: 2397
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 107

agagaaagaa gcgtctccag ctgaagccaa tgcagccctc cggctctccg

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cgaagaagtt ccctgccccg atgagcccc gccgtgcgtc cccgactatc	100
cccaggcggg cgtggggcac cgggcccagc gccgacgac gctgccgttt	150
tgcccttggg agtaggatgt ggtgaaagga tggggcttct cccttacggg	200
gctcacaatg gccagagaag attccgtgaa gtgtctgcgc tgcctgctct	250
acgcccccaa tctgctcttt tggttaatgt ccatcagtgt gttggcagtt	300
tctgcttgga tgagggacta cctaaataat gttctcactt taactgcaga	350
aacgagggta gaggaagcag tcattttgac ttactttcct gtggttcac	400
cggtcacgat tgctgtttgc tgtttcccta tcattgtggg gatgttagga	450
tattgtggaa cggtgaaaaa aaatctgttg cttcttgcat ggtactttgg	500
aagtttgctt gtcatTTTTt gtgtagaact ggcttgtggc gtttgacat	550
atgaacagga acttatgggt ccagtacaat ggtcagatat ggtcactttg	600
aaagccagga tgacaaatta tggattacct agatatcggg ggcttactca	650
tgcttggaat ttttttcaga gagagtttaa gtgctgtgga gtagtatatt	700
tcactgactg gttgaaatg acagagatgg actggcccc agattcctgc	750
tgtgttagag aattcccagg atgttccaaa caggcccacc aggaagatct	800
cagtacactt tatcaagagg gttgtgggaa gaaaaatgtat tcctttttga	850
gaggaaccaa acaactgcag gtgctgaggt ttctgggaat ctccattggg	900
gtgacacaaa tcctggccat gattctcacc attactctgc tctgggctct	950
gtattatgat agaagggagc ctgggacaga ccaaatgatg tccttgaaga	1000
atgacaactc tcagcacctg tcatgtccct cagtagaact gttgaaacca	1050
agcctgtcaa gaatctttga acacacatcc atggcaaaca gctttaatac	1100
acactttgag atggaggagt tataaaaaga aatgtcacag aagaaaacca	1150
caaacttggt ttattggact tgtgaatttt tgagtacata ctatgtgttt	1200
cagaaatatg tagaaataaa aatgttgcca taaaataaca cctaagcata	1250
tactattcta tgctttaaaa tgaggatgga aaagtttcat gtcataagtc	1300
accacttgga caataattga tgccttaaa atgctgaaga cagatgtcat	1350
accactgtg tagcctgtgt atgactttta ctgaacacag ttatgttttg	1400
aggcagcatg gtttgattag catttccgca tccatgcaa cgagtccat	1450
atggtgggac tggagccata gtaaagggtg atttacttct accaactagt	1500
atataaagta ctaattaaat gctaacatag gaagttagaa aataactaata	1550
acttttatta ctacgacatc tattcttctg atgctaaata aattatatat	1600
cagaaaactt tcaatatggg tgactaccta aatgtgattt ttgctgggta	1650
ctaaaaatatt cttaccactt aaaagagcaa gctaacacat tgtcttaagc	1700
tgatcaggga ttttttgat ataagtctgt gttaaactctg tataattcag	1750
tcgatttcag ttctgataat gtaagaata accattatga aaaggaaaat	1800
ttgtcctgta tagcatcatt attttagcc ttctctgtta ataaagcttt	1850
actattctgt cctgggctta tattacacat ataactgtta tttaaatact	1900
taaccactaa ttttgaaaat taccagtgtg atacatagga atcattattc	1950

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agaatgtagt ctggtcttta ggaagtatta ataagaaaat ttgcacataa      2000
cttagttgat tcagaaagga cttgtatgct gtttttctcc caaatgaaga      2050
ctctttttga cactaaacac tttttaaaaa gcttatcttt gccttctcca      2100
aacaagaagc aatagtctcc aagtcaatat aaattctaca gaaaatagtg      2150
ttctttttct ccagaaaaat gcttgtgaga atcattaaaa catgtgacaa      2200
tttagagatt ctttgtttta ttctactgat taatatactg tggcaaatta      2250
cacagattat taaatTTTTT tacaagagta tagtatattt atttgaaatg      2300
ggaaaagtgc attttactgt attttgtgta ttttgtttat ttctcagaat      2350
atggaagaa aattaaaatg tgtcaataaa tattttctag agagtaa      2397

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<210> SEQ ID NO 108

<211> LENGTH: 305

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 108

```

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

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	245		250		255
Gly Thr Asp Gln Met Met Ser Leu Lys Asn Asp Asn Ser Gln His					
	260		265		270
Leu Ser Cys Pro Ser Val Glu Leu Leu Lys Pro Ser Leu Ser Arg					
	275		280		285
Ile Phe Glu His Thr Ser Met Ala Asn Ser Phe Asn Thr His Phe					
	290		295		300
Glu Met Glu Glu Leu					
	305				

<210> SEQ ID NO 109
<211> LENGTH: 2339
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 109

ccaaggccag agctgtggac accttatccc actcatcctc atcctcttcc	50
tctgataaag cccctaccag tgctgataaa gtctttctcg tgagagccta	100
gaggccttaa aaaaaaagt gcttgaaaga gaaggggaca aaggaacacc	150
agtattaaga ggattttcca gtgtttctgg cagttggtcc agaaggatgc	200
ctccattcct gcttctcacc tgctcttcca tcacaggcac ctccgtgtca	250
cccgtggccc tagatccttg ttctgcttac atcagcctga atgagccctg	300
gaggaacct gaccaccagt tggatgagtc tcaaggtcct cctctatgtg	350
acaacctatg gaatggggag tggatccact tcacgggcat ggcgggagat	400
gccatgccta ccttctgcat accagaaaac cactgtggaa cccacgcacc	450
tgtctggctc aatggcagcc accccctaga aggcgacggc attgtgcaac	500
gccaggcttg tgccagcttc aatgggaact gctgtctctg gaacaccacg	550
gtggaagtca aggcttgccc tggaggctac tatgtgtatc gtctgaccaa	600
gcccagcgtc tgcttccacg tctactgtgg tcatttttat gacatctgcg	650
acgaggactg ccatggcagc tgctcagata ccagcgagtg cacatgcgct	700
ccaggaactg tgctaggccc tgacaggcag acatgctttg atgaaaatga	750
atgtgagcaa aacaacggtg gctgcagtga gatctgtgtg aacctcaaaa	800
actcctaccg ctgtgagtgt ggggttggcc gtgtgctaag aagtgatggc	850
aagacttgtg aagacgttga aggatgccac aataacaatg gtggctgcag	900
ccactcttgc cttggatctg agaaaggcta ccagtgtgaa tgtccccggg	950
gcctgggtgct gtctgaggat aaccacactt gccaaagccc tgtgttgtgc	1000
aaatcaaatg ccattgaagt gaacatcccc agggagctgg ttgggtggcct	1050
ggagctcttc ctgaccaaca cctcctgccg aggagtgtcc aacggcacc	1100
atgtcaacat cctcttctct ctcaagacat gtggtacagt ggtcgatgtg	1150
gtgaatgaca agattgtggc cagcaacctc gtgacagtc taccgaagca	1200
gaccccgggg agcagcgggg acttcatcat ccgaaccagc aagctgctga	1250
tcccggtgac ctgcgagttt ccacgcctgt acaccatttc tgaaggatac	1300
gttcccaacc ttcgaaactc cccactggaa atcatgagcc gaaatcatgg	1350

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gatcttccca ttcactctgg agatcttcaa ggacaatgag ttggaagagc	1400
cttaccggga agctctgccc accctcaagc ttcgtgactc cctctacttt	1450
ggcattgagc ccgtggtgca cgtgagcggc ttggaaagct tggtgagagc	1500
ctgctttgcc acccccacct ccaagatcga cgaggctctg aaatactacc	1550
tcatccggga tggctgtggt tcagatgact cggtaaagca gtacacatcc	1600
cgggatcacc tagcaaagca cttccaggtc cctgtcttca agtttgtggg	1650
caaagaccac aaggaagtgt ttctgcactg ccgggttctt gtctgtggag	1700
tgttggacga gcgttccgc tgtgccagg gttgccaccg gcgaatgcgt	1750
cgtggggcag gaggagagga ctacagccgt ctacagggcc agacgctaac	1800
aggcgggccg atccgcacgc actgggagga ctagtctgta gccatactc	1850
gagtcctgc attggacggc tctgtctttt ggagcttctc cccccaccgc	1900
cctctaagaa catctgcaa cagctgggtt cagacttcac actgtgagtt	1950
cagactccca gcaccaactc actctgattc tggtcattc agtgggcaca	2000
ggtcacagca ctgctgaaca atgtggcctg ggtggggttt catctttcta	2050
gggttgaaaa ctaaactgtc caccagaaa gacactcacc ccatttcct	2100
cattttcttc ctacacttaa atacctcgtg tatggtgcaa tcagaccaca	2150
aaatcagaag ctgggtataa tatttcaagt taaaaccct agaaaaatta	2200
aacagt tact gaaattatga cttaaatacc caatgactcc ttaaatatgt	2250
aaattatagt tataccttga aatttcaatt caaatgcaga ctaattatag	2300
ggaatttgga agtgtatcaa taaaacagta tataatttt	2339

<210> SEQ ID NO 110
<211> LENGTH: 545
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 110

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

-continued

140					145					150				
Cys	Asp	Glu	Asp	Cys	His	Gly	Ser	Cys	Ser	Asp	Thr	Ser	Glu	Cys
				155					160					165
Thr	Cys	Ala	Pro	Gly	Thr	Val	Leu	Gly	Pro	Asp	Arg	Gln	Thr	Cys
				170					175					180
Phe	Asp	Glu	Asn	Glu	Cys	Glu	Gln	Asn	Asn	Gly	Gly	Cys	Ser	Glu
				185					190					195
Ile	Cys	Val	Asn	Leu	Lys	Asn	Ser	Tyr	Arg	Cys	Glu	Cys	Gly	Val
				200					205					210
Gly	Arg	Val	Leu	Arg	Ser	Asp	Gly	Lys	Thr	Cys	Glu	Asp	Val	Glu
				215					220					225
Gly	Cys	His	Asn	Asn	Asn	Gly	Gly	Cys	Ser	His	Ser	Cys	Leu	Gly
				230					235					240
Ser	Glu	Lys	Gly	Tyr	Gln	Cys	Glu	Cys	Pro	Arg	Gly	Leu	Val	Leu
				245					250					255
Ser	Glu	Asp	Asn	His	Thr	Cys	Gln	Val	Pro	Val	Leu	Cys	Lys	Ser
				260					265					270
Asn	Ala	Ile	Glu	Val	Asn	Ile	Pro	Arg	Glu	Leu	Val	Gly	Gly	Leu
				275					280					285
Glu	Leu	Phe	Leu	Thr	Asn	Thr	Ser	Cys	Arg	Gly	Val	Ser	Asn	Gly
				290					295					300
Thr	His	Val	Asn	Ile	Leu	Phe	Ser	Leu	Lys	Thr	Cys	Gly	Thr	Val
				305					310					315
Val	Asp	Val	Val	Asn	Asp	Lys	Ile	Val	Ala	Ser	Asn	Leu	Val	Thr
				320					325					330
Gly	Leu	Pro	Lys	Gln	Thr	Pro	Gly	Ser	Ser	Gly	Asp	Phe	Ile	Ile
				335					340					345
Arg	Thr	Ser	Lys	Leu	Leu	Ile	Pro	Val	Thr	Cys	Glu	Phe	Pro	Arg
				350					355					360
Leu	Tyr	Thr	Ile	Ser	Glu	Gly	Tyr	Val	Pro	Asn	Leu	Arg	Asn	Ser
				365					370					375
Pro	Leu	Glu	Ile	Met	Ser	Arg	Asn	His	Gly	Ile	Phe	Pro	Phe	Thr
				380					385					390
Leu	Glu	Ile	Phe	Lys	Asp	Asn	Glu	Phe	Glu	Glu	Pro	Tyr	Arg	Glu
				395					400					405
Ala	Leu	Pro	Thr	Leu	Lys	Leu	Arg	Asp	Ser	Leu	Tyr	Phe	Gly	Ile
				410					415					420
Glu	Pro	Val	Val	His	Val	Ser	Gly	Leu	Glu	Ser	Leu	Val	Glu	Ser
				425					430					435
Cys	Phe	Ala	Thr	Pro	Thr	Ser	Lys	Ile	Asp	Glu	Val	Leu	Lys	Tyr
				440					445					450
Tyr	Leu	Ile	Arg	Asp	Gly	Cys	Val	Ser	Asp	Asp	Ser	Val	Lys	Gln
				455					460					465
Tyr	Thr	Ser	Arg	Asp	His	Leu	Ala	Lys	His	Phe	Gln	Val	Pro	Val
				470					475					480
Phe	Lys	Phe	Val	Gly	Lys	Asp	His	Lys	Glu	Val	Phe	Leu	His	Cys
				485					490					495
Arg	Val	Leu	Val	Cys	Gly	Val	Leu	Asp	Glu	Arg	Ser	Arg	Cys	Ala
				500					505					510
Gln	Gly	Cys	His	Arg	Arg	Met	Arg	Arg	Gly	Ala	Gly	Gly	Glu	Asp
				515					520					525

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Ser Ala Gly Leu Gln Gly Gln Thr Leu Thr Gly Gly Pro Ile Arg
530 535 540
Ile Asp Trp Glu Asp
545

<210> SEQ ID NO 111
<211> LENGTH: 2063
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 111

gagagagggca gcagcttgct cagcggacaa ggatgctggg cgtgagggac	50
caaggcctgc cctgcactcg ggccctctcc agccagtgct gaccagggac	100
ttctgacctg ctggccagcc aggacctgtg tggggaggcc ctctgtctgc	150
cttgggggtg caatctcagc tccaggctac agggagaccg ggaggatcac	200
agagccagca tgttacagga tcctgacagt gatcaacctc tgaacagcct	250
cgatgtcaaa cccctgcgca aaccccgat ccccatggag accttcagaa	300
aggtggggat ccccatcatc atagcactac tgagcctggc gagtatcatc	350
attgtggttg tcctcatcaa ggtgattctg gataaatact acttcctctg	400
cgggcagcct ctccacttca tcccaggaa gcagctgtgt gacggagagc	450
tggactgtcc cttgggggag gacgaggagc actgtgtcaa gagottcccc	500
gaagggcctg cagtggcagt ccgcctctcc aaggaccgat ccacactgca	550
ggtgctggac tcggccacag ggaactggtt ctctgcctgt ttcgacaact	600
tcacagaagc tctcgctgag acagcctgta ggcagatggg ctacagcaga	650
gctgtggaga ttggcccaga ccaggatctg gatgtgttg aaatcacaga	700
aaacagccag gagcttcgca tgcggaactc aagtggggcc tgtctctcag	750
gtccctcgtt ctccctgcac tgtcttcgct gtgggaagag cctgaagacc	800
ccccgtgtgg tgggtgggga ggaggcctct gtggattctt ggccttgga	850
ggtcagcatc cagtacgaca aacagcacgt ctgtggaggg agcatcctgg	900
acccccactg ggtcctcacg gcagccact gcttcaggaa acataccgat	950
gtgttcaact ggaaggtgcg ggcaggctca gacaaactgg gcagcttccc	1000
atccctggct gtggccaaga tcatcatcat tgaattcaac cccatgtacc	1050
ccaaagacaa tgacatcgcc ctcatgaagc tgcagttccc actcactttc	1100
tcaggcacag tcaggcccat ctgtctgccc ttctttgatg aggagctcac	1150
tccagccacc ccaactctga tcattggatg gggctttacg aagcagaatg	1200
gagggaagat gtctgacata ctgctgcagg cgtcagtcca ggtcattgac	1250
agcacacggt gcaatgcaga cgatgcgtac cagggggaag tcaccgagaa	1300
gatgatgtgt gcaggcatcc cggaaggggg tgtggacacc tgccaggggtg	1350
acagtgggtg gccctgatg taccaatctg accagtggca tgtggtgggc	1400
atcgttagct ggggctatgg ctgcgggggc ccgagcacc caggagtata	1450
caccaaggtc tcagcctatc tcaactggat ctacaatgtc tggaaggctg	1500
agctgtaatg ctgctgcccc ttgcagtgc tgggagccgc ttccttctctg	1550

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ccctgcccac ctggggatcc cccaaagtca gacacagagc aagagtcccc      1600
ttgggtacac ccctctgccc acagcctcag catttcttg agcagcaaag      1650
ggcctcaatt cctgtaagag accctcgag cccagaggcg cccagaggaa      1700
gtcagcagcc ctgctcggc cacacttggt gctcccagca tcccaggag      1750
agacacagcc cactgaacaa ggtctcaggg gtattgctaa gccaagaagg      1800
aactttccca cactactgaa tggaagcagg ctgtcttgta aaagcccaga      1850
tcactgtggg ctggagagga gaaggaaagg gtctgcgcca gccctgtccg      1900
tcttcaccca tcccgaagcc tactagagca agaaaccagt tgtaataataa      1950
aatgcactgc cctactgttg gtatgactac cgttacctac tgttgtcatt      2000
gttattacag ctatggccac tattattaaa gagctgtgta acatctctgg      2050
caaaaaaaaaaaa aaa      2063

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<210> SEQ ID NO 112

<211> LENGTH: 432

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 112

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

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Gly	Gly	Ser	Ile	Leu	Asp	Pro	His	Trp	Val	Leu	Thr	Ala	Ala	His
				230					235					240
Cys	Phe	Arg	Lys	His	Thr	Asp	Val	Phe	Asn	Trp	Lys	Val	Arg	Ala
				245					250					255
Gly	Ser	Asp	Lys	Leu	Gly	Ser	Phe	Pro	Ser	Leu	Ala	Val	Ala	Lys
				260					265					270
Ile	Ile	Ile	Ile	Glu	Phe	Asn	Pro	Met	Tyr	Pro	Lys	Asp	Asn	Asp
				275					280					285
Ile	Ala	Leu	Met	Lys	Leu	Gln	Phe	Pro	Leu	Thr	Phe	Ser	Gly	Thr
				290					295					300
Val	Arg	Pro	Ile	Cys	Leu	Pro	Phe	Phe	Asp	Glu	Glu	Leu	Thr	Pro
				305					310					315
Ala	Thr	Pro	Leu	Trp	Ile	Ile	Gly	Trp	Gly	Phe	Thr	Lys	Gln	Asn
				320					325					330
Gly	Gly	Lys	Met	Ser	Asp	Ile	Leu	Leu	Gln	Ala	Ser	Val	Gln	Val
				335					340					345
Ile	Asp	Ser	Thr	Arg	Cys	Asn	Ala	Asp	Asp	Ala	Tyr	Gln	Gly	Glu
				350					355					360
Val	Thr	Glu	Lys	Met	Met	Cys	Ala	Gly	Ile	Pro	Glu	Gly	Gly	Val
				365					370					375
Asp	Thr	Cys	Gln	Gly	Asp	Ser	Gly	Gly	Pro	Leu	Met	Tyr	Gln	Ser
				380					385					390
Asp	Gln	Trp	His	Val	Val	Gly	Ile	Val	Ser	Trp	Gly	Tyr	Gly	Cys
				395					400					405
Gly	Gly	Pro	Ser	Thr	Pro	Gly	Val	Tyr	Thr	Lys	Val	Ser	Ala	Tyr
				410					415					420
Leu	Asn	Trp	Ile	Tyr	Asn	Val	Trp	Lys	Ala	Glu	Leu			
				425					430					

<210> SEQ ID NO 113
<211> LENGTH: 1768
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 113

ggctggactg	gaactcctgg	tcccaagtga	tccacccgcc	tcagcctccc	50
aaggtgctgt	gattataggt	gtaagccacc	gtgtctggcc	tctgaacaac	100
tttttcagca	actaaaaaag	ccacaggagt	tgaactgcta	ggattctgac	150
tatgctgtgg	tggctagtgc	tcctactcct	acctacatta	aaatctgttt	200
tttgttctct	tgtaactagc	ctttaccctc	ctaacacaga	ggatctgtca	250
ctgtggctct	ggcccaaacc	tgaccttcac	tctggaacga	gaacagaggt	300
ttctaccac	accgtcccct	cgaagccggg	gacagcctca	ccttgctggc	350
ctctcgctgg	agcagtggcc	tcaccaactg	tctcacgtct	ggaggcactg	400
actcgggcag	tgcaggtagc	tgagcctctt	ggtagctgcg	gctttcaagg	450
tgggccttgc	cctggccgta	gaagggattg	acaagccoga	agatttcata	500
ggcgatggct	cccactggcc	aggcatcagc	cttgctgtag	tcaatcactg	550
ccctggggcc	aggacggggc	gtggacacct	gctcagaagc	agtgggtgag	600
acatcacgct	gccgcgccat	ctaacctttt	catgtcctgc	acatcacctg	650

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atccatgggc taatctgaac tctgtcccaa ggaaccacaga gcttgagtga	700
gctgtggctc agaccacaga ggggtctgct tagaccacct ggtttatgtg	750
acaggacttg cattctcctg gaacatgagg gaacgccgga ggaagcaaaa	800
gtggcaggga aggaacttgt gccaaattat gggtcagaaa agatggaggt	850
gttgggttat cacaaggcat cgagtctcct gcattcagtg gacatgtggg	900
ggaagggtcg ccgatggcgc atgacacact cgggactcac ctctggggcc	950
atcacagacg cgtttccgcc ccgatccacg taccagctgc tgaagggcaa	1000
ctgcaggccg atgctctcat cagccaggca gcagccaaaa tctgcgatca	1050
ccagccaggg gcagccgtct gggaaggagc aagcaaagt accatttctc	1100
ctccctctct tccctctgag aggcctctct atgtccctac taaagccacc	1150
agcaagacat agctgacagg ggctaattgg tcagtgttgg cccaggaggt	1200
cagcaaggcc tgagagctga tcagaagggc ctgctgtgcg aacacggaaa	1250
tgctctcagt aagcacaggc tgcaaatcc ccaggcaaag gactgtgtgg	1300
ctcaatttaa atcatgttct agtaattgga gctgtcccca agaccaaagg	1350
agctagagct tggttcaaat gatctccaag ggccttata ccccaggaga	1400
ctttgatttg aatttgaaac cccaaatcca aacctaagaa ccagggtgat	1450
taagaatcag ttattgccg gtgtggtggc ctgtaatgcc aacattttgg	1500
gaggccgagg cgggtagatc acctgaggtc aggagttaa gaccagcctg	1550
gccaacatgg tgaaaccctt gtctctacta aaaatacaaa aaaactagcc	1600
aggcatggtg gtgtgtgcct gtatccacg tactcgggag gctgagacag	1650
gagaattact tgaacctggg aggtgaagga ggctgagaca ggagaatcac	1700
ttcagcctga gcaacacagc gagactctgt ctcagaaaaa ataaaaaaag	1750
aattatgggt atttgtaa	1768

<210> SEQ ID NO 114
<211> LENGTH: 109
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 114

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

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<210> SEQ ID NO 115
<211> LENGTH: 1197
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 115

cagcagtggt ctctcagtc tctcaaagca aggaaagagt actgtgtgct	50
gagagaccat ggcaaagaat cctccagaga attgtgaaga ctgtcacatt	100
ctaaatgcag aagcttttaa atccaagaaa atatgtaaat cacttaagat	150
ttgtggactg gtgtttggt tcttgccct aactctaatt gtcctgtttt	200
gggggagcaa gcacttctgg ccggagggtac ccaaaaaagc ctatgacatg	250
gagcacactt tctacagcaa tggagagaag aagaagattt acatggaaat	300
tgatcctgtg accagaactg aaatattcag aagcggaaat ggcactgatg	350
aaacattgga agtgcacgac tttaaaaacg gatacactgg catctacttc	400
gtgggtcttc aaaaatgttt tatcaaaact cagattaaag tgattcctga	450
attttctgaa ccagaagagg aaatagatga gaatgaagaa attaccacaa	500
ctttcttga acagtcatg atttgggtcc cagcagaaaa gcctattgaa	550
aaccgagatt ttcttaaaaa ttccaaaatt ctggagattt gtgataacgt	600
gaccatgtat tggatcaatc cactctaata atcagtttct gagttacaag	650
actttgagga ggaggagaa gatcttcaat ttcttgccaa cgaaaaaaaa	700
gggattgaac aaaaatgaaca gtgggtggtc cctcaagtga aagtagagaa	750
gacccgtcac gccagacaag caagtgagga agaacttcca ataatgact	800
atactgaaaa tggaatagaa ttgatccca tgctggatga gagaggttat	850
tgttgtattt actgccgtcg aggcaaccgc tattgccgcc gcgtctgtga	900
acctttacta ggctactacc catatccata ctgctaccaa ggaggacgag	950
tcatctgtcg tgtcatcatg ccttgtaact ggtgggtggc ccgcatgctg	1000
gggagggtct aataggaggt ttgagctcaa atgcttaaac tgctggcaac	1050
atataataaa tgcattctat tcaatgaatt tctgcctatg aggcattctg	1100
cccctggtag ccagctctcc agaattactt gtaggtaatt cctctcttca	1150
tgttctaata aacttctaca ttatcaccaa aaaaaaaaaa aaaaaaa	1197

<210> SEQ ID NO 116
<211> LENGTH: 317
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 116

Met Ala Lys Asn Pro Pro Glu Asn Cys Glu Asp Cys His Ile Leu	
1 5 10 15	
Asn Ala Glu Ala Phe Lys Ser Lys Lys Ile Cys Lys Ser Leu Lys	
20 25 30	
Ile Cys Gly Leu Val Phe Gly Ile Leu Ala Leu Thr Leu Ile Val	
35 40 45	
Leu Phe Trp Gly Ser Lys His Phe Trp Pro Glu Val Pro Lys Lys	
50 55 60	

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Ala	Tyr	Asp	Met	Glu	His	Thr	Phe	Tyr	Ser	Asn	Gly	Glu	Lys	Lys	
			65						70					75	
Lys	Ile	Tyr	Met	Glu	Ile	Asp	Pro	Val	Thr	Arg	Thr	Glu	Ile	Phe	
			80						85					90	
Arg	Ser	Gly	Asn	Gly	Thr	Asp	Glu	Thr	Leu	Glu	Val	His	Asp	Phe	
			95						100					105	
Lys	Asn	Gly	Tyr	Thr	Gly	Ile	Tyr	Phe	Val	Gly	Leu	Gln	Lys	Cys	
			110						115					120	
Phe	Ile	Lys	Thr	Gln	Ile	Lys	Val	Ile	Pro	Glu	Phe	Ser	Glu	Pro	
			125						130					135	
Glu	Glu	Glu	Ile	Asp	Glu	Asn	Glu	Glu	Ile	Thr	Thr	Thr	Phe	Phe	
			140						145					150	
Glu	Gln	Ser	Val	Ile	Trp	Val	Pro	Ala	Glu	Lys	Pro	Ile	Glu	Asn	
			155						160					165	
Arg	Asp	Phe	Leu	Lys	Asn	Ser	Lys	Ile	Leu	Glu	Ile	Cys	Asp	Asn	
			170						175					180	
Val	Thr	Met	Tyr	Trp	Ile	Asn	Pro	Thr	Leu	Ile	Ser	Val	Ser	Glu	
			185						190					195	
Leu	Gln	Asp	Phe	Glu	Glu	Glu	Gly	Glu	Asp	Leu	His	Phe	Pro	Ala	
			200						205					210	
Asn	Glu	Lys	Lys	Gly	Ile	Glu	Gln	Asn	Glu	Gln	Trp	Val	Val	Pro	
			215						220					225	
Gln	Val	Lys	Val	Glu	Lys	Thr	Arg	His	Ala	Arg	Gln	Ala	Ser	Glu	
			230						235					240	
Glu	Glu	Leu	Pro	Ile	Asn	Asp	Tyr	Thr	Glu	Asn	Gly	Ile	Glu	Phe	
			245						250					255	
Asp	Pro	Met	Leu	Asp	Glu	Arg	Gly	Tyr	Cys	Cys	Ile	Tyr	Cys	Arg	
			260						265					270	
Arg	Gly	Asn	Arg	Tyr	Cys	Arg	Arg	Val	Cys	Glu	Pro	Leu	Leu	Gly	
			275						280					285	
Tyr	Tyr	Pro	Tyr	Pro	Tyr	Cys	Tyr	Gln	Gly	Gly	Arg	Val	Ile	Cys	
			290						295					300	
Arg	Val	Ile	Met	Pro	Cys	Asn	Trp	Trp	Val	Ala	Arg	Met	Leu	Gly	
			305						310					315	
Arg	Val														

<210> SEQ ID NO 117
<211> LENGTH: 2121
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 117

gagctccctc caggagcgcg ttagcttcac accttcgcca gcaggagggc	50
ggcagcttct cgcaggcggc agggcgggcg gccaggatca tgtccaccac	100
cacatgccaa gtggtggcgt tcctcctgtc catcctgggg ctggccggct	150
gcacgcggc caccgggatg gacatgtgga gcaccagga cctgtacgac	200
aaccccgta cctccgtgtt ccagtacgaa gggctctgga ggagctgcgt	250
gaggcagagt tcaggcttca ccgaatgcag gccctatttc accatcctgg	300
gacttcagc catgctgcag gcagtgcgag ccctgatgat cgtaggcac	350

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gtcctgggtg ccatggcct cctggtatcc atctttgccc tgaatgcat	400
ccgcattggc agcatggagg actctgcaa agccaacatg aactgacct	450
ccggatcat gttcattgtc tcaggctctt gtgcaattgc tggagtgtct	500
gtgtttgcca acatgctggt gactaacttc tggatgtcca cagctaact	550
gtacaccggc atgggtggga tgggtgcagac tgttcagacc aggtacacat	600
ttggtgcggc tctgttcgtg ggctgggtcg ctggaggcct cacactaatt	650
gggggtgtga tgatgtgcat cgctgccgg ggctggcac cagaagaaac	700
caactacaaa gccgtttctt atcatgcctc aggccacagt gttgcctaca	750
agcctggagg cttcaaggcc agcactggct ttgggtccaa caccaaaaac	800
aagaagatat acgatggagg tgccgcaca gaggacgagg tacaatctta	850
tccttccaag cactgactatg tgtaatgctc taagacctct cagcacgggc	900
ggaagaaact cccggagagc tcacccaaaa aacaaggaga tcccatctag	950
atttcttctt gcttttgact cacagctgga agttagaaaa gcctcgattt	1000
catctttgga gaggccaaat ggtcttagcc tcagtctctg tctctaaata	1050
ttccaccata aaacagctga gttatttatg aattagaggc tatagctcac	1100
attttcaatc ctctatttct ttttttaaat ataactttct actctgatga	1150
gagaatgtgg ttttaatctc tctctcacat tttgatgatt tagacagact	1200
ccccctcttc ctctagtgca ataaacccat tgatgatcta tttcccagct	1250
tatccccaag aaaacttttg aaaggaaaga gtagacccaa agatgttatt	1300
ttctgctgtt tgaattttgt ctccccaccc ccaacttggc tagtaataaa	1350
cacttactga agaagaagca ataagagaaa gatatttgta atctctccag	1400
cccagtatct cggttttctt aactgtgat cttaaaagtt accaaaccaa	1450
agtcattttc agtttgaggc aaccaaacct ttctactgct gttgacatct	1500
tcttattaca gcaacacccat tctaggagtt tcctgagctc tccactggag	1550
tcctctttct gtcgcggtc agaaattgtc cctagatgaa tgagaaaatt	1600
atttttttta atttaagtcc taaatatagt taaaataaat aatgttttag	1650
taaaatgata cactatctct gtgaaatagc ctcccccta catgtggata	1700
gaaggaaatg aaaaaataat tgctttgaca ttgtctatat ggtactttgt	1750
aaagtcatgc ttaagtacaa attccatgaa aagctcacac ctgtaatcct	1800
agcactttgg gaggtgagg aggaaggatc acttgagccc agaagttcga	1850
gactagcctg ggcaacatgg agaagccctg tctctacaaa atacagagag	1900
aaaaaatcag ccagtcatgg tggcatacac ctgtagtccc agcattccgg	1950
gaggctgagg tgggaggatc acttgagccc agggagggtg gggctgcagt	2000
gagccatgat cacaccactg cactccagcc aggtgacata gcgagatcct	2050
gtctaaaaaa ataaaaata aataatggaa cacagcaagt cctaggaaat	2100
aggttaaaac taattcttta a	2121

<210> SEQ ID NO 118

<211> LENGTH: 261

<212> TYPE: PRT

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<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 118

Met Ser Thr Thr Thr Cys Gln Val Val Ala Phe Leu Leu Ser Ile
1 5 10 15

Leu Gly Leu Ala Gly Cys Ile Ala Ala Thr Gly Met Asp Met Trp
20 25 30

Ser Thr Gln Asp Leu Tyr Asp Asn Pro Val Thr Ser Val Phe Gln
35 40 45

Tyr Glu Gly Leu Trp Arg Ser Cys Val Arg Gln Ser Ser Gly Phe
50 55 60

Thr Glu Cys Arg Pro Tyr Phe Thr Ile Leu Gly Leu Pro Ala Met
65 70 75

Leu Gln Ala Val Arg Ala Leu Met Ile Val Gly Ile Val Leu Gly
80 85 90

Ala Ile Gly Leu Leu Val Ser Ile Phe Ala Leu Lys Cys Ile Arg
95 100 105

Ile Gly Ser Met Glu Asp Ser Ala Lys Ala Asn Met Thr Leu Thr
110 115 120

Ser Gly Ile Met Phe Ile Val Ser Gly Leu Cys Ala Ile Ala Gly
125 130 135

Val Ser Val Phe Ala Asn Met Leu Val Thr Asn Phe Trp Met Ser
140 145 150

Thr Ala Asn Met Tyr Thr Gly Met Gly Gly Met Val Gln Thr Val
155 160 165

Gln Thr Arg Tyr Thr Phe Gly Ala Ala Leu Phe Val Gly Trp Val
170 175 180

Ala Gly Gly Leu Thr Leu Ile Gly Gly Val Met Met Cys Ile Ala
185 190 195

Cys Arg Gly Leu Ala Pro Glu Glu Thr Asn Tyr Lys Ala Val Ser
200 205 210

Tyr His Ala Ser Gly His Ser Val Ala Tyr Lys Pro Gly Gly Phe
215 220 225

Lys Ala Ser Thr Gly Phe Gly Ser Asn Thr Lys Asn Lys Lys Ile
230 235 240

Tyr Asp Gly Gly Ala Arg Thr Glu Asp Glu Val Gln Ser Tyr Pro
245 250 255

Ser Lys His Asp Tyr Val
260

<210> SEQ ID NO 119
<211> LENGTH: 2010
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 119

ggaaaaactg ttctcttctg tggcacagag aacctgctt caaagcagaa	50
gtagcagttc cggagtccag ctggctaaaa ctcatcccag aggataatgg	100
caacccatgc cttagaaatc gctgggctgt ttcttggtgg tgttggaatg	150
gtgggcacag tggctgtcac tgtcatgcct cagtggagag tgtcggcctt	200
cattgaaaac aacatcgtgg tttttgaaaa cttctgggaa ggactgtgga	250

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tgaattgcgt gaggcaggct aacatcagga tgcagtgcaa aatctatgat	300
tccctgctgg ctctttctcc ggacctacag gcagccagag gactgatgtg	350
tgctgcttcc gtgatgtcct tcttgcttt catgatggcc atccttgcca	400
tgaaatgcac caggtgcacg ggggacaatg agaaggtgaa ggctcacatt	450
ctgctgacgg ctggaatcat ctctcatcgc acgggcatgg tggtgctcat	500
ccctgtgagc tgggttgcca atgccatcat cagagatttc tataactcaa	550
tagtgaatgt tgcccaaaaa cgtgagcttg gagaagctct ctacttagga	600
tggaccacgg cactggtgct gattgttgga ggagctctgt tctgctgcgt	650
ttttgttgc aacgaaaaga gcagtagcta cagatactcg atacctccc	700
atcgcacaca ccaaaaaagt tatcacaccg gaaagaagtc accgagcgtc	750
tactccagaa gtcagtatgt gtagttgtgt atgttttttt aactttacta	800
taaagccatg caaatgacaa aaatctatat tactttctca aaatggaccc	850
caaagaaact ttgatttact gttcttaact gctaactctt aattacagga	900
actgtgcac agctatttat gattctataa gctatttcag cagaatgaga	950
tattaaaccc aatgctttga ttgttctaga aagtatagta atttgttttc	1000
taaggtggtt caagcatcta ctctttttat catttacttc aaaatgacat	1050
tgctaagagc tgcattattt tactactgta atttctccac gacatagcat	1100
tatgtacata gatgagtgtg acatttatat ctacataga gacatgctta	1150
tatggtttta tttaaaatga aatgccagtc cattacactg aataaataga	1200
actcaactat tgcttttcag ggaaatcatg gatagggttg aagaaggtta	1250
ctattaattg tttaaaaaca gcttagggat taatgtcctc cattttataat	1300
gaagattaaa atgaaggctt taatcagcat tgtaaaggaa attgaatggc	1350
tttctgatat gctgtttttt agcctaggag ttagaaatcc taacttcottt	1400
atcctcttct cccagaggct ttttttttct tgtgtattaa attaacattt	1450
ttaaaacgca gatattttgt caaggggctt tgcattcaaa ctgcttttcc	1500
agggtatata tcagaagaaa gataaaagtg tgatctaaga aaaagtgatg	1550
gttttaggaa agtgaaaata tttttgtttt tgtatttgaa gaagaatgat	1600
gcattttgac aagaatcat atatgtatgg atatatttta ataagtattt	1650
gagtacagac tttgaggttt catcaatata aataaaagag cagaaaaata	1700
tgtcttggtt ttcatattgct taccaaaaaa acaacaacaa aaaaagttgt	1750
cctttgagaa cttcacctgc tcctatgtgg gtacctgagt caaaattgtc	1800
atttttgttc tgtgaaaaat aaatttcctt cttgtacctt ttctgtttag	1850
ttttactaaa atctgtaaat actgtatttt tctgtttatt ccaaatttga	1900
tgaaactgac aatccaattt gaaagtttgt gtcgacgtct gtctagctta	1950
aatgaatgtg ttctatttgc tttatacatt tatattaata aattgtacat	2000
ttttctaatt	2010

<210> SEQ ID NO 120

<211> LENGTH: 225

<212> TYPE: PRT

-continued

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 120

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

<210> SEQ ID NO 121

<211> LENGTH: 1257

<212> TYPE: DNA

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 121

ggagagaggg gcgcgggtga aaggcgcatg gatgcagcct gcggcggcct 50
cggagcgcgg cggagccaga cgctgaccac gttcctctcc tcggtctcct 100
ccgcctccag ctccgcgctg cccggcagcc gggagccatg cgaccccgag 150
gccccgcgcg ctccccgcag cggtccgcg gcctcctgct gctcctgctg 200
ctgcagctgc ccgcgcgctc gagcgctct gagatcccca aggggaagca 250
aaaggcgag ctccggcaga gggaggtggt ggacctgtat aatggaatgt 300
gcttacaagg gccagcagga gtgcctggtc gagacgggag ccctggggcc 350
aatgttattc cgggtacacc tgggatccca ggtcgggatg gattcaaagg 400
agaaaagggg gaatgtctga gggaaagctt tgaggagtcc tggacacca 450
actacaagca gtgttcattg agttcattga attatggcat agatcttggg 500

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aaaattgcgg agtgtacatt tacaaagatg cgttcaaata gtgctctaag      550
agttttgttc agtggtcac ttcggctaaa atgcagaaat gcatgctgtc      600
agcgttggtta ttccacattc aatggagctg aatgttcagg acctcttccc      650
attgaagcta taattttatt ggaccaagga agccctgaaa tgaattcaac      700
aattaatatt catcgcaactt cttctgtgga aggactttgt gaaggaattg      750
gtgctgggatt agtggatggt gctatctggg ttggcacttg ttcagattac      800
ccaaaaggag atgcttctac tggatggaat tcagtttctc gcatcattat      850
tgaagaacta ccaaataaaa tgctttaatt ttcatttgct acctcttttt      900
ttattatgcc ttggaatggt tcacttaaat gacattttaa ataagtttat      950
gtatacatct gaatgaaaag caaagctaaa tatgtttaca gaccaaagtg     1000
tgatttcaca ctgtttttta atctagcatt attcattttg cttcaatcaa     1050
aagtggtttc aatatttttt ttagttggtt agaatacttt cttcatagtc     1100
acattctctc aacctataat ttggaatatt gttgtggtct tttgtttttt     1150
ctcttagtat agcattttta aaaaaatata aaagctacca atctttgtac     1200
aatttgtaaa tgtaagaat tttttttata tctgttaaat aaaaattatt     1250
tccaaca                                           1257

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<210> SEQ ID NO 122

<211> LENGTH: 243

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 122

```

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

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Gly Ser Pro Glu Met Asn Ser Thr Ile Asn Ile His Arg Thr Ser
185 190 195
Ser Val Glu Gly Leu Cys Glu Gly Ile Gly Ala Gly Leu Val Asp
200 205 210
Val Ala Ile Trp Val Gly Thr Cys Ser Asp Tyr Pro Lys Gly Asp
215 220 225
Ala Ser Thr Gly Trp Asn Ser Val Ser Arg Ile Ile Ile Glu Glu
230 235 240
Leu Pro Lys

<210> SEQ ID NO 123
<211> LENGTH: 2379
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 123

gctgagcgtg tgcgcggtac ggggctctcc tgccttctgg gctccaacgc 50
agctctgtgg ctgaactggg tgctcatcac gggaactgct gggctatgga 100
atacagatgt ggcagctcag gtagcccaa attgctgga agaatacatc 150
atgtttttctg ataagaagaa attgtaggat ccagtttttt ttttaaccgc 200
ccccctccca ccccccaaaa aaactgtaaa gatgcaaaaa cgtaatatcc 250
atgaagatcc tattacctag gaagattttg atgttttgct gcgaatgcgg 300
tgttgggatt tatttgttct tggagtgttc tgcgtggctg gcaaagaata 350
atgttccaaa atcgggtccat ctccaaggg gtccaatttt tcttcctggg 400
tgtcagcgag ccctgactca ctacagtgc gctgacaggg gctgtcatgc 450
aactggcccc taagccaaag caaaagacct aaggacgacc tttgaacaat 500
acaaaggatg ggtttcaatg taattaggct actgagcgga tcagctgtag 550
cactgggttat agccccact gtcttactga caatgctttc tctgcgaa 600
cgaggatgcc ctaagggctg taggtgtgaa ggcaaatgg tatattgtga 650
atctcagaaa ttacaggaga taccctcaag tatactctgct ggttgcttag 700
gtttgtccct tcgctataac agccttcaaa aacttaagta taatcaattt 750
aaagggtcca accagctcac ctggctatac cttgaccata accatatcag 800
caatattgac gaaaatgctt ttaatggaat acgcagactc aaagagctga 850
ttcttagttc caatagaatc tcctattttc ttaacaatac cttcagacct 900
gtgacaaatt tacggaactt ggatctgtcc tataatcagc tgcattctct 950
gggatctgaa cagtttcggg gcttgcgga gctgctgagt ttacatttac 1000
ggctaaactc cctgagaacc atccctgtgc gaattattcca agactgccgc 1050
aacctggaac ttttgacct gggatataac cggatccgaa gtttagccag 1100
gaatgtcttt gctggcatga tcagactcaa agaacttcac ctggagcaca 1150
atcaattttc caagctcaac ctggcccttt ttccaagggt ggtcagcctt 1200
cagaaccttt acttgacgtg gaataaaatc agtgtcatag gacagacct 1250
gtcctggacc tggagctcct tacaaaggct tgatttatca ggcaatgaga 1300
tcgaagcttt cagtggacct agtgttttcc agtgtgtccc gaatctgcag 1350

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cgccccaacc tggattccaa caagctcaca tttattggtc aagagatttt      1400
ggattcttgg atatccctca atgacatcag tcttgctggg aatatatggg      1450
aatgcagcag aaatatattgc tcccttgtaa actggctgaa aagttttaa      1500
ggtctaaggg agaatacaat tatctgtgcc agtcccaaag agctgcaagg      1550
agtaaatgtg atcgatgcag tgaagaacta cagcatctgt ggcaaaagta      1600
ctacagagag gtttgatctg gccaggggctc tcccaaagcc gacgtttaag      1650
cccaagctcc ccaggccgaa gcatgagagc aaacccctt tgccccgac      1700
ggtgaggagc acagagcccg gccagagac cgatgctgac gccgagcaca      1750
tctctttcca taaatcatc gcgggcagcg tggcgctttt cctgtccgtg      1800
ctcgtcatcc tgctggttat ctacgtgtca tggaagcggg accctgcgag      1850
catgaagcag ctgcagcagc gtcctctcat gcgaaggcac aggaaaaaga      1900
aaagacagtc cctaaagcaa atgactccca gcaccagga attttatgta      1950
gattataaac ccaccaacac ggagaccagc gagatgctgc tgaatgggac      2000
gggaccctgc acctataaca aatcgggctc cagggagtgt gaggtatgaa      2050
ccattgtgat aaaaagagct cttaaagct gggaaataag tggtgcttta      2100
ttgaactctg gtgactatca agggaacgcg atgccccccc tccccttccc      2150
tctccctctc actttggtgg caagatcctt cctgtccgt ttagtgcat      2200
tcataatact ggtcattttc ctctcataca taatcaaccc attgaaattt      2250
aaataccaca atcaatgtga agcttgaact ccggtttaat ataataccta      2300
ttgtataaga ccctttactg attccattaa tgcgcattt gttttaagat      2350
aaaacttctt tcataggtaa aaaaaaaaa      2379
```

<210> SEQ ID NO 124

<211> LENGTH: 513

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 124

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

125					130					135				
Arg	Asn	Leu	Asp	Leu	Ser	Tyr	Asn	Gln	Leu	His	Ser	Leu	Gly	Ser
				140					145					150
Glu	Gln	Phe	Arg	Gly	Leu	Arg	Lys	Leu	Leu	Ser	Leu	His	Leu	Arg
				155					160					165
Ser	Asn	Ser	Leu	Arg	Thr	Ile	Pro	Val	Arg	Ile	Phe	Gln	Asp	Cys
				170					175					180
Arg	Asn	Leu	Glu	Leu	Leu	Asp	Leu	Gly	Tyr	Asn	Arg	Ile	Arg	Ser
				185					190					195
Leu	Ala	Arg	Asn	Val	Phe	Ala	Gly	Met	Ile	Arg	Leu	Lys	Glu	Leu
				200					205					210
His	Leu	Glu	His	Asn	Gln	Phe	Ser	Lys	Leu	Asn	Leu	Ala	Leu	Phe
				215					220					225
Pro	Arg	Leu	Val	Ser	Leu	Gln	Asn	Leu	Tyr	Leu	Gln	Trp	Asn	Lys
				230					235					240
Ile	Ser	Val	Ile	Gly	Gln	Thr	Met	Ser	Trp	Thr	Trp	Ser	Ser	Leu
				245					250					255
Gln	Arg	Leu	Asp	Leu	Ser	Gly	Asn	Glu	Ile	Glu	Ala	Phe	Ser	Gly
				260					265					270
Pro	Ser	Val	Phe	Gln	Cys	Val	Pro	Asn	Leu	Gln	Arg	Leu	Asn	Leu
				275					280					285
Asp	Ser	Asn	Lys	Leu	Thr	Phe	Ile	Gly	Gln	Glu	Ile	Leu	Asp	Ser
				290					295					300
Trp	Ile	Ser	Leu	Asn	Asp	Ile	Ser	Leu	Ala	Gly	Asn	Ile	Trp	Glu
				305					310					315
Cys	Ser	Arg	Asn	Ile	Cys	Ser	Leu	Val	Asn	Trp	Leu	Lys	Ser	Phe
				320					325					330
Lys	Gly	Leu	Arg	Glu	Asn	Thr	Ile	Ile	Cys	Ala	Ser	Pro	Lys	Glu
				335					340					345
Leu	Gln	Gly	Val	Asn	Val	Ile	Asp	Ala	Val	Lys	Asn	Tyr	Ser	Ile
				350					355					360
Cys	Gly	Lys	Ser	Thr	Thr	Glu	Arg	Phe	Asp	Leu	Ala	Arg	Ala	Leu
				365					370					375
Pro	Lys	Pro	Thr	Phe	Lys	Pro	Lys	Leu	Pro	Arg	Pro	Lys	His	Glu
				380					385					390
Ser	Lys	Pro	Pro	Leu	Pro	Pro	Thr	Val	Gly	Ala	Thr	Glu	Pro	Gly
				395					400					405
Pro	Glu	Thr	Asp	Ala	Asp	Ala	Glu	His	Ile	Ser	Phe	His	Lys	Ile
				410					415					420
Ile	Ala	Gly	Ser	Val	Ala	Leu	Phe	Leu	Ser	Val	Leu	Val	Ile	Leu
				425					430					435
Leu	Val	Ile	Tyr	Val	Ser	Trp	Lys	Arg	Tyr	Pro	Ala	Ser	Met	Lys
				440					445					450
Gln	Leu	Gln	Gln	Arg	Ser	Leu	Met	Arg	Arg	His	Arg	Lys	Lys	Lys
				455					460					465
Arg	Gln	Ser	Leu	Lys	Gln	Met	Thr	Pro	Ser	Thr	Gln	Glu	Phe	Tyr
				470					475					480
Val	Asp	Tyr	Lys	Pro	Thr	Asn	Thr	Glu	Thr	Ser	Glu	Met	Leu	Leu
				485					490					495
Asn	Gly	Thr	Gly	Pro	Cys	Thr	Tyr	Asn	Lys	Ser	Gly	Ser	Arg	Glu
				500					505					510

-continued

Cys Glu Val

<210> SEQ ID NO 125
<211> LENGTH: 998
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 125

ccgttatcgt cttgcgctac tgctgaatgt ccgtcccgga ggaggaggag	50
aggccttttc cgctgaccca gagatggccc cgagcgagca aattcctact	100
gtccggctgc gcggctaccg tggccgagct agcaaccttt cccctggatc	150
tcacaaaaac tcgactccaa atgcaaggag aagcagctct tgctcggttg	200
ggagacggtg caagagaatc tgcccctat aggggaatgg tgcgcacagc	250
cctagggatc attgaagagg aaggctttct aaagctttgg caaggagtga	300
cacccgccat ttacagacac gtagtgtatt ctggaggtcg aatggtcaca	350
tatgaacatc tccgagaggt tgtgtttggc aaaagtgaag atgagcatta	400
tccccttttg aaatcagtca ttggagggat gatggctggt gttattggcc	450
agtttttagc caatccaact gacctagtga aggttcagat gcaaatggaa	500
ggaaaaagga aactggaagg aaaaccattg cgatttcgtg gtgtacatca	550
tgcatattga aaaaatcttag ctgaaggagg aatacagagg ctttgggcag	600
gctgggtacc caatatacaa agagcagcac tggatgaatat gggagattta	650
accacttatg atacagtga acactacttg gtattgaata caccacttga	700
ggacaatatc atgactcacg gtttatcaag tttatgttct ggactggtag	750
cttctattct gggaacacca gccgatgtca tcaaaagcag aataatgaat	800
caaccacgag ataaacaagg aaggggactt ttgtataaat catcgactga	850
ctgcttgatt caggctgttc aaggtgaagg attcatgagt ctatataaag	900
gctttttacc atcttggtg agaatgacct cttggccaat ggtgttctgg	950
cttacttatg aaaaaatcag agagatgagt ggagtcagtc cattttaa	998

<210> SEQ ID NO 126
<211> LENGTH: 323
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 126

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

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Val Thr Pro Ala Ile Tyr Arg His Val Val Tyr Ser Gly Gly Arg
95 100 105
Met Val Thr Tyr Glu His Leu Arg Glu Val Val Phe Gly Lys Ser
110 115 120
Glu Asp Glu His Tyr Pro Leu Trp Lys Ser Val Ile Gly Gly Met
125 130 135
Met Ala Gly Val Ile Gly Gln Phe Leu Ala Asn Pro Thr Asp Leu
140 145 150
Val Lys Val Gln Met Gln Met Glu Gly Lys Arg Lys Leu Glu Gly
155 160 165
Lys Pro Leu Arg Phe Arg Gly Val His His Ala Phe Ala Lys Ile
170 175 180
Leu Ala Glu Gly Gly Ile Arg Gly Leu Trp Ala Gly Trp Val Pro
185 190 195
Asn Ile Gln Arg Ala Ala Leu Val Asn Met Gly Asp Leu Thr Thr
200 205 210
Tyr Asp Thr Val Lys His Tyr Leu Val Leu Asn Thr Pro Leu Glu
215 220 225
Asp Asn Ile Met Thr His Gly Leu Ser Ser Leu Cys Ser Gly Leu
230 235 240
Val Ala Ser Ile Leu Gly Thr Pro Ala Asp Val Ile Lys Ser Arg
245 250 255
Ile Met Asn Gln Pro Arg Asp Lys Gln Gly Arg Gly Leu Leu Tyr
260 265 270
Lys Ser Ser Thr Asp Cys Leu Ile Gln Ala Val Gln Gly Glu Gly
275 280 285
Phe Met Ser Leu Tyr Lys Gly Phe Leu Pro Ser Trp Leu Arg Met
290 295 300
Thr Pro Trp Ser Met Val Phe Trp Leu Thr Tyr Glu Lys Ile Arg
305 310 315
Glu Met Ser Gly Val Ser Pro Phe
320

<210> SEQ ID NO 127
<211> LENGTH: 1505
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 127

cgcggtatcgg acccaagcag gtcggcgggc gcggcaggag agcgggccggg 50
cgtcagctcc tcgacccccg tgtcgggcta gtccagcgag gcggacgggc 100
ggcgtggggc catggccagg cccggcatgg agcgggtggcg cgaccggctg 150
gcgctgtgta cggggggcctc ggggggcctc ggcgcggccg tggccccggc 200
cctggtccag cagggactga aggtggtggg ctgcgccgc actgtgggca 250
acatcgagga gctggctgct gaatgtaaga gtgcaggcta ccccgggact 300
ttgatccct acagatgta cctatcaaat gaagaggaca tcctctccat 350
gttctcagct atccgttctc agcacagcg tgtagacatc tgcataaca 400
atgctggctt ggcccggcct gacaccctgc tctcaggcag caccagtgg 450
tggaaggaca tgttcaatgt gaacgtgctg gccctcagca tctgcacacg 500

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ggaagcctac cagtcctatga aggagcggaa tgtggacgat gggcacatca	550
ttaacatcaa tagcatgtct ggcacccgag tgttaccctt gtctgtgacc	600
cactttctata gtgccaccaa gtatgccgtc actgcgctga cagagggact	650
gaggcaagag cttcgggagg ccagaccca catccgagcc acgtgcatct	700
ctccaggtgt ggtggagaca caattcgctt tcaaactcca cgacaaggac	750
cctgagaagg cagctgccac ctatgagcaa atgaagtgtc tcaaaccgca	800
ggatgtggcc gaggtgttta tctacgtcct cagcaccccc gcacacatcc	850
agattggaga catccagatg aggcccacg agcaggtgac ctagtgactg	900
tgggagctcc tccttccttc ccaccccttc atggcttgcc tcctgcctct	950
ggattttagg tgttgatttc tggatcacgg gataccactt cctgtccaca	1000
ccccgaccag gggctagaaa atttgtttga gatttttata tcatcttgtc	1050
aaattgcttc agttgtaaat gtgaaaaatg ggctggggaa aggaggtggt	1100
gtccctaatt gttttacttg ttaactgttt cttgtgcccc tgggcacttg	1150
gcctttgtct gctctcagtg tcttcctttt gacatgggaa aggagtgtg	1200
gccaaaatcc ccatcttctt gcacctcaac gtctgtggct cagggctggg	1250
gtggcagagg gaggccttca ccttatatct gtgttggtat ccagggctcc	1300
agacttcctc ctctgcctgc cccactgcac cctctccccc ttatctatct	1350
ccttctcggc tcccagccc agtcttggtt tctgtcccc tcctggggtc	1400
atccctccac tctgactctg actatggcag cagaacacca gggcctggcc	1450
cagtggattt catggtgatc attaaaaaag aaaaatcgca accaaaaaaa	1500
aaaaa	1505

<210> SEQ ID NO 128
<211> LENGTH: 260
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 128

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

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Asn	Val	Leu	Ala	Leu	Ser	Ile	Cys	Thr	Arg	Glu	Ala	Tyr	Gln	Ser
		125							130					135
Met	Lys	Glu	Arg	Asn	Val	Asp	Asp	Gly	His	Ile	Ile	Asn	Ile	Asn
				140					145					150
Ser	Met	Ser	Gly	His	Arg	Val	Leu	Pro	Leu	Ser	Val	Thr	His	Phe
				155					160					165
Tyr	Ser	Ala	Thr	Lys	Tyr	Ala	Val	Thr	Ala	Leu	Thr	Glu	Gly	Leu
				170					175					180
Arg	Gln	Glu	Leu	Arg	Glu	Ala	Gln	Thr	His	Ile	Arg	Ala	Thr	Cys
				185					190					195
Ile	Ser	Pro	Gly	Val	Val	Glu	Thr	Gln	Phe	Ala	Phe	Lys	Leu	His
				200					205					210
Asp	Lys	Asp	Pro	Glu	Lys	Ala	Ala	Ala	Thr	Tyr	Glu	Gln	Met	Lys
				215					220					225
Cys	Leu	Lys	Pro	Glu	Asp	Val	Ala	Glu	Ala	Val	Ile	Tyr	Val	Leu
				230					235					240
Ser	Thr	Pro	Ala	His	Ile	Gln	Ile	Gly	Asp	Ile	Gln	Met	Arg	Pro
				245					250					255
Thr	Glu	Gln	Val	Thr										
				260										

<210> SEQ ID NO 129
<211> LENGTH: 1177
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 129

aacttctaca tgggcctcct gctgctggtg ctcttctccta gcctcctgcc	50
ggtggcctac accatcatgt ccctcccacc ctcccttgac tgcggggccgt	100
tcaggtgcag agtctcagtt gcccgaggc acctcccctc ccgaggcagt	150
ctgctcagag ggctcggcc cagaattcca gttctggttt catgccagcc	200
tgtaaaaggc catggaactt tgggtgaatc accgatgcca tttaagaggg	250
ttttctgcca ggatggaaat gttaggctgt tctgtgtctg cgctgttcat	300
ttcagtagcc accagccacc tgtggccgtt gagtgctga aatgaggaac	350
tgagaaaatt aatttctcat gtatttttct catttattta ttaattttta	400
actgatagtt gtacatatTT gggggtacat gtgatatttg gatacatgta	450
tacaatatat aatgatcaaa tcagggtaac tgggatatcc atcacatcaa	500
acatttattt tttattcttt ttagacagag tctcactctg tcaccagggc	550
tggagtgcag tggtgccatc tcagcttact gcaacctctg cctgccaggt	600
tcaagcgatt ctcatgcctc cacctcccaa gtactgga ctacaggcat	650
gcaccacaat gcccaactaa tttttgtatt tttagtagag acgggggtttt	700
gccatgttgc ccaggctggc cttgaactcc tggcctcaa caatccactt	750
gcctcggcct cccaaagtgt tatgattaca ggcgtgagcc accgtgcctg	800
gcctaaacat ttatcttttc tttgtgttg gaaacttgaa attatacaat	850
gaattattgt taactgtcat ctccctgctg tgctatggaa cactgggact	900
tcttcctct atctaactgt atattgtac cagttaacca accgtacttc	950

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atccccactc ctctctatcc ttccaacct ctgatcacct cattctactc	1000
tctacctcca tgagatccac ttttttagct cccacatgtg agtaagaaaa	1050
tgcaatatatt gtctttctgt gcctggctta ttccacttaa cataatgact	1100
tcctgttcca tccatgttgc tgcaaatgac aggatttcgt tcttaatttc	1150
aattaaaata accacacatg gcaaaaa	1177

<210> SEQ ID NO 130

<211> LENGTH: 111

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 130

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

<210> SEQ ID NO 131

<211> LENGTH: 2061

<212> TYPE: DNA

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 131

ttctgaagta acggaagcta ccttgtataa agacctcaac actgctgacc	50
atgatcagcg cagcctggag catcttcctc atcgggacta aaattgggct	100
gttccttcaa gtagcacctc tatcagttat ggctaaatcc tgtccattctg	150
tgtgtcgtcg cgatgcgggt ttcatttact gtaatgatcg ctttctgaca	200
tccattccaa caggaatacc agaggatgct acaactctct accttcagaa	250
caaccaaata aataatgctg ggattccttc agatttgaaa aacttgctga	300
aagtagaaag aatataccta taccacaaca gtttagatga atttcctacc	350
aacctcccaa agtatgtaaa agagttacat ttgcaagaaa ataacataag	400
gactatcact tatgattcac ttcaaaaat tocctatctg gaagaattac	450
atttagatga caactctgtc tctgcagtta gcatagaaga gggagcattc	500
cgagacagca actatctccg actgcttttc ctgtcccgta atcaccttag	550
cacaattccc tggggtttgc ccaggactat agaagaacta cgcttggtatg	600
ataatcgcat atccactatt tcatcaccat ctcttcaagg tctcactagt	650

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ctaaaacgcc tggttctaga tggaaacctg ttgaacaatc atggtttagg	700
tgacaaagtt ttcttcaacc tagttaattt gacagagctg tccctggtgc	750
ggaattccct gactgctgca ccagtaaacc ttccaggcac aaacctgag	800
aagctttatc ttcaagataa ccacatcaat cgggtgcccc caaatgcttt	850
ttcttatcta aggcagctct atcgactgga tatgtccaat aataacctaa	900
gtaatttacc tcagggtatc ttgatgatt tggacaatat aacacaactg	950
attcttcgca acaatccctg gtattgctgg tgcaagatga aatgggtacg	1000
tgactgggta caatcactac ctgtgaaggc caacgtgcgt gggctcatgt	1050
gccaaagccc agaaaagggt cgtgggatgg ctattaagga tctcaatgca	1100
gaactgtttg attgtaagga cagtgggatt gtaagcacca ttcagataac	1150
cactgcaata cccaacacag tgtatcctgc ccaaggacag tggccagctc	1200
cagtgcacaa acagccagat attaagaacc ccaagctcac taaggatcaa	1250
caaaccacag ggagtccctc aagaaaaaca attacaatta ctgtgaagtc	1300
tgtcacctct gataccattc atatctcttg gaaacttgct ctacctatga	1350
ctgctttgag actcagctgg cttaaactgg gccatagccc ggcatttgga	1400
tctataacag aaacaattgt aacaggggaa cgcagtgagt acttggtcac	1450
agccctggag cctgattcac cctataaagt atgcatgggt cccatggaaa	1500
ccagcaacct ctacctattt gatgaaactc ctgtttgtat tgagactgaa	1550
actgcacccc ttcgaatgta caaccctaca accaccctca atcgagagca	1600
agagaaagaa ccttacaaaa accccaattt acctttggct gccatcattg	1650
gtggggctgt ggccctgggt accattgccc ttcttgcttt agtgtgttg	1700
tatgttcata ggaatggatc gctcttctca aggaactgtg catatagcaa	1750
agggaggaga agaaaggatg actatgcaga agctggcact aagaaggaca	1800
actctatcct ggaaatcagg gaaacttctt ttcagatggt accaataagc	1850
aatgaaccca tctcgaagga ggagtttgta atacacacca tatttcctcc	1900
taatggaatg aatctgtaca aaaacaatca cagtgaagc agtagtaacc	1950
gaagctacag agacagtgtt attccagact cagatcactc acactcatga	2000
tgctgaagga ctcacagcag acttgtgttt tgggtttttt aaacctaagg	2050
gaggtgatgg t	2061

<210> SEQ ID NO 132

<211> LENGTH: 649

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 132

Met	Ile	Ser	Ala	Ala	Trp	Ser	Ile	Phe	Leu	Ile	Gly	Thr	Lys	Ile
1				5					10					15
Gly	Leu	Phe	Leu	Gln	Val	Ala	Pro	Leu	Ser	Val	Met	Ala	Lys	Ser
			20						25					30
Cys	Pro	Ser	Val	Cys	Arg	Cys	Asp	Ala	Gly	Phe	Ile	Tyr	Cys	Asn
			35						40					45
Asp	Arg	Phe	Leu	Thr	Ser	Ile	Pro	Thr	Gly	Ile	Pro	Glu	Asp	Ala

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50					55					60				
Thr	Thr	Leu	Tyr	Leu	Gln	Asn	Asn	Gln	Ile	Asn	Asn	Ala	Gly	Ile
				65					70					75
Pro	Ser	Asp	Leu	Lys	Asn	Leu	Leu	Lys	Val	Glu	Arg	Ile	Tyr	Leu
				80					85					90
Tyr	His	Asn	Ser	Leu	Asp	Glu	Phe	Pro	Thr	Asn	Leu	Pro	Lys	Tyr
				95					100					105
Val	Lys	Glu	Leu	His	Leu	Gln	Glu	Asn	Asn	Ile	Arg	Thr	Ile	Thr
				110					115					120
Tyr	Asp	Ser	Leu	Ser	Lys	Ile	Pro	Tyr	Leu	Glu	Glu	Leu	His	Leu
				125					130					135
Asp	Asp	Asn	Ser	Val	Ser	Ala	Val	Ser	Ile	Glu	Glu	Gly	Ala	Phe
				140					145					150
Arg	Asp	Ser	Asn	Tyr	Leu	Arg	Leu	Leu	Phe	Leu	Ser	Arg	Asn	His
				155					160					165
Leu	Ser	Thr	Ile	Pro	Trp	Gly	Leu	Pro	Arg	Thr	Ile	Glu	Glu	Leu
				170					175					180
Arg	Leu	Asp	Asp	Asn	Arg	Ile	Ser	Thr	Ile	Ser	Ser	Pro	Ser	Leu
				185					190					195
Gln	Gly	Leu	Thr	Ser	Leu	Lys	Arg	Leu	Val	Leu	Asp	Gly	Asn	Leu
				200					205					210
Leu	Asn	Asn	His	Gly	Leu	Gly	Asp	Lys	Val	Phe	Phe	Asn	Leu	Val
				215					220					225
Asn	Leu	Thr	Glu	Leu	Ser	Leu	Val	Arg	Asn	Ser	Leu	Thr	Ala	Ala
				230					235					240
Pro	Val	Asn	Leu	Pro	Gly	Thr	Asn	Leu	Arg	Lys	Leu	Tyr	Leu	Gln
				245					250					255
Asp	Asn	His	Ile	Asn	Arg	Val	Pro	Pro	Asn	Ala	Phe	Ser	Tyr	Leu
				260					265					270
Arg	Gln	Leu	Tyr	Arg	Leu	Asp	Met	Ser	Asn	Asn	Asn	Leu	Ser	Asn
				275					280					285
Leu	Pro	Gln	Gly	Ile	Phe	Asp	Asp	Leu	Asp	Asn	Ile	Thr	Gln	Leu
				290					295					300
Ile	Leu	Arg	Asn	Asn	Pro	Trp	Tyr	Cys	Gly	Cys	Lys	Met	Lys	Trp
				305					310					315
Val	Arg	Asp	Trp	Leu	Gln	Ser	Leu	Pro	Val	Lys	Val	Asn	Val	Arg
				320					325					330
Gly	Leu	Met	Cys	Gln	Ala	Pro	Glu	Lys	Val	Arg	Gly	Met	Ala	Ile
				335					340					345
Lys	Asp	Leu	Asn	Ala	Glu	Leu	Phe	Asp	Cys	Lys	Asp	Ser	Gly	Ile
				350					355					360
Val	Ser	Thr	Ile	Gln	Ile	Thr	Thr	Ala	Ile	Pro	Asn	Thr	Val	Tyr
				365					370					375
Pro	Ala	Gln	Gly	Gln	Trp	Pro	Ala	Pro	Val	Thr	Lys	Gln	Pro	Asp
				380					385					390
Ile	Lys	Asn	Pro	Lys	Leu	Thr	Lys	Asp	Gln	Gln	Thr	Thr	Gly	Ser
				395					400					405
Pro	Ser	Arg	Lys	Thr	Ile	Thr	Ile	Thr	Val	Lys	Ser	Val	Thr	Ser
				410					415					420
Asp	Thr	Ile	His	Ile	Ser	Trp	Lys	Leu	Ala	Leu	Pro	Met	Thr	Ala
				425					430					435

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Leu	Arg	Leu	Ser	Trp	Leu	Lys	Leu	Gly	His	Ser	Pro	Ala	Phe	Gly
				440					445					450
Ser	Ile	Thr	Glu	Thr	Ile	Val	Thr	Gly	Glu	Arg	Ser	Glu	Tyr	Leu
				455					460					465
Val	Thr	Ala	Leu	Glu	Pro	Asp	Ser	Pro	Tyr	Lys	Val	Cys	Met	Val
				470					475					480
Pro	Met	Glu	Thr	Ser	Asn	Leu	Tyr	Leu	Phe	Asp	Glu	Thr	Pro	Val
				485					490					495
Cys	Ile	Glu	Thr	Glu	Thr	Ala	Pro	Leu	Arg	Met	Tyr	Asn	Pro	Thr
				500					505					510
Thr	Thr	Leu	Asn	Arg	Glu	Gln	Glu	Lys	Glu	Pro	Tyr	Lys	Asn	Pro
				515					520					525
Asn	Leu	Pro	Leu	Ala	Ala	Ile	Ile	Gly	Gly	Ala	Val	Ala	Leu	Val
				530					535					540
Thr	Ile	Ala	Leu	Leu	Ala	Leu	Val	Cys	Trp	Tyr	Val	His	Arg	Asn
				545					550					555
Gly	Ser	Leu	Phe	Ser	Arg	Asn	Cys	Ala	Tyr	Ser	Lys	Gly	Arg	Arg
				560					565					570
Arg	Lys	Asp	Asp	Tyr	Ala	Glu	Ala	Gly	Thr	Lys	Lys	Asp	Asn	Ser
				575					580					585
Ile	Leu	Glu	Ile	Arg	Glu	Thr	Ser	Phe	Gln	Met	Leu	Pro	Ile	Ser
				590					595					600
Asn	Glu	Pro	Ile	Ser	Lys	Glu	Glu	Phe	Val	Ile	His	Thr	Ile	Phe
				605					610					615
Pro	Pro	Asn	Gly	Met	Asn	Leu	Tyr	Lys	Asn	Asn	His	Ser	Glu	Ser
				620					625					630
Ser	Ser	Asn	Arg	Ser	Tyr	Arg	Asp	Ser	Gly	Ile	Pro	Asp	Ser	Asp
				635					640					645
His	Ser	His	Ser											

<210> SEQ ID NO 133
<211> LENGTH: 1882
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 133

ccgtcatccc cctgcagcca cccttcccag agtcctttgc ccaggccacc	50
ccaggcttct tggcagccct gccgggccac ttgtcttcat gtctgccagg	100
gggaggtggg aaggaggtgg gaggagggcg tgcagaggca gtctgggctt	150
ggccagagct cagggtgctg agcgtgtgac cagcagtgag cagaggccgg	200
ccatggccag cctggggctg ctgctcctgc tcttactgac agcactgcca	250
ccgctgtggt cctcctcact gcctgggctg gacactgctg aaagtaaagc	300
caccattgca gacctgatcc tgtctgcgct ggagagagcc accgtcttcc	350
tagaacagag gctgcctgaa atcaacctgg atggcatggt gggggtcgga	400
gtgtgtggaag agcagctaaa aagtgtccgg gagaagtggg cccaggagcc	450
cctgtctgac ccgctgagcc tgcgcgtggg gatgctgggg gagaagctgg	500
aggctgccat ccagagatcc ctccactacc tcaagctgag tgatcccaag	550
tacctaagag agttccagct gaccctccag cccgggtttt ggaagctccc	600

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acatgcctgg atccacactg atgcctcctt ggtgtacccc acgttcgggc	650
cccaggactc attctcagag gagagaagtg acgtgtgcct ggtgcagctg	700
ctgggaaccg ggacggacag cagcgagccc tgcggcctct cagacctctg	750
caggagcctc atgaccaagc ccggctgctc aggctactgc ctgtcccacc	800
aactgctctt cttcctctgg gccagaatga ggggatgcac acagggacca	850
ctccaacaga gccaggacta tatcaacctc ttctgcgcca acatgatgga	900
cttgaaccgc agagctgagg ccatcgata cgcctaccct acccgggaca	950
tcttcatgga aaacatcatg ttctgtggaa tgggcggtt ctccgacttc	1000
tacaagctcc ggtggctgga ggccattctc agctggcaga aacagcagga	1050
aggatgcttc ggggagcctg atgctgaaga tgaagaatta tctaaagcta	1100
ttcaatatca gcagcathtt tcgaggagag tgaagaggcg aaaaaacaa	1150
tttccagatt ctcgctctgt tgctcaggct ggagtacagt ggcgcaatct	1200
cggctcactg caacctttgc ctctgggtt caagcaattc tcttgctca	1250
tctcccagag tagctgggac tacaggagcg tgccaccata cctggcta	1300
ttttatattt ttttagtaga gacagggtt catcatgtt ctcatgctg	1350
tctcgaactc ctgatctcaa gagatccgcc cacctcaggc tcccaaagt	1400
tgggattata ggtgtgagcc accgtgtctg gctgaaaagc actttcaaag	1450
agactgtgtt gaataaagg ccaaggttct tgccaccag cactcatggg	1500
ggctctctcc cctagatggc tgctcctccc acaacacagc cacagcagtg	1550
gcagccctgg gtggcttcct atacatcctg gcagaatacc cccagcaaa	1600
cagagagcca caccatcca caccgccacc accaagcagc cgctgagacg	1650
gacggttcca tgccagctgc ctggaggagg aacagacccc tttagtctc	1700
atcccttaga tcctggaggg caccgatcac atcctgggaa gaaggcatct	1750
ggagataaag caaagccacc ccgacacca atcttgaag ccctgagtag	1800
gcagggccag ggtagggtgg gcccgggagg gaccaggtg tgaacggatg	1850
aataaagttc aactgcaact gaaaaaaaa aa	1882

<210> SEQ ID NO 134
<211> LENGTH: 440
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 134

Met Ser Ala Arg Gly Arg Trp Glu Gly Gly Gly Arg Arg Ala Cys	1	5	10	15
Arg Gly Ser Leu Gly Leu Ala Arg Ala Gln Gly Ala Glu Arg Val	20	25	30	
Thr Ser Ser Glu Gln Arg Pro Ala Met Ala Ser Leu Gly Leu Leu	35	40	45	
Leu Leu Leu Leu Leu Thr Ala Leu Pro Pro Leu Trp Ser Ser Ser	50	55	60	
Leu Pro Gly Leu Asp Thr Ala Glu Ser Lys Ala Thr Ile Ala Asp	65	70	75	

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

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<210> SEQ ID NO 135
<211> LENGTH: 884
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 135

ggctctgagtg cagagctgct gtcattggcgg ccgctctgtg gggcttcttt 50
cccgtcctgc tgctgctgct gctatcgggg gatgtccaga gtcggagggt 100
gcccggggct gctgctgagg gatcgggagg gaggggggtc ggcataggag 150
atcgcttcaa gattgagggg cgtgcagttg ttccaggggg gaagcctcag 200
gactggatct cggcggcccg agtgcgtgta gacggagaag agcacgtcgg 250
tttcttaag acagatggga gttttgtggt tcatgatata ctttctggat 300
cttatgtagt ggaagttgta tctccagctt acagatttga tccggttcga 350
gtggatatca cttcgaaagg aaaaatgaga gcaagatatg tgaattacat 400
caaaacatca gaggttgta gactgcccta tctctccaa atgaaatctt 450
caggtccacc ttcttacttt attaaaaggg aatcgtgggg ctggacagac 500
tttctaatag acccaatggt tatgatgatg gttcttcctt tattgatatt 550
tgtgtcttctg cctaaagtgg tcaacacaag tgatcctgac atgagacggg 600
aaatggagca gtcaatgaat atgtgaatt ccaaccatga gttgcctgat 650
gtttctgagt tcatgacaag actcttctct tcaaaatcat ctggcaaata 700
tagcagcggc agcagtaaaa caggcaaaag tggggctggc aaaaggagggt 750
agtcaggccg tccagagctg gcatttgcac aaacacggca aactgggtg 800
gcatccaagt cttgaaaaac cgtgtgaagc aactactata aacttgagtc 850
atccccgacgt tgatctctta caactgtgta tggt 884

<210> SEQ ID NO 136
<211> LENGTH: 242
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 136

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

-continued

					125					130					135
Tyr	Pro	Leu	Gln	Met	Lys	Ser	Ser	Gly	Pro	Pro	Ser	Tyr	Phe	Ile	
				140					145					150	
Lys	Arg	Glu	Ser	Trp	Gly	Trp	Thr	Asp	Phe	Leu	Met	Asn	Pro	Met	
				155					160					165	
Val	Met	Met	Met	Val	Leu	Pro	Leu	Leu	Ile	Phe	Val	Leu	Leu	Pro	
				170					175					180	
Lys	Val	Val	Asn	Thr	Ser	Asp	Pro	Asp	Met	Arg	Arg	Glu	Met	Glu	
				185					190					195	
Gln	Ser	Met	Asn	Met	Leu	Asn	Ser	Asn	His	Glu	Leu	Pro	Asp	Val	
				200					205					210	
Ser	Glu	Phe	Met	Thr	Arg	Leu	Phe	Ser	Ser	Lys	Ser	Ser	Gly	Lys	
				215					220					225	
Ser	Ser	Ser	Gly	Ser	Ser	Lys	Thr	Gly	Lys	Ser	Gly	Ala	Gly	Lys	
				230					235					240	

Arg Arg

<210> SEQ ID NO 137
<211> LENGTH: 1571
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 137

gatggcgcag ccacagcttc tgtgagattc gatttctccc cagttcccct	50
gtgggtctga ggggaccaga agggtagact acgttggtt tctggaagg	100
gaggctatat gcgtcaattc cccaaaacaa gttttgacat ttcccctgaa	150
atgtcattct ctatctattc actgcaagtg cctgctgttc caggccttac	200
ctgctgggca ctaacggcgg agccaggatg gggacagaat aaaggagcca	250
cgacctgtgc caccaactcg cactcagact ctgaactcag acctgaaatc	300
ttctcttcac gggaggcttg gcagtttttc ttactcctgt ggtctccaga	350
tttcaggcct aagatgaaag cctctagtct tgccttcagc cttctctctg	400
ctgcgtttta tctcctatgg actccttcca ctggactgaa gacactcaat	450
ttgggaagct gtgtgatcgc cacaaacctt caggaaatac gaaatggatt	500
ttctgagata cggggcagtg tgcaagccaa agatggaaac attgacatca	550
gaatcttaag gaggactgag tctttgcaag acacaaagcc tgcgaatcga	600
tgctgcctcc tgcgccatth gctaagactc tatctggaca gggtatthaa	650
aaactaccag acccctgacc attatactct ccggaagatc agcagcctcg	700
ccaattcctt tcttaccatc aagaaggacc tccggctctc tcatgcccac	750
atgacatgcc attgtgggga ggaagcaatg aagaaatata gccagattct	800
gagtcacttt gaaaagctgg aacctcaggc agcagttgtg aaggctttgg	850
gggaactaga cattcttctg caatggatgg aggagacaga ataggaggaa	900
agtgtagtct ctgctaagaa tattcagagt caagagctcc agtcttcaat	950
acctgcagag gaggcagtac cccaaaccac catctcttta ctgtactagt	1000
cttgtgtctg tcacagtgtg tcttatttat gcattacttg ctctcttgca	1050
tgattgtctt tatgcatccc caatcttaat tgagaccata cttgtataag	1100

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```
atTTTTgtaa tatctttctg ctattggata tatttattag ttaatatatt      1150
tattttatTTT ttgctatttta atgtatttat ttttttactt ggacatgaaa      1200
ctttaaaaaa attcacagat tatatttata acctgactag agcaggtgat      1250
gtatttttat acagtaaaaa aaaaaaacct tgtaaattct agaagagtgg      1300
ctaggggggt tattcatttg tattcaacta aggacatatt tactcatgct      1350
gatgctctgt gagatatttg aaattgaacc aatgactact taggatgggt      1400
tgtggaataa gttttgatgt ggaattgcac atctacctta caattactga      1450
ccatccccag tagactcccc agtcccataa ttgtgtatct tccagccagg      1500
aatcctacac ggcagcatg tatttctaca aataaagttt tctttgcata      1550
ccaaaaaaaa aaaaaaaaaa a      1571
```

<210> SEQ ID NO 138

<211> LENGTH: 261

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 138

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

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Ala Ala Val Val Lys Ala Leu Gly Glu Leu Asp Ile Leu Leu Gln
245 250 255

Trp Met Glu Glu Thr Glu
260

<210> SEQ ID NO 139
<211> LENGTH: 2395
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 139

cctggagcgcg gaagcgcggc tgcagcaggg cgaggctcca ggtgggggtcg	50
gttcgcgcac cagcctagcg tgtccacgat gcggctgggc tccgggactt	100
tcgtacacctg ttgcgtagcg atcgagggtgc tagggatcgc ggtcttcctt	150
cggggattct tcccggctcc cgttcgttcc tctgccagag cggaacacgg	200
agcggagccc ccagcgcccg aaccctcggc tggagccagt tctaactgga	250
ccacgctgcc accacctctc ttcagtaaag ttgttattgt tctgatagat	300
gccttgagag atgattttgt gtttgggtca aaggggtgtga aatttatgcc	350
ctacacaact taccttgtagg aaaaaggagc atctcacagt tttgtggctg	400
aagcaaagcc acctacagtt actatgcctc gaatcaaggc attgatgacg	450
gggagccttc ctggctttgt cgacgtcatc aggaacctca attctcctgc	500
actgctggaa gacagtgtga taagacaagc aaaagcagct ggaaaaagaa	550
tagtctttta tggagatgaa acctgggtta aattattccc aaagcatttt	600
gtggaatatg atggaacaac ctcatTTTTt gtgtcagatt acacagaggt	650
ggataataat gtcacgaggc atttgataa agtattaaaa agaggagatt	700
gggacataatt aatcctccac tacctggggc tggaccacat tggccacatt	750
tcagggccca acagccccct gattgggcag aagctgagcg agatggacag	800
cgtgctgatg aagatccaca cctcactgca gtcgaaggag agagagacgc	850
ctttacccaa tttgctggtt ctttgtggtg accatggcat gtctgaaaca	900
ggaagtcaag gggcctcctc caccgaggag gtgaatacac ctctgatttt	950
aatcagttct cgttttgaaa ggaaaccccg tgatatccga catccaaagc	1000
acgtccaata gacggatgtg gctgcgacac tggcgatagc acttggttta	1050
ccgattccaa aagacagtgt agggagcctc ctattcccag ttgtggaagg	1100
aagaccaatg agagagcagt tgagattttt acatttgaat acagtgcagc	1150
ttagtaaaact gttgcaagag aatgtgccgt catatgaaaa agatcctggg	1200
tttgagcagt ttaaaatgtc agaaagattg catgggaact ggatcagact	1250
gtacttgagg gaaaagcatt cagaagtctc attcaacctg ggctccaagg	1300
ttctcaggca gtacctggat gctctgaaga cgctgagctt gtccctgagt	1350
gcacaagtgg ccagttctc accctgctcc tgctcagcgt ccacaggca	1400
ctgcacagaa aggctgagct ggaagtccca ctgtcatctc ctgggttttc	1450
tctgtctctt tatttggtga tcctggttct ttcggccggt cacgtcattg	1500
tgtgcacctc agctgaaagt tcgtgtact tctgtggcct ctctggctg	1550

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gcggcaggct gcctttcggt taccagactc tgggtgaaca cctgggtgtgt 1600
gccaaagtgt ggcagtgtcc tggacagggg gcctcaggga aggacgtgga 1650
gcagccttat ccagggcctc tgggtgtccc gacacaggtg ttcacatctg 1700
tgctgtcagg tcagatgcct cagttcttgg aaagctaggt tcctgcgact 1750
gttaccaagg tgattgtaaa gagctggcgg tcacagagga acaagccccc 1800
cagctgaggg ggtgtgtgaa tcggacagcc tcccagcaga ggtgtgggag 1850
ctgcagctga gggaagaaga gacaatcggc ctggacactc aggagggtca 1900
aaaggagact tggctgcacc actcatcctg ccacccccag aatgcacatc 1950
gcctcatcag gtccagattt ctttccaagg cggacgtttt ctgttggaat 2000
tcttagtcct tggcctcgga caccttcatt cgtagctgg ggagtgggtg 2050
tgaggcagtg aagaagaggc ggatggtcac actcagatcc acagagccca 2100
ggatcaaggg acccactgca gtggcagcag gactgttggg cccccacccc 2150
aacctgtcac agccctcatc ccctcttggc ttgagccgtc agaggccctg 2200
tgctgagtg ctgaccgaga cactcacagc tttgtcatca gggcacaggc 2250
ttcctcggag ccaggatgat ctgtgccacg cttgcacctc gggcccatct 2300
gggctcatgc tctctctcct gctattgaat tagtacctag ctgcacacag 2350
tatgtagtta caaaagaat aaacggcaat aattgagaaa aaaaa 2395

<210> SEQ ID NO 140

<211> LENGTH: 310

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 140

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

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Trp	Val	Lys	Leu	Phe	Pro	Lys	His	Phe	Val	Glu	Tyr	Asp	Gly	Thr
			170						175					180
Thr	Ser	Phe	Phe	Val	Ser	Asp	Tyr	Thr	Glu	Val	Asp	Asn	Asn	Val
			185						190					195
Thr	Arg	His	Leu	Asp	Lys	Val	Leu	Lys	Arg	Gly	Asp	Trp	Asp	Ile
			200						205					210
Leu	Ile	Leu	His	Tyr	Leu	Gly	Leu	Asp	His	Ile	Gly	His	Ile	Ser
			215						220					225
Gly	Pro	Asn	Ser	Pro	Leu	Ile	Gly	Gln	Lys	Leu	Ser	Glu	Met	Asp
			230						235					240
Ser	Val	Leu	Met	Lys	Ile	His	Thr	Ser	Leu	Gln	Ser	Lys	Glu	Arg
			245						250					255
Glu	Thr	Pro	Leu	Pro	Asn	Leu	Leu	Val	Leu	Cys	Gly	Asp	His	Gly
			260						265					270
Met	Ser	Glu	Thr	Gly	Ser	His	Gly	Ala	Ser	Ser	Thr	Glu	Glu	Val
			275						280					285
Asn	Thr	Pro	Leu	Ile	Leu	Ile	Ser	Ser	Ala	Phe	Glu	Arg	Lys	Pro
			290						295					300
Gly	Asp	Ile	Arg	His	Pro	Lys	His	Val	Gln					
			305						310					

<210> SEQ ID NO 141
<211> LENGTH: 754
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 141

ggcacgaggc aagccttcca ggttatcgtg acgcaccttg aaagtctgag	50
agctactgcc ctacagaaag ttactagtgc cctaaagctg gcgctggcac	100
tgatgttact gctgctgttg gagtacaact tccctataga aaacaactgc	150
cagcacctta agaccactca caccttcaga gtgaagaact taaacccgaa	200
gaaattcagc attcatgacc aggatcacaa agtactggtc ctggactctg	250
ggaatctcat agcagttcca gataaaaact acatacgccc agagatcttc	300
tttgattag cctcatcctt gagctcagcc tctgcggaga aaggaagtcc	350
gattctcctg ggggtctcta aaggggagtt ttgtctctac tgtgacaagg	400
ataaaggaca aagtcattcca tcccttcagc tgaagaagga gaaactgatg	450
aagctggctg cccaaaagga atcagcacgc cggcccttca tcttttatag	500
ggctcaggtg ggctcctgga acatgctgga gtcggcggct caccctcgat	550
ggttcattctg cacctcctgc aattgtaatg agcctgttgg ggtgacagat	600
aaatttgaga acaggaaaca cattgaattt tcatttcaac cagtttgcaa	650
agctgaaatg agccccagtg aggtcagcga ttaggaaact gccccattga	700
acgccttcct cgctaatttg aactaattgt ataaaaacac caaacctgct	750
cact	754

<210> SEQ ID NO 142
<211> LENGTH: 193
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

-continued

<400> SEQUENCE: 142

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

<210> SEQ ID NO 143
<211> LENGTH: 961
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 143

ctagagagta tagggcagaa ggatggcaga tgagtgactc cacatccaga	50
gctgcctccc tttaatccag gatcctgtcc ttcctgtcct gtaggagtgc	100
ctgttgccag tgtggggtga gacaagtttg tcccacaggg ctgtctgagc	150
agataagatt aagggtctggg tctgtgctca attaaactcct gtgggcacgg	200
gggctgggaa gagcaaagtc agcggcgcct acagtcagca ccatgctggg	250
cctgccgtgg aagggaggtc tgcctgggc gctgctgctg cttctcttag	300
gctcccagat cctgctgata tatgcctggc atttccacga gcaaaggac	350
tgtgatgaac acaatgtcat ggctcgttac ctccctgcc agtggagtt	400
tgctgtccac acattcaacc aacagagcaa ggactactat gcctacagac	450
tggggcacat cttgaattcc tggaaggagc aggtggagtc caagactgta	500
ttctcaatgg agctactgct ggggagaact aggtgtggga aatttgaaga	550
cgacattgac aactgccatt tccaagaaag cacagagctg aacaatactt	600
tcacctgctt cttcaccatc agcaccaggc cctggatgac tcagttcagc	650
ctcctgaaca agacctgctt ggagggatc cactgagtga aaccactca	700

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caggcttgctc catgtgctgc tcccacattc cgtggacatc agcactactc	750
tcctgaggac tcttcagtgg ctgagcagct ttggacttgt ttgttatcct	800
attttgcatg tgtttgagat ctgagatcag tgttttagaa aatccacaca	850
tcttgagcct aatcatgtag tntagatcat taaacatcag cattttaaga	900
aaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa	950
aaaaaaaaa a	961

<210> SEQ ID NO 144
<211> LENGTH: 147
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 144

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

<210> SEQ ID NO 145
<211> LENGTH: 1157
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 145

ctgtgcagct cgaggctcca gaggcacact ccagagagag ccaaggttct	50
gacgcgatga ggaagcacct gagctggtgg tggctggcca ctgtctgcat	100
gtgctctctc agccacctct ctgcggtcca gacgaggggc atcaagcaca	150
gaatcaagtg gaaccggaag gccctgccca gcaactgccca gatcactgag	200
gcccaggtgg ctgagaaccg cccgggagcc ttcatacaagc aaggccgcaa	250
gtcgacatt gacttcggag ccgagggcaa cagggtactac gaggccaact	300
actggcagtt ccccgatggc atccactaca acggctgctc tgaggctaatt	350
gtgaccaagg aggcatttgt caccggtgc atcaatgccca cccaggcggc	400
gaaccagggg gaggttccaga agccagacaa caagctccac cagcaggtgc	450

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tctggcggct ggtccaggag ctctgctccc tcaagcattg cgagttttgg      500
ttggagaggg gcgcaggact tcgggtcacc atgcaccagc cagtgtctct      550
ctgccttctg gctttgatct ggctcatggt gaaataagct tgccaggagg      600
ctggcagtac agagcgcagc agcgagcaaa tcctggcaag tgaccagct      650
cttctcccc aaacccacgc gtgttctgaa ggtgcccagg agcggcgatg      700
cactcgcact gcaaatgccg ctcccacgta tgcgccctgg tatgtgcctg      750
cgttctgata gatggggggac tgtggcttct ccgtcactcc attctcagcc      800
cctagcagag cgtctggcac actagattag tagtaaatgc ttgatgagaa      850
gaacacatca ggcaactgcg cacctgcttc acagtacttc ccaacaactc      900
ttagaggtag gtgtattccc gttttacaga taaggaaact gaggcccaga      950
gagctgaagt actgcaccca gcatcaccag ctagaaagtg gcagagccag     1000
gattcaacco tggcttgtct aaccccaggt tttctgctct gtccaattcc     1050
agagctgtct ggtgatcact ttatgtctca cagggacca catccaaaca     1100
tgtatctcta atgaaattgt gaaagctcca tgtttagaaa taaatgaaaa     1150
cacctga                                             1157
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<210> SEQ ID NO 146
<211> LENGTH: 176
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 146

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

-continued

<210> SEQ ID NO 147
<211> LENGTH: 333
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 147

gccttggcct cccaaagggc tgggattata ggcgtgacca ccatgtctgg 50
tccagagtct catttcctga tgatttatag actcaaagaa aactcatgtt 100
cagaagctct cttctcttct ggccctcctct ctgtcttctt tccctctttc 150
ttcttatttt aattagtagc atctactcag agtcatgcaa gctggaaatc 200
tttcattttg cttgtcagtg gggtaggtca ctgagtctta gtttttattt 250
tttgaaattt caactttcag attcaggggg tacatgtgaa ggtttgtttt 300
atgagtatat tgcatgatgc tgaggtttgg ggt 333

<210> SEQ ID NO 148
<211> LENGTH: 73
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 148

Met Phe Arg Ser Ser Leu Leu Phe Trp Pro Pro Leu Cys Leu Leu
1 5 10 15
Ser Leu Phe Leu Leu Ile Leu Ile Ser Ser Ile Tyr Ser Glu Ser
20 25 30
Cys Lys Leu Glu Ile Phe His Phe Ala Cys Gln Trp Gly Arg Ser
35 40 45
Leu Ser Leu Ser Phe Tyr Phe Leu Lys Phe Gln Leu Ser Asp Ser
50 55 60
Gly Gly Thr Cys Glu Gly Leu Phe Tyr Glu Tyr Ile Ala
65 70

<210> SEQ ID NO 149
<211> LENGTH: 1893
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 149

gtctccgcgt cacaggaact tcagcaccca cagggcggac agcgctcccc 50
tctacctgga gacttgactc ccgcgcgccc caaccctgct tatcccttga 100
ccgtcgagtg tcagagatcc tgcagccgcc cagtcccggc ccctctcccg 150
ccccacaccc accctcctgg ctcttcctgt ttttactcct ccttttcatt 200
cataacaaaa gctacagctc caggagccca gcgcgggct gtgacccaag 250
ccgagcgtgg aagaatgggg ttccctcgga ccggcacttg gattctggtg 300
ttagtgctcc cgattcaagc tttcccaaa cctggaggaa gccaagacaa 350
atctctacat aatagagaat taagtgcaga aagaccttg aatgaacaga 400
ttgtgaagc agaagaagac aagattaaaa aaacatatcc tccagaaaac 450
aagccaggtc agagcaacta ttcttttgtt gataacttga acctgctaaa 500
ggcaataaca gaaaaggaaa aaattgagaa agaaagacaa tctataagaa 550
gtccccact tgataataag ttgaatgtgg aagatgttga ttcaaccaag 600

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aatcgaaaac tgatcgatga ttatgactct actaagagtg gattggatca	650
taaatttcaa gatgatccag atggctttca tcaactagac gggactcctt	700
taaccgctga agacattgtc cataaaatcg ctgccaggat ttatgaagaa	750
aatgacagag ccgtgtttga caagattgtt tctaaactac ttaatctcgg	800
ccttatcaca gaaagccaag cacatacact ggaagatgaa gtagcagagg	850
ttttacaaaa attaattctca aaggaagcca acaattatga ggaggatccc	900
aataagccca caagctggac tgagaatcag gctggaaaaa taccagagaa	950
agtgactcca atggcagcaa ttcaagatgg tcttgctaag ggagaaaacg	1000
atgaaacagt atctaacaca ttaaccttga caaatggctt ggaaggaga	1050
actaaaacct acagtgaaga caactttgag gaactccaat atttccaaa	1100
tttctatgcg ctactgaaaa gtattgattc agaaaaagaa gcaaaagaga	1150
aagaaacact gattactatc atgaaaacac tgattgactt tgtgaagatg	1200
atggtgaaat atggaacaat atctccagaa gaaggtgttt cctaccttga	1250
aaacttgatg gaaatgattg ctcttcagac caaaaacaag ctagaaaaaa	1300
atgctactga caatataagc aagcttttcc cagcaccatc agagaagagt	1350
catgaagaaa cagacagtac caaggaagaa gcagctaaga tggaaaagga	1400
atatggaagc ttgaaggatt ccacaaaaga tgataactcc aaccaggag	1450
gaaagacaga tgaaccctaa ggaaaaacag aagcctatctt ggaagccatc	1500
agaaaaata ttgaatgggt gaagaacat gacaaaaagg gaaataaaga	1550
agattatgac ctttcaaaga tgagagactt catcaataaa caagctgatg	1600
cttatgtgga gaaaggcatc ctgacaagg aagaagccga ggccatcaag	1650
cgcattttata gcagcctgta aaaatggcaa aagatccagg agtctttcaa	1700
ctgtttcaga aaacataata tagcttaaaa cacttctaatt tctgtgatta	1750
aaatTTTTTg acccaaggtt tattagaaag tgctgaattt acagtagtta	1800
acctttttaca agtgggttaa acatagcttt cttcccgtaa aaactatctg	1850
aaagtaaagt tgtatgtaag ctgaaaaaaa aaaaaaaaaa aaa	1893

<210> SEQ ID NO 150

<211> LENGTH: 468

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 150

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

-continued

80					85					90				
Arg	Gln	Ser	Ile	Arg	Ser	Ser	Pro	Leu	Asp	Asn	Lys	Leu	Asn	Val
				95					100					105
Glu	Asp	Val	Asp	Ser	Thr	Lys	Asn	Arg	Lys	Leu	Ile	Asp	Asp	Tyr
				110					115					120
Asp	Ser	Thr	Lys	Ser	Gly	Leu	Asp	His	Lys	Phe	Gln	Asp	Asp	Pro
				125					130					135
Asp	Gly	Leu	His	Gln	Leu	Asp	Gly	Thr	Pro	Leu	Thr	Ala	Glu	Asp
				140					145					150
Ile	Val	His	Lys	Ile	Ala	Ala	Arg	Ile	Tyr	Glu	Glu	Asn	Asp	Arg
				155					160					165
Ala	Val	Phe	Asp	Lys	Ile	Val	Ser	Lys	Leu	Leu	Asn	Leu	Gly	Leu
				170					175					180
Ile	Thr	Glu	Ser	Gln	Ala	His	Thr	Leu	Glu	Asp	Glu	Val	Ala	Glu
				185					190					195
Val	Leu	Gln	Lys	Leu	Ile	Ser	Lys	Glu	Ala	Asn	Asn	Tyr	Glu	Glu
				200					205					210
Asp	Pro	Asn	Lys	Pro	Thr	Ser	Trp	Thr	Glu	Asn	Gln	Ala	Gly	Lys
				215					220					225
Ile	Pro	Glu	Lys	Val	Thr	Pro	Met	Ala	Ala	Ile	Gln	Asp	Gly	Leu
				230					235					240
Ala	Lys	Gly	Glu	Asn	Asp	Glu	Thr	Val	Ser	Asn	Thr	Leu	Thr	Leu
				245					250					255
Thr	Asn	Gly	Leu	Glu	Arg	Arg	Thr	Lys	Thr	Tyr	Ser	Glu	Asp	Asn
				260					265					270
Phe	Glu	Glu	Leu	Gln	Tyr	Phe	Pro	Asn	Phe	Tyr	Ala	Leu	Leu	Lys
				275					280					285
Ser	Ile	Asp	Ser	Glu	Lys	Glu	Ala	Lys	Glu	Lys	Glu	Thr	Leu	Ile
				290					295					300
Thr	Ile	Met	Lys	Thr	Leu	Ile	Asp	Phe	Val	Lys	Met	Met	Val	Lys
				305					310					315
Tyr	Gly	Thr	Ile	Ser	Pro	Glu	Glu	Gly	Val	Ser	Tyr	Leu	Glu	Asn
				320					325					330
Leu	Asp	Glu	Met	Ile	Ala	Leu	Gln	Thr	Lys	Asn	Lys	Leu	Glu	Lys
				335					340					345
Asn	Ala	Thr	Asp	Asn	Ile	Ser	Lys	Leu	Phe	Pro	Ala	Pro	Ser	Glu
				350					355					360
Lys	Ser	His	Glu	Glu	Thr	Asp	Ser	Thr	Lys	Glu	Glu	Ala	Ala	Lys
				365					370					375
Met	Glu	Lys	Glu	Tyr	Gly	Ser	Leu	Lys	Asp	Ser	Thr	Lys	Asp	Asp
				380					385					390
Asn	Ser	Asn	Pro	Gly	Gly	Lys	Thr	Asp	Glu	Pro	Lys	Gly	Lys	Thr
				395					400					405
Glu	Ala	Tyr	Leu	Glu	Ala	Ile	Arg	Lys	Asn	Ile	Glu	Trp	Leu	Lys
				410					415					420
Lys	His	Asp	Lys	Lys	Gly	Asn	Lys	Glu	Asp	Tyr	Asp	Leu	Ser	Lys
				425					430					435
Met	Arg	Asp	Phe	Ile	Asn	Lys	Gln	Ala	Asp	Ala	Tyr	Val	Glu	Lys
				440					445					450
Gly	Ile	Leu	Asp	Lys	Glu	Glu	Ala	Glu	Ala	Ile	Lys	Arg	Ile	Tyr
				455					460					465

-continued

Ser Ser Leu

<210> SEQ ID NO 151
<211> LENGTH: 2598
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 151

cggctcgagg ctcccgccag gagaaaggaa cattctgagg ggagtctaca	50
ccctgtggag ctcaagatgg tcctgagtgg ggcgctgtgc ttccgaatga	100
aggactcggc attgaaggtg ctttatctgc ataataacca gcttctagct	150
ggagggctgc atgcagggaa ggtcattaaa ggtgaagaga tcagcgtggt	200
ccccaatcgg tggctggatg ccagcctgtc ccccgtcac cttgggtgtcc	250
aggggtgga cagtgccctg tcatgtgggg tggggcagga gccgactcta	300
acactagagc cagtgaacat catggagctc tatcttggtg ccaaggaatc	350
caagagcttc accttctacc ggccggacat ggggctcacc tccagcttcg	400
agtcggctgc ctacccgggc tgggtcctgt gcacggtgcc tgaagccgat	450
cagcctgtca gactcaccca gcttcccag aatggtggct ggaatgcccc	500
catcacagac ttctacttcc agcagtgtga ctagggcaac gtgcccccca	550
gaactccctg ggcagagcca gctcgggtga ggggtgagtg gaggagaccc	600
atggcggaca atcactctct ctgctctcag gacccccacg tctgacttag	650
tgggcacctg accactttgt cttctggttc ccagtttgga taaattctga	700
gatttgagc tcagtccacg gtccctcccc actggatggt gctactgctg	750
tggaaccttg taaaaacccat gtggggtaaa ctgggaataa catgaaaaga	800
tttctgtggg ggtgggggtg gggagtgggt ggaatcattc ctgcttaatg	850
gtaactgaca agtgttaccc tgagccccgc aggccaaacc atccccagtt	900
gagccttata gggtcagtag ctctccacat gaagtcctgt cactcaccac	950
tgtgcaggag agggaggtgg tcatagagtc agggatctat ggcccttggc	1000
ccagccccac ccccttccct ttaatcctgc cactgtcata tgctaccttt	1050
cctatctctt ccctcatcat cttgttgtgg gcatgaggag gtgggtgatgt	1100
cagaagaaat ggctcgagct cagaagataa aagataagta gggtatgctg	1150
atcctctttt aaaaacccaa gatacaatca aaatcccaga tgctggtctc	1200
tattcccatg aaaaagtgt catgacatat tgagaagacc tacttacaaa	1250
gtggcatata ttgcaattta ttttaattaa aagataccta tttatatatt	1300
tctttataga aaaaagtctg gaagagttta cttcaattgt agcaatgtca	1350
gggtgggtgg agtatagggt atttttcttt taattctggt aatttatctg	1400
tatttcctaa tttttctaca atgaagatga attccttgta taaaaataag	1450
aaaagaaatt aatcttgagg taagcagagc agacatcatc tctgattgtc	1500
ctcagcctcc acttccccag agtaaatca aattgaatcg agctctgctg	1550
ctctggttgg ttgtagtagt gatcaggaaa cagatctcag caaagccact	1600
gaggaggagg ctgtgctgag tttgtgtggc tggaatctct gggtaaggaa	1650

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cttaaagaac aaaatcatc tggtaatctt ttcctagaag gatcacagcc      1700
cctggggattc caaggcattg gatccagtct ctaagaaggc tgctgtactg      1750
gttgaattgt gtccccctca aattcacatc cttcttgga tctcagtcgt      1800
tgagttttatt tggagataag gtctctgcag atgtagttag ttaagacaag      1850
gtcatgctgg atgaaggtag acctaaattc aatatgactg gtttccttgt      1900
atgaaaagga gaggacacag agacagagga gacgcgggga agactatgta      1950
aagatgaagg cagagatcgg agttttgcag ccacaagcta agaaacacca      2000
aggattgtgg caaccatcag aagcttgga gaggcaaaga agaattcttc      2050
cctagaggct ttagagggat aacggctctg ctgaaacctt aatctcagac      2100
ttccagcctc ctgaacgaag aaagaataaa tttcggctgt tttaagccac      2150
caaggataat tggttacagc agctctagga aactaatata gctgctaaaa      2200
tgatccctgt ctctctgtgt ttacattctg tgtgtgtccc ctcccacaat      2250
gtaccaaagt tgtctttgtg accaatagaa tatggcagaa gtgatggcat      2300
gccacttcca agattagggt ataaaagaca ctgcagcttc tacttgagcc      2350
ctctctctct gccaccacc gccccaatc tatcttggt cactcgctct      2400
gggggaagct agctgccatg ctatgagcag gcctataaag agacttacgt      2450
ggtaaaaaat gaagtctcct gccacagcc acattagtga acctagaagc      2500
agagactctg tgagataatc gatgtttgtt gttttaagtt gctcagtttt      2550
gggtctaactt gttatgcagc aatagataaa taatatgcag agaaagag      2598
```

<210> SEQ ID NO 152

<211> LENGTH: 155

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 152

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

-continued

Phe Gln Gln Cys Asp
155

<210> SEQ ID NO 153
<211> LENGTH: 1152
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 153

cttcagaaca ggttctcctt ccccgagtcac cagttgctcg agttagaatt 50
gtctgcaatg gccgccctgc agaaatctgt gagctctttc cttatgggga 100
ccctggccac cagctgcctc cttctcttgg ccctcttggt acagggagga 150
gcagctgcgc ccatcagctc cactgcagc cttgacaagt ccaacttcca 200
gcagccctat atcaccaacc gcaccttcac gctggctaag gaggctagct 250
tggctgataa caacacagac gttcgtctca ttggggagaa actgttccac 300
ggagtcaagta tgagtgcgc ctgctatctg atgaagcagg tgctgaactt 350
cacccttgaa gaagtgtgt tccctcaatc tgataggctc cagccctata 400
tgaggagagt ggtgcccttc ctggccagc tcagcaacag gctaagcaca 450
tgtcatattg aaggatga cctgcatac cagaggaatg tgcaaaagct 500
gaaggacaca gtgaaaaagc ttggagagag tggagagatc aaagcaattg 550
gagaactgga tttgtgttt atgtctctga gaaatgcctg catttgacca 600
gagcaaaagt gaaaaatgaa taactaaccc ccttccctg ctgaaataa 650
caattagatg ccccaaagc atttttttta accaaaagga agatgggaag 700
ccaaactcca tcatgatggg tggattccaa atgaaccct gcgttagtta 750
caaaggaaac caatgccact ttgtttata agaccagaag gtagactttc 800
taagcataga tatttattga taacatttca ttgtaactgg tgttctatac 850
acagaaaaca atttattttt taaataattg tctttttcca taaaaagat 900
tactttccat tcctttaggg gaaaaaaccc ctaaatagct tcatgtttcc 950
ataatcagta ctttatattt ataaatgtat ttattattat tataagactg 1000
cattttattt atatcatttt attaatatgg atttatttat agaaacatca 1050
ttcgatattg ctacttgagt gtaaggctaa tattgatatt tatgacaata 1100
attatagagc tataacatgt ttatttgacc tcaataaaca cttggatatc 1150
cc 1152

<210> SEQ ID NO 154
<211> LENGTH: 179
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 154

Met Ala Ala Leu Gln Lys Ser Val Ser Ser Phe Leu Met Gly Thr
1 5 10 15
Leu Ala Thr Ser Cys Leu Leu Leu Leu Ala Leu Leu Val Gln Gly
20 25 30
Gly Ala Ala Ala Pro Ile Ser Ser His Cys Arg Leu Asp Lys Ser
35 40 45

-continued

Asn	Phe	Gln	Gln	Pro	Tyr	Ile	Thr	Asn	Arg	Thr	Phe	Met	Leu	Ala
				50					55					60
Lys	Glu	Ala	Ser	Leu	Ala	Asp	Asn	Asn	Thr	Asp	Val	Arg	Leu	Ile
			65						70					75
Gly	Glu	Lys	Leu	Phe	His	Gly	Val	Ser	Met	Ser	Glu	Arg	Cys	Tyr
			80						85					90
Leu	Met	Lys	Gln	Val	Leu	Asn	Phe	Thr	Leu	Glu	Glu	Val	Leu	Phe
			95						100					105
Pro	Gln	Ser	Asp	Arg	Phe	Gln	Pro	Tyr	Met	Gln	Glu	Val	Val	Pro
			110						115					120
Phe	Leu	Ala	Arg	Leu	Ser	Asn	Arg	Leu	Ser	Thr	Cys	His	Ile	Glu
			125						130					135
Gly	Asp	Asp	Leu	His	Ile	Gln	Arg	Asn	Val	Gln	Lys	Leu	Lys	Asp
			140						145					150
Thr	Val	Lys	Lys	Leu	Gly	Glu	Ser	Gly	Glu	Ile	Lys	Ala	Ile	Gly
			155						160					165
Glu	Leu	Asp	Leu	Leu	Phe	Met	Ser	Leu	Arg	Asn	Ala	Cys	Ile	
			170						175					

<210> SEQ ID NO 155
<211> LENGTH: 1320
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 155

ggcttgctga aaataaaatc aggactccta acctgctcca gtcagcctgc	50
ttccacgagg cctgtcagtc agtgcccac ttgtgactga gtgtgcagtg	100
cccagcatgt accaggtcag tgcagagggc tgctgaggg ctgtgctgag	150
agggagagga gcagagatgc tgctgagggt ggagggaggc caagctgcca	200
ggtttggggc tgggggccaa gtggagttag aaactgggat cccaggggga	250
gggtgcagat gaggagcgca cccagattag gtgaggacag ttctctcatt	300
agccttttcc tacagtggtg tgcattcttg gcaatggtca tgggaaccca	350
cacctacagc cactggccca gctgctgccc cagcaaaggg caggacacct	400
ctgaggagct gctgaggtgg agcactgtgc ctgtgcctcc cctagagcct	450
gctaggccca accgccccc agagtctgt agggccagt aagatggacc	500
cctcaacagc agggccatct ccccctggag atatgagttg gacagagact	550
tgaaccggct ccccaggac ctgtaccacg ccggttgct gtgccgcac	600
tgcgtcagcc tacagacagg ctcccacatg gacccccgg gcaactcgga	650
gctgctctac cacaaccaga ctgtcttcta caggcgccca tgccatggcg	700
agaagggcac ccacaaggc tactgcctgg agcgcaggct gtaccgtgtt	750
tccttagctt gtgtgtgtgt gcggccccgt gtgatgggct agccggacct	800
gctggaggct ggtccctttt tgggaaacct ggagccaggt gtacaaccac	850
ttgccatgaa gggccaggat gccagatgc ttggcccctg tgaagtgtg	900
tctggagcag caggatccc ggacaggatg gggggctttg gggaaaacct	950
gcacttctgc acattttgaa aagagcagct gctgcttagg gccgccggaa	1000

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gctggtgtcc tgtcattttc tctcaggaaa ggttttcaaa gttctgccca	1050
ttttcggagg ccaccactcc tgtctcttcc tcttttccca tcccctgcta	1100
ccctggccca gcacaggcac tttctagata tttccccctt gctggagaag	1150
aaagagcccc tggttttatt tgtttgttta ctcactctc agtgagcatc	1200
tactttgggt gcattctagt gtagttacta gtcttttgac atggatgatt	1250
ctgaggagga agctgttatt gaatgtatag agatttatcc aaataaatat	1300
ctttatttaa aaatgaaaaa	1320

<210> SEQ ID NO 156
<211> LENGTH: 177
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 156

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

<210> SEQ ID NO 157
<211> LENGTH: 1515
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 157

ccggcgatgt cgctcgtgct gctaagcctg gccgcgctgt gcaggagcgc	50
cgtaccgccga gagccgaccg ttcaatgtgg ctctgaaact gggccatctc	100
cagagtggat gctacaacat gatctaatacc ccggagactt gagggacctc	150
cgagtagaac ctgttacaac tagtgttgca acaggggact attcaatttt	200
gatgaatgta agctgggtac tccgggcaga tgccagcatc cgcttggtga	250

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agggccaccaa gatttgtgtg acggggcaaaa gcaacttcca gtcctacagc	300
tgtgtgaggt gcaattacac agaggccttc cagactcaga ccagaccctc	350
tggtggtaaa tggacatttt cctacatcgg cttccctgta gagctgaaca	400
cagtctatatt cattggggcc cataatatcc ctaatgcaaa tatgaatgaa	450
gatggccctt ccatgtctgt gaatttcacc tcaccaggct gcctagacca	500
cataatgaaa tataaaaaaa agtgtgtcaa ggccggaagc ctgtgggatc	550
cgaacatcac tgcttgaag aagaatgagg agacagtaga agtgaacttc	600
acaaccactc ccctgggaaa cagatacatg gctcttatcc aacacagcac	650
tatcatcggg ttttctcagg tgtttgagcc acaccagaag aaacaaacgc	700
gagcttcagt ggtgattcca gtgactgggg atagtgaagg tgctacggtg	750
cagctgactc catattttcc tacttgtggc agcgactgca tccgacataa	800
aggaacagtt gtgctctgcc cacaaacagg cgtccctttc cctctggata	850
acaacaaaag caagccggga ggctggctgc ctctctcctc gctgtctctg	900
ctggtggcca catgggtgct ggtggcaggg atctatctaa tgtggaggca	950
cgaaaggatc aagaagactt ccttttctac caccacacta ctgcccccca	1000
ttaaggttct tgtggtttac ccactgaaa tatgtttcca tcacacaatt	1050
tgttacttca ctgaatttct tcaaaaccat tgcagaagtg aggtcatcct	1100
tgaaaagtgg cagaaaaaga aaatagcaga gatgggtcca gtgcagtggc	1150
ttgccactca aaagaaggca gcagacaaag tcgtcttcct tctttccaat	1200
gacgtcaaca gtgtgtgcga tggtagctgt ggcaagagcg agggcagtc	1250
cagtgagaac tctcaagacc tcttccccct tgcctttaac cttttctgca	1300
gtgatctaag aagccagatt catctgcaca aatacgtggt ggtctacttt	1350
agagagattg atacaaaaga cgattacaat gctctcagtg tctgccccaa	1400
gtaccacctc atgaaggatg ccaactgcttt ctgtgcagaa cttctccatg	1450
tcaagcagca ggtgtcagca ggaaaaagat cacaagcctg ccacgatggc	1500
tgctgctcct tgtag	1515

<210> SEQ ID NO 158

<211> LENGTH: 502

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 158

Met	Ser	Leu	Val	Leu	Leu	Ser	Leu	Ala	Ala	Leu	Cys	Arg	Ser	Ala
1			5					10						15
Val	Pro	Arg	Glu	Pro	Thr	Val	Gln	Cys	Gly	Ser	Glu	Thr	Gly	Pro
			20						25					30
Ser	Pro	Glu	Trp	Met	Leu	Gln	His	Asp	Leu	Ile	Pro	Gly	Asp	Leu
			35						40					45
Arg	Asp	Leu	Arg	Val	Glu	Pro	Val	Thr	Thr	Ser	Val	Ala	Thr	Gly
			50						55					60
Asp	Tyr	Ser	Ile	Leu	Met	Asn	Val	Ser	Trp	Val	Leu	Arg	Ala	Asp
			65						70					75
Ala	Ser	Ile	Arg	Leu	Leu	Lys	Ala	Thr	Lys	Ile	Cys	Val	Thr	Gly

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80					85					90				
Lys	Ser	Asn	Phe	Gln	Ser	Tyr	Ser	Cys	Val	Arg	Cys	Asn	Tyr	Thr
				95					100					105
Glu	Ala	Phe	Gln	Thr	Gln	Thr	Arg	Pro	Ser	Gly	Gly	Lys	Trp	Thr
				110					115					120
Phe	Ser	Tyr	Ile	Gly	Phe	Pro	Val	Glu	Leu	Asn	Thr	Val	Tyr	Phe
				125					130					135
Ile	Gly	Ala	His	Asn	Ile	Pro	Asn	Ala	Asn	Met	Asn	Glu	Asp	Gly
				140					145					150
Pro	Ser	Met	Ser	Val	Asn	Phe	Thr	Ser	Pro	Gly	Cys	Leu	Asp	His
				155					160					165
Ile	Met	Lys	Tyr	Lys	Lys	Lys	Cys	Val	Lys	Ala	Gly	Ser	Leu	Trp
				170					175					180
Asp	Pro	Asn	Ile	Thr	Ala	Cys	Lys	Lys	Asn	Glu	Glu	Thr	Val	Glu
				185					190					195
Val	Asn	Phe	Thr	Thr	Thr	Pro	Leu	Gly	Asn	Arg	Tyr	Met	Ala	Leu
				200					205					210
Ile	Gln	His	Ser	Thr	Ile	Ile	Gly	Phe	Ser	Gln	Val	Phe	Glu	Pro
				215					220					225
His	Gln	Lys	Lys	Gln	Thr	Arg	Ala	Ser	Val	Val	Ile	Pro	Val	Thr
				230					235					240
Gly	Asp	Ser	Glu	Gly	Ala	Thr	Val	Gln	Leu	Thr	Pro	Tyr	Phe	Pro
				245					250					255
Thr	Cys	Gly	Ser	Asp	Cys	Ile	Arg	His	Lys	Gly	Thr	Val	Val	Leu
				260					265					270
Cys	Pro	Gln	Thr	Gly	Val	Pro	Phe	Pro	Leu	Asp	Asn	Asn	Lys	Ser
				275					280					285
Lys	Pro	Gly	Gly	Trp	Leu	Pro	Leu	Leu	Leu	Leu	Ser	Leu	Leu	Val
				290					295					300
Ala	Thr	Trp	Val	Leu	Val	Ala	Gly	Ile	Tyr	Leu	Met	Trp	Arg	His
				305					310					315
Glu	Arg	Ile	Lys	Lys	Thr	Ser	Phe	Ser	Thr	Thr	Thr	Leu	Leu	Pro
				320					325					330
Pro	Ile	Lys	Val	Leu	Val	Val	Tyr	Pro	Ser	Glu	Ile	Cys	Phe	His
				335					340					345
His	Thr	Ile	Cys	Tyr	Phe	Thr	Glu	Phe	Leu	Gln	Asn	His	Cys	Arg
				350					355					360
Ser	Glu	Val	Ile	Leu	Glu	Lys	Trp	Gln	Lys	Lys	Lys	Ile	Ala	Glu
				365					370					375
Met	Gly	Pro	Val	Gln	Trp	Leu	Ala	Thr	Gln	Lys	Lys	Ala	Ala	Asp
				380					385					390
Lys	Val	Val	Phe	Leu	Leu	Ser	Asn	Asp	Val	Asn	Ser	Val	Cys	Asp
				395					400					405
Gly	Thr	Cys	Gly	Lys	Ser	Glu	Gly	Ser	Pro	Ser	Glu	Asn	Ser	Gln
				410					415					420
Asp	Leu	Phe	Pro	Leu	Ala	Phe	Asn	Leu	Phe	Cys	Ser	Asp	Leu	Arg
				425					430					435
Ser	Gln	Ile	His	Leu	His	Lys	Tyr	Val	Val	Val	Tyr	Phe	Arg	Glu
				440					445					450
Ile	Asp	Thr	Lys	Asp	Asp	Tyr	Asn	Ala	Leu	Ser	Val	Cys	Pro	Lys
				455					460					465

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Tyr His Leu Met Lys Asp Ala Thr Ala Phe Cys Ala Glu Leu Leu
470 475 480
His Val Lys Gln Gln Val Ser Ala Gly Lys Arg Ser Gln Ala Cys
485 490 495
His Asp Gly Cys Cys Ser Leu
500

<210> SEQ ID NO 159
<211> LENGTH: 535
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 159
agccaccagc gcaacatgac agtgaagacc ctgcatggcc cagccatggt 50
caagtacttg ctgctgtcga tattggggct tgcctttctg agtgaggcgg 100
cagctcggaa aatccccaaa gtaggacata cttttttcca aaagcctgag 150
agttgccgcg ctgtgccagg aggtagtatg aagcttgaca ttggcatcat 200
caatgaaaac cagcgcgttt ccatgtcacg taacatcgag agccgctcca 250
cctccccctg gaattacact gtcacttggg accccaaccg gtaccctctg 300
gaagttgtac aggcccagtg taggaacttg ggctgcatca atgctcaagg 350
aaaggaagac atctccatga attccgttcc catccagcaa gagaccctgg 400
tcgtccggag gaagcaccaa ggctgctctg tttctttcca gttggagaag 450
gtgctggtga ctgttggtg cactgcgtgc acccctgtca tccaccatgt 500
gcagtaagag gtgcatatcc actcagctga agaag 535

<210> SEQ ID NO 160
<211> LENGTH: 163
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 160
Met Thr Val Lys Thr Leu His Gly Pro Ala Met Val Lys Tyr Leu
1 5 10 15
Leu Leu Ser Ile Leu Gly Leu Ala Phe Leu Ser Glu Ala Ala Ala
20 25 30
Arg Lys Ile Pro Lys Val Gly His Thr Phe Phe Gln Lys Pro Glu
35 40 45
Ser Cys Pro Pro Val Pro Gly Gly Ser Met Lys Leu Asp Ile Gly
50 55 60
Ile Ile Asn Glu Asn Gln Arg Val Ser Met Ser Arg Asn Ile Glu
65 70 75
Ser Arg Ser Thr Ser Pro Trp Asn Tyr Thr Val Thr Trp Asp Pro
80 85 90
Asn Arg Tyr Pro Ser Glu Val Val Gln Ala Gln Cys Arg Asn Leu
95 100 105
Gly Cys Ile Asn Ala Gln Gly Lys Glu Asp Ile Ser Met Asn Ser
110 115 120
Val Pro Ile Gln Gln Glu Thr Leu Val Val Arg Arg Lys His Gln
125 130 135
Gly Cys Ser Val Ser Phe Gln Leu Glu Lys Val Leu Val Thr Val

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tggtggcct gcctactctt tgccgtgcg ctttccctca tcctccttct	1650
caaaaaggat cacgcgaaa ggtggctgag gctcttgaaa caggacgtcc	1700
gctcgggggc ggccgccagg ggccgcgcgg ctctgctcct ctactcagcc	1750
gatgactcgg gtttcgagcg cctgggtggc gccctggcgt cggccctgtg	1800
ccagctgccg ctgcgcgtgg ccgtagacct gtggagccgt cgtgaactga	1850
gcgcgcaggg gcccggtggc tggtttcacg cgcagcggcg ccagaccctg	1900
caggaggggc gcgtggtggt ctgtctcttc tctcccgtg cgtggcgct	1950
gtgcagcgag tggctacagg atgggggtgc cgggcccggg gcgcacggcc	2000
cgcacgacgc cttccgcgc tcgctcagct gcgtgctgcc cgacttcttg	2050
cagggccggg cgcccgccag ctacgtgggg gcctgcttcg acaggctgct	2100
ccaccggcag gccgtaccgg cccttttccg caccgtgccc gtcttcacac	2150
tgccctccca actgccagac ttctggggg ccctgcagca gcctcgcgcc	2200
ccgcgttccg ggcggctcca agagagagcg gagcaagtgt cccggggcct	2250
tcagccagcc ctggatagct acttccatcc cccggggact cccgcgccgg	2300
gacgcggggg gggaccaggg gcgggacctg gggcggggga cgggacttaa	2350
ataaaggcag acgtgtttt tctaaaaaaa	2380

<210> SEQ ID NO 162

<211> LENGTH: 705

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 162

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

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Tyr	Glu	Lys	Glu	Leu	Asn	His	Thr	Gln	Gln	Leu	Pro	Ala	Leu	Pro	
				185					190					195	
Trp	Leu	Asn	Val	Ser	Ala	Asp	Gly	Asp	Asn	Val	His	Leu	Val	Leu	
				200					205					210	
Asn	Val	Ser	Glu	Glu	Gln	His	Phe	Gly	Leu	Ser	Leu	Tyr	Trp	Asn	
				215					220					225	
Gln	Val	Gln	Gly	Pro	Pro	Lys	Pro	Arg	Trp	His	Lys	Asn	Leu	Thr	
				230					235					240	
Gly	Pro	Gln	Ile	Ile	Thr	Leu	Asn	His	Thr	Asp	Leu	Val	Pro	Cys	
				245					250					255	
Leu	Cys	Ile	Gln	Val	Trp	Pro	Leu	Glu	Pro	Asp	Ser	Val	Arg	Thr	
				260					265					270	
Asn	Ile	Cys	Pro	Phe	Arg	Glu	Asp	Pro	Arg	Ala	His	Gln	Asn	Leu	
				275					280					285	
Trp	Gln	Ala	Ala	Arg	Leu	Arg	Leu	Leu	Thr	Leu	Gln	Ser	Trp	Leu	
				290					295					300	
Leu	Asp	Ala	Pro	Cys	Ser	Leu	Pro	Ala	Glu	Ala	Ala	Leu	Cys	Trp	
				305					310					315	
Arg	Ala	Pro	Gly	Gly	Asp	Pro	Cys	Gln	Pro	Leu	Val	Pro	Pro	Leu	
				320					325					330	
Ser	Trp	Glu	Asn	Val	Thr	Val	Asp	Lys	Val	Leu	Glu	Phe	Pro	Leu	
				335					340					345	
Leu	Lys	Gly	His	Pro	Asn	Leu	Cys	Val	Gln	Val	Asn	Ser	Ser	Glu	
				350					355					360	
Lys	Leu	Gln	Leu	Gln	Glu	Cys	Leu	Trp	Ala	Asp	Ser	Leu	Gly	Pro	
				365					370					375	
Leu	Lys	Asp	Asp	Val	Leu	Leu	Leu	Glu	Thr	Arg	Gly	Pro	Gln	Asp	
				380					385					390	
Asn	Arg	Ser	Leu	Cys	Ala	Leu	Glu	Pro	Ser	Gly	Cys	Thr	Ser	Leu	
				395					400					405	
Pro	Ser	Lys	Ala	Ser	Thr	Arg	Ala	Ala	Arg	Leu	Gly	Glu	Tyr	Leu	
				410					415					420	
Leu	Gln	Asp	Leu	Gln	Ser	Gly	Gln	Cys	Leu	Gln	Leu	Trp	Asp	Asp	
				425					430					435	
Asp	Leu	Gly	Ala	Leu	Trp	Ala	Cys	Pro	Met	Asp	Lys	Tyr	Ile	His	
				440					445					450	
Lys	Arg	Trp	Ala	Leu	Val	Trp	Leu	Ala	Cys	Leu	Leu	Phe	Ala	Ala	
				455					460					465	
Ala	Leu	Ser	Leu	Ile	Leu	Leu	Leu	Lys	Lys	Asp	His	Ala	Lys	Gly	
				470					475					480	
Trp	Leu	Arg	Leu	Leu	Lys	Gln	Asp	Val	Arg	Ser	Gly	Ala	Ala	Ala	
				485					490					495	
Arg	Gly	Arg	Ala	Ala	Leu	Leu	Leu	Tyr	Ser	Ala	Asp	Asp	Ser	Gly	
				500					505					510	
Phe	Glu	Arg	Leu	Val	Gly	Ala	Leu	Ala	Ser	Ala	Leu	Cys	Gln	Leu	
				515					520					525	
Pro	Leu	Arg	Val	Ala	Val	Asp	Leu	Trp	Ser	Arg	Arg	Glu	Leu	Ser	
				530					535					540	
Ala	Gln	Gly	Pro	Val	Ala	Trp	Phe	His	Ala	Gln	Arg	Arg	Gln	Thr	
				545					550					555	

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Leu	Gln	Glu	Gly	Gly	Val	Val	Val	Leu	Leu	Phe	Ser	Pro	Gly	Ala
			560						565					570
Val	Ala	Leu	Cys	Ser	Glu	Trp	Leu	Gln	Asp	Gly	Val	Ser	Gly	Pro
			575						580					585
Gly	Ala	His	Gly	Pro	His	Asp	Ala	Phe	Arg	Ala	Ser	Leu	Ser	Cys
			590						595					600
Val	Leu	Pro	Asp	Phe	Leu	Gln	Gly	Arg	Ala	Pro	Gly	Ser	Tyr	Val
			605						610					615
Gly	Ala	Cys	Phe	Asp	Arg	Leu	Leu	His	Pro	Asp	Ala	Val	Pro	Ala
			620						625					630
Leu	Phe	Arg	Thr	Val	Pro	Val	Phe	Thr	Leu	Pro	Ser	Gln	Leu	Pro
			635						640					645
Asp	Phe	Leu	Gly	Ala	Leu	Gln	Gln	Pro	Arg	Ala	Pro	Arg	Ser	Gly
			650						655					660
Arg	Leu	Gln	Glu	Arg	Ala	Glu	Gln	Val	Ser	Arg	Ala	Leu	Gln	Pro
			665						670					675
Ala	Leu	Asp	Ser	Tyr	Phe	His	Pro	Pro	Gly	Thr	Pro	Ala	Pro	Gly
			680						685					690
Arg	Gly	Val	Gly	Pro	Gly	Ala	Gly	Pro	Gly	Ala	Gly	Asp	Gly	Thr
			695						700					705

<210> SEQ ID NO 163
<211> LENGTH: 2478
<212> TYPE: DNA
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 163

gtcagtgcgg gaggccggtc agccaccaag atgactgaca gggttcagctc	50
tctgcagcac actaccctca agccacctga tgtgacctgt atctccaaag	100
tgagatcgat tcagatgatt gttcatccta cccccacgcc aatccgtgca	150
ggcgatggcc accggctaac cctggaagac atcttccatg acctgttcta	200
ccacttagag ctccaggtca accgcaccta ccaaatgcac cttggaggga	250
agcagagaga atatgagttc ttccgcctga cccctgacac agagttcctt	300
ggcaccatca tgatttgcgt tcccacctgg gccaaggaga gtgcccccta	350
catgtgccga gtgaagacac tgccagaccg gacatggacc tactccttct	400
ccggagcctt cctgtttctcc atgggcttcc tcgtcgagct actctgctac	450
ctgagctaca gatatgtcac caagccgcct gcacctccca actccctgaa	500
cgtccagcga gtcctgactt tccagccgct gcgcttcac caggagcacg	550
tctgatccc tgtctttgac ctccagcgcc ccagcagctt ggcccagcct	600
gtccagtact ccagatcag ggtgtctgga cccaggagc ccgcaggagc	650
tccacagcgg catagcctgt ccgagatcac ctacttaggg cagccagaca	700
tctccatcct ccagccctcc aacgtgccac ctcccagat cctctcccca	750
ctgtcctatg ccccaaacgc tgccctgag gtcgggcccc catcctatgc	800
acctcaggtg acccccgaag ctcaattccc attctacgcc ccacaggcca	850
tctctaaggt ccagccttcc tcctatgccc ctcaagccac tccggacagc	900
tggcctccct cctatggggg atgcatggaa gggtctggca aagactcccc	950

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cactggggaca ctttctagtc ctaaacacct taggcctaaa ggtcagcttc	1000
agaaagagcc accagctgga agctgcatgt taggtggcct ttctctgcag	1050
gaggtgacct ccttggtctat ggaggaatcc caagaagcaa aatcattgca	1100
ccagcccctg gggatttgca cagacagAAC atctgaccca aatgtgctac	1150
acagtgggga ggaagggaca ccacagtacc taaagggccA gctccccctc	1200
ctctcctcag tccagatcga gggccacccc atgtccctcc ctttgcaacc	1250
tccttcctggt ccatgttccc cctcggacca aggtccaagt ccttggggcc	1300
tgctggagtc ccttgtgtgt cccaaggatg aagccaagag cccagcccct	1350
gagacctcag acctggagca gcccacagaa ctggattctc ttttcagagg	1400
cctggccctg actgtgcagt gggagtccctg aggggaatgg gaaaggcttg	1450
gtgcttcctc cctgtcccta cccagtgtca catccttggc tgtcaatccc	1500
atgcctgccc atgccacaca ctctgcgacg tggcctcaga cgggtgccct	1550
tgagagaagc agagggagtg gcatgcaggg cccctgccat ggggtcgctc	1600
ctcaccggaA caaagcagca tgataaggac tgcagcgggg gagctctggg	1650
gagcagcttg tgtagacaag cgcgtgctcg ctgagccctg caaggcagaa	1700
atgacagtgc aaggaggaaa tgcagggaaa ctcccgaggt ccagagcccc	1750
acctcctaac accatggatt caaagtgtcA agggaatttg cctctccttg	1800
ccccattcct ggccagtttc acaatctagc tcgacagagc atgaggcccc	1850
tgctctctct gtcatgtttc aaagtgggga agagagcctg gaaaagaacc	1900
aggcctggaa aagaaccaga aggaggctgg gcagaaccag aacaacctgc	1950
acttctgccA aggccagggc cagcaggacg gcaggactct agggaggggt	2000
gtggcctgca gctcattccc agccagggca actgcctgac gttgcacgat	2050
ttcagcttca ttctctgat agaacaaagc gaaatgcagg tccaccaggg	2100
agggagacac acaagccttt tctgcaggca ggagtttcag acctatcct	2150
gagaatgggg tttgaaagga aggtgagggc tgtggcccct ggacgggtac	2200
aataacacac tgtactgatg tcacaacttt gcaagctctg ccttgggttc	2250
agcccactct ggctcaaatt ccagcctcac cactcacaaA ctgtgtgact	2300
tcaaacaaat gaaatcagtg cccagaacct cggtttcctc atctgtaatg	2350
tggggatcat aacacctacc tcatggagtt gtggtgaaga tgaaatgaag	2400
tcatgtcttt aaagtgttA atagtgcctg gtacatgggc agtgcccaat	2450
aaacggtagc tatttaaaaa aaaaaaaa	2478

<210> SEQ ID NO 164
<211> LENGTH: 574
<212> TYPE: PRT
<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 164

Met Arg Thr Leu Leu Thr Ile Leu Thr Val Gly Ser Leu Ala Ala
1 5 10 15

His Ala Pro Glu Asp Pro Ser Asp Leu Leu Gln His Val Lys Phe
20 25 30

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

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				410				415				420			
Ser	Pro	Lys	His	Leu	Arg	Pro	Lys	Gly	Gln	Leu	Gln	Lys	Glu	Pro	
				425					430					435	
Pro	Ala	Gly	Ser	Cys	Met	Leu	Gly	Gly	Leu	Ser	Leu	Gln	Glu	Val	
				440					445					450	
Thr	Ser	Leu	Ala	Met	Glu	Glu	Ser	Gln	Glu	Ala	Lys	Ser	Leu	His	
				455					460					465	
Gln	Pro	Leu	Gly	Ile	Cys	Thr	Asp	Arg	Thr	Ser	Asp	Pro	Asn	Val	
				470					475					480	
Leu	His	Ser	Gly	Glu	Glu	Gly	Thr	Pro	Gln	Tyr	Leu	Lys	Gly	Gln	
				485					490					495	
Leu	Pro	Leu	Leu	Ser	Ser	Val	Gln	Ile	Glu	Gly	His	Pro	Met	Ser	
				500					505					510	
Leu	Pro	Leu	Gln	Pro	Pro	Ser	Gly	Pro	Cys	Ser	Pro	Ser	Asp	Gln	
				515					520					525	
Gly	Pro	Ser	Pro	Trp	Gly	Leu	Leu	Glu	Ser	Leu	Val	Cys	Pro	Lys	
				530					535					540	
Asp	Glu	Ala	Lys	Ser	Pro	Ala	Pro	Glu	Thr	Ser	Asp	Leu	Glu	Gln	
				545					550					555	
Pro	Thr	Glu	Leu	Asp	Ser	Leu	Phe	Arg	Gly	Leu	Ala	Leu	Thr	Val	
				560					565					570	
Gln	Trp	Glu	Ser												

<210> SEQ ID NO 165

<211> LENGTH: 1060

<212> TYPE: DNA

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 165

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gtggccacaa	catggctgcg	gcgcgggggc	tgctcttctg	gctgttcgtg	100
ctggggggcg	tctggtgggt	ccggggccag	tcggatctca	gccacggacg	150
gcgtttctcg	gacctcaaa	tgctcgggga	cgaagagtgc	agcatgttaa	200
tgtaccgtgg	gaaagctctt	gaagacttca	cgggccttga	ttgtcgtttt	250
gtgaatttta	aaaaagggtg	cgatgtatat	gtctactaca	aactggcagg	300
gggatccctt	gaactttggg	ctggaagtgt	tgaacacagt	tttgatattt	350
ttccaaaaga	tttgatcaag	gtacttcata	aatacacgga	agaagagcta	400
catattccag	cagatgagac	agactttgtc	tgctttgaag	gaggaagaga	450
tgattttaat	agttataatg	tagaagagct	tttaggatct	ttggaactgg	500
aggactctgt	acctgaagag	tcgaagaaa	ctgaagaagt	ttctcagcac	550
agagagaaat	ctcctgagga	gtctcggggg	cgtgaacttc	accctgtgcc	600
tgagcccag	gcattcagag	ctgattcaga	ggatggagaa	ggtgctttct	650
cagagagcac	cgaggggctg	cagggacagc	cctcagctca	ggagagccac	700
cctcacacca	gcggtcctcg	ggctaacgct	cagggagtgc	agtcttcgtt	750
ggacactttt	gaagaaattc	tgacagataa	attgaaagtg	cggggaagcg	800
aaqcagaac	tggcaataqt	tctcctgcct	cqgtggaqcg	ggagaagaca	850

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gatgcttaca aagtcctgaa aacagaaatg agtcagagag gaagtggaca	900
gtgcgttatt cattacagca aaggatttcg ttggcatcaa aatctaagtt	950
tgttttacaa agattgtttt tagtactaag ctgccttggc agtttgcatt	1000
tttgagccaa acaaaaatat attattttcc cttctaagta aaaaaaaaaa	1050
aaaaaaaaaa	1060

<210> SEQ ID NO 166

<211> LENGTH: 303

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 166

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

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290	295	300
Asp Cys Phe		
<210> SEQ ID NO 167		
<211> LENGTH: 2570		
<212> TYPE: DNA		
<213> ORGANISM: Homo Sapien		
<400> SEQUENCE: 167		
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agagaagcaa agcgcaacgg tgtgtgtccaa gccgggggctt ctgcttcgcc		100
tctaggacat acacgggacc ccctaacttc agtcccccaa acgcgcaccc		150
tcgaagtctt gaactccagc cccgcacatc cagcgcgggc acaggcgcg		200
caggcgggag gtcccggccg aaggcgatgc gcgcaggggg tcgggcagct		250
gggctcgggc ggcgggagta gggcccgga gggaggcagg gaggtgcat		300
attcagagtc gcgggctgcg ccctgggcag aggcgcacct cgctccacgc		350
aacacctgct gctgccaccg cgcgcgatg agccgcgtgg tctcgtgct		400
gctggggccc gcgctgctct gcggccacgg agccttctgc cgcgcgtgg		450
tcagcgccaa aaaggtgtgt tttgtgact tcaagcatcc ctgctacaaa		500
atggcctact tccatgaact gtccagccga gtgagcttcc aggaggcacg		550
cctggtctgt gagagtgagg gaggagtcct cctcagcctt gagaatgaag		600
cagaacagaa gttaatagag agcatgttgc aaaacctgac aaaaccggg		650
acagggattt ctgatggtga tttctggata gggctttgga ggaatggaga		700
tgggcaaaac tctggtgcct gccagatct ctaccagtgg tctgatggaa		750
gcaattccca gtaccgaaac tggtagacag atgaaccttc ctgcggaagt		800
gaaaagtgtg ttgtgatgta tcaccaacca actgccaatc ctggccttgg		850
gggtccctac ctttaccagt ggaatgatga caggtgtaac atgaagcaca		900
attatatttg caagtatgaa ccagagatta atccaacagc ccctgtagaa		950
aagccttacc ttacaaatca accaggagac acccatcaga atgtggtgt		1000
tactgaagca ggtataattc ccaatctaata ttatgttgtt ataccaacaa		1050
taccctgct cttactgata ctggttgctt ttggaacctg ttgtttccag		1100
atgtgcgata aaagtaaagg aagaacaaaa actagtccaa accagtctac		1150
actgtggatt tcaaagagta ccagaaaaga aagtggcatg gaagtataat		1200
aactcattga cttgggtcca gaattttgta attctggatc tgtataagga		1250
atggcatcag aacaatagct tggaatggct tgaaatcaca aaggatctgc		1300
aagatgaact gtaagctccc ccttgaggca aatattaaag taatttttat		1350
atgtctatta ttccatttaa agaatatgct gtgctaataa tggagtgaga		1400
catgcttatt ttgctaaagg atgcacccaa acttcaaact tcaagcaaat		1450
gaaatggaca atgcagataa agttgttatc aacacgtcgg gagtatgtgt		1500
gttagaagca attcctttta tttctttcac ctttcataag ttgttatcta		1550
gtcaatgtaa tgtatatgtt attgaaattt acagtgtgca aaagtatttt		1600

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acctttgcat aagtgtttga taaaaatgaa ctgttctaatt atttattttt	1650
atggcatctc atttttcaat acatgctctt ttgattaaag aaacttatta	1700
ctgttgtcaa ctgaattcac acacacacaa atatagtacc atagaaaaag	1750
tttgtttttc cgaataaatt catctttcag cttctctgct ttgtgtcaat	1800
gtctaggaaa tctcttcaga aataagaagc tatttcatta agtgtgatat	1850
aaacctctc aaacatttta cttagaggca aggattgtct aatttcaatt	1900
gtgcaagaca tgtgccttat aattattttt agcttaaaat taaacagatt	1950
ttgtaataat gtaactttgt taatagggtc ataaacacta atgcagtcaa	2000
tttgaacaaa agaagtgaca tacacaatat aaatcatatg tcttcacacg	2050
ttgcctatat aatgagaagc agctctctga gggttctgaa atcaatgtgg	2100
tccctctctt gccactaaa caaagatggg tgttcggggg ttgggattga	2150
cactggaggc agatagtgtc aaagttagtc taaggtttcc ctgctgtat	2200
ttagcctctg actatattag tatacaaaga ggtcatgtgg ttgagaccag	2250
gtgaatagtc actatcagtg tggagacaag cacagcacac agacatttta	2300
ggaaggaaa gaactacgaa atcgtgtgaa aatgggttgg aaccatcag	2350
tgatcgcata ttcatgtatg agggtttgct tgagatagaa aatggtggct	2400
cctttctgtc ttatctccta gtttcttcaa tgcttacgcc ttgttcttct	2450
caagagaaa ttgtaactct ctggtcttca tatgtccctg tgcctctttt	2500
aaccaaataa agagtctctg tttctggggg aaaaaaaaaa aaaaaaaaaa	2550
aaaaaaaaa aaaaaaaaaa	2570

<210> SEQ ID NO 168

<211> LENGTH: 273

<212> TYPE: PRT

<213> ORGANISM: Homo Sapien

<400> SEQUENCE: 168

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

-continued														
140				145				150						
Pro	Thr	Ala	Asn	Pro	Gly	Leu	Gly	Gly	Pro	Tyr	Leu	Tyr	Gln	Trp
155				160				165						
Asn	Asp	Asp	Arg	Cys	Asn	Met	Lys	His	Asn	Tyr	Ile	Cys	Lys	Tyr
170				175				180						
Glu	Pro	Glu	Ile	Asn	Pro	Thr	Ala	Pro	Val	Glu	Lys	Pro	Tyr	Leu
185				190				195						
Thr	Asn	Gln	Pro	Gly	Asp	Thr	His	Gln	Asn	Val	Val	Val	Thr	Glu
200				205				210						
Ala	Gly	Ile	Ile	Pro	Asn	Leu	Ile	Tyr	Val	Val	Ile	Pro	Thr	Ile
215				220				225						
Pro	Leu	Leu	Leu	Leu	Ile	Leu	Val	Ala	Phe	Gly	Thr	Cys	Cys	Phe
230				235				240						
Gln	Met	Leu	His	Lys	Ser	Lys	Gly	Arg	Thr	Lys	Thr	Ser	Pro	Asn
245				250				255						
Gln	Ser	Thr	Leu	Trp	Ile	Ser	Lys	Ser	Thr	Arg	Lys	Glu	Ser	Gly
260				265				270						
Met Glu Val														
<210> SEQ ID NO 169														
<211> LENGTH: 43														
<212> TYPE: DNA														
<213> ORGANISM: Artificial Sequence														
<220> FEATURE:														
<223> OTHER INFORMATION: Synthetic oligonucleotide probe														
<400> SEQUENCE: 169														
tgtaaaacga cggccagtta aatagacctg caattattaa tct										43				
<210> SEQ ID NO 170														
<211> LENGTH: 41														
<212> TYPE: DNA														
<213> ORGANISM: Artificial Sequence														
<220> FEATURE:														
<223> OTHER INFORMATION: Synthetic oligonucleotide probe														
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caggaaacag ctatgaccac ctgcacacct gcaaatccat t										41				

1. An isolated polypeptide having at least 80% amino acid sequence identity to:

(a) the amino acid sequence of the polypeptide shown in **FIG. 6** (SEQ ID NO:6);

(b) the amino acid sequence of the polypeptide shown in **FIG. 6** (SEQ ID NO:6), lacking its associated signal peptide;

(c) the amino acid sequence of the extracellular domain of the polypeptide shown in **FIG. 6** (SEQ ID NO:6);

(d) the amino acid sequence of the extracellular domain of the polypeptide shown in **FIG. 6** (SEQ ID NO:6), lacking its associated signal peptide; or

(e) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 209399.
2. The isolated polypeptide of claim 1 having at least 85% amino acid sequence identity to:

(a) the amino acid sequence of the polypeptide shown in **FIG. 6** (SEQ ID NO:6);

(b) the amino acid sequence of the polypeptide shown in **FIG. 6** (SEQ ID NO:6), lacking its associated signal peptide;

(c) the amino acid sequence of the extracellular domain of the polypeptide shown in **FIG. 6** (SEQ ID NO:6);

(d) the amino acid sequence of the extracellular domain of the polypeptide shown in **FIG. 6** (SEQ ID NO:6), lacking its associated signal peptide; or

(e) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 209399.
3. The isolated polypeptide of claim 1 having at least 90% amino acid sequence identity to:

- (a) the amino acid sequence of the polypeptide shown in **FIG. 6** (SEQ ID NO:6);
 - (b) the amino acid sequence of the polypeptide shown in **FIG. 6** (SEQ ID NO:6), lacking its associated signal peptide;
 - (c) the amino acid sequence of the extracellular domain of the polypeptide shown in **FIG. 6** (SEQ ID NO:6);
 - (d) the amino acid sequence of the extracellular domain of the polypeptide shown in **FIG. 6** (SEQ ID NO:6), lacking its associated signal peptide; or
 - (e) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 209399.
4. The isolated polypeptide of claim 1 having at least 95% amino acid sequence identity to:
- (a) the amino acid sequence of the polypeptide shown in **FIG. 6** (SEQ ID NO:6);
 - (b) the amino acid sequence of the polypeptide shown in **FIG. 6** (SEQ ID NO:6), lacking its associated signal peptide;
 - (c) the amino acid sequence of the extracellular domain of the polypeptide shown in **FIG. 6** (SEQ ID NO:6);
 - (d) the amino acid sequence of the extracellular domain of the polypeptide shown in **FIG. 6** (SEQ ID NO:6), lacking its associated signal peptide; or
 - (e) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 209399.
5. The isolated polypeptide of claim 1 having at least 99% amino acid sequence identity to:
- (a) the amino acid sequence of the polypeptide shown in **FIG. 6** (SEQ ID NO:6);
 - (b) the amino acid sequence of the polypeptide shown in **FIG. 6** (SEQ ID NO:6), lacking its associated signal peptide;
 - (c) the amino acid sequence of the extracellular domain of the polypeptide shown in **FIG. 6** (SEQ ID NO:6);
 - (d) the amino acid sequence of the extracellular domain of the polypeptide shown in **FIG. 6** (SEQ ID NO:6), lacking its associated signal peptide; or
 - (e) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 209399.
6. An isolated polypeptide comprising:
- (a) the amino acid sequence of the polypeptide shown in **FIG. 6** (SEQ ID NO:6);
 - (b) the amino acid sequence of the polypeptide shown in **FIG. 6** (SEQ ID NO:6), lacking its associated signal peptide;
 - (c) the amino acid sequence of the extracellular domain of the polypeptide shown in **FIG. 6** (SEQ ID NO:6);
 - (d) the amino acid sequence of the extracellular domain of the polypeptide shown in **FIG. 6** (SEQ ID NO:6), lacking its associated signal peptide; or
 - (e) the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 209399.
7. The isolated polypeptide of claim 6 comprising the amino acid sequence of the polypeptide shown in **FIG. 6** (SEQ ID NO:6).
8. The isolated polypeptide of claim 6 comprising the amino acid sequence of the polypeptide shown in **FIG. 6** (SEQ ID NO:6), lacking its associated signal peptide.
9. The isolated polypeptide of claim 6 comprising the amino acid sequence of the extracellular domain of the polypeptide shown in **FIG. 6** (SEQ ID NO:6).
10. The isolated polypeptide of claim 6 comprising the amino acid sequence of the extracellular domain of the polypeptide shown in **FIG. 6** (SEQ ID NO:6), lacking its associated signal peptide.
11. The isolated polypeptide of claim 6 comprising the amino acid sequence of the polypeptide encoded by the full-length coding sequence of the cDNA deposited under ATCC accession number 209399.
12. A chimeric polypeptide comprising a polypeptide according to claim 1 fused to a heterologous polypeptide.
13. The chimeric polypeptide of claim 12, wherein said heterologous polypeptide is an epitope tag or an Fc region of an immunoglobulin.

* * * * *