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(54) Title: CHLAMYDIA ANTIGENS

(57) Abstract: The invention provides identifies Chlamydia antigens for use in the treatment, prevention and/or diagnosis of Chlamydia infection. In particular, the invention provides antigens CT733, CT1 53, CT601, CT279, CT443, CT372, CT456, CT381, CT255, CT341, CT716, CT745, CT387, CT812, CT869, CT166, CT175, CT163, CT214, CT721, CT127, CT043, CT823 and/or CT600 from *C. trachomatis* for the treatment, prevention or diagnosis of Chlamydia infection.

CHLAMYDIA ANTIGENS**TECHNICAL FIELD**

This invention is in the field of *Chlamydia trachomatis* proteins and their uses.

BACKGROUND ART

5 Vaccine development has been identified as essential to controlling infection with *C. trachomatis*. Vaccines against *C. trachomatis* appear to elicit protective T-cell and/or B-cell immunity in the genital tract mucosa.

Protective immunity to *C. trachomatis* seems to depend on a Th1-polarized cell-mediated immune response, in particular on CD4⁺ lymphocytes secreting IFN γ . For example, depletion of CD4⁺ T
10 cells in mice results in loss of protective immunity, and adoptive transfer of Chlamydia-specific CD4⁺ T cells confers protection against challenge with *C. trachomatis*. Furthermore, recent studies report that *C. trachomatis* infection in mice induces a CD4-Th1 protective immune response, indicating that critical Chlamydia antigens are processed and presented via the MHC class II pathway (Brunham RC and Rey-Ladino J (2005), Nat Rev Immunol 5: 149-1611; Su H and Caldwell HD
15 (1995), Infect Immun 63: 3302-3308).

Although B-cells and antibodies do not have a decisive role in resolution of primary infection, they are likely to be important for enhancing the protective effector T-cell response and to be required to control re-infection with various mechanisms such as antibody-mediated neutralization and opsonization.

20 Because immune protection against infection with *C. trachomatis* is likely to be mediated by immunization with *C. trachomatis* proteins that are targets of CD4⁺ T cells and that are capable of inducing B-cell responses, identification of such proteins is particularly important. It is therefore an object of the invention to provide further antigens for use in Chlamydia vaccines.

DISCLOSURE OF THE INVENTION

25 The invention provides identifies Chlamydia antigens for use in the treatment, prevention and/or diagnosis of Chlamydia infection. In particular, the invention provides one or more of the following antigens (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 20, 21, 22, 23, 24, 25, 26, 27, 28, 29 or 30) from *C. trachomatis* for the treatment, prevention or diagnosis of Chlamydia infection (and, in particular, *C. trachomatis* infection): CT733, CT153, CT601, CT279, CT443,
30 CT372, CT456, CT381, CT255, CT341, CT716, CT745, CT812, CT869, CT387, CT166, CT175, CT163, CT214, CT721, CT127, CT043, CT823, CT600, CT711, CT114, CT480, CT089, CT734 and CT016 for example, one or more of CT733, CT153, CT601, CT279, CT443, CT372, CT456, CT381, CT255, CT341, CT716 and CT745.

In particular, the invention provides proteins for use in the treatment, prevention and/or diagnosis of Chlamydia infection (and, in particular, *C. trachomatis* infection). Immunisation with the proteins is preferably able to induce a specific CD4+ Th1 cell mediated response against Chlamydia.

In one embodiment, the nucleic acid sequence and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:1 and SEQ ID NO:2 respectively. This protein is also known as "CT733" and is annotated as a hypothetical protein from *C. trachomatis*. In another embodiment, the nucleic acid sequence and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:3 and SEQ ID NO:4 respectively. This protein is also known as "CT153" and is annotated as MACPF/ membrane-attack complex (MAC)/ perforin from *C. trachomatis*. In another embodiment, the nucleic acid sequence and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:5 and SEQ ID NO:6 respectively. This protein is also known as "CT601" from *C. trachomatis*. In another embodiment, the nucleic acid sequence and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:7 and SEQ ID NO:8 respectively. This protein is also known as "CT279" from *C. trachomatis*. In another embodiment, the nucleic acid sequence and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:9 and SEQ ID NO:10 respectively. This protein is also known as "CT443" from *C. trachomatis*. In another embodiment, the nucleic acid sequence and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:11 and SEQ ID NO:12 respectively. This protein is also known as "CT372" from *C. trachomatis*. In another embodiment, the nucleic acid sequence and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:13 and SEQ ID NO:14 respectively. This protein is also known as "CT456" from *C. trachomatis*. In another embodiment, the nucleic acid sequence and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:15 and SEQ ID NO:16 respectively. This protein is also known as "CT381" from *C. trachomatis*. In another embodiment, the nucleic acid sequence and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:39 and SEQ ID NO:40 respectively. This protein is also known as "CT255" from *C. trachomatis*. In another embodiment, the nucleic acid sequence and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:41 and SEQ ID NO:42 respectively. This protein is also known as "CT341" from *C. trachomatis*. In another embodiment, the nucleic acid sequence and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:43 and SEQ ID NO:44 respectively. This protein is also known as "CT716" from *C. trachomatis*. In another embodiment, the nucleic acid sequence and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:45 and SEQ ID NO:46 respectively. This protein is also known as "CT745" from *C. trachomatis*. In another embodiment, the nucleic acid sequence and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:47 and SEQ ID NO:48, respectively. This protein is also known as "CT387" from *C. trachomatis* and is annotated as a hypothetical protein. In another embodiment, the nucleic acid and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:49 and

SEQ ID NO:50, respectively. This protein is also known as “CT812” from *C. trachomatis* and is annotated as a polymorphic outer membrane protein. In another embodiment, the nucleic acid and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:51 and SEQ ID NO:52, respectively. This protein is also known as “CT869” from *C. trachomatis* and is annotated as a polymorphic outer membrane protein. In another embodiment, the nucleic acid and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:53 and SEQ ID NO:54, respectively. This protein is also known as “CT166” from *C. trachomatis*. In another embodiment, the nucleic acid and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:55 and SEQ ID NO:56, respectively. This protein is also known as “CT175” from *C. trachomatis*. In another embodiment, the nucleic acid and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:155 and SEQ ID NO:156, respectively. This protein is also known as “CT163” from *C. trachomatis*. In another embodiment, the nucleic acid and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:159 and SEQ ID NO:160, respectively. This protein is also known as “CT214” from *C. trachomatis*. In another embodiment, the nucleic acid and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:163 and SEQ ID NO:164, respectively. This protein is also known as “CT721” from *C. trachomatis*. In another embodiment, the nucleic acid and/or amino acid sequence of the protein comprises the sequence presented in SEQ ID NO:167 and SEQ ID NO:168, respectively. This protein is also known as “CT127” from *C. trachomatis*.

In some embodiments, the protein is a variant of a protein as described above. For example, the protein may comprise one or more mutations (for example, 1, 2, 3, 4, 5, 6, 7, 8, 9, 10 or more mutations) in the sequence of SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 18, 19, 20, 21, 22, 23, 24, 40, 42, 44, 46, 48, 50, 52, 54, 56, 136, 140, 156, 160, 164 or 168, for example, in the sequence of SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 40, 42, 44, or 46. Preferred mutations are those which do not cause a significant conformational change in the protein such that the protein of the invention retains the ability to elicit an immune response against the wild-type Chlamydia protein. The proteins having the sequences presented in SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 40, 42, 44, 46, 48, 50, 52, 54 and 56 are the wild-type proteins.

In some embodiments, the one or more mutations are present in the N-terminal portion of the protein, for example, between residues 1 and 20 of the protein, between residues 21 and 40, between residues 41 and 60, between residues 1 and 60 or between residues 1 and 40 of the protein. In some embodiments, the one or more mutations are present in the C-terminal portion of the protein, for example, between the C-terminal 20 residues of the protein, between residues 21 and 40 from the C-terminus, between residues 41 and 60 from the C-terminus; between residues 1 and 60 from the C-terminus or between residues 1 and 40 from the C-terminus of the protein.

Preferably, the amino acid sequences contain fewer than twenty mutations (e.g. 19, 18, 17, 16, 15, 14, 13, 12, 11, 10, 9, 8, 7, 6, 5, 4, 3, 2 or 1). Each mutation preferably involves a single amino acid

and is preferably a point mutation. The mutations may each independently be a substitution, an insertion or a deletion. Preferred mutations are single amino acid substitutions. The proteins may also include one or more (*e.g.* 1, 2, 3, 4, 5, 6, 7, 8, 9, *etc.*) single amino acid deletions relative to the Chlamydia sequences. The proteins may also include one or more (*e.g.* 1, 2, 3, 4, 5, 6, 7, 8, 9, *etc.*) insertions (*e.g.* each of 1, 2, 3, 4 or 5 or more amino acids) relative to the Chlamydia sequences. Deletions, substitutions or insertions may be at the N-terminus and/or C-terminus, or may be between the two termini. Thus a truncation is an example of a deletion. Truncations may involve deletion of up to 40 (or more) amino acids at the N-terminus and/or C-terminus (for example, 1-10, 11-40, 41-70, 71-100 or more amino acids).

Amino acid substitutions may be to any one of the other nineteen naturally occurring amino acids. Preferably, a substitution mutation is a conservative substitution. Alternatively, a substitution mutation is a non-conservative substitution. A conservative substitution is commonly defined as a substitution introducing an amino acid having sufficiently similar chemical properties, *e.g.* having a related side chain (*e.g.* a basic, positively charged amino acid should be replaced by another basic, positively charged amino acid), in order to preserve the structure and the biological function of the molecule. Genetically-encoded amino acids are generally divided into four families: (1) acidic *i.e.* aspartate, glutamate; (2) basic *i.e.* lysine, arginine, histidine; (3) non-polar *i.e.* alanine, valine, leucine, isoleucine, proline, phenylalanine, methionine, tryptophan; and (4) uncharged polar *i.e.* glycine, asparagine, glutamine, cysteine, serine, threonine, tyrosine. Phenylalanine, tryptophan, and tyrosine are sometimes classified jointly as aromatic amino acids. In general, substitution of single amino acids within these families does not have a major effect on the biological activity. Further examples of conservative substitutions that may be used in the invention are presented in Table 1.

TABLE 1

Amino Acid	Synonymous Groups	More Preferred Synonymous Groups
Ser	Gly, Ala, Ser, Thr, Pro	Thr, Ser
Arg	Asn, Lys, Gln, Arg, His	Arg, Lys, His
Leu	Phe, Ile, Val, Leu, Met	Ile, Val, Leu, Met
Pro	Gly, Ala, Ser, Thr, Pro	Pro
Thr	Gly, Ala, Ser, Thr, Pro	Thr, Ser
Ala	Gly, Thr, Pro, Ala, Ser	Gly, Ala
Val	Met, Phe, Ile, Leu, Val	Met, Ile, Val, Leu
Gly	Ala, Thr, Pro, Ser, Gly	Gly, Ala
Ile	Phe, Ile, Val, Leu, Met	Ile, Val, Leu, Met
Phe	Trp, Phe, Tyr	Tyr, Phe
Tyr	Trp, Phe, Tyr	Phe, Tyr
Cys	Ser, Thr, Cys	Cys
His	Asn, Lys, Gln, Arg, His	Arg, Lys, His
Gln	Glu, Asn, Asp, Gln	Asn, Gln
Asn	Glu, Asn, Asp, Gln	Asn, Gln
Lys	Asn, Lys, Gln, Arg, His	Arg, Lys, His

Asp	Glu, Asn, Asp, Gln	Asp, Glu
Glu	Glu, Asn, Asp, Gln	Asp, Glu
Met	Phe, Ile, Val, Leu, Met	Ile, Val, Leu, Met
Trp	Trp, Phe, Tyr	Trp

5 Examples of non-conservative substitutions that may be used in the invention include the substitution of an uncharged polar amino acid with a nonpolar amino acid, the substitution of a nonpolar amino acid with an uncharged polar amino acid, the substitution of an acidic amino acid with a basic amino acid and the substitution of a basic amino acid with an acidic amino acid.

Mutations may also be introduced to improve stability, *e.g.*, the insertion of disulphide bonds (van den Akker et al. Protein Sci., 1997, 6:2644-2649). For example, the protein may comprise an amino acid sequence having sequence identity to the amino acid sequence of any one of SEQ ID NOs: 2, 4, 6, 8, 10, 12, 14, 16, 18, 19, 20, 21, 22, 23, 24, 40, 42, 44, 46, 48, 50, 52, 54, 56, 136, 140, 156, 160, 164 and 168, for example, of any one of SEQ ID NOs: 2, 4, 6, 8, 10, 12, 14, 16, 40, 42, 44 and 46. The degree of sequence identity is preferably greater than 50% (*e.g.* 60%, 70%, 80%, 90%, 95%, 97%, 98%, 99% or more). These proteins include homologs, orthologs, allelic variants and functional mutants. Identity between proteins is preferably determined by the Smith-Waterman homology search algorithm as implemented in the MPSRCH program (Oxford Molecular), using an affine gap search with parameters gap open penalty = 12 and gap extension penalty = 1.

20 The Chlamydia protein of the invention may comprise one or more amino acid derivatives. By "amino acid derivative" is intended an amino acid or amino acid-like chemical entity other than one of the 20 genetically encoded naturally occurring amino acids. In particular, the amino acid derivative may contain substituted or non-substituted, linear, branched, or cyclic alkyl moieties, and may include one or more heteroatoms. The amino acid derivatives can be made *de novo* or obtained from commercial sources (Calbiochem-Novabiochem AG; Bachem).

In some embodiments, the variant protein is a homologous protein from *C. pneumoniae*, *C. psittaci*, *C. pecorum*, *C. muridarum* or *C. suis*.

25 The invention further provides a protein comprising or consisting of a fragment of a protein comprising or consisting of the amino acid sequence of any of SEQ ID NOs: 2, 4, 6, 8, 10, 12, 14, 16, 18, 19, 20, 21, 22, 23, 24, 40, 42, 44, 46, 48, 50, 52, 54, 56, 136, 140, 156, 160, 164 or 168, for example, of any one of SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 40, 42, 44 or 46, or a fragment of a variant thereof. The fragment should comprise at least *n* consecutive amino acids from the protein and, depending on the particular sequence, *n* is 6 or more (*e.g.* 8, 11, 16, 31, 51, 76, 121, 181, 231, 281, 331, 381, 431, 440, 445, 446, 481, 531, 581, 631, 681, 731, 781, 801, 806, 808 or more). The fragment is *n*-1 amino acids or less in length, wherein *n* = the number of amino acids in the full length protein (*e.g.* *n*-5, *n*-20, *n*-50, *n*-110, *n*-180, *n*-240, *n*-310, *n*-380, *n*-445, *n*-515, *n*-595, *n*-675, *n*-745, *n*-785, *n*-800 amino acids or less in length). Preferably the fragment comprises one or more

epitopes from the protein. Preferably, one or more of the epitopes is an MHC class II epitope, for example, a CD4+ T cell epitope. In some embodiments, the fragment comprises or consists of the amino acid sequence of any of SEQ ID NOs 64, 66, 68, 70, 72, 74, 76, 78, 80, 82, 84, 86, 88, 90, 92, 94, 96, 98, 100, 102, 104, 106, 108, 110, 112, 114, 116, 118, 120, 122, 124, 126, 128, 130, 132, 134, 138, 142, 146, 150, 154, 158, 162, 166 and 170. In some embodiments, the invention provides a protein comprising or consisting of a fragment of a protein comprising or consisting of the amino acid sequence recited in SEQ ID NO: 122. Table 3 below shows which fragments correspond to which full length sequences.

TABLE 3

Annotation	SEQ ID NO. for full length sequence	SEQ ID NO. for fragment
CT733	1	63
CT733	2	64
CT153	3	65
CT153	4	66
CT601	5	67
CT601	6	68
CT279	7	69
CT279	8	70
CT443	9	71
CT443	10	72
CT372	11	73
CT372	12	74
CT456	13	75
CT456	14	76
CT381	15	77
CT381	16	78
CT043	17	79
CT043	18	80
CT711	19	81 (nucleotide); 82 (protein)
CT114	20	83 (nucleotide); 84 (protein)
CT480	21	85 (nucleotide); 86 (protein)
CT089	22	87 (nucleotide); 88 (protein)
CT734	23	89 (nucleotide); 90 (protein)
CT016	24	91 (nucleotide); 92 (protein)
TC0551 (CT279)	25	93

TC0551 (CT279)	26	94
TC0651 (CT372)	27	95
TC0651 (CT372)	28	96
TC0727 (CT443)	29	97
TC0727 (CT443)	30	98
TC0313 (CT043)	31	99
TC0313 (CT043)	32	100
TC0890 (CT601)	33	101
TC0890 (CT601)	34	102
TC0741 (CT456)	35	103
TC0741 (CT456)	36	104
TC0660 (CT381)	37	105
TC0660 (CT381)	38	106
CT255	39	107
CT255	40	108
CT341	41	109
CT341	42	110
CT716	43	111
CT716	44	112
CT745	45	113
CT745	46	114
CT387	47	115
CT387	48	116
CT812	49	117 (mature full length); 119 (N-terminal fragment); 121 (C-terminal fragment)
CT812	50	118 (mature full length) 120 (N-terminal fragment) 122 (C-terminal fragment)
CT869	51	123
CT869	52	124
CT166	53	125
CT166	54	126
CT175	55	127
CT175	56	128
TC0666 (CT387)	57	129
TC0666 (CT387)	58	130

TC0197	59	131
TC0197	60	132
TC0261	61	133
TC0261	62	134
CT600	135	137
CT600	136	138
CT823	139	141
CT823	140	142
TC0106	143	145
TC0106	144	146
TC0431	147	149
TC0431	148	150
TC0210	151	153
TC0210	152	154
CT163	155	157
CT163	156	158
CT214	159	161
CT214	160	162
CT721	163	165
CT721	164	166
CT127	167	169
CT127	168	170

The protein of the invention, for example the variant protein or the fragment, is preferably immunogenic.

The term “immunogenic” in the context of “an immunogenic variant” and “immunogenic fragment”,
5 is used to mean that the protein is capable of eliciting an immune response, such as a cell-mediated and/or an antibody response, against the wild-type Chlamydia protein from which it is derived, for example, when used to immunise a subject (preferably a mammal, more preferably a human or a mouse). For example, the protein of the invention (for example, the variant or fragment) is preferably capable of stimulating *in vitro* CD4⁺ IFN γ ⁺ cells in splenocytes purified from mice
10 infected with live *C. trachomatis* to a level comparable with the wild-type Chlamydia protein. The protein of the invention preferably retains the ability to elicit antibodies that recognise the wild-type protein. For example, the protein of the invention preferably elicits antibodies that can bind to, and preferably neutralise the activity of, the wild-type protein. In a further embodiment, the protein of the invention is capable of eliciting antibodies that are capable of neutralising Chlamydia infectivity
15 and/or virulence. In some embodiments, the antibodies are able to cross-react with the protein of the

invention and the wild-type protein, but with no other homologous protein (e.g. from another Chlamydia species). In other embodiments, the antibodies are cross-reactive with the wild-type protein and with homologous proteins from other Chlamydia species. In some embodiments, the antibodies are cross-reactive with the wild-type protein and with homologous protein from other organisms (for example from *E.coli* or *H.influenzae*). Mice immunized with the protein of the invention and the wild-type Chlamydia protein preferably show similar antigen-specific antibody titers. Antibody titres and specificities can be measured using standard methods available in the art. Other methods of testing the immunogenicity of proteins are also well known in the art.

For example, the variant or fragment is preferably capable of eliciting an immune response, such as a cell-mediated and/or an antibody response, against the wild-type Chlamydia protein. In one embodiment the fragment is capable of stimulating *in vitro* CD4+ IFN γ + cells in splenocytes purified from mice infected with live *C. trachomatis* to a level comparable with the wild-type Chlamydia protein and/or retains the ability to elicit antibodies that recognise the wild-type protein.

Preferably, the variant or the fragment is capable of inducing a specific CD4-Th1 cell mediated response against the wild type Chlamydia protein.

The proteins of the invention can, of course, be prepared by various means (e.g. recombinant expression, purification from native host, purification from cell culture, chemical synthesis *etc.*) and in various forms (e.g. native, fusions, glycosylated, non-glycosylated, lipidated, non-lipidated, phosphorylated, non-phosphorylated, myristoylated, non-myristoylated, monomeric, multimeric, particulate, denatured, *etc.*). Generally, the recombinant fusion proteins of the present invention are prepared as a GST-fusion protein and/or a His-tagged fusion protein.

The proteins of the invention are preferably prepared in purified or substantially pure form (*i.e.* substantially free from host cell proteins and/or other *Chlamydia* proteins), and are generally at least about 50% pure (by weight), and usually at least about 90% pure, *i.e.* less than about 50%, and more preferably less than about 10% (e.g. 5%) of a composition is made up of other expressed polypeptides. Thus the antigens in the compositions are separated from the whole organism with which the molecule is expressed.

Whilst expression of the proteins of the invention may take place in *Chlamydia*, the invention preferably utilises a heterologous host. The heterologous host may be prokaryotic (e.g. a bacterium) or eukaryotic. It is preferably *E.coli*, but other suitable hosts include *Bacillus subtilis*, *Vibrio cholerae*, *Salmonella typhi*, *Salmonella typhimurium*, *Neisseria lactamica*, *Neisseria cinerea*, *Mycobacteria* (e.g. *M.tuberculosis*), yeasts, *etc.*

The term "polypeptide" or "protein" refers to amino acid polymers of any length. The polymer may be linear or branched, it may comprise modified amino acids, and it may be interrupted by non-amino acids. The terms also encompass an amino acid polymer that has been modified naturally or by intervention; for example, disulfide bond formation, glycosylation, lipidation, acetylation,

phosphorylation, or any other manipulation or modification, such as conjugation with a labeling component. Also included are, for example, polypeptides containing one or more analogs of an amino acid (including, for example, unnatural amino acids, *etc.*), as well as other modifications known in the art. Polypeptides can occur as single chains or associated chains.

- 5 The invention provides polypeptides comprising a sequence -P-Q- or -Q-P-, wherein: -P- is an amino acid sequence as defined above and -Q- is not a sequence as defined above *i.e.* the invention provides fusion proteins. Where the N-terminus codon of -P- is not ATG, but this codon is not present at the N-terminus of a polypeptide, it will be translated as the standard amino acid for that codon rather than as a Met. Where this codon is at the N-terminus of a polypeptide, however, it will be translated
10 as Met. Examples of -Q- moieties include, but are not limited to, histidine tags (*i.e.* His_n where *n* = 3, 4, 5, 6, 7, 8, 9, 10 or more), maltose-binding protein, or glutathione-S-transferase (GST).

Proteins of the invention may be attached to a solid support. They may comprise a detectable label (*e.g.* a radioactive or fluorescent label, or a biotin label).

Antibodies

- 15 The proteins of the invention induce antibodies that may be used as a vaccine capable of neutralising the activity of infectious EB. The antibodies may alternatively be used for the diagnosis of Chlamydia infection. Thus, the invention provides antibodies for use in the treatment, prevention or diagnosis of Chlamydia infection. Preferably, the infection is by *C. trachomatis*, but may alternatively be by *C. psittaci*, *C. pecorum*, *C. muridarum* or *C. suis*.
- 20 The term "antibody" includes intact immunoglobulin molecules, as well as fragments thereof which are capable of binding an antigen. These include hybrid (chimeric) antibody molecules (Winter *et al.*, (1991) *Nature* 349:293-99; US 4,816,567); F(ab')₂ and F(ab) fragments and Fv molecules; non-covalent heterodimers (Inbar *et al.*, (1972) *Proc. Natl. Acad. Sci. U.S.A.* 69:2659-62; Ehrlich *et al.*, (1980) *Biochem* 19:4091-96); single-chain Fv molecules (sFv) (Huston *et al.*, (1988) *Proc. Natl.*
25 *Acad. Sci. U.S.A.* 85:5897-83); dimeric and trimeric antibody fragment constructs; minibodies Pack *et al.*, (1992) *Biochem* 31, 1579-84; Cumber *et al.*, (1992) *J. Immunology* 149B, 120-26); humanized antibody molecules (Riechmann *et al.*, (1988) *Nature* 332, 323-27; Verhoeyan *et al.*, (1988) *Science* 239, 1534-36; and GB 2,276,169); and any functional fragments obtained from such molecules, as well as antibodies obtained through non-conventional processes such as phage display. Preferably,
30 the antibodies are monoclonal antibodies. Methods of obtaining monoclonal antibodies are well known in the art. Humanised or fully-human antibodies are preferred.

The antibodies may be polyclonal or monoclonal and may be produced by any suitable means. The antibody may include a detectable label.

- 35 Also provided is a method for preparing antibodies comprising immunising a mammal (such as a mouse or a rabbit) with a protein of the invention and obtaining polyclonal antibodies or monoclonal antibodies by conventional techniques. For example, polyclonal antisera may be

obtained by bleeding the immunized animal into a glass or plastic container, incubating the blood at 25°C for one hour, followed by incubating at 4°C for 2-18 hours. The serum is recovered by centrifugation (eg. 1,000g for 10 minutes). Monoclonal antibodies may be prepared using the standard method of Kohler & Milstein [*Nature* (1975) 256:495-96], or a modification thereof, or by any other suitable method.

Nucleic acids

According to a further aspect, the invention provides a nucleic acid encoding a protein or antibody of the invention. In some embodiments, the nucleic acid sequence encoding a protein of the invention preferably comprises or consists of any one of SEQ ID NOs:1, 3, 5, 7, 9, 11, 13, 15, 17, 39, 41, 43, 45, 47, 49, 51, 53, 55, 135, 139, 155, 159, 163 or 167, for example, of any one of SEQ ID NOs:1, 3, 5, 7, 9, 11, 13, 15, 39, 41, 43 or 45. In some embodiments, the nucleic acid sequence encoding a protein of the invention comprises or consists of any one of SEQ ID NOs: 63, 65, 67, 69, 71, 73, 75, 77, 79, 81, 83, 85, 87, 89, 91, 93, 95, 97, 99, 101, 103, 105, 107, 109, 111, 113, 115, 117, 119, 121, 123, 125, 127, 129, 131 and 133.

The invention also provides nucleic acid comprising nucleotide sequences having sequence identity to such nucleotide sequences. Identity between sequences is preferably determined by the Smith-Waterman homology search algorithm as described above. Such nucleic acids include those using alternative codons to encode the same amino acid.

The invention also provides nucleic acid which can hybridize to these nucleic acids. Hybridization reactions can be performed under conditions of different "stringency". Conditions that increase stringency of a hybridization reaction of widely known and published in the art (e.g. page 7.52 of Kaplitt, *Nature Genetics* (1994) 6:148). Examples of relevant conditions include (in order of increasing stringency): incubation temperatures of 25°C, 37°C, 50°C, 55°C and 68°C; buffer concentrations of 10 x SSC, 6 x SSC, 1 x SSC, 0.1 x SSC (where SSC is 0.15 M NaCl and 15 mM citrate buffer) and their equivalents using other buffer systems; formamide concentrations of 0%, 25%, 50%, and 75%; incubation times from 5 minutes to 24 hours; 1, 2, or more washing steps; wash incubation times of 1, 2, or 15 minutes; and wash solutions of 6 x SSC, 1 x SSC, 0.1 x SSC, or de-ionized water. Hybridization techniques and their optimization are well known in the art (e.g. see US patent 5,707,829, *Current Protocols in Molecular Biology* (F.M. Ausubel *et al.* eds., 1987) Supplement 30, Kaplitt, *Nature Genetics* (1994) 6:148, and WO 94/03622, *etc.*).

The nucleic acid may be used in hybridisation reactions (e.g. Northern or Southern blots, or in nucleic acid microarrays or 'gene chips') or in amplification reactions (e.g. PCR, SDA, SSSR, LCR, NASBA, TMA) *etc.*

The invention also provides a nucleic acid comprising sequences complementary to those described above (e.g. for antisense or probing, or for use as primers). In one embodiment, the nucleic acid is complementary to the full length of the nucleic acid described above.

Nucleic acid according to the invention may be labelled *e.g.* with a radioactive or fluorescent label. This is particularly useful where the nucleic acid is to be used as a primer or probe *e.g.* in PCR, LCR or TMA.

5 The term “nucleic acid” includes in general means a polymeric form of nucleotides of any length, which contain deoxyribonucleotides, ribonucleotides, and/or their analogs. It includes DNA, RNA, DNA/RNA hybrids. It also includes DNA or RNA analogs, such as those containing modified backbones (*e.g.* peptide nucleic acids (PNAs) or phosphorothioates) or modified bases. Thus the invention includes mRNA, ribozymes, DNA, cDNA, recombinant nucleic acids, branched nucleic acids, plasmids, vectors, probes, primers, *etc.*. Where nucleic acid of the invention takes the form of
10 RNA, it may or may not have a 5' cap.

Nucleic acids of the invention can take various forms (*e.g.* single stranded, double stranded, vectors, primers, probes *etc.*). Unless otherwise specified or required, any embodiment of the invention that utilizes a nucleic acid may utilize both the double-stranded form and each of two complementary single-stranded forms which make up the double-stranded form. Primers and probes are generally
15 single-stranded, as are antisense nucleic acids.

Nucleic acids of the invention are preferably prepared in substantially pure form (*i.e.* substantially free from naturally-occurring nucleic acids, particularly from chlamydial or other host cell nucleic acids), generally being at least about 50% pure (by weight), and usually at least about 90% pure.

20 Nucleic acids of the invention may be prepared in many ways *e.g.* by chemical synthesis (*e.g.* phosphoramidite synthesis of DNA) in whole or in part, by digesting longer nucleic acids using nucleases (*e.g.* restriction enzymes), by joining shorter nucleic acids or nucleotides (*e.g.* using ligases or polymerases), from genomic or cDNA libraries, *etc.*

The invention provides vectors comprising nucleotide sequences of the invention (*e.g.* cloning or expression vectors) and host cells transformed with such vectors. Nucleic acids of the invention may
25 be part of a vector *i.e.* part of a nucleic acid construct designed for transduction/transfection of one or more cell types. Vectors may be, for example, “cloning vectors” which are designed for isolation, propagation and replication of inserted nucleotides, “expression vectors” which are designed for expression of a nucleotide sequence in a host cell, “viral vectors” which are designed to result in the production of a recombinant virus or virus-like particle, or “shuttle vectors”, which comprise the
30 attributes of more than one type of vector. Preferred vectors are plasmids.

Also provided is a host cell comprising a nucleic acid of the invention. A “host cell” includes an individual cell or cell culture which can be or has been a recipient of exogenous nucleic acid. Host cells include progeny of a single host cell, and the progeny may not necessarily be completely identical (in morphology or in total DNA complement) to the original parent cell due to natural,
35 accidental, or deliberate mutation and/or change. Host cells include cells transfected or infected *in vivo* or *in vitro* with nucleic acid of the invention, for example, with a vector of the invention.

Where a nucleic acid is DNA, it will be appreciated that “U” in a RNA sequence will be replaced by “T” in the DNA. Similarly, where a nucleic acid is RNA, it will be appreciated that “T” in a DNA sequence will be replaced by “U” in the RNA.

5 The term “complement” or “complementary” when used in relation to nucleic acids refers to Watson-Crick base pairing. Thus the complement of C is G, the complement of G is C, the complement of A is T (or U), and the complement of T (or U) is A. It is also possible to use bases such as I (the purine inosine) *e.g.* to complement pyrimidines (C or T).

10 Nucleic acids of the invention can be used, for example: to produce polypeptides; as hybridization probes for the detection of nucleic acid in biological samples; to generate additional copies of the nucleic acids; to generate ribozymes or antisense oligonucleotides; as single-stranded DNA primers or probes; or as triple-strand forming oligonucleotides.

The invention provides a process for producing nucleic acid of the invention, wherein the nucleic acid is synthesised in part or in whole using chemical means.

15 For certain embodiments of the invention, nucleic acids are preferably at least 24 nucleotides in length (*e.g.* 60, 120, 240, 390, 540, 720, 900, 1200, 1320, 1500, 1800, 2100, 2400, 2415 nucleotides or longer).

For certain embodiments of the invention, nucleic acids are preferably at most 2430 nucleotides in length (*e.g.* 2427, 2394, 2250, 2034, 1450, 1300, 1150, 1000, 850, 700, 500 nucleotides or shorter).

20 Primers and probes of the invention, and other nucleic acids used for hybridization, are preferably between 10 and 30 nucleotides in length (*e.g.* 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 20, 21, 22, 23, 24, 25, 26, 27, 28, 29, or 30 nucleotides).

Immunogenic compositions and medicaments

25 The protein, antibody, and/or nucleic acid or medicament may be in the form of a composition. These compositions may be suitable as immunogenic compositions (*e.g.* vaccines), or as diagnostic reagents.

Preferably, the composition is an immunogenic composition. It is particularly advantageous to use a protein of the invention in an immunogenic composition such as a vaccine. It is also envisaged that the immunogenic composition may comprise a nucleic acid which encodes a protein of the invention such that the protein is generated *in vivo*.

30 An immunogenic composition of the invention comprises a protein, antibody, nucleic acid, vector and/or host cell according to the invention. Immunogenic compositions according to the invention may either be prophylactic (*i.e.* to prevent infection) or therapeutic (*i.e.* to treat infection), but will typically be prophylactic. Where the immunogenic composition is for prophylactic use, the human is preferably a child (*e.g.* a toddler or infant) or a teenager; where the immunogenic composition is for

therapeutic use, the human is preferably a teenager or an adult. An immunogenic composition intended for children may also be administered to adults *e.g.* to assess safety, dosage, immunogenicity, *etc.*

In some embodiments, the immunogenic composition is for treatment or prevention of Chlamydia infection or an associated condition (*e.g.* trachoma, blindness, cervicitis, pelvic inflammatory disease, infertility, ectopic pregnancy, chronic pelvic pain, salpingitis, urethritis, epididymitis, infant pneumonia, patients infected with cervical squamous cell carcinoma, and/or HIV infection, *etc.*), preferably, *C. trachomatis* infection. The immunogenic composition may be effective against *C. pneumoniae*.

10 Immunogenic compositions used as vaccines comprise an immunologically effective amount of the protein of the invention, as well as any other components, as needed. By 'immunologically effective amount', it is meant that the administration of that amount to an individual, either in a single dose or as part of a series, is effective for treatment or prevention. This amount varies depending upon the health and physical condition of the individual to be treated, age, the taxonomic group of the individual to be treated (*e.g.* non-human primate, primate, *etc.*), the capacity of the individual's immune system to synthesise antibodies, the degree of protection desired, the formulation of the vaccine, the treating doctor's assessment of the medical situation, and other relevant factors. It is expected that the amount will fall in a relatively broad range that can be determined through routine trials.

20 Antigens in the composition will typically be present at a concentration of at least 1 µg/ml each.

In general, the concentration of any given antigen will be sufficient to elicit an immune response against that antigen.

Dosage treatment can be a single dose schedule or a multiple dose schedule. Multiple doses may be used in a primary immunisation schedule and/or in a booster immunisation schedule. In a multiple dose schedule the various doses may be given by the same or different routes *e.g.* a parenteral prime and mucosal boost, a mucosal prime and parenteral boost, *etc.* Multiple doses will typically be administered at least 1 week apart (*e.g.* about 2 weeks, about 3 weeks, about 4 weeks, about 6 weeks, about 8 weeks, about 10 weeks, about 12 weeks, about 16 weeks, *etc.*). In some embodiments, three or more doses are provided (for example, three, four or five) doses. In some embodiments, three doses are given intramuscularly at 2 week-intervals, for example, three doses of 10-20 µg of each protein, at 2 week-intervals, given intramuscularly.

The pH of an immunogenic composition is preferably between 6 and 8, preferably about 7. pH may be maintained by the use of a buffer. The composition may be sterile and/or pyrogen-free. The composition may be isotonic with respect to humans.

Immunogenic compositions of the invention will generally be administered directly to a patient. Direct delivery may be accomplished by parenteral injection (*e.g.* subcutaneously, intraperitoneally, intravenously, intramuscularly, or to the interstitial space of a tissue), or mucosally, such as by rectal, oral (*e.g.* tablet, spray), vaginal, topical, transdermal (*See e.g.* WO99/27961) or transcutaneous (*See e.g.* WO02/074244 and WO02/064162), intranasal (*See e.g.* WO03/028760), ocular, aural, pulmonary or other mucosal administration.

Chlamydia infections affect various areas of the body and so the immunogenic compositions of the invention may be prepared in various forms. For example, the compositions may be prepared as injectables, either as liquid solutions or suspensions. Solid forms suitable for solution in, or suspension in, liquid vehicles prior to injection can also be prepared (*e.g.* a lyophilised composition). The composition may be prepared for topical administration *e.g.* as an ointment, cream or powder. The composition may be prepared for oral administration *e.g.* as a tablet or capsule, or as a syrup (optionally flavoured). The composition may be prepared for pulmonary administration *e.g.* as an inhaler, using a fine powder or a spray. The composition may be prepared as a suppository or pessary. The composition may be prepared for nasal, aural or ocular administration *e.g.* as drops.

The invention also provides a delivery device pre-filled with an immunogenic composition of the invention.

The invention also provides a kit comprising a first component and a second component wherein neither the first component nor the second component is a composition of the invention as described herein, but wherein the first component and the second component can be combined to provide a composition of the invention as described herein. The kit may further include a third component comprising one or more of the following: instructions, syringe or other delivery device, adjuvant, or pharmaceutically acceptable formulating solution.

A composition as described above may alternatively and/or additionally be used for diagnosis of chlamydia infection.

Combinations with other antigens

The therapeutic or diagnostic efficiency of a *Chlamydia* antigen may be improved by combination with a different *Chlamydia* antigen. For example, the immunogenicity of a protein of the invention may be improved by combination with another protein of the invention or with another known *Chlamydia* antigen. The invention thus includes an immunogenic composition comprising a combination of *Chlamydia* antigens, said combination comprising a protein of the invention in combination with one or more additional *Chlamydia* antigens. The one or more additional *Chlamydia* antigens that are present in the composition may be in the form of a protein or nucleic acid or any other suitable form. A protein of the invention may be combined with one or more (*e.g.* 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20 or more) different proteins of the invention and/or with one or more (*e.g.* 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20 or more) other known *Chlamydia* antigens. For example, an

immunogenic composition is provided comprising two or more (e.g. 3, 4, 5, 6, 7, 8, 9, 10, 15, 20 or more) proteins of the invention. The proteins of the invention may alternatively and/or additionally be provided in the composition in the form of their corresponding nucleic acids, vectors, host cells, etc. Also provided is a protein or nucleic acid of the invention for a use as described above, wherein
5 the protein or nucleic acid is for use in combination with one or more additional Chlamydia antigens (or their encoding nucleic acids). The one or more additional antigens (e.g. 2, 3, 4, 5, 6, 7 or more additional antigens) may be administered simultaneously, separately or sequentially with the protein or nucleic acid of the invention, for example as a combined preparation.

Likewise, the antibodies of the invention may be used in combination with one or more antibodies
10 specific for one or more additional Chlamydia antigens for use in diagnosis of Chlamydia infections.

In one embodiment, one or more of the additional Chlamydia antigens is selected from the antigens presented in Table 2, or their variants. For example, one or more (for example, all) of the additional antigens are selected from the *Chlamydia trachomatis* antigens listed in Table 2, but may alternatively or additionally be selected from the *Chlamydia pneumoniae* antigens listed in Table 2.

15 In some embodiments, the one or more (for example, all) of the additional antigens are selected from the *Chlamydia trachomatis* antigens and/or *Chlamydia pneumoniae* antigens listed in Table 2 and CT387, CT812, CT869, CT166, CT175, CT163, CT214, CT721 and CT127. In one embodiment, one or more of the one or more additional antigens are selected from CT372, CT443, CT043, CT153, CT279, CT601, CT711, CT114, CT480, CT456, CT381, CT089, CT734, CT016, CT600, CT823,
20 CT387, CT812, CT869, CT166, CT175, CT163, CT214, CT721 and CT127 (or their variants), for example, from CT372, CT443, CT043, CT153, CT279, CT601, CT711, CT114, CT480, CT456, CT381, CT089, CT734, CT016, CT600 and CT823. These additional antigens are listed in Table 2 and their sequences are set out in the "Sequences" section that follows Table 2. In one embodiment, one or more proteins of the invention is combined with CT089. In another embodiment, one or more
25 proteins of the invention is combined with CT089 and CT381 (or their variants). In some embodiments, the C-terminal fragment of CT812 "CT812C" (for example, a protein comprising or consisting of the amino acid sequence set out in SEQ ID NO:122 or a fragment or variant thereof) is used instead of full length CT812.

In some embodiments, the following combinations of antigens (or their variants) are used:
30 CT733+CT601, CT733+CT279, CT733+CT443, CT733+CT372, CT733+CT456, CT733+CT381, CT153+CT601, CT153+CT279, CT153+CT443, CT153+CT372, CT153+CT456, CT153+CT381, CT601+CT443, CT601+CT372, CT601+CT456, CT601+CT381, CT279+CT443, CT279+CT372, CT279+CT456, CT279+CT381, CT443+CT372, CT443+CT456, CT443+CT381, CT372+CT456, CT372+CT381, CT387+CT812+CT869, CT387+CT812C+CT869. These combinations may be
35 used in the absence of any other chlamydia antigens or in the presence of one or more additional chlamydia antigens. Particularly preferred combinations are: (i) CT279+CT601; (ii) CT372+CT443; (iii) CT733+CT153; (iv) CT456+CT381; (v) CT279+ CT601+CT733+CT153; (vi)

CT279+CT601+CT372+CT443; (vii) CT823+CT733+CT043+ CT456; (viii) CT387+CT812+CT869; and (ix) CT387+CT812C+CT869 (or their variants).

The human serovariants (“serovars”) of *C. trachomatis* are divided into two biovariants (“biovars”). Serovars A-K elicit epithelial infections primarily in the ocular tissue (A-C) or urogenital tract (D-K). Serovars L1, L2 and L3 are the agents of invasive lymphogranuloma venereum (LGV). In some embodiments, one or more of the additional Chlamydial antigens may, for example, be of any of Serovars A-K or L1, L2 or L3. One or more of the additional Chlamydia antigens is preferably from *C. trachomatis* serovar D, or from another epidemiologically prevalent serotype.

In some embodiments, one or more of the additional Chlamydia antigens is a homologous antigen from *C. pneumoniae*, *C. psittaci*, *C. pecorum*, *C. muridarum* or *C. suis*.

In some embodiments, TC0551 (the *C. muridarum* homologue of CT279) is used in place of the *C. trachomatis* protein. *C. muridarum* is the mouse adapted strain of *Chlamydia trachomatis*. Although *C. muridarum* is not a human pathogen, infection of mice with *C. muridarum* phenotypically mimics many aspects of *C. trachomatis* infection in humans and is frequently used to measure immunoprotective responses against *C. trachomatis*. In some embodiments, TC0890 (the *C. muridarum* homologue of CT601) is used in place of the *C. trachomatis* protein. In some embodiments, TC0651 (the *C. muridarum* homologue of CT372) is used in place of the *C. trachomatis* protein. In some embodiments, TC0727 (the *C. muridarum* homologue of CT443) is used in place of the *C. trachomatis* protein. In some embodiments, TC0106 (the *C. muridarum* homologue of CT733) is used in place of the *C. trachomatis* protein. In some embodiments, TC0431 (the *C. muridarum* homologue of CT153) is used in place of the *C. trachomatis* protein. In some embodiments, TC0660 (the *C. muridarum* homologue of CT381) is used in place of the *C. trachomatis* protein. In some embodiments, TC0741 (the *C. muridarum* homologue of CT456) is used in place of the *C. trachomatis* protein. In some embodiments, TC0210 (the *C. muridarum* homologue of CT823) is used in place of the *C. trachomatis* protein. In some embodiments, TC0666 (the *C. muridarum* homologue of CT387) is used in place of the *C. trachomatis* protein. TC0666 is annotated as a hypothetical protein. In some embodiments, TC0197 (the *C. muridarum* homologue of CT812) is used in place of the *C. trachomatis* protein. TC0197 is annotated as polymorphic membrane protein D family protein. In some embodiments, TC0261 (the *C. muridarum* homologue of CT869) is used in place of the *C. trachomatis* protein. TC0261 is annotated as polymorphic membrane protein E/F family protein. In some embodiments, TC0313 (the *C. muridarum* homologue of CT043) is used in place of the *C. trachomatis* protein. In some embodiments, TC0889 (the *C. muridarum* homologue of CT600) is used in place of the *C. trachomatis* protein. In some embodiments, TC0210 (the *C. muridarum* homologue of CT823) is used in place of the *C. trachomatis* protein. In some embodiments in which the composition comprises a single Chlamydia antigen, the *C. muridarum* homologue is used in place of the single *C. trachomatis* antigen. In some embodiments in which the composition comprises a combination of Chlamydia antigens, the *C.*

muridarum homologue is used in place of one or more (for example, 1, 2, 3, 4, 5, 6, 7, 8, 9, 10) or all *C. trachomatis* antigens.

Advantageous combinations of the invention are those in which two or more antigens (for example, two, three or four antigens) act synergistically. Thus, the protection against Chlamydia achieved by their combined administration exceeds that expected by mere addition of their individual protective efficacy.

In some embodiments, the one or more additional Chlamydia antigens may comprise an amino acid sequence: (a) which is a variant of a Table 2 antigen (i.e. has 50% or more identity (e.g. 60%, 65%, 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, 99.5% or more) to a sequence presented in Table 2); and/or (b) comprising a fragment of at least 'n' consecutive amino acids of a sequence presented in Table 2 or of a variant of a Table 2 antigen, wherein 'n' is 7 or more (e.g. 8, 10, 12, 14, 16, 18, 20, 25, 30, 35, 40, 50, 60, 70, 80, 90, 100, 150, 200, 250, 350, 450, 550, 650, 750, 780, 800 or more). Preferred fragments of (b) comprise an epitope from a sequence presented in Table 2. Preferably, the epitope is a MHC class II epitope, for example, a CD4+ T cell epitope. Other preferred fragments lack one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the C-terminus and/or one or more amino acids (e.g. 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 20, 25 or more) from the N-terminus of a sequence presented in Table 2, while retaining at least one epitope of a sequence presented in Table 2. Other fragments omit one or more protein domains. When an additional Chlamydia antigen comprises a sequence that is not identical to a complete sequence from Table 2 (e.g. when it comprises a sequence with less than 100% sequence identity thereto, or when it comprises a fragment thereof), it is preferred in each individual instance that the additional Chlamydia antigen can elicit an antibody that recognises a protein having the complete sequence from the Table 2 antigen from which it is derived.

In some embodiments, the combination of two or more chlamydia antigens is provided as a combined preparation for simultaneous, separate or sequential administration. The invention also provides a kit comprising a protein of the invention and one or more additional antigens for simultaneous, separate or sequential administration.

The *Chlamydia* antigens used in the invention may be present in the composition as individual separate polypeptides. Alternatively, the combination may be present as a hybrid polypeptide in which two or more (i.e. 2, 3, 4, 5, 6, 7, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19 or 20 or more) of the antigens are expressed as a single polypeptide chain. Hybrid polypeptides offer two principal advantages: first, a polypeptide that may be unstable or poorly expressed on its own can be assisted by adding a suitable hybrid partner that overcomes the problem; second, commercial manufacture is simplified as only one expression and purification need be employed in order to produce two polypeptides which are both antigenically useful. Different hybrid polypeptides may be mixed together in a single formulation. Within such combinations, a *Chlamydia trachomatis* antigen may

be present in more than one hybrid polypeptide and/or as a non-hybrid polypeptide. It is preferred, however, that an antigen is present either as a hybrid or as a non-hybrid, but not as both.

Hybrid polypeptides can be represented by the formula $\text{NH}_2\text{-A-}\{-\text{X-L}\}_n\text{-B-COOH}$, wherein: at least one X is an amino acid sequence of a Chlamydia protein according to the invention as described above; L is an optional linker amino acid sequence; A is an optional N-terminal amino acid sequence; B is an optional C-terminal amino acid sequence; n is an integer of 2 or more (*e.g.* 2, 3, 4, 5, 6, *etc.*). Usually n is 2 or 3.

If a -X- moiety has a leader peptide sequence in its wild-type form, this may be included or omitted in the hybrid protein. In some embodiments, the leader peptides will be deleted except for that of the -X- moiety located at the N-terminus of the hybrid protein *i.e.* the leader peptide of X_1 will be retained, but the leader peptides of $X_2 \dots X_n$ will be omitted. This is equivalent to deleting all leader peptides and using the leader peptide of X_1 as moiety -A-.

For each n instances of $\{-\text{X-L}\}$, linker amino acid sequence -L- may be present or absent. For instance, when $n=2$ the hybrid may be $\text{NH}_2\text{-X}_1\text{-L}_1\text{-X}_2\text{-L}_2\text{-COOH}$, $\text{NH}_2\text{-X}_1\text{-X}_2\text{-COOH}$, $\text{NH}_2\text{-X}_1\text{-L}_1\text{-X}_2\text{-COOH}$, $\text{NH}_2\text{-X}_1\text{-X}_2\text{-L}_2\text{-COOH}$, *etc.* Linker amino acid sequence(s) -L- will typically be short (*e.g.* 20 or fewer amino acids *i.e.* 20, 19, 18, 17, 16, 15, 14, 13, 12, 11, 10, 9, 8, 7, 6, 5, 4, 3, 2, 1). Examples comprise short peptide sequences which facilitate cloning, poly-glycine linkers (*i.e.* comprising Gly_n where $n = 2, 3, 4, 5, 6, 7, 8, 9, 10$ or more), and histidine tags (*i.e.* His_n where $n = 3, 4, 5, 6, 7, 8, 9, 10$ or more). Other suitable linker amino acid sequences will be apparent to those skilled in the art. A useful linker is GSGGGG, with the Gly-Ser dipeptide being formed from a *Bam*HI restriction site, thus aiding cloning and manipulation, and the $(\text{Gly})_4$ tetrapeptide being a typical poly-glycine linker.

-A- is an optional N-terminal amino acid sequence. This will typically be short (*e.g.* 40 or fewer amino acids *i.e.* 40, 39, 38, 37, 36, 35, 34, 33, 32, 31, 30, 29, 28, 27, 26, 25, 24, 23, 22, 21, 20, 19, 18, 17, 16, 15, 14, 13, 12, 11, 10, 9, 8, 7, 6, 5, 4, 3, 2, 1). Examples include leader sequences to direct protein trafficking, or short peptide sequences which facilitate cloning or purification (*e.g.* histidine tags *i.e.* His_n where $n = 3, 4, 5, 6, 7, 8, 9, 10$ or more). Other suitable N-terminal amino acid sequences will be apparent to those skilled in the art. If X_1 lacks its own N-terminus methionine, -A- is preferably an oligopeptide (*e.g.* with 1, 2, 3, 4, 5, 6, 7 or 8 amino acids) which provides a N-terminus methionine.

-B- is an optional C-terminal amino acid sequence. This will typically be short (*e.g.* 40 or fewer amino acids *i.e.* 39, 38, 37, 36, 35, 34, 33, 32, 31, 30, 29, 28, 27, 26, 25, 24, 23, 22, 21, 20, 19, 18, 17, 16, 15, 14, 13, 12, 11, 10, 9, 8, 7, 6, 5, 4, 3, 2, 1). Examples include sequences to direct protein trafficking, short peptide sequences which facilitate cloning or purification (*e.g.* comprising histidine tags *i.e.* His_n where $n = 3, 4, 5, 6, 7, 8, 9, 10$ or more), or sequences which enhance protein stability. Other suitable C-terminal amino acid sequences will be apparent to those skilled in the art.

Where hybrid polypeptides are used, the individual antigens within the hybrid (*i.e.* individual -X-moieties) may be from one or more strains. Where $n=2$, for instance, X_2 may be from the same strain as X_1 or from a different strain. Where $n=3$, the strains might be (i) $X_1=X_2=X_3$ (ii) $X_1=X_2\neq X_3$ (iii) $X_1\neq X_2=X_3$ (iv) $X_1\neq X_2\neq X_3$ or (v) $X_1=X_3\neq X_2$, *etc.*

- 5 The invention also provides a nucleic acid encoding a hybrid polypeptide of the invention. Furthermore, the invention provides a nucleic acid which can hybridise to this nucleic acid, preferably under “high stringency” conditions (*e.g.* 65°C in a 0.1 x SSC, 0.5% SDS solution).

Further components of the composition

10 Compositions may thus be pharmaceutically acceptable. They will usually include components in addition to the antigens *e.g.* they typically include one or more pharmaceutical carrier(s) and/or excipient(s). A thorough discussion of such components is available in *Remington The Science and Practice of Pharmacy*.

15 Compositions will generally be administered to a mammal in aqueous form. Prior to administration, however, the composition may have been in a non-aqueous form. For instance, although some vaccines are manufactured in aqueous form, then filled and distributed and administered also in aqueous form, other vaccines are lyophilised during manufacture and are reconstituted into an aqueous form at the time of use. Thus a composition of the invention may be dried, such as a lyophilised formulation.

20 The composition may include preservatives such as thiomersal or 2-phenoxyethanol. It is preferred, however, that the vaccine should be substantially free from (*i.e.* less than 5µg/ml) mercurial material *e.g.* thiomersal-free. Vaccines containing no mercury are more preferred. Preservative-free vaccines are particularly preferred.

25 To control tonicity, it is preferred to include a physiological salt, such as a sodium salt. Sodium chloride (NaCl) is preferred, which may be present at between 1 and 20 mg/ml *e.g.* about 10±2mg/ml NaCl. Other salts that may be present include potassium chloride, potassium dihydrogen phosphate, disodium phosphate dehydrate, magnesium chloride, calcium chloride, *etc.*

Compositions will generally have an osmolality of between 200 mOsm/kg and 400 mOsm/kg, preferably between 240-360 mOsm/kg, and will more preferably fall within the range of 290-310 mOsm/kg.

30 Compositions may include one or more buffers. Typical buffers include: a phosphate buffer; a Tris buffer; a borate buffer; a succinate buffer; a histidine buffer (particularly with an aluminum hydroxide adjuvant); or a citrate buffer. Buffers will typically be included in the 5-20mM range.

The pH of a composition will generally be between 5.0 and 8.1, and more typically between 6.0 and 8.0 *e.g.* 6.5 and 7.5, or between 7.0 and 7.8.

The composition is preferably sterile. The composition is preferably non-pyrogenic *e.g.* containing <1 EU (endotoxin unit, a standard measure) per dose, and preferably <0.1 EU per dose. The composition is preferably gluten free.

5 The composition may include material for a single immunisation, or may include material for multiple immunisations (*i.e.* a ‘multidose’ kit). The inclusion of a preservative is preferred in multidose arrangements. As an alternative (or in addition) to including a preservative in multidose compositions, the compositions may be contained in a container having an aseptic adaptor for removal of material.

10 Human vaccines are typically administered in a dosage volume of about 0.5ml, although a half dose (*i.e.* about 0.25ml) may be administered to children.

Immunogenic compositions of the invention may also comprise one or more immunoregulatory agents. Preferably, one or more of the immunoregulatory agents include one or more adjuvants. The adjuvants may include a TH1 adjuvant and/or a TH2 adjuvant, further discussed below.

Adjuvants which may be used in compositions of the invention include, but are not limited to:

15 A. *Mineral-containing compositions*

Mineral containing compositions suitable for use as adjuvants in the invention include mineral salts, such as aluminium salts and calcium salts (or mixtures thereof). Calcium salts include calcium phosphate (*e.g.* the “CAP” particles disclosed in US patent 6355271). Aluminum salts include hydroxides, phosphates, sulfates, *etc.*, with the salts taking any suitable form (*e.g.* gel, crystalline, 20 amorphous, *etc.*). Adsorption to these salts is preferred. The mineral containing compositions may also be formulated as a particle of metal salt [WO00/23105].

The adjuvants known as aluminum hydroxide and aluminum phosphate may be used. These names are conventional, but are used for convenience only, as neither is a precise description of the actual chemical compound which is present (*e.g.* see chapter 9 of *Vaccine Design...* (1995) eds. Powell & Newman. ISBN: 030644867X. Plenum). The invention can use any of the “hydroxide” or 25 “phosphate” adjuvants that are in general use as adjuvants. The adjuvants known as “aluminium hydroxide” are typically aluminium oxyhydroxide salts, which are usually at least partially crystalline. The adjuvants known as “aluminium phosphate” are typically aluminium hydroxyphosphates, often also containing a small amount of sulfate (*i.e.* aluminium hydroxyphosphate sulfate). They may be obtained by precipitation, and the reaction conditions and 30 concentrations during precipitation influence the degree of substitution of phosphate for hydroxyl in the salt.

A fibrous morphology (*e.g.* as seen in transmission electron micrographs) is typical for aluminium hydroxide adjuvants. The pI of aluminium hydroxide adjuvants is typically about 11 *i.e.* the adjuvant 35 itself has a positive surface charge at physiological pH. Adsorptive capacities of between 1.8-2.6 mg protein per mg Al⁺⁺⁺ at pH 7.4 have been reported for aluminium hydroxide adjuvants.

Aluminium phosphate adjuvants generally have a PO_4/Al molar ratio between 0.3 and 1.2, preferably between 0.8 and 1.2, and more preferably 0.95 ± 0.1 . The aluminium phosphate will generally be amorphous, particularly for hydroxyphosphate salts. A typical adjuvant is amorphous aluminium hydroxyphosphate with PO_4/Al molar ratio between 0.84 and 0.92, included at $0.6 \text{ mg Al}^{3+}/\text{ml}$. The aluminium phosphate will generally be particulate (*e.g.* plate-like morphology as seen in transmission electron micrographs). Typical diameters of the particles are in the range $0.5\text{-}20 \mu\text{m}$ (*e.g.* about $5\text{-}10 \mu\text{m}$) after any antigen adsorption. Adsorptive capacities of between $0.7\text{-}1.5 \text{ mg protein per mg Al}^{+++}$ at pH 7.4 have been reported for aluminium phosphate adjuvants.

The point of zero charge (PZC) of aluminium phosphate is inversely related to the degree of substitution of phosphate for hydroxyl, and this degree of substitution can vary depending on reaction conditions and concentration of reactants used for preparing the salt by precipitation. PZC is also altered by changing the concentration of free phosphate ions in solution (more phosphate = more acidic PZC) or by adding a buffer such as a histidine buffer (makes PZC more basic). Aluminium phosphates used according to the invention will generally have a PZC of between 4.0 and 7.0, more preferably between 5.0 and 6.5 *e.g.* about 5.7.

Suspensions of aluminium salts used to prepare compositions of the invention may contain a buffer (*e.g.* a phosphate or a histidine or a Tris buffer), but this is not always necessary. The suspensions are preferably sterile and pyrogen-free. A suspension may include free aqueous phosphate ions *e.g.* present at a concentration between 1.0 and 20 mM, preferably between 5 and 15 mM, and more preferably about 10 mM. The suspensions may also comprise sodium chloride.

The invention can use a mixture of both an aluminium hydroxide and an aluminium phosphate. In this case there may be more aluminium phosphate than hydroxide *e.g.* a weight ratio of at least 2:1 *e.g.* $\geq 5:1, \geq 6:1, \geq 7:1, \geq 8:1, \geq 9:1, \text{ etc.}$

The concentration of Al^{+++} in a composition for administration to a patient is preferably less than 10 mg/ml *e.g.* $\leq 5 \text{ mg/ml}, \leq 4 \text{ mg/ml}, \leq 3 \text{ mg/ml}, \leq 2 \text{ mg/ml}, \leq 1 \text{ mg/ml}, \text{ etc.}$ A preferred range is between 0.3 and 1 mg/ml . A maximum of 0.85 mg/dose is preferred.

Aluminium phosphates are particularly preferred, particularly in compositions which include a *H. influenzae* saccharide antigen, and a typical adjuvant is amorphous aluminium hydroxyphosphate with PO_4/Al molar ratio between 0.84 and 0.92, included at $0.6 \text{ mg Al}^{3+}/\text{ml}$. Adsorption with a low dose of aluminium phosphate may be used *e.g.* between 50 and $100 \mu\text{g Al}^{3+}$ per conjugate per dose. Where there is more than one conjugate in a composition, not all conjugates need to be adsorbed.

B. Oil Emulsions

Oil emulsion compositions suitable for use as adjuvants in the invention include squalene-water emulsions, such as MF59 [Chapter 10 of *Vaccine Design...* (1995) eds. Powell & Newman. ISBN: 030644867X. Plenum; see also WO90/14837] (5% Squalene, 0.5% Tween 80, and 0.5% Span 85,

formulated into submicron particles using a microfluidizer). Complete Freund's adjuvant (CFA) and incomplete Freund's adjuvant (IFA) may also be used.

5 Various oil-in-water emulsion adjuvants are known, and they typically include at least one oil and at least one surfactant, with the oil(s) and surfactant(s) being biodegradable (metabolisable) and biocompatible. The oil droplets in the emulsion are generally less than 5µm in diameter, and ideally have a sub-micron diameter, with these small sizes being achieved with a microfluidiser to provide stable emulsions. Droplets with a size less than 220nm are preferred as they can be subjected to filter sterilization.

10 The emulsion can comprise oils such as those from an animal (such as fish) or vegetable source. Sources for vegetable oils include nuts, seeds and grains. Peanut oil, soybean oil, coconut oil, and olive oil, the most commonly available, exemplify the nut oils. Jojoba oil can be used *e.g.* obtained from the jojoba bean. Seed oils include safflower oil, cottonseed oil, sunflower seed oil, sesame seed oil and the like. In the grain group, corn oil is the most readily available, but the oil of other cereal grains such as wheat, oats, rye, rice, teff, triticale and the like may also be used. 6-10 carbon fatty acid esters of glycerol and 1,2-propanediol, while not occurring naturally in seed oils, may be prepared by hydrolysis, separation and esterification of the appropriate materials starting from the nut and seed oils. Fats and oils from mammalian milk are metabolizable and may therefore be used in the practice of this invention. The procedures for separation, purification, saponification and other means necessary for obtaining pure oils from animal sources are well known in the art. Most fish contain metabolizable oils which may be readily recovered. For example, cod liver oil, shark liver oils, and whale oil such as spermaceti exemplify several of the fish oils which may be used herein. A number of branched chain oils are synthesized biochemically in 5-carbon isoprene units and are generally referred to as terpenoids. Shark liver oil contains a branched, unsaturated terpenoids known as squalene, 2,6,10,15,19,23-hexamethyl-2,6,10,14,18,22-tetracosahexaene, which is particularly preferred herein. Squalane, the saturated analog to squalene, is also a preferred oil. Fish oils, including squalene and squalane, are readily available from commercial sources or may be obtained by methods known in the art. Other preferred oils are the tocopherols (see below). Mixtures of oils can be used.

30 Surfactants can be classified by their 'HLB' (hydrophile/lipophile balance). Preferred surfactants of the invention have a HLB of at least 10, preferably at least 15, and more preferably at least 16. The invention can be used with surfactants including, but not limited to: the polyoxyethylene sorbitan esters surfactants (commonly referred to as the Tweens), especially polysorbate 20 and polysorbate 80; copolymers of ethylene oxide (EO), propylene oxide (PO), and/or butylene oxide (BO), sold under the DOWFAX™ tradename, such as linear EO/PO block copolymers; octoxynols, which can vary in the number of repeating ethoxy (oxy-1,2-ethanediyl) groups, with octoxynol-9 (Triton X-100, or t-octylphenoxy polyethoxyethanol) being of particular interest; (octylphenoxy)polyethoxyethanol (IGEPAL CA-630/NP-40); phospholipids such as phosphatidylcholine (lecithin); nonylphenol

ethoxylates, such as the Tergitol™ NP series; polyoxyethylene fatty ethers derived from lauryl, cetyl, stearyl and oleyl alcohols (known as Brij surfactants), such as triethyleneglycol monolauryl ether (Brij 30); and sorbitan esters (commonly known as the SPANs), such as sorbitan trioleate (Span 85) and sorbitan monolaurate. Non-ionic surfactants are preferred. Preferred surfactants for including in the emulsion are Tween 80 (polyoxyethylene sorbitan monooleate), Span 85 (sorbitan trioleate), lecithin and Triton X-100.

Mixtures of surfactants can be used e.g. Tween 80/Span 85 mixtures. A combination of a polyoxyethylene sorbitan ester such as polyoxyethylene sorbitan monooleate (Tween 80) and an octoxynol such as t-octylphenoxypolyoxyethanol (Triton X-100) is also suitable. Another useful combination comprises laureth 9 plus a polyoxyethylene sorbitan ester and/or an octoxynol.

Preferred amounts of surfactants (% by weight) are: polyoxyethylene sorbitan esters (such as Tween 80) 0.01 to 1%, in particular about 0.1 %; octyl- or nonylphenoxy polyoxyethanols (such as Triton X-100, or other detergents in the Triton series) 0.001 to 0.1 %, in particular 0.005 to 0.02%; polyoxyethylene ethers (such as laureth 9) 0.1 to 20 %, preferably 0.1 to 10 % and in particular 0.1 to 1 % or about 0.5%.

Preferred emulsion adjuvants have an average droplets size of $<1\mu\text{m}$ e.g. $\leq 750\text{nm}$, $\leq 500\text{nm}$, $\leq 400\text{nm}$, $\leq 300\text{nm}$, $\leq 250\text{nm}$, $\leq 220\text{nm}$, $\leq 200\text{nm}$, or smaller. These droplet sizes can conveniently be achieved by techniques such as microfluidisation.

Specific oil-in-water emulsion adjuvants useful with the invention include, but are not limited to:

- A submicron emulsion of squalene, Tween 80, and Span 85. The composition of the emulsion by volume can be about 5% squalene, about 0.5% polysorbate 80 and about 0.5% Span 85. In weight terms, these ratios become 4.3% squalene, 0.5% polysorbate 80 and 0.48% Span 85. This adjuvant is known as 'MF59' (WO90/14837, Podda & Del Giudice (2003) *Expert Rev Vaccines* 2:197-203, Podda (2001) *Vaccine* 19: 2673-2680; as described in more detail in Chapter 10 of *Vaccine Design: The Subunit and Adjuvant Approach* (eds. Powell & Newman) Plenum Press 1995 (ISBN 0-306-44867-X) and chapter 12 of *Vaccine Adjuvants: Preparation Methods and Research Protocols* (Volume 42 of *Methods in Molecular Medicine* series). ISBN: 1-59259-083-7. Ed. O'Hagan). The MF59 emulsion advantageously includes citrate ions e.g. 10mM sodium citrate buffer.
- An emulsion of squalene, a tocopherol, and Tween 80. The emulsion may include phosphate buffered saline. It may also include Span 85 (e.g. at 1%) and/or lecithin. These emulsions may have from 2 to 10% squalene, from 2 to 10% tocopherol and from 0.3 to 3% Tween 80, and the weight ratio of squalene:tocopherol is preferably ≤ 1 as this provides a more stable emulsion. Squalene and Tween 80 may be present volume ratio of about 5:2. One such emulsion can be made by dissolving Tween 80 in PBS to give a 2% solution, then mixing 90ml of this solution with a mixture of (5g of DL- α -tocopherol and 5ml squalene), then microfluidising the mixture.

The resulting emulsion may have submicron oil droplets *e.g.* with an average diameter of between 100 and 250nm, preferably about 180nm.

- An emulsion of squalene, a tocopherol, and a Triton detergent (*e.g.* Triton X-100). The emulsion may also include a 3d-MPL (see below). The emulsion may contain a phosphate buffer.
- 5 • An emulsion comprising a polysorbate (*e.g.* polysorbate 80), a Triton detergent (*e.g.* Triton X-100) and a tocopherol (*e.g.* an α -tocopherol succinate). The emulsion may include these three components at a mass ratio of about 75:11:10 (*e.g.* 750 μ g/ml polysorbate 80, 110 μ g/ml Triton X-100 and 100 μ g/ml α -tocopherol succinate), and these concentrations should include any contribution of these components from antigens. The emulsion may also include squalene. The
10 emulsion may also include a 3d-MPL (see below). The aqueous phase may contain a phosphate buffer.
- An emulsion of squalane, polysorbate 80 and poloxamer 401 (“Pluronic™ L121”). The emulsion can be formulated in phosphate buffered saline, pH 7.4. This emulsion is a useful delivery vehicle for muramyl dipeptides, and has been used with threonyl-MDP in the “SAF-1” adjuvant
15 (Allison & Byars (1992) *Res Immunol* 143:519-25) (0.05-1% Thr-MDP, 5% squalane, 2.5% Pluronic L121 and 0.2% polysorbate 80). It can also be used without the Thr-MDP, as in the “AF” adjuvant (Hariharan *et al.* (1995) *Cancer Res* 55:3486-9) (5% squalane, 1.25% Pluronic L121 and 0.2% polysorbate 80). Microfluidisation is preferred.
- An emulsion comprising squalene, an aqueous solvent, a polyoxyethylene alkyl ether hydrophilic
20 nonionic surfactant (*e.g.* polyoxyethylene (12) cetostearyl ether) and a hydrophobic nonionic surfactant (*e.g.* a sorbitan ester or mannide ester, such as sorbitan monooleate or ‘Span 80’). The emulsion is preferably thermoreversible and/or has at least 90% of the oil droplets (by volume) with a size less than 200 nm (US-2007/014805.). The emulsion may also include one or more of: alditol; a cryoprotective agent (*e.g.* a sugar, such as dodecylmaltoside and/or sucrose); and/or an
25 alkylpolyglycoside. Such emulsions may be lyophilized.
- An emulsion of US-2007/014805.f squalene, poloxamer 105 and Abil-Care (Suli *et al.* (2004) *Vaccine* 22(25-26):3464-9). The final concentration (weight) of these components in adjuvanted vaccines are 5% squalene, 4% poloxamer 105 (pluronic polyol) and 2% Abil-Care 85 (Bis-PEG/PPG-16/16 PEG/PPG-16/16 dimethicone; caprylic/capric triglyceride).
- 30 • An emulsion having from 0.5-50% of an oil, 0.1-10% of a phospholipid, and 0.05-5% of a non-ionic surfactant. As described in WO95/11700, preferred phospholipid components are phosphatidylcholine, phosphatidylethanolamine, phosphatidylserine, phosphatidylinositol, phosphatidylglycerol, phosphatidic acid, sphingomyelin and cardiolipin. Submicron droplet sizes are advantageous.

- A submicron oil-in-water emulsion of a non-metabolisable oil (such as light mineral oil) and at least one surfactant (such as lecithin, Tween 80 or Span 80). Additives may be included, such as QuilA saponin, cholesterol, a saponin-lipophile conjugate (such as GPI-0100, described in US patent 6,080,725, produced by addition of aliphatic amine to desacylsaponin via the carboxyl group of glucuronic acid), dimethyldioctadecylammonium bromide and/or N,N-dioctadecyl-N,N-bis (2-hydroxyethyl)propanediamine.
- An emulsion in which a saponin (*e.g.* QuilA or QS21) and a sterol (*e.g.* a cholesterol) are associated as helical micelles (WO2005/097181).
- An emulsion comprising a mineral oil, a non-ionic lipophilic ethoxylated fatty alcohol, and a non-ionic hydrophilic surfactant (*e.g.* an ethoxylated fatty alcohol and/or polyoxyethylene-polyoxypropylene block copolymer) (WO2006/113373).
- An emulsion comprising a mineral oil, a non-ionic hydrophilic ethoxylated fatty alcohol, and a non-ionic lipophilic surfactant (*e.g.* an ethoxylated fatty alcohol and/or polyoxyethylene-polyoxypropylene block copolymer) (Wu *et al.* (2004) *Antiviral Res.* 64(2):79-83).

In some embodiments an emulsion may be mixed with antigen extemporaneously, at the time of delivery, and thus the adjuvant and antigen may be kept separately in a packaged or distributed vaccine, ready for final formulation at the time of use. In other embodiments an emulsion is mixed with antigen during manufacture, and thus the composition is packaged in a liquid adjuvanted form. The antigen will generally be in an aqueous form, such that the vaccine is finally prepared by mixing two liquids. The volume ratio of the two liquids for mixing can vary (*e.g.* between 5:1 and 1:5) but is generally about 1:1. Where concentrations of components are given in the above descriptions of specific emulsions, these concentrations are typically for an undiluted composition, and the concentration after mixing with an antigen solution will thus decrease. Where a composition is to be prepared extemporaneously prior to use (*e.g.* where a component is presented in lyophilised form) and is presented as a kit, the kit may comprise two vials, or it may comprise one ready-filled syringe and one vial, with the contents of the syringe being used to reactivate the contents of the vial prior to injection.

Where a composition includes a tocopherol, any of the α , β , γ , δ , ϵ or ξ tocopherols can be used, but α -tocopherols are preferred. The tocopherol can take several forms *e.g.* different salts and/or isomers.

Salts include organic salts, such as succinate, acetate, nicotinate, *etc.* D- α -tocopherol and DL- α -tocopherol can both be used. Tocopherols are advantageously included in vaccines for use in elderly patients (*e.g.* aged 60 years or older) because vitamin E has been reported to have a positive effect on the immune response in this patient group (Han *et al.* (2005) *Impact of Vitamin E on Immune Function and Infectious Diseases in the Aged at Nutrition, Immune functions and Health EuroConference, Paris, 9-10 June 2005*). They also have antioxidant properties that may help to stabilize the emulsions (US-6630161). A preferred α -tocopherol is DL- α -tocopherol, and the

preferred salt of this tocopherol is the succinate. The succinate salt has been found to cooperate with TNF-related ligands *in vivo*.

C. *Saponin formulations (chapter 22 of Vaccine Design... (1995) eds. Powell & Newman. ISBN: 030644867X. Plenum)*

5 Saponin formulations may also be used as adjuvants in the invention. Saponins are a heterogeneous group of sterol glycosides and triterpenoid glycosides that are found in the bark, leaves, stems, roots and even flowers of a wide range of plant species. Saponin from the bark of the *Quillaia saponaria* Molina tree have been widely studied as adjuvants. Saponin can also be commercially obtained from *Smilax ornata* (sarsapilla), *Gypsophilla paniculata* (brides veil), and *Saponaria officinalis* (soap
10 root). Saponin adjuvant formulations include purified formulations, such as QS21, as well as lipid formulations, such as ISCOMs. QS21 is marketed as Stimulon™.

Saponin compositions have been purified using HPLC and RP-HPLC. Specific purified fractions using these techniques have been identified, including QS7, QS17, QS18, QS21, QH-A, QH-B and QH-C. Preferably, the saponin is QS21. A method of production of QS21 is disclosed in US
15 5,057,540. Saponin formulations may also comprise a sterol, such as cholesterol (WO96/33739).

Combinations of saponins and cholesterol can be used to form unique particles called immunostimulating complexes (ISCOMs) (chapter 23 of *Vaccine Design... (1995) eds. Powell & Newman. ISBN: 030644867X. Plenum*). ISCOMs typically also include a phospholipid such as phosphatidylethanolamine or phosphatidylcholine. Any known saponin can be used in ISCOMs.
20 Preferably, the ISCOM includes one or more of Quila, QHA & QHC. ISCOMs are further described in Podda & Del Giudice (2003) *Expert Rev Vaccines* 2:197-203; Podda (2001) *Vaccine* 19: 2673-2680; *Vaccine Design: The Subunit and Adjuvant Approach* (eds. Powell & Newman) Plenum Press 1995 (ISBN 0-306-44867-X); *Vaccine Adjuvants: Preparation Methods and Research Protocols* (Volume 42 of *Methods in Molecular Medicine* series). ISBN: 1-59259-083-7. Ed. O'Hagan; Allison
25 & Byars (1992) *Res Immunol* 143:519-25; Hariharan *et al.* (1995) *Cancer Res* 55:3486-9; US-2007/014805; Suli *et al.* (2004) *Vaccine* 22(25-26):3464-9; WO95/11700; US patent 6,080,725; WO2005/097181; WO2006/113373; Han *et al.* (2005) *Impact of Vitamin E on Immune Function and Infectious Diseases in the Aged at Nutrition, Immune functions and Health EuroConference, Paris, 9-10 June 2005*; US- 6630161; US 5,057,540; WO96/33739; EP-A-0109942; and WO96/11711.
30 Optionally, the ISCOMS may be devoid of additional detergent (WO00/07621).

A review of the development of saponin based adjuvants can be found in Barr *et al.* (1998) *Advanced Drug Delivery Reviews* 32:247-271 and Sjolanderet *et al.* (1998) *Advanced Drug Delivery Reviews* 32:321-338.

D. *Virosomes and virus-like particles*

35 Virosomes and virus-like particles (VLPs) can also be used as adjuvants in the invention. These structures generally contain one or more proteins from a virus optionally combined or formulated with a phospholipid. They are generally non-pathogenic, non-replicating and generally do not contain

any of the native viral genome. The viral proteins may be recombinantly produced or isolated from whole viruses. These viral proteins suitable for use in virosomes or VLPs include proteins derived from influenza virus (such as HA or NA), Hepatitis B virus (such as core or capsid proteins), Hepatitis E virus, measles virus, Sindbis virus, Rotavirus, Foot-and-Mouth Disease virus, Retrovirus, Norwalk virus, human Papilloma virus, HIV, RNA-phages, Q β -phage (such as coat proteins), GA-phage, fr-phage, AP205 phage, and Ty (such as retrotransposon Ty protein p1). VLPs are discussed further in Niikura *et al.* (2002) *Virology* 293:273-280; Lenz *et al.* (2001) *J Immunol* 166:5346-5355; Pinto *et al.* (2003) *J Infect Dis* 188:327-338; Gerber *et al.* (2001) *J Virol* 75:4752-4760; WO03/024480 and WO03/024481. Virosomes are discussed further in, for example, Gluck *et al.* (2002) *Vaccine* 20:B10-B16.

E. Bacterial or microbial derivatives

Adjuvants suitable for use in the invention include bacterial or microbial derivatives such as non-toxic derivatives of enterobacterial lipopolysaccharide (LPS), Lipid A derivatives, immunostimulatory oligonucleotides and ADP-ribosylating toxins and detoxified derivatives thereof.

Non-toxic derivatives of LPS include monophosphoryl lipid A (MPL) and 3-O-deacylated MPL (3dMPL). 3dMPL is a mixture of 3 de-O-acylated monophosphoryl lipid A with 4, 5 or 6 acylated chains. A preferred "small particle" form of 3 De-O-acylated monophosphoryl lipid A is disclosed in EP-A-0689454. Such "small particles" of 3dMPL are small enough to be sterile filtered through a 0.22 μ m membrane (US- 6630161). Other non-toxic LPS derivatives include monophosphoryl lipid A mimics, such as aminoalkyl glucosaminide phosphate derivatives *e.g.* RC-529 (Johnson *et al.* (1999) *Bioorg Med Chem Lett* 9:2273-2278; and Evans *et al.* (2003) *Expert Rev Vaccines* 2:219-229).

Lipid A derivatives include derivatives of lipid A from *Escherichia coli* such as OM-174. OM-174 is described for example in Meraldi *et al.* (2003) *Vaccine* 21:2485-2491 and Pajak *et al.* (2003) *Vaccine* 21:836-842.

Immunostimulatory oligonucleotides suitable for use as adjuvants in the invention include nucleotide sequences containing a CpG motif (a dinucleotide sequence containing an unmethylated cytosine linked by a phosphate bond to a guanosine). Double-stranded RNAs and oligonucleotides containing palindromic or poly(dG) sequences have also been shown to be immunostimulatory.

The CpG's can include nucleotide modifications/analogs such as phosphorothioate modifications and can be double-stranded or single-stranded. Kandimalla *et al.* (2003) *Nucleic Acids Research* 31:2393-2400, WO02/26757 and WO99/62923 disclose possible analog substitutions *e.g.* replacement of guanosine with 2'-deoxy-7-deazaguanosine. The adjuvant effect of CpG oligonucleotides is further discussed in Krieg (2003) *Nature Medicine* 9:831-835; McCluskie *et al.* (2002) *FEMS Immunology and Medical Microbiology* 32:179-185; WO98/40100; US 6,207,646; US 6,239,116 and US 6,429,199.

The CpG sequence may be directed to TLR9, such as the motif GTCGTT or TTCGTT (Kandimalla *et al.* (2003) *Biochemical Society Transactions* 31 (part 3):654-658). The CpG sequence may be

specific for inducing a Th1 immune response, such as a CpG-A ODN, or it may be more specific for inducing a B cell response, such as a CpG-B ODN. CpG-A and CpG-B ODNs are discussed in Blackwell *et al.* (2003) *J Immunol* 170:4061-4068; Krieg (2002) *Trends Immunol* 23:64-65; and WO01/95935. Preferably, the CpG is a CpG-A ODN.

5 Preferably, the CpG oligonucleotide is constructed so that the 5' end is accessible for receptor recognition. Optionally, two CpG oligonucleotide sequences may be attached at their 3' ends to form "immunomers". See, for example, Gluck *et al.* (2002) *Vaccine* 20:B10-B16; Kandimalla *et al.* (2003) *BBRC* 306:948-953; Bhagat *et al.* (2003) *BBRC* 300:853-861; and WO03/035836.

A useful CpG adjuvant is CpG7909, also known as ProMune™ (Coley Pharmaceutical Group, Inc.).
10 Another is CpG1826. As an alternative, or in addition, to using CpG sequences, TpG sequences can be used (WO01/22972), and these oligonucleotides may be free from unmethylated CpG motifs. The immunostimulatory oligonucleotide may be pyrimidine-rich. For example, it may comprise more than one consecutive thymidine nucleotide (*e.g.* TTTT, as disclosed in Pajak *et al.* (2003) *Vaccine* 21:836-842), and/or it may have a nucleotide composition with >25% thymidine (*e.g.* >35%, >40%,
15 >50%, >60%, >80%, *etc.*). For example, it may comprise more than one consecutive cytosine nucleotide (*e.g.* CCCC, as disclosed in Pajak *et al.* (2003) *Vaccine* 21:836-842), and/or it may have a nucleotide composition with >25% cytosine (*e.g.* >35%, >40%, >50%, >60%, >80%, *etc.*). These oligonucleotides may be free from unmethylated CpG motifs. Immunostimulatory oligonucleotides will typically comprise at least 20 nucleotides. They may comprise fewer than 100 nucleotides.

20 A particularly useful adjuvant based around immunostimulatory oligonucleotides is known as IC-31™ (Schellack *et al.* (2006) *Vaccine* 24:5461-72). Thus an adjuvant used with the invention may comprise a mixture of (i) an oligonucleotide (*e.g.* between 15-40 nucleotides) including at least one (and preferably multiple) CpI motifs (*i.e.* a cytosine linked to an inosine to form a dinucleotide), and
25 (ii) a polycationic polymer, such as an oligopeptide (*e.g.* between 5-20 amino acids) including at least one (and preferably multiple) Lys-Arg-Lys tripeptide sequence(s). The oligonucleotide may be a deoxynucleotide comprising 26-mer sequence 5'-(IC)₁₃-3'. The polycationic polymer may be a peptide comprising 11-mer amino acid sequence KKLKLLLLLKLK.

Bacterial ADP-ribosylating toxins and detoxified derivatives thereof may be used as adjuvants in the invention. Preferably, the protein is derived from *E.coli* (*E.coli* heat labile enterotoxin "LT"), cholera
30 ("CT"), or pertussis ("PT"). The use of detoxified ADP-ribosylating toxins as mucosal adjuvants is described in WO95/17211 and as parenteral adjuvants in WO98/42375. The toxin or toxoid is preferably in the form of a holotoxin, comprising both A and B subunits. Preferably, the A subunit contains a detoxifying mutation; preferably the B subunit is not mutated. Preferably, the adjuvant is a detoxified LT mutant such as LT-K63, LT-R72, and LT-G192. The use of ADP-ribosylating toxins
35 and detoxified derivatives thereof, particularly LT-K63 and LT-R72, as adjuvants can be found in Beignon *et al.* (2002) *Infect Immun* 70:3012-3019; Pizza *et al.* (2001) *Vaccine* 19:2534-2541; Pizza *et al.* (2000) *Int J Med Microbiol* 290:455-461; Scharton-Kersten *et al.* (2000) *Infect Immun*

68:5306-5313; Ryan *et al.* (1999) *Infect Immun* 67:6270-6280; Partidos *et al.* (1999) *Immunol Lett* 67:209-216; Peppoloni *et al.* (2003) *Expert Rev Vaccines* 2:285-293; and Pine *et al.* (2002) *J Control Release* 85:263-270.

5 A useful CT mutant is or CT-E29H (Tebbey *et al.* (2000) *Vaccine* 18:2723-34). Numerical reference for amino acid substitutions is preferably based on the alignments of the A and B subunits of ADP-ribosylating toxins set forth in Domenighini *et al.* (1995) *Mol Microbiol* 15:1165-1167, specifically incorporated herein by reference in its entirety.

F. Human immunomodulators

10 Human immunomodulators suitable for use as adjuvants in the invention include cytokines, such as interleukins (*e.g.* IL-1, IL-2, IL-4, IL-5, IL-6, IL-7, IL-12 (WO99/40936), *etc.*) (WO99/44636), interferons (*e.g.* interferon- γ), macrophage colony stimulating factor, and tumor necrosis factor. A preferred immunomodulator is IL-12.

G. Bioadhesives and Mucoadhesives

15 Bioadhesives and mucoadhesives may also be used as adjuvants in the invention. Suitable bioadhesives include esterified hyaluronic acid microspheres (Singh *et al.* (2001) *J Cont Release* 70:267-276) or mucoadhesives such as cross-linked derivatives of poly(acrylic acid), polyvinyl alcohol, polyvinyl pyrrolidone, polysaccharides and carboxymethylcellulose. Chitosan and derivatives thereof may also be used as adjuvants in the invention (WO99/27960).

H. Microparticles

20 Microparticles may also be used as adjuvants in the invention. Microparticles (*i.e.* a particle of ~100nm to ~150 μ m in diameter, more preferably ~200nm to ~30 μ m in diameter, and most preferably ~500nm to ~10 μ m in diameter) formed from materials that are biodegradable and non-toxic (*e.g.* a poly(α -hydroxy acid), a polyhydroxybutyric acid, a polyorthoester, a polyanhydride, a polycaprolactone, *etc.*), with poly(lactide-co-glycolide) are preferred, optionally treated to have a negatively-charged surface (*e.g.* with SDS) or a positively-charged surface (*e.g.* with a cationic detergent, such as CTAB).

I. Liposomes (Chapters 13 & 14 of Vaccine Design... (1995) eds. Powell & Newman. ISBN: 030644867X. Plenum.)

30 Examples of liposome formulations suitable for use as adjuvants are described in US 6,090,406; US 5,916,588; and EP-A-0626169.

J. Polyoxyethylene ether and polyoxyethylene ester formulations

35 Adjuvants suitable for use in the invention include polyoxyethylene ethers and polyoxyethylene esters (WO99/52549). Such formulations further include polyoxyethylene sorbitan ester surfactants in combination with an octoxynol (WO01/21207) as well as polyoxyethylene alkyl ethers or ester surfactants in combination with at least one additional non-ionic surfactant such as an octoxynol (WO01/21152). Preferred polyoxyethylene ethers are selected from the following group:

polyoxyethylene-9-lauryl ether (laureth 9), polyoxyethylene-9-stearyl ether, polyoxyethylene-8-stearyl ether, polyoxyethylene-4-lauryl ether, polyoxyethylene-35-lauryl ether, and polyoxyethylene-23-lauryl ether.

K. Phosphazenes

- 5 A phosphazene, such as poly[di(carboxylatophenoxy)phosphazene] ("PCPP") as described, for example, in Andrianov *et al.* (1998) *Biomaterials* 19:109-115 and Payne *et al.* (1998) *Adv Drug Delivery Review* 31:185-196, may be used.

L. Muramyl peptides

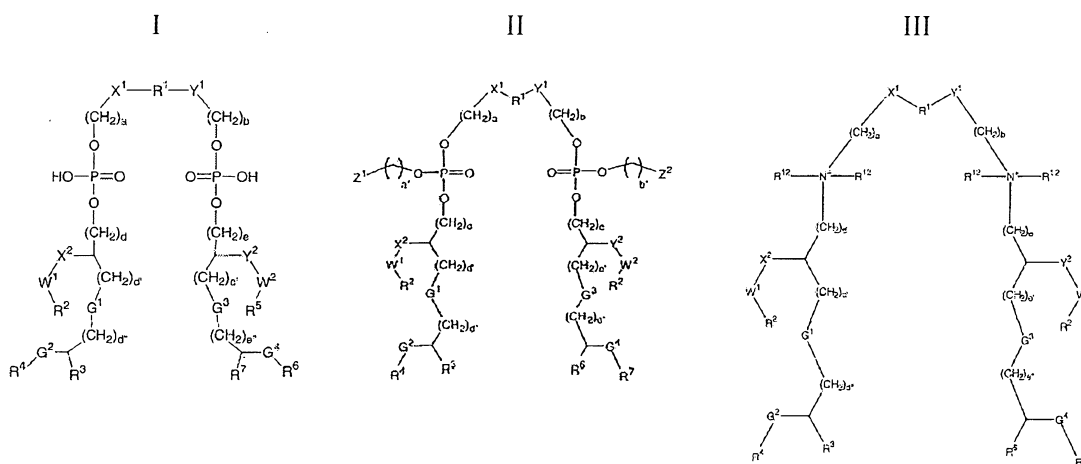
- 10 Examples of muramyl peptides suitable for use as adjuvants in the invention include N-acetyl-muramyl-L-threonyl-D-isoglutamine (thr-MDP), N-acetyl-normuramyl-L-alanyl-D-isoglutamine (nor-MDP), and N-acetylmuramyl-L-alanyl-D-isoglutaminyl-L-alanine-2-(1'-2'-dipalmitoyl-*sn*-glycero-3-hydroxyphosphoryloxy)-ethylamine MTP-PE).

M. Imidazoquinolone Compounds.

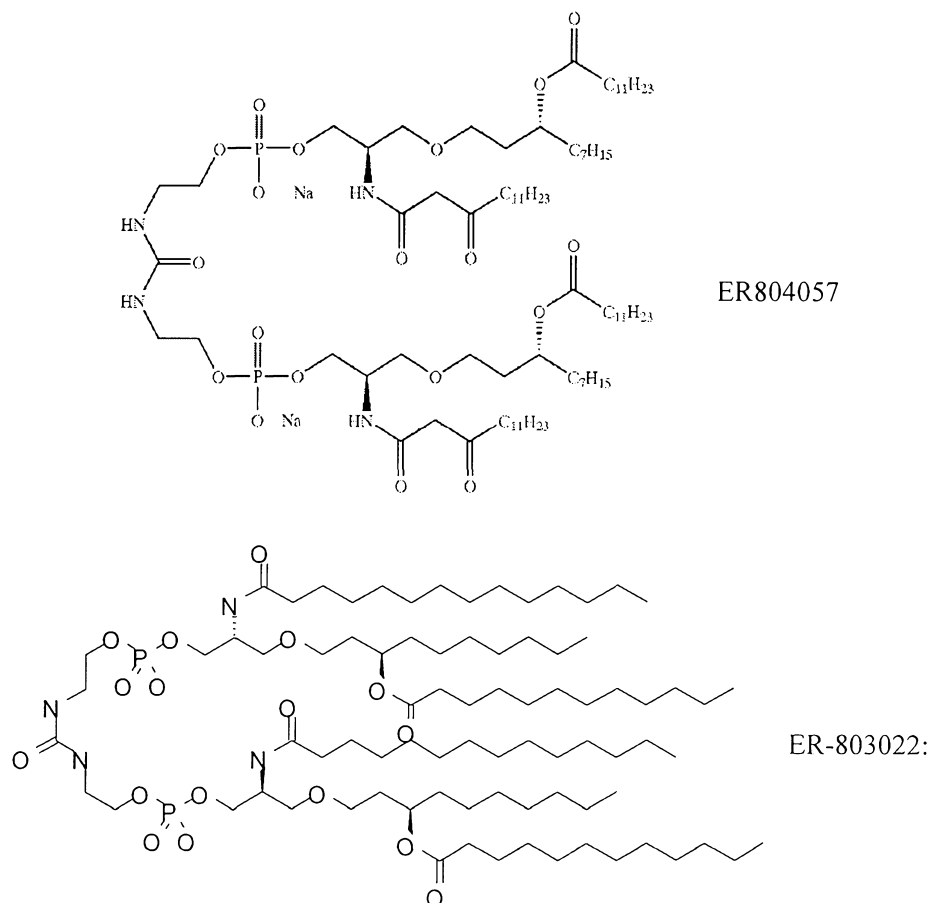
- 15 Examples of imidazoquinolone compounds suitable for use adjuvants in the invention include Imiquimod ("R-837") (US 4,680,338; US 4,988,815), Resiquimod ("R-848") (WO92/15582), and their analogs; and salts thereof (*e.g.* the hydrochloride salts). Further details about immunostimulatory imidazoquinolines can be found in Stanley (2002) *Clin Exp Dermatol* 27:571-577; Wu *et al.* (2004) *Antiviral Res.* 64(2):79-83; Vasilakos *et al.* (2000) *Cell Immunol.* 204(1):64-74; US patents 4689338, 4929624, 5238944, 5266575, 5268376, 5346905, 5352784, 5389640, 20 5395937, 5482936, 5494916, 5525612, 6083505, 6440992, 6627640, 6656938, 6660735, 6660747, 6664260, 6664264, 6664265, 6667312, 6670372, 6677347, 6677348, 6677349, 6683088, 6703402, 6743920, 6800624, 6809203, 6888000 and 6924293; and Jones (2003) *Curr Opin Investig Drugs* 4:214-218.

N. Substituted ureas

- 25 Substituted ureas useful as adjuvants include compounds of formula I, II or III, or salts thereof:



as defined in WO03/011223, such as ‘ER 803058’, ‘ER 803732’, ‘ER 804053’, ‘ER 804058’, ‘ER 804059’, ‘ER 804442’, ‘ER 804680’, ‘ER 804764’, ER 803022 or ‘ER 804057’ e.g.:

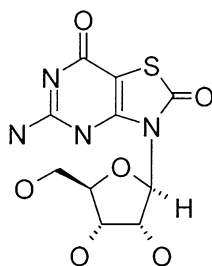


5 O. Further adjuvants

Further adjuvants that may be used with the invention include:

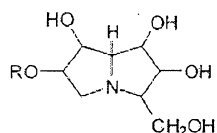
- An aminoalkyl glucosaminide phosphate derivative, such as RC-529 (Johnson *et al.* (1999) *Bioorg Med Chem Lett* 9:2273-2278; Evans *et al.* (2003) *Expert Rev Vaccines* 2:219-229).

- A thiosemicarbazone compound, such as those disclosed in WO2004/060308. Methods of formulating, manufacturing, and screening for active compounds are also described in Bhagat *et al.* (2003) *BBRC* 300:853-861. The thiosemicarbazones are particularly effective in the stimulation of human peripheral blood mononuclear cells for the production of cytokines, such as TNF- α .
- A tryptanthrin compound, such as those disclosed in WO2004/064759. Methods of formulating, manufacturing, and screening for active compounds are also described in WO03/035836. The thiosemicarbazones are particularly effective in the stimulation of human peripheral blood mononuclear cells for the production of cytokines, such as TNF- α .
- A nucleoside analog, such as: (a) Isatorabine (ANA-245; 7-thia-8-oxoguanosine):



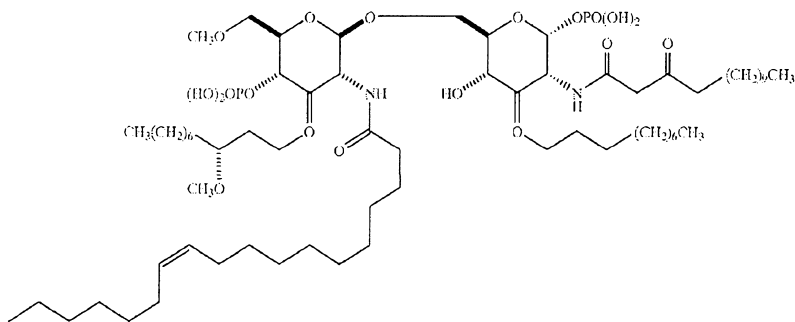
and prodrugs thereof; (b) ANA975; (c) ANA-025-1; (d) ANA380; (e) the compounds disclosed in US 6,924,271, US2005/0070556 and US 5,658,731, oxoribine (7-allyl-8-oxoguanosine) (US patent 5,011,828).

- Compounds disclosed in WO2004/87153, including: Acylpiperazine compounds, Indoleione compounds, Tetrahydroisoquinoline (THIQ) compounds, Benzocyclodione compounds, Aminoazavinyl compounds, Aminobenzimidazole quinolinone (ABIQ) compounds (US 6,605,617, WO02/18383), Hydraphtalamide compounds, Benzophenone compounds, Isoxazole compounds, Sterol compounds, Quinazolinone compounds, Pyrrole compounds (WO2004/018455), Anthraquinone compounds, Quinoxaline compounds, Triazine compounds, Pyrazalopyrimidine compounds, and Benzazole compounds (WO03/082272).
- Compounds containing lipids linked to a phosphate-containing acyclic backbone, such as the TLR4 antagonist E5564 (Wong *et al.* (2003) *J Clin Pharmacol* 43(7):735-42; US2005/0215517).
- A polyoxidonium polymer (Dyakonova *et al.* (2004) *Int Immunopharmacol* 4(13):1615-23; FR-2859633) or other N-oxidized polyethylene-piperazine derivative.
- Methyl inosine 5'-monophosphate ("MIMP") (Signorelli & Hadden (2003) *Int Immunopharmacol* 3(8):1177-86).
- A polyhydroxylated pyrrolizidine compound (WO2004/064715), such as one having formula:



where R is selected from the group comprising hydrogen, straight or branched, unsubstituted or substituted, saturated or unsaturated acyl, alkyl (*e.g.* cycloalkyl), alkenyl, alkynyl and aryl groups, or a pharmaceutically acceptable salt or derivative thereof. Examples include, but are not limited to: casuarine, casuarine-6- α -D-glucopyranose, 3-*epi*-casuarine, 7-*epi*-casuarine, 3,7-*diepi*-casuarine, *etc.*

- A CD1d ligand, such as an α -glycosylceramide (De Libero *et al*, *Nature Reviews Immunology*, 2005, 5: 485-496; US patent 5,936,076 ; Oki *et al*, *J. Clin. Investig.*, 113: 1631-1640 ; US2005/0192248; Yang *et al*, *Angew. Chem. Int. Ed.*, 2004, 43: 3818-3822; WO2005/102049; Goff *et al*, *J. Am. Chem., Soc.*, 2004, 126: 13602-13603; WO03/105769) *e.g.* α -galactosylceramide), phytosphingosine-containing α -glycosylceramides, OCH, KRN7000 [(2S,3S,4R)-1-O-(α -D-galactopyranosyl)-2-(N-hexacosanoylamino)-1,3,4-octadecanetriol], CRONY-101, 3''-O-sulfo-galactosylceramide, *etc.*
- A gamma inulin (Cooper (1995) *Pharm Biotechnol* 6:559-80) or derivative thereof, such as algammulin.



Adjuvant combinations

The invention may also comprise combinations of aspects of one or more of the adjuvants identified above. For example, the following adjuvant compositions may be used in the invention: (1) a saponin and an oil-in-water emulsion (WO99/11241); (2) a saponin (*e.g.* QS21) + a non-toxic LPS derivative (*e.g.* 3dMPL) (WO94/00153); (3) a saponin (*e.g.* QS21) + a non-toxic LPS derivative (*e.g.* 3dMPL) + a cholesterol; (4) a saponin (*e.g.* QS21) + 3dMPL + IL-12 (optionally + a sterol) (WO98/57659); (5) combinations of 3dMPL with, for example, QS21 and/or oil-in-water emulsions (European patent applications 0835318, 0735898 and 0761231); (6) SAF, containing 10% squalane, 0.4% Tween 80TM, 5% pluronic-block polymer L121, and thr-MDP, either microfluidized into a submicron emulsion or vortexed to generate a larger particle size emulsion. (7) RibiTM adjuvant system (RAS), (Ribi Immunochem) containing 2% squalene, 0.2% Tween 80, and one or more bacterial cell wall

components from the group consisting of monophosphorylipid A (MPL), trehalose dimycolate (TDM), and cell wall skeleton (CWS), preferably MPL + CWS (Detox™); and (8) one or more mineral salts (such as an aluminum salt) + a non-toxic derivative of LPS (such as 3dMPL). In some embodiments a combination of a toxin (e.g. LTK63) and an immunostimulatory oligonucleotide (e.g. CpG) is used. In some embodiments, a combination of an emulsion (e.g. montanide) and an immunostimulatory oligonucleotide (e.g. CpG) is used.

Other substances that act as immunostimulating agents are disclosed in chapter 7 of *Vaccine Design*, (1995) eds. Powell & Newman. ISBN: 030644867X. Plenum.

The use of an aluminium hydroxide and/or aluminium phosphate adjuvant is particularly preferred, and antigens are generally adsorbed to these salts. Calcium phosphate is another preferred adjuvant. Other preferred adjuvant combinations include combinations of Th1 and Th2 adjuvants such as CpG & alum or resiquimod & alum. A combination of aluminium phosphate and 3dMPL may be used.

To improve thermal stability, a composition may include a temperature protective agent. This component may be particularly useful in adjuvanted compositions (particularly those containing a mineral adjuvant, such as an aluminium salt). As described in WO2006/110603, a liquid temperature protective agent may be added to an aqueous vaccine composition to lower its freezing point e.g. to reduce the freezing point to below 0°C. Thus the composition can be stored below 0°C, but above its freezing point, to inhibit thermal breakdown. The temperature protective agent also permits freezing of the composition while protecting mineral salt adjuvants against agglomeration or sedimentation after freezing and thawing, and may also protect the composition at elevated temperatures e.g. above 40°C. A starting aqueous vaccine and the liquid temperature protective agent may be mixed such that the liquid temperature protective agent forms from 1-80% by volume of the final mixture. Suitable temperature protective agents should be safe for human administration, readily miscible/soluble in water, and should not damage other components (e.g. antigen and adjuvant) in the composition. Examples include glycerin, propylene glycol, and/or polyethylene glycol (PEG). Suitable PEGs may have an average molecular weight ranging from 200-20,000 Da. In a preferred embodiment, the polyethylene glycol can have an average molecular weight of about 300 Da ('PEG-300').

The invention provides an immunogenic composition comprising: (i) one or more proteins of the invention; and (ii) a temperature protective agent. This composition may be formed by mixing (i) an aqueous composition comprising one or more proteins of the invention, with (ii) a temperature protective agent. The mixture may then be stored e.g. below 0°C, from 0-20°C, from 20-35°C, from 35-55°C, or higher. It may be stored in liquid or frozen form. The mixture may be lyophilised. The composition may alternatively be formed by mixing (i) a dried composition comprising one or more proteins of the invention, with (ii) a liquid composition comprising the temperature protective agent. Thus component (ii) can be used to reconstitute component (i).

The compositions of the invention may elicit either or both of a cell mediated immune response and a humoral immune response. This immune response will preferably induce long lasting (*e.g.* neutralising) antibodies and a cell mediated immunity that can quickly respond upon exposure to chlamydia.

5 Two types of T cells, CD4 and CD8 cells, are generally thought necessary to initiate and/or enhance cell mediated immunity and humoral immunity. CD8 T cells can express a CD8 co-receptor and are commonly referred to as Cytotoxic T lymphocytes (CTLs). CD8 T cells are able to recognized or interact with antigens displayed on MHC Class I molecules.

CD4 T cells can express a CD4 co-receptor and are commonly referred to as T helper cells. CD4 T
10 cells are able to recognize antigenic peptides bound to MHC class II molecules. Upon interaction with a MHC class II molecule, the CD4 cells can secrete factors such as cytokines. These secreted cytokines can activate B cells, cytotoxic T cells, macrophages, and other cells that participate in an immune response. Helper T cells or CD4+ cells can be further divided into two functionally distinct subsets: TH1 phenotype and TH2 phenotypes which differ in their cytokine and effector function.

15 Activated TH1 cells enhance cellular immunity (including an increase in antigen-specific CTL production) and are therefore of particular value in responding to intracellular infections. Activated TH1 cells may secrete one or more of IL-2, IFN γ , and TNF- β . A TH1 immune response may result in local inflammatory reactions by activating macrophages, NK (natural killer) cells, and CD8 cytotoxic T cells (CTLs). A TH1 immune response may also act to expand the immune response by stimulating
20 growth of B and T cells with IL-12. TH1 stimulated B cells may secrete IgG2a.

Activated TH2 cells enhance antibody production and are therefore of value in responding to extracellular infections. Activated TH2 cells may secrete one or more of IL-4, IL-5, IL-6, and IL-10. A TH2 immune response may result in the production of IgG1, IgE, IgA and memory B cells for future protection.

25 An enhanced immune response may include one or more of an enhanced TH1 immune response and a TH2 immune response.

A TH1 immune response may include one or more of an increase in CTLs, an increase in one or more of the cytokines associated with a TH1 immune response (such as IL-2, IFN γ , and TNF- β), an increase in activated macrophages, an increase in NK activity, or an increase in the production of
30 IgG2a. Preferably, the enhanced TH1 immune response will include an increase in IgG2a production.

A TH1 immune response may be elicited using a TH1 adjuvant. A TH1 adjuvant will generally elicit increased levels of IgG2a production relative to immunization of the antigen without adjuvant. TH1 adjuvants suitable for use in the invention may include for example saponin formulations, virosomes and virus like particles, non-toxic derivatives of enterobacterial lipopolysaccharide (LPS),

immunostimulatory oligonucleotides. Immunostimulatory oligonucleotides, such as oligonucleotides containing a CpG motif, are preferred TH1 adjuvants for use in the invention.

5 A TH2 immune response may include one or more of an increase in one or more of the cytokines associated with a TH2 immune response (such as IL-4, IL-5, IL-6 and IL-10), or an increase in the production of IgG1, IgE, IgA and memory B cells. Preferably, the enhanced TH2 immune response will include an increase in IgG1 production.

10 A TH2 immune response may be elicited using a TH2 adjuvant. A TH2 adjuvant will generally elicit increased levels of IgG1 production relative to immunization of the antigen without adjuvant. TH2 adjuvants suitable for use in the invention include, for example, mineral containing compositions, oil-emulsions, and ADP-ribosylating toxins and detoxified derivatives thereof. Mineral containing compositions, such as aluminium salts are preferred TH2 adjuvants for use in the invention.

15 Preferably, the invention includes a composition comprising a combination of a TH1 adjuvant and a TH2 adjuvant. Preferably, such a composition elicits an enhanced TH1 and an enhanced TH2 response, i.e., an increase in the production of both IgG1 and IgG2a production relative to immunization without an adjuvant. Still more preferably, the composition comprising a combination of a TH1 and a TH2 adjuvant elicits an increased TH1 and/or an increased TH2 immune response relative to immunization with a single adjuvant (*i.e.*, relative to immunization with a TH1 adjuvant alone or immunization with a TH2 adjuvant alone).

20 The immune response may be one or both of a TH1 immune response and a TH2 response. Preferably, immune response provides for one or both of an enhanced TH1 response and an enhanced TH2 response. Preferably, the immune response includes an increase in the production of IgG1 and/or IgG2 and/or IgGA.

25 The invention is preferably used to elicit systemic and/or mucosal immunity. The enhanced immune response may be one or both of a systemic and a mucosal immune response. Preferably, the immune response provides for one or both of an enhanced systemic and an enhanced mucosal immune response. Preferably the mucosal immune response is a TH2 immune response. Preferably, the mucosal immune response includes an increase in the production of IgA.

Methods of treatment, and administration of the vaccine

30 The invention also provides a method for raising an immune response in a mammal comprising the step of administering an effective amount of a protein, antibody, nucleic acid, vector, host cell or composition of the invention. The immune response is preferably protective and preferably involves antibodies and/or cell-mediated immunity. The method may raise a booster response.

The invention also provides a protein or combination, as defined above, for use as a medicament *e.g.* for use in raising an immune response in a mammal.

The invention also provides the use of a protein or combination of the invention in the manufacture of a medicament for raising an immune response in a mammal. By raising an immune response in the mammal by these uses and methods, the mammal can be protected against Chlamydia infection. More particularly, the mammal may be protected against *Chlamydia trachomatis*. The invention is effective against Chlamydia of various different serotypes, but can be particularly useful in protecting against disease resulting from Chlamydia infection by strains in serovar D.

Thus, according to a further aspect, the invention also provides a nucleic acid, protein, antibody, vector or host cell according to the invention for use as a medicament (*e.g.* a vaccine) or a diagnostic reagent. In one embodiment, the protein, nucleic acid or antibody is used for treatment, prevention or diagnosis of Chlamydia infection (preferably *C. trachomatis*) in a mammal. The invention also provides a method of treating, preventing or diagnosing Chlamydia infection (preferably, *C. trachomatis* infection) in a patient (preferably a mammal), comprising administering a therapeutically effective amount of a nucleic acid, protein or antibody of the invention.

Preferably, the nucleic acid, protein or antibody according to the invention is for treatment or prevention of Chlamydia infection or an associated condition (*e.g.* trachoma, blindness, cervicitis, pelvic inflammatory disease, infertility, ectopic pregnancy, chronic pelvic pain, salpingitis, urethritis, epididymitis, infant pneumonia, cervical squamous cell carcinoma, *etc.*), preferably, *C. trachomatis* infection. The immunogenic composition may additionally or alternatively be effective against *C. pneumoniae*.

The mammal is preferably a human. Where the vaccine is for prophylactic use, the human is preferably a child (*e.g.* a toddler or infant) or a teenager; where the vaccine is for therapeutic use, the human is preferably a teenager or an adult. A vaccine intended for children may also be administered to adults *e.g.* to assess safety, dosage, immunogenicity, *etc.* Thus a human patient may be less than 1 year old, 1-5 years old, 5-15 years old, 15-55 years old, or at least 55 years old. Preferred patients for receiving the vaccines are people going through puberty, teenagers, sexually active people, the elderly (*e.g.* ≥ 50 years old, ≥ 60 years old, and preferably ≥ 65 years), the young (*e.g.* ≤ 5 years old), hospitalised patients, healthcare workers, armed service and military personnel, pregnant women, the chronically ill, or immunodeficient patients. The vaccines are not suitable solely for these groups, however, and may be used more generally in a population.

Vaccines produced by the invention may be administered to patients at substantially the same time as (*e.g.* during the same medical consultation or visit to a healthcare professional or vaccination centre) other vaccines *e.g.* at substantially the same time as a human papillomavirus vaccine such as Cervarix™ or Gardasil™; a tetanus, diphtheria and acellular pertussis vaccine such as TDaP, DTaP or Boostrix™; a rubella vaccine such as MMR; or a tuberculosis vaccine such as the BCG. Examples of other vaccines that the vaccine produced by the invention may be administered at substantially the same time as are a measles vaccine, a mumps vaccine, a varicella vaccine, a MMRV vaccine, a diphtheria vaccine, a tetanus vaccine, a pertussis vaccine, a DTP vaccine, a conjugated *H. influenzae*

type b vaccine, an inactivated poliovirus vaccine, a hepatitis B virus vaccine, a meningococcal conjugate vaccine (such as a tetravalent A-C-W135-Y vaccine), a respiratory syncytial virus vaccine, etc.

5 In a preferred embodiment, the protein of the invention is used to elicit antibodies that are capable of neutralising the activity of the wild type Chlamydia protein, for example, of one or more of wild-type Chlamydia CT733, CT153, CT601, CT279, CT443, CT372, CT456, CT381, CT255, CT341, CT716, CT745, CT387, CT812, CT869, CT166, CT175, CT163, CT214, CT721, CT127, CT043, CT600 and/or CT823 for example, of one or more of wild-type Chlamydia CT733, CT153, CT601, CT279, CT443, CT372, CT456 and/or CT381. Neutralizing antibodies may be used as a vaccine capable of
10 neutralising the activity of a native Chlamydia protein expressed by infectious EB. In one embodiment, the protein of the invention is used to elicit antibodies that are capable of neutralising Chlamydia infectivity and/or virulence. Thus, the invention also provides the antibodies of the invention for neutralising wild-type Chlamydia proteins and/or Chlamydia infectivity and/or virulence.

15 The invention also provides the use of a nucleic acid, protein, or antibody of the invention in the manufacture of: (i) a medicament for treating or preventing bacterial infection; (ii) a diagnostic reagent for detecting the presence of bacteria or of antibodies raised against bacteria; and/or (iii) a reagent which can raise antibodies against bacteria. Said bacteria is preferably a *Chlamydia*, e.g. *Chlamydia trachomatis* or *Chlamydia pneumoniae*, but is preferably *Chlamydia trachomatis*.

20 Also provided is a method for diagnosing Chlamydia infection, comprising:

- (a) raising an antibody against a protein of the invention;
- (b) contacting the antibody of step (a) with a biological sample suspected of being infected with Chlamydia under conditions suitable for the formation of antibody-antigen complexes; and
25 (c) detecting said complexes, wherein detection of said complex is indicative of Chlamydia infection.

Also provided is a method for diagnosing Chlamydia infection, comprising: (a) contacting an antibody which was raised against a protein of the invention with a biological sample suspected of being infected with Chlamydia under conditions suitable for the formation of antibody-antigen
30 complexes; and (b) detecting said complexes, wherein detection of said complex is indicative of Chlamydia infection.

Proteins of the invention can be used in immunoassays to detect antibody levels (or, conversely, antibodies of the invention can be used to detect protein levels). Immunoassays based on well defined, recombinant antigens can be developed to replace invasive diagnostics methods. Antibodies
35 to proteins within biological samples, including for example, blood or serum samples, can be

detected. Design of the immunoassays is subject to a great deal of variation, and a variety of these are known in the art. Protocols for the immunoassay may be based, for example, upon competition, or direct reaction, or sandwich type assays. Protocols may also, for example, use solid supports, or may be by immunoprecipitation. Most assays involve the use of labeled antibody or polypeptide; the labels may be, for example, fluorescent, chemiluminescent, radioactive, or dye molecules. Assays which amplify the signals from the probe are also known; examples of which are assays which utilize biotin and avidin, and enzyme-labeled and mediated immunoassays, such as ELISA assays.

Kits suitable for immunodiagnosis and containing the appropriate labeled reagents are constructed by packaging the appropriate materials, including the compositions of the invention, in suitable containers, along with the remaining reagents and materials (for example, suitable buffers, salt solutions, *etc.*) required for the conduct of the assay, as well as suitable set of assay instructions.

Testing efficacy of compositions

The efficacy of the immunogenic compositions of the present invention can be evaluated in *in vitro* and *in vivo* animal models prior to host, e.g., human, administration. For example, *in vitro* neutralization by Peterson et al (1988) is suitable for testing vaccine compositions directed toward *Chlamydia trachomatis*.

One way of checking efficacy of therapeutic treatment involves monitoring *C. trachomatis* infection after administration of the compositions of the invention. One way of checking efficacy of prophylactic treatment involves monitoring immune responses both systemically (such as monitoring the level of IgG1 and IgG2a production) and mucosally (such as monitoring the level of IgA production) against the *Chlamydia trachomatis* antigens in the compositions of the invention after administration of the composition. Typically, serum *Chlamydia* specific antibody responses are determined post-immunisation but pre-challenge whereas mucosal *Chlamydia* specific antibody body responses are determined post-immunisation and post-challenge.

One example of such an *in vitro* test is described as follows. Hyper-immune antisera is diluted in PBS containing 5% guinea pig serum, as a complement source. *Chlamydia trachomatis* (10^4 IFU; inclusion forming units) are added to the antisera dilutions. The antigen-antibody mixtures are incubated at 37°C for 45 minutes and inoculated into duplicate confluent Hep-2 or HeLa cell monolayers contained in glass vials (e.g., 15 by 45 mm), which have been washed twice with PBS prior to inoculation. The monolayer cells are infected by centrifugation at 1000X g for 1 hour followed by stationary incubation at 37°C for 1 hour. Infected monolayers are incubated for 48 or 72 hours, fixed and stained with Chlamydia specific antibody, such as anti-MOMP. Inclusion-bearing cells are counted in ten fields at a magnification of 200X. Neutralization titer is assigned on the dilution that gives 50% inhibition as compared to control monolayers/IFU.

Another way of assessing the immunogenicity of the compositions of the present invention is to express the proteins recombinantly for screening patient sera or mucosal secretions by immunoblot

and/or microarrays. A positive reaction between the protein and the patient sample indicates that the patient has mounted an immune response to the protein in question. This method may also be used to identify immunodominant antigens and/or epitopes within antigens.

The efficacy of vaccine compositions can also be determined *in vivo* by challenging animal models of *Chlamydia trachomatis* infection, e.g., guinea pigs or mice, with the vaccine compositions. For example, *in vivo* vaccine composition challenge studies in the guinea pig model of *Chlamydia trachomatis* infection can be performed. A description of one example of this type of approach follows. Female guinea pigs weighing 450 – 500 g are housed in an environmentally controlled room with a 12 hour light-dark cycle and immunized with vaccine compositions via a variety of immunization routes. Post-vaccination, guinea pigs are infected in the genital tract with the agent of guinea pig inclusion conjunctivitis (GPIC), which has been grown in HeLa or McCoy cells (Rank et al. (1988)). Each animal receives approximately 1.4×10^7 inclusion forming units (IFU) contained in 0.05 ml of sucrose-phosphate-glutamate buffer, pH 7.4 (Schacter, 1980). The course of infection monitored by determining the percentage of inclusion-bearing cells by indirect immunofluorescence with GPIC specific antisera, or by Giemsa-stained smear from a scraping from the genital tract (Rank et al 1988). Antibody titers in the serum is determined by an enzyme-linked immunosorbent assay.

Alternatively, *in vivo* vaccine compositions challenge studies can be performed in the murine model of *Chlamydia trachomatis* (Morrison et al 1995). A description of one example of this type of approach is as follows. Female mice 7 to 12 weeks of age receive 2.5 mg of depo-provera subcutaneously at 10 and 3 days before vaginal infection. Post-vaccination, mice are infected in the genital tract with 1,500 inclusion-forming units of *Chlamydia trachomatis* contained in 5ml of sucrose-phosphate-glutamate buffer, pH 7.4. The course of infection is monitored by determining the percentage of inclusion-bearing cells by indirect immunofluorescence with *Chlamydia trachomatis* specific antisera, or by a Giemsa-stained smear from a scraping from the genital tract of an infected mouse. The presence of antibody titers in the serum of a mouse is determined by an enzyme-linked immunosorbent assay.

Nucleic acid immunisation

The immunogenic compositions described above include Chlamydia antigens. In all cases, however, the polypeptide antigens can be replaced by nucleic acids (typically DNA) encoding those polypeptides, to give compositions, methods and uses based on nucleic acid immunisation. Nucleic acid immunisation is now a developed field (e.g. see Donnelly *et al.* (1997) *Annu Rev Immunol* 15:617-648; Strugnell *et al.* (1997) *Immunol Cell Biol* 75(4):364-369; Cui (2005) *Adv Genet* 54:257-89; Robinson & Torres (1997) *Seminars in Immunol* 9:271-283; Brunham *et al.* (2000) *J Infect Dis* 181 Suppl 3:S538-43; Svanholm *et al.* (2000) *Scand J Immunol* 51(4):345-53; *DNA Vaccination - Genetic Vaccination* (1998) eds. Koprowski *et al.* (ISBN 3540633928); *Gene Vaccination : Theory and Practice* (1998) ed. Raz (ISBN 3540644288), etc.).

The nucleic acid encoding the immunogen is expressed *in vivo* after delivery to a patient and the expressed immunogen then stimulates the immune system. The active ingredient will typically take the form of a nucleic acid vector comprising: (i) a promoter; (ii) a sequence encoding the immunogen, operably linked to the promoter; and optionally (iii) a selectable marker. Preferred
5 vectors may further comprise (iv) an origin of replication; and (v) a transcription terminator downstream of and operably linked to (ii). In general, (i) & (v) will be eukaryotic and (iii) & (iv) will be prokaryotic.

Preferred promoters are viral promoters *e.g.* from cytomegalovirus (CMV). The vector may also include transcriptional regulatory sequences (*e.g.* enhancers) in addition to the promoter and which
10 interact functionally with the promoter. Preferred vectors include the immediate-early CMV enhancer/promoter, and more preferred vectors also include CMV intron A. The promoter is operably linked to a downstream sequence encoding an immunogen, such that expression of the immunogen-encoding sequence is under the promoter's control.

Where a marker is used, it preferably functions in a microbial host (*e.g.* in a prokaryote, in a bacteria,
15 in a yeast). The marker is preferably a prokaryotic selectable marker (*e.g.* transcribed under the control of a prokaryotic promoter). For convenience, typical markers are antibiotic resistance genes.

The vector of the invention is preferably an autonomously replicating episomal or extrachromosomal vector, such as a plasmid.

The vector of the invention preferably comprises an origin of replication. It is preferred that the
20 origin of replication is active in prokaryotes but not in eukaryotes.

Preferred vectors thus include a prokaryotic marker for selection of the vector, a prokaryotic origin of replication, but a *eukaryotic* promoter for driving transcription of the immunogen-encoding sequence. The vectors will therefore (a) be amplified and selected in prokaryotic hosts without polypeptide expression, but (b) be expressed in eukaryotic hosts without being amplified. This
25 arrangement is ideal for nucleic acid immunization vectors.

The vector of the invention may comprise a eukaryotic transcriptional terminator sequence downstream of the coding sequence. This can enhance transcription levels. Where the coding sequence does not have its own, the vector of the invention preferably comprises a polyadenylation sequence. A preferred polyadenylation sequence is from bovine growth hormone.

30 The vector of the invention may comprise a multiple cloning site.

In addition to sequences encoding the immunogen and a marker, the vector may comprise a second eukaryotic coding sequence. The vector may also comprise an IRES upstream of said second sequence in order to permit translation of a second eukaryotic polypeptide from the same transcript as the immunogen. Alternatively, the immunogen-coding sequence may be downstream of an IRES.

The vector of the invention may comprise unmethylated CpG motifs *e.g.* unmethylated DNA sequences which have in common a cytosine preceding a guanosine, flanked by two 5' purines and two 3' pyrimidines. In their unmethylated form these DNA motifs have been demonstrated to be potent stimulators of several types of immune cell.

5 Vectors may be delivered in a targeted way. Receptor-mediated DNA delivery techniques are described in, for example, Findeis *et al.*, *Trends Biotechnol.* (1993) 11:202; Chiou *et al.* (1994) *Gene Therapeutics: Methods And Applications Of Direct Gene Transfer*. ed. Wolff; Wu *et al.*, *J. Biol. Chem.* (1988) 263:621; Wu *et al.*, *J. Biol. Chem.* (1994) 269:542; Zenke *et al.*, *Proc. Natl. Acad. Sci. (USA)* (1990) 87:3655; and Wu *et al.*, *J. Biol. Chem.* (1991) 266:338.

10 Therapeutic compositions containing a nucleic acid are administered in a range of about 100ng to about 200mg of DNA for local administration in a gene therapy protocol. Concentration ranges of about 500 ng to about 50 mg, about 1µg to about 2 mg, about 5µg to about 500µg, and about 20µg to about 100µg of DNA can also be used during a gene therapy protocol. Factors such as method of action (*e.g.* for enhancing or inhibiting levels of the encoded gene product) and efficacy of
15 transformation and expression are considerations which will affect the dosage required for ultimate efficacy. Where greater expression is desired over a larger area of tissue, larger amounts of vector or the same amounts re-administered in a successive protocol of administrations, or several administrations to different adjacent or close tissue portions may be required to effect a positive therapeutic outcome. In all cases, routine experimentation in clinical trials will determine specific
20 ranges for optimal therapeutic effect.

Vectors can be delivered using gene delivery vehicles. The gene delivery vehicle can be of viral or non-viral origin (see generally Jolly, *Cancer Gene Therapy* (1994) 1:51; Kimura, *Human Gene Therapy* (1994) 5:845; Connelly, *Human Gene Therapy* (1995) 1:185; and Kaplitt, *Nature Genetics* (1994) 6:148).

25 Viral-based vectors for delivery of a desired nucleic acid and expression in a desired cell are well known in the art. Exemplary viral-based vehicles include, but are not limited to, recombinant retroviruses (*e.g.* WO 90/07936; WO 94/03622; WO 93/25698; WO 93/25234; US patent 5,219,740; WO 93/11230; WO 93/10218; US patent 4,777,127; GB Patent No. 2,200,651; EP-A-0345242; and WO 91/02805), alphavirus-based vectors (*e.g.* Sindbis virus vectors, Semliki forest virus (ATCC
30 VR-67; ATCC VR-1247), Ross River virus (ATCC VR-373; ATCC VR-1246) and Venezuelan equine encephalitis virus (ATCC VR-923; ATCC VR-1250; ATCC VR 1249; ATCC VR-532); hybrids or chimeras of these viruses may also be used), poxvirus vectors (*e.g.* vaccinia, fowlpox, canarypox, modified vaccinia Ankara, *etc.*), adenovirus vectors, and adeno-associated virus (AAV)
35 vectors (*e.g.* see WO 90/07936; WO 94/03622; WO 93/25698; WO 93/25234; US patent 5,219,740; WO 93/11230; WO 93/10218; US patent 4,777,127; GB Patent No. 2,200,651; EP-A-0345242; WO 91/02805; WO 94/12649; WO 93/03769; WO 93/19191; WO 94/28938; WO 95/11984; and WO

95/00655). Administration of DNA linked to killed adenovirus (Curiel, *Hum. Gene Ther.* (1992) 3:147) can also be employed.

Non-viral delivery vehicles and methods can also be employed, including, but not limited to, polycationic condensed DNA linked or unlinked to killed adenovirus alone (e.g. De Libero *et al.*, *Nature Reviews Immunology*, 2005, 5: 485-496), ligand-linked DNA (Wu, *J. Biol. Chem.* (1989) 264:16985), eukaryotic cell delivery vehicles cells (US patent 5,814,482; WO 95/07994; WO 96/17072; WO 95/30763; and WO 97/42338) and nucleic charge neutralization or fusion with cell membranes. Naked DNA can also be employed. Exemplary naked DNA introduction methods are described in WO 90/11092 and US patent 5,580,859. Liposomes (e.g. immunoliposomes) that can act as gene delivery vehicles are described in US patent 5,422,120; WO 95/13796; WO 94/23697; WO 91/14445; and EP-0524968. Additional approaches are described in Philip, *Mol. Cell Biol.* (1994) 14:2411 and Woffendin, *Proc. Natl. Acad. Sci.* (1994) 91:11581.

Further non-viral delivery suitable for use includes mechanical delivery systems such as the approach described in Donnelly *et al.* (1997) *Annu Rev Immunol* 15:617-648. Moreover, the coding sequence and the product of expression of such can be delivered through deposition of photopolymerized hydrogel materials or use of ionizing radiation (e.g. US patent 5,206,152 and WO 92/11033). Other conventional methods for gene delivery that can be used for delivery of the coding sequence include, for example, use of hand-held gene transfer particle gun (US patent 5,149,655) or use of ionizing radiation for activating transferred genes (Strugnell *et al.* (1997) *Immunol Cell Biol* 75(4):364-369 and Cui (2005) *Adv Genet* 54:257-89).

Delivery DNA using PLG {poly(lactide-co-glycolide)} microparticles is a particularly preferred method e.g. by adsorption to the microparticles, which are optionally treated to have a negatively-charged surface (e.g. treated with SDS) or a positively-charged surface (e.g. treated with a cationic detergent, such as CTAB).

25 ***Antibody immunisation***

The antibodies of the invention may be used, for example, for neutralising the activity of the wild-type Chlamydia protein. Antibodies against Chlamydia antigens can be used for passive immunisation (Brandt *et al.* (2006) *J Antimicrob Chemother.* 58(6):1291-4. Epub 2006 Oct 26). Thus the invention provides the use of antibodies of the invention in therapy. The invention also provides the use of such antibodies in the manufacture of a medicament. The invention also provides a method for treating a mammal comprising the step of administering an effective amount of an antibody of the invention. As described above for immunogenic compositions, these methods and uses allow a mammal to be protected against Chlamydia infection.

Processes

35 According to further aspects, the invention provides various processes.

A process for producing a protein of the invention is provided, comprising the step of culturing a host cell of the invention under conditions which induce protein expression.

A process for producing protein or nucleic acid of the invention is provided, wherein the protein or nucleic acid is synthesised in part or in whole using chemical means.

- 5 A process for detecting Chlamydia (preferably *C. trachomatis*) in a biological sample is also provided, comprising the step of contacting a nucleic acid according to the invention with the biological sample under hybridising conditions. The process may involve nucleic acid amplification (*e.g.* PCR, SDA, SSSR, LCR, TMA *etc.*) or hybridisation (*e.g.* microarrays, blots, hybridisation with probe in solution *etc.*).
- 10 A process for detecting wild-type Chlamydia (preferably, *C. trachomatis*) is provided, comprising the steps of: (a) contacting an antibody of the invention with a biological sample under conditions suitable for the formation of an antibody-antigen complex(es); and (b) detecting said complex(es). This process may advantageously be used to diagnose Chlamydia infection.

General

- 15 The practice of the present invention will employ, unless otherwise indicated, conventional methods of chemistry, biochemistry, molecular biology, immunology and pharmacology, within the skill of the art. Such techniques are explained fully in the literature. See, *e.g.*, Gennaro (2000) *Remington: The Science and Practice of Pharmacy*. 20th edition, ISBN: 0683306472; *Methods In Enzymology* (S. Colowick and N. Kaplan, eds., Academic Press, Inc.); *Handbook of Experimental Immunology*,
20 Vols. I-IV (D.M. Weir and C.C. Blackwell, eds, 1986, Blackwell Scientific Publications); Sambrook *et al.* (2001) *Molecular Cloning: A Laboratory Manual*, 3rd edition (Cold Spring Harbor Laboratory Press); *Handbook of Surface and Colloidal Chemistry* (Birdi, K.S. ed., CRC Press, 1997); Ausubel *et al.* (eds) (2002) *Short protocols in molecular biology*, 5th edition (Current Protocols); *Molecular Biology Techniques: An Intensive Laboratory Course*, (Ream *et al.*, eds., 1998, Academic Press); and
25 *PCR (Introduction to Biotechniques Series)*, 2nd ed. (Newton & Graham eds., 1997, Springer Verlag) *etc.*

“GI” numbering is used herein. A GI number, or “GenInfo Identifier”, is a series of digits assigned consecutively to each sequence record processed by NCBI when sequences are added to its databases. The GI number bears no resemblance to the accession number of the sequence record.

- 30 When a sequence is updated (*e.g.* for correction, or to add more annotation or information) then it receives a new GI number. Thus the sequence associated with a given GI number is never changed.

- Where the invention concerns an “epitope”, this epitope may be a B-cell epitope and/or a T-cell epitope. Such epitopes can be identified empirically (*e.g.* using PEPSCAN (Geysen *et al.* (1984) *PNAS USA* 81:3998-4002; Carter (1994) *Methods Mol Biol* 36:207-23) or similar methods), or they
35 can be predicted (*e.g.* using the Jameson-Wolf antigenic index (Jameson, BA *et al.* 1988, *CABIOS* 4(1):181-186), matrix-based approaches (Raddrizzani & Hammer (2000) *Brief Bioinform* 1(2):179-

89), MAPITOPE (Bublil *et al.* (2007) *Proteins* 68(1):294-304), TEPITOPE (De Lalla *et al.* (1999) *J. Immunol.* 163:1725-29; Kwok *et al.* (2001) *Trends Immunol* 22:583-88), neural networks (Brusic *et al.* (1998) *Bioinformatics* 14(2):121-30), OptiMer & EpiMer (Meister *et al.* (1995) *Vaccine* 13(6):581-91; Roberts *et al.* (1996) *AIDS Res Hum Retroviruses* 12(7):593-610), ADEPT (Maksyutov & Zagrebelnaya (1993) *Comput Appl Biosci* 9(3):291-7), Tsites (Feller & de la Cruz (1991) *Nature* 349(6311):720-1), hydrophilicity (Hopp (1993) *Peptide Research* 6:183-190), antigenic index (Welling *et al.* (1985) *FEBS Lett.* 188:215-218) or the methods disclosed in Davenport *et al.* (1995) *Immunogenetics* 42:392-297; Tsurui & Takahashi (2007) *J Pharmacol Sci.* 105(4):299-316; Tong *et al.* (2007) *Brief Bioinform.* 8(2):96-108 ; Schirle *et al.* (2001) *J Immunol Methods.* 257(1-2):1-16; and Chen *et al.* (2007) *Amino Acids* 33(3):423-8, *etc.*). Epitopes are the parts of an antigen that are recognised by and bind to the antigen binding sites of antibodies or T-cell receptors, and they may also be referred to as “antigenic determinants”.

Where an antigen “domain” is omitted, this may involve omission of a signal peptide, of a cytoplasmic domain, of a transmembrane domain, of an extracellular domain, *etc.*

15 The term “comprising” encompasses “including” as well as “consisting” *e.g.* a composition “comprising” X may consist exclusively of X or may include something additional *e.g.* X + Y.

The term “about” in relation to a numerical value x is optional and means, for example, $x \pm 10\%$.

References to a percentage sequence identity between two amino acid sequences means that, when aligned, that percentage of amino acids are the same in comparing the two sequences. This alignment and the percent homology or sequence identity can be determined using software programs known in the art, for example those described in section 7.7.18 of *Current Protocols in Molecular Biology* (F.M. Ausubel *et al.*, eds., 1987) Supplement 30. A preferred alignment is determined by the Smith-Waterman homology search algorithm using an affine gap search with a gap open penalty of 12 and a gap extension penalty of 2, BLOSUM matrix of 62. The Smith-Waterman homology search algorithm is disclosed in Smith & Waterman (1981) *Adv. Appl. Math.* 2: 482-489.

BRIEF DESCRIPTION OF DRAWINGS

Figure 1 is a graph which shows the ability of 20 selected *C. trachomatis* antigens to induce IFN γ production by CD4+ T cells.

30 Figure 2a shows the bacterial shedding (IFUs recovered from lungs) after Chlamydia challenge in mice to whom EB-CM CD4+ T cells had been adoptively transferred. Figure 2b shows the ability of various *C. muridarum* antigens to stimulate the protective EB-CD4+ T cell line to produce IFN γ .

Figure 3 is a histogram which shows the number of CD4+ T cells that produce IFN γ , upon specific stimulation with *C. trachomatis* recombinant antigens CT153 and CT733.

35 Figure 4 shows the protective activity of TC0106 (*C. muridarum* homologue of CT733) and TC0431 (*C. muridarum* homologue of CT153) as single antigens. The graph shows mean IFU/ml in BALB/C

mice immunised with the two antigens and then challenged with *C. muridarum*. The three bars are, from left to right: adjuvant alone; TC0106 as immunogen; and TC0431 as immunogen.

Figure 5 shows the protective activity of the combination TC0106+TC0431. The graph shows mean IFU per lung (Log10) recovered from infected lungs of mice immunised with the combination. The three bars are, from left to right: 10³ live Ebs; adjuvant alone; antigen combination.

Figure 6 shows CD4 T cells producing IFN γ in PBMC of mice immunized with TC0106+TC0431, TC0106, TC0431 and LTK 63+CpG. From left to right, the bars represent stimulation with 1) LTK 63, TC0106+TC0431, TC0106, TC0431 (all EB-immunized mice); 2) LTK 63, TC0106+TC0431, TC0106 (all TC 0106-immunized mice); 3) LTK63, TC0106+TC0431, TC0431 (all TC0431-immunized mice); and 4) LTK63 and TC0106+TC0431 (both TC0106+TC0431-immunized mice). It shows that immunization with TC0106 (*C. muridarum* homologue of CT733) and TC0431 (*C. muridarum* homologue of CT153) elicits a significant frequency of specific CD4+/IFN γ + cells. The Y axis shows frequency on 10⁶ CD4.

Figure 7 is a summary of protection results for various combinations and single antigens in the mouse model of *C. muridarum* intranasal challenge. It shows the mean IFU/lung of mice immunised intramuscularly with single antigens, or antigen combinations, adjuvanted with LTK63 and CpG, then challenged intranasally with 10³ *C. muridarum* IFU.

Figure 8 is a summary of protection results for various combinations of antigens in the mouse model of *C. muridarum* intranasal challenge. It shows mean IFU/lung (log 10) of *C. muridarum* recovered from infected lungs of immunised mice.

Figure 9 shows the results of the combination TC0551+TC0651+TC0727+TC0890 in the mouse model of ovarian bursa challenge with *C. muridarum*. The Y axis shows IFU/swab (log10). The three groups, from left to right, are for different immunizing antigens: ovalbumin; the combination; and nMOMP.

Figure 10A shows the protection results achieved with various antigens combinations in the mouse model of *C. muridarum* intranasal challenge. Figure 10B shows the frequency of IFN γ -producing CD4+ T cells induced by vaccination with the antigen combination TC0890+TC0551. From left to right, the bars represent stimulation with 1) TC0551, TC0890, TC0551+TC0890 (for adjuvant-immunized mice) and 2) TC0551, TC0890, TC0551+TC0890 (for MIX TC0890+TC0551-immunized mice). Figure 10C shows CD4 T cells producing IFN γ and IL2/TNF in PBMC of mice immunized with TC0106+TC0431 with Ltk63 + CpG. From left to right, the bars represent stimulation with a) TC0106, TC0431, TC0106+TC0431, CT153+CT733 (all adjuvant-immunized mice); 2) TC0106, TC0431, TC0106+TC0431, CT153+CT733 (all MIX TC0106+TC0431-immunized mice).

Figure 11 shows an immunoblot analysis of CT601, CT279, CT153 and CT733 in Ct-EBs and *C. trachomatis*-infected HeLa cells using their specific mouse immune antisera.

Figure 12 shows protective activity of antigens TC0313, TC0741, TC0106 and TC0210 given singly or in combination. In 12A to 12D the bars show mean IFU/lung (Log10), with the left-hand bar being adjuvant alone (LTK61+CpG) and the right-hand bar being the TC antigen. Figure 12E shows the IFU reduction over time (Mean IFU/lung against days post-challenge) using the combination (squares) or adjuvant alone (diamonds).

Figure 13A and 13B are histograms showing the antigen specific CD4 Th1 response in BALB/c mice after a primary *C. trachomatis* (CT) infection. Results are the mean of 4 independent experiments. Two results are shown for each experiment: non-infected mice (left hand bar) and primary infected mice (right hand bar). From left to right in Figure 13A, the results relate to stimulation with CT812C, CT387, CT869, CT166 and CT175. From left to right in Figure 13B, the results relate to stimulation with MOMP, CT163, CT812, CT812C, CT166, CT869, CT163, CT812, CT214, CT387, CT721, CT127 and CT175. The frequency on 10^5 CD4 T cells is shown on the Y axis.

Figure 14 is a histogram showing *C. muridarum* IFUs recovered from infected lungs of immunised mice (Day 12 post I.N. challenge with 10^3 IFUs). The immunisation group is shown on the X axis: the left hand bar relates to mice immunised with LTK63+CpG; the right hand bar relates to mice immunised with TC0197+TC0261+TC0666+LTK63+CpG. Mean IFU/lung (Log10) is shown on the Y axis.

Figure 15 is a histogram showing *C. muridarum* IFUs recovered from infected lungs of immunised mice (Day 12 post I.N. challenge with 10^3 IFUs). The immunisation group is shown on the X axis: from left to right, the results relate to mice immunised with i) LTK63+CpG, ii) TC0261+LTK63+CpG, iii) TC0197+LTK63+CpG, and iv) TC0666+LTK63+CpG. Mean IFU/lung (Log10) is shown on the Y axis.

25 MODES FOR CARRYING OUT THE INVENTION

Example 1: Induction of population of CD4+ T cells to produce IFN γ

20 antigens have been found which induce a population of CD4+ T cells to produce IFN γ (see Figure 1). 17 of these are newly discovered (CT016, CT043, CT114, CT153, CT255, CT279, CT341, CT372, CT480, CT600, CT601, CT711, CT716, CT733, CT734, CT745, CT823), while three antigens (CT681-MOMP, CT396-Hsp60 and CT587-Enolase) have already been described as targets of CD4+ T cells (Goodall JC et al. 2001; Hassell AB et al. 1993). Significantly, some antigens were able to induce a frequency of antigen-specific CD4+ responding T cells at least comparable to what observed with the positive control antigen MOMP.

The 17 new antigens are as follows:

Antigen	Annotation	Gene name
CT016	Hypothetical protein	
CT043	Hypothetical protein	
CT114	Hypothetical protein	
CT153	Hypothetical protein	
CT255	Hypothetical protein	
CT279	Na(+)-translocating NADH-quinone reductase subunit C	nqr3
CT341	Heat shock protein J (Hsp-J)	dnaJ
CT372	Hypothetical protein	
CT480	Oligopeptide Binding Lipoprotein	oppA_4
CT600		
CT601	Invasin repeat family phosphatase	papQ
CT711	Hypothetical protein	
CT716	Hypothetical protein	
CT733	Hypothetical protein	
CT734	Hypothetical protein	
CT745	protoporphyrinogen oxidase	hemG
CT823	DO serine protease	htrA

Of these 17 new antigens, CT341 may be the least suitable for use in immunization because it is a heat shock protein.

5 **Example 2: Characterization of the antigen-specificity of protective *Chlamydia* specific CD4+ Th1 cell lines**

The relevance of the newly discovered antigens for protective immunity to *Chlamydia* was further corroborated by showing that they were recognized by T cells belonging to a *Chlamydia*-specific CD4+/IFN γ + cell line, conferring protection when adoptively transferred to naïve recipient mice. To this aim we have derived a short-term CD4+ T cell line, produced against the extracellular EB form of *C. muridarum* that showed a high capacity to protect adoptively transferred naïve mice from *C. muridarum* challenge. The protective CD4+ cell line, which had undergone only a few cycles of expansion, maintained a polyclonal cell population with broad specificity that should correlate more closely to the *in vivo* protective response than long-term lines or clones. The polyclonal cell line was analysed for its antigen recognition profile versus the *C. muridarum* antigens, homologous to the *C. trachomatis* CD4-Th1 inducing proteins. The dissection of the antigen specificity of the protective CD4+ T cell polyclonal population demonstrated that the *Chlamydia* CD4+/IFN γ + inducing-antigens identified during an infection are also targets of CD4+ T cells that play a part in the rapid clearance of the bacterium in a protective response to the infection, in the absence of antibodies.

Chlamydia T cell lines were derived from Balb/c infected mice and their protective activity was verified in naïve mice against *C. muridarum* challenge. Subsequently, the antigen recognition profile of the *C. muridarum* CD4+ T cell line was characterized to define the possible contribution of each *C. muridarum* antigen in inducing protective CD4+ T cells. For the preparation of Chlamydia –

5 specific CD4+ T cells, splenic CD4+ T lymphocytes were purified from donor Balb/c mice that had previously been infected intranasally with 10^3 viable Elementary Bodies (EBs) of *C. muridarum*. An EB-responding CD4+ T cell line was derived (referred as EB-CD4+ cell line) and expanded *in vitro* with a short term stimulation with heat inactivated EBs. The line showed the capacity to respond to *C. muridarum* EBs by producing IFN γ with a high frequency (data not shown). To determine the

10 efficacy of the EB-CD4+ cell line in resolving an infection, 10^7 CD4+ T cells were adoptively transferred into 4 Balb/c recipient naïve mice. Mice were challenged intranasally 24 hours after i.v. infusion of CD4+ T cells with 10^3 IFUs of *C. muridarum*. The protective effect of adoptive immunization was evaluated by quantitating the number of IFUs recovered from lungs taken 10 days after Chlamydia challenge. As shown in Figure 2a, naïve mice adoptively transferred with EB-CM

15 CD4+ T cells shed 3 Log $_{10}$ fewer IFUs in the lungs 10 days after intranasal challenge with 10^3 IFUs of *C. muridarum*, as compared to either non treated mice (p value: 0.008) or mice receiving an unrelated CD4+ T cell line. Similarly, splenic CD4+ T cells isolated from mice that resolved an intravaginal primary infection with 10^5 IFUs of *C. trachomatis* conferred significant IFU reduction in adoptively transferred mice (data not shown).

20 To characterize the antigen recognition profile of the *C. muridarum* CD4+ T cells, most of the *C. muridarum* proteins, homolog of the proteins identified as CD4+ Th1 inducers during *C. trachomatis* infection (Figure 1), were obtained in recombinant form and tested for their ability to stimulate the protective EB-CD4+ T cell line to produce IFN γ . In this analysis we excluded both the proteins which after purification did not reach the purity /endotoxin level required for the cytokine stimulation

25 assay, or those which, due to their homology with human bacterial proteins were not suitable for developing a vaccine (e.g. heat shock proteins, enolase). The protective EB-CD4+ T cell line was stimulated *in vitro* with a panel of 19 *C. muridarum* recombinant proteins, including MOMP. Fourteen of them were homologs of *C. trachomatis* CD4+ Th1 inducing antigens identified in the primary screening in infected mice, and 5 were negative controls. As shown in Figure 2b, all the 14

30 CD4+-inducing antigens tested were found also to be targets of the protective EB-CM CD4+ T cell line, and able to induce IFN γ production in a percentage of CD4+ T cells at least 3 times higher than the frequency of negative control antigens. Therefore the pattern of T cell antigens recognized by the protective Chlamydia EB-CM T cell line is comparable to the recognition profile of T cells identified in the *C. trachomatis* infected mice.

Example 3: CT733 and CT153 specific CD4⁺ Th1 response in BALB/c mice after a primary *C. trachomatis* infection.

Splenocytes of primary infected BALB/c mice and non infected controls were collected 10 days after infection and stimulated with LPS-free recombinant antigens CT733 and CT153 (20mg/ml). After 4
5 hours of stimulation, 5mg/ml of Brefeldin A were added to the cells for the following 12 hrs to block cytokine secretion. Afterwards, cells were fixed, permeabilized and stained. Intracellular IFN γ and IL-5 expression were analyzed versus CD4 surface expression of the gated viable cells and assessed by flow cytometry.

The histogram in Figure 3 shows the number of CD4⁺ T cells per 10⁵ CD4⁺ T splenocytes of
10 primary infected (dark gery bars) and non-infected (light grey bars) mice that produce IFN γ upon specific stimulation with the *C. trachomatis* recombinant antigens CT153 and CT733. The data were confirmed in several further experiments using the same protocol.

The results indicate that CT733 and CT153 are able to induce significant frequencies of specific CD4⁺/IFN γ ⁺ cells in splenocytes from Balb/c mice that were infected intravaginally with *C.*
15 *trachomatis*, suggesting a potential role as antigen candidates for these proteins.

Example 4: Protective activity of single antigens TC0106 and TC0431 against *C. muridarum* challenge.

CT733 and CT153 were tested in a mouse model of chlamydial infection to evaluate their protective properties. This was done by adopting the mouse model of lung infection with the species
20 *Chlamydia muridarum*.

The *C. muridarum* proteins TC0106 and TC0431, homologous to CT733 and CT153, respectively, were cloned and purified, and used for the mouse model.

Groups of BALB/c mice were immunized with either TC0106 or TC0431 recombinant antigens formulated with LTK63+CpG adjuvant (3 doses of 15 ug protein, at 2 week interval, given
25 intramuscularly). As negative control, mice were immunized with the adjuvant only. Four weeks after the last immunization animals were infected intranasally with 10³ IFU of infectious *C. muridarum*. After 10 days, the protective activity conferred by the two antigens was measured by counting the infectious IFU in the lung of challenge animals.

As shown in Figure 4, each of the two antigens (middle and right hand columns of the histogram)
30 was able to reduce significantly the number of IFU/lung in challenged mice as compared to adjuvant immunized mice (left hand column of the histogram), indicating that both TC0106 and TC0431 (and therefore CT733 and CT153) confer protective immunity to Chlamydia infection

Example 5: Protective activity of the combination of TC0106+TC0431 against C. muridarum challenge.

Groups of BALB/c mice (10 to 15 mice) were immunized with the combination of TC0106+TC0431 recombinant antigens formulated with LTK63+CpG adjuvant (3 doses of 10 ug of each protein at 2 week-interval, given intramuscularly). As negative control, mice were immunized with the adjuvant only. Four weeks after the last immunization, animals were infected intranasally with 10^3 IFU of infectious *C. muridarum*. After 10 days, the protective activity conferred by the two antigens was measured by counting the infectious IFU in the lung of challenge animals. As positive control, a group of mice receiving a primary and a secondary *C. muridarum* infection was also included (left column in the histogram of Figure 5).

As shown in Figure 5, the antigen combination (right hand column of histogram) was able to significantly reduce the number of IFU/lung in challenged mice as compared to adjuvant immunized mice (middle column of histogram).

Thus, immunization with the CT733 and CT153, either alone or in combination, was able to significantly reduce the bacterial load in the lungs of challenged mice (see Figures 4 and 5).

Example 6: Elicitation of CD4+ Th1 cells in BALB/c mice after immunization with TC0431 and TC0106 recombinant antigens, alone or in combination.

Groups of BALB/c mice (10 to 15 mice) were immunized with the recombinant antigens TC0431 and TC0106 as single antigens or in combination (i.m., 10-15 micrograms/dose, 3 doses at 2 week-intervals) using LTK63+CpG adjuvant. Ten days after the third immunization dose, splenocytes were collected and stimulated with LPS-free recombinant antigens (20mg/ml). As negative control, splenocytes of adjuvant immunized mice were included. After 4 hours of stimulation, 5mg/ml of Brefeldin A was added to the cells for the following 12 hrs to block cytokine secretion. Afterwards, cells were fixed, permeabilized and stained. The intracellular IFN γ was analyzed versus CD4 surface expression of the gated viable cells and assessed by flow cytometry. The histogram in Figure 6 shows the number of CD4+ T cells per 10^5 CD4+ T splenocytes that produce IFN γ upon specific stimulation with the recombinant antigens in mice immunized with TC0106, TC0431, the combination of TC0106+TC0431 and adjuvant immunized mice.

The results indicate that immunization with these antigens elicits a high frequency of CD4+ Th1 cells.

Example 7: Evaluation of the protective effect of the chlamydial antigen(s) against C. muridarum challenge.

The protective effect of combinations of two antigens selected from *C. trachomatis* CT279, CT601, CT372, CT443, CT733, CT153, CT456 and CT381 was tested in the *C. muridarum* mouse model using their *C. muridarum* homologues TC0551 (CT279), TC0651 (CT372), TC0727 (CT443),

TC0890 (CT601), TC0106 (CT733), TC0431 (CT153), TC0660 (CT381) and TC0741 (CT456). The protective effect of CT733 and CT153 individually was also tested.

BALB/c mice were immunized three times intramuscularly with a combination of two antigens or single antigens with LTK63+CpG as adjuvant. Twenty-four days post last immunization mice were challenged intranasally with 10^3 IFU *C. muridarum*. After 10 days, lungs were collected, homogenized and the number of viable chlamydiae (IFU/lung) was measured. The data in Figure 7 shows the mean IFU/lung counts in antigen-immunized mice and adjuvant-immunized control. From left to right, the lanes relate to (a) adjuvant only; (b) TC0551+TC0890 (CT279+CT601); (c) TC0651+TC0727 (CT372+CT443); (d) TC0106+TC0431 (CT733+CT153); (e) TC0660+TC0741 (CT456+CT381); (f) TC0106 (CT733); (g) TC0431 (CT153). For each antigen formulation, the numbers of infected mice out of the total immunized are reported in the form "Inf X/Y", wherein X is the number of infected mice and Y is the total number of mice challenged. The statistical significance of immunizing antigen/s (P), was determined by Student t-test.

Four combinations of two antigens have been identified as capable of conferring protection against *C. muridarum* intranasal challenge. For three of them (TC0431+TC0106; TC0727+TC0651; TC0551+TC0890; homologs of CT733+CT153; CT443+CT372; CT279+CT601) protection has been confirmed in a high number of mice using LTK63+CpG adjuvant (Figure 7). Immunization experiments with TC0431 and TC0106 (CT153 and CT733) as single antigens indicate that the two antigens are both immunogenic individually and that either of the two antigens contributes to protection of the CT153+CT733 combination (Figure 7). A fourth antigen combination has been recently identified (TC660+TC0741; homologs of CT456 and CT381) showing protection in an immunization experiment (15 mice) (Figure 7).

The experiments were repeated where the protocol differed from that described above in that the mice were challenged intranasally with 10^3 IFUs of *C. muridarum* three weeks after the last immunization. Since differences in the duration of infections in the animals may occur, the presence of infectious *Chlamydiae* in the lungs was determined in each mouse at days 10 and 12 after challenge. Immunization experiments were repeated at least three times so as to generate data from a statistically significant number of mice. Figure 8 reports the mean number of infectious chlamydiae recovered from lungs of mice immunized with each antigen formulation, in which data obtained at days 10 and 12 were averaged. As shown in Figure 8, two of the four combinations tested in the mouse model, namely TC0551 (CT279 homolog, 82.6 % identity) + TC0890 (CT0601 homolog, 87.6% identity) and TC0106 (CT733 homolog, 84.8% identity) +TC0431 (CT153 homolog, 64.6% identity), showed a statistically significant protective effect in the immunized groups with an IFU reduction of more than 1 Log as compared to the adjuvant-injected mice ($P < 0.001$). Moreover, 20-25% of the animals immunized with either of the two combinations resolved completely the infection by days 10-12, as compared to 9% of the adjuvant group.

Example 8: Evaluation of the protective activity of the combination TC0551+TC0890+TC0106+TC0431 against challenge with *C. muridarum*.

On the basis of the result discussed in the preceding Example, groups of mice were immunized with a combination of four antigens TC0551+TC0890+TC0106+TC0431 using the same immunization regimen as in the Example above. As shown in Figure 8, the 4-antigen combination appeared to have an additive protective effect over the 2-antigen combinations, showing 2.2 Logs reduction of bacterial shedding in the lung (P:0.0003). Moreover, 39% of animals totally resolved the infection, indicating a higher efficacy of this antigen combination in accelerating the bacterial clearance.

The remarkable reduction observed in the number of viable Chlamydiae recovered from the lungs of immunized mice is the first demonstration of a high level of protection induced by systemic immunization with recombinant Chlamydia proteins. It has also to be pointed out that, since denatured forms of the recombinant antigens were used, further optimization of antigen conformation could maximize their protective activity.

Preliminary data aimed at defining whether any of the 4 recombinant antigens were protective when given as single antigens, indicated that a lower level of IFU reduction was observed (less than 1 log) was obtained with any of them (data not shown). This is in agreement with the notion that, in general, combinatorial vaccination approaches are more effective in conferring protective immunity against a given pathogen than single vaccine approaches, since elicited immune responses target different aspects of the bacterial developmental cycle.

Example 9: Evaluation of the protective activity of the combination TC0551+TC0651+TC0727+TC0890 against intraovarian bursa challenge with *C. muridarum*.

The protective effect of the combination TC0551+TC0651+TC0727+TC0890 (homologs of *C. trachomatis* CT279+CT372+CT443+CT601) was tested in the mouse model of ovarian bursa challenge with *C. muridarum* using the Montanide+CpG adjuvant. This model has previously been described to assess the protective activity of native MOMP (nMOMP), the chlamydial major outer membrane protein (Pal S et al, Infect Immun., 73:8153, 2005). In this model, the protective activity of the antigens is assessed against progression of infection by counting the chlamydia shedding in vaginal swabs.

BALB/c mice were immunized three times intranasally with a combination of the four antigens or with MOMP, with LTK63+CpG as adjuvant. As negative control, a group of mice immunized with ovalbumin was also included. Four weeks after the last immunization, mice received a *C. muridarum* challenge in the ovarian bursa and chlamydial shedding was measured by counting the IFU in the vaginal swabs of infected animals.

The results shown in Figure 9 represent the number of IFU/vaginal swab at two weeks post challenge. As shown in Figure 9, mice receiving the combination of all four antigens show a reduced bacterial shedding as compared to the negative control group (Ovalbumin). Thus, the combination

reduced the progression of infection. Interestingly, the protection level obtained with the combination does not differ significantly from that obtained with nMOMP, which is the most protective antigen that has been identified so far. Thus, this combination of four antigens is a particularly immunogenic combination.

5 ***Example 10: Antigen-specific cytokine profiles of protective CD4+ T cells***

Given the importance of the CD4-Th1 response in mediating protection from Chlamydia infection, the type of immune response induced by vaccination with two antigen combinations that elicited protection in mice was analysed (TC0551+TC0890 and TC0106+TC0431). In particular, we measured the simultaneous production from antigen-specific CD4+ T cells of IFN γ , TNF- α and IL-2, considering this as an indication of optimal effector functions of CD4+ T cells, possibly improving protection for vaccines aimed at targeting T-cell responses. The assessment of the cytokine profile induced in a single antigen specific CD4+ T cell by vaccination was performed by multiparametric flow cytometric analysis (Perfetto SP et al., Nat.Rev.Immunol. 4, 648-655, 2004) in immunized mice. Peripheral blood was collected 12 days after the last immunization with antigen combinations TC0551+TC0890 and TC0106+TC0431. PBMC were prepared and the frequency of CD3+, CD4+ antigen-specific IFN γ , IL-2 and TNF-producing cells was assessed by intracellular cytokine staining and flow cytometric determination. As shown in Figure 10B, vaccination with the antigen combination TC0551-TC0890 induced a high frequency of TC0551-responding CD4+ T cells producing IFN γ (93 TC0551 specific CD4+ T cells on 10⁵ CD4+ cells), while the response to TC0890 was very low, with a frequency of 16 IFN γ + responding T cells on 10⁵ CD4+ cells. The response to the antigen combination used for immunization showed an increased response compared to single antigens, with 132 IFN γ producing T cells on 10⁵ CD4+ cells. Furthermore, there was a predominant frequency of multifunctional CD4+ T cells, producing either IFN γ and TNF- α or IFN γ /TNF- α /IL-2 simultaneously. In the control group of mock immunized mice there was no cytokine secretion in response to any recombinant antigen used for stimulation, indicating the specificity of the response observed in the vaccinated mice. As far as the CD4+ response to the antigen combination TC0106-TC0431 is concerned (Figure 10C) both antigens, TC0106 and TC0431 induced a similar response with a frequency respectively of 120 and 98 IFN γ antigen-specific T cells on 10⁵ CD4+, while the antigen combination showed a frequency of 145 IFN γ + responding T cells on 10⁵ CD4+ cells. The further analysis of cytokines produced simultaneously with IFN γ showed that about 50% of IFN γ + cells produced also TNF- α and IL-2, while about 30% of them produced TNF- α . Overall these data underline that the Th1 cytokines produced by antigen-specific CD4+ T cells induced by vaccination showed a functional difference that could reflect differences in the capacity to clear the infection.

Example: 11. Expression analysis of CD4+ inducing Chlamydia antigens.

We then investigated the expression of CT279 (subunit C of Na(+)-translocating NADH-quinone reductase), CT601 (Invasin repeat family phosphatase), CT733 (-Hypothetical protein) and CT153 (MAC-Perforin Protein) by immunoblot analysis both in Ct-EBs and within *C. trachomatis* infected HeLa cells, using their specific mouse immune antisera (Figure 11A). Total protein lysates of renografin-purified EBs (corresponding to approximately 10⁷ EBs per lane) showed that each tested antiserum was able to react with a protein band of the expected molecular weight in both EB samples, showing in general a higher reactivity against CM EBs. For analysis of antigen expression in Chlamydia-infected cells, total protein extracts were prepared from Hela 229 cells at different time points after infection (24 – 48 - 72 h) and tested by immunoblot.

The amount of Chlamydial proteins loaded on the gel was normalized on the basis of MOMP expression as described. As shown in Figure 11B, the four antigens appeared to be expressed at different phases of the Chlamydia development.

Finally, we also investigated antigen cellular localization within infected HeLa cells by confocal microscopy in infected Hela cells at 6, 24, 48 and 72 h post infection. As shown in Figure 11B, expression of all antigens was clearly detected within the inclusions at 24h post infection and was still visible at 72h. Interestingly, CT153 staining appeared to accumulate at the inclusion membrane while the other proteins were homogeneously distributed. Since CT153 encodes a MAC-Perforin protein, belonging to a protein family capable of disrupting the cell membrane, the ammassing of this protein at the inclusion membrane might anticipate its involvement in the Chlamydia exit from infected cells.

The analysis of the immune response after vaccination with the combinations has shown that all the recombinant antigens induced a robust humoral response, with the production of IgG2a antibody titers higher than IgG1, as expected for a Th1 driven immune response. Since the resolution of a Chlamydia infection requires a Th1 type of cellular immune response, the regulation of CD4+ Th1 effector and memory cells after vaccination has also been investigated. Differences in the type of cytokines produced by individual cells have important implications for their capacity to mediate effector functions, be sustained as memory T cells or both. CD4+ T cells that secrete only IFN γ have limited capacity to develop into memory T cells as compared with IL-2-IFN γ double positive cells (Hayashi N. et al. 2002). Therefore vaccines eliciting high frequency of single-positive IFN γ producing cells may be limited in their ability to provide long-lasting protection. Furthermore the majority of CD4+ T cells that produce IL-2, IFN γ and TNF are classified as effector memory cells, playing an essential role for mediating protection against intracellular pathogens (Darrah PA et al. 2007). We demonstrated that antigen-specific CD4+ T cells induced by immunization with the protective combinations were predominantly multifunctional, being differentiated to ensure a population of Th1 cells that included either effectors and memory cells. An appropriate balance of

Th1 lineage cells that can be maintained and those with immediate protective functions might be the successful formula for an effective vaccine.

Example 12: Combination of CT823+CT733+CT043+CT456

To evaluate the protective activity of antigens TC0106, TC0313, TC0210, TC0741 and their combination, groups of mice were immunized with the 4 antigens either as single or in a 4 antigen-combination, using the same immunization regimen described in Example 7. The protective activity of the single antigens was assessed by measuring the IFU/Lung at day 12 post infection. The protective activity of the 4-ag combination was measured at days 10, 12, 14 post infection, to evaluate the kinetics of the infection clearance. As shown in Figure 12, the single antigens conferred approximately 0.5-1log IFU reduction in the lung of infected animals.

The four antigens combination showed a highest protective property, indicating a synergic activity of the four antigens in conferring protection, eliciting approximately 4 logs reduction of bacterial shedding in the lung ($P < 0.0001$) at day 12 and showing the tendency to resolve the infection at day 12. Moreover a high number of mice (42 %) totally resolved the infection, indicating the efficacy of the antigen combination in accelerating the bacterial clearance.

Example 13: Evaluation of antigenicity of CT812, CT387, CT869, CT166 and CT175

Antigen specific CD4 Tg1 response in BALB/c mice after a primary C.trachomatis (CT infection

The antigen specific CD4 Th1 response in BALB/c mice after a primary *C. trachomatis* (CT) infection was evaluated. *C. trachomatis* antigens identified by the proteomic characterization of the membrane fraction of CT infected HeLa cells were tested for their capability to induce specific CD4+ Th1 response in mice that received an experimental CT infection. Splenocytes of primary infected BALB/c mice and non infected controls were collected 10 days after infection and stimulated with LPS-free recombinant antigens (20µg/ml). After 4 hours of stimulation, 5µg/ml of Brefeldin A was added to the cells for the following 12 hrs, to block cytokine secretion. Afterwards, cells were fixed, permeabilized and stained. The intracellular IFN-γ expression was analyzed versus CD4 surface expression of the gated viable cells, and assessed by flow cytometry. The histogram in Figure 13A and figure 13B show the number of CD4+ T cells that produce IFNγ, upon specific stimulation with CT recombinant antigens per 10⁵ CD4+ T splenocytes of primary infected (right hand bars) and not-infected (left hand bars) mice. Data are representative of 4 different experiments. As shown in Figure 13A, CT812C, CT387, CT869 and CT166 induced a significant frequency of CD4⁺-IFNγ⁺ cells in splenocytes of infected animals ($P_{val} < 0.05$). As shown in Figure 13B, CT812C (a C-terminal fragment of CT812) surprisingly induced a higher frequency of CD4⁺-IFNγ⁺ cells in splenocytes of infected animals than did the full length CT812 sequence.

Protective activity of the combination of TC0197+TC0261+TC0666 against C. muridarum challenge

The protective effect of the combination of the three *C. trachomatis* antigens CT387+CT812+CT869 was tested in the *C. muridarum* mouse model using their *C. muridarum* orthologues TC0666, TC0197 and TC0261, respectively. TC0197, TC0261 and TC0666 were cloned and purified for protection studies in the mouse model of intranasal infection with *C. muridarum*. Groups of BALB/c mice (16 mice per group) were immunized with the combination of the three recombinant antigens TC0197+TC0261+TC0666 formulated with LTK63+CpG adjuvant (3 doses of 10 µg of each protein, at 2 week-interval, given intramuscularly). As a negative control, mice were immunized with the adjuvant only. Four weeks after the last immunization, animals were infected intranasally with 10³ IFU of infectious *C. muridarum*. After 12 days, the protective activity conferred by the two antigens was measured by counting the infectious IFU in the lung of challenge animals. As shown in Figure 14, the antigen combination TC0197+TC0261+TC0666 was able to reduce significantly the number of IFU/lung in challenged mice as compared to adjuvant immunized mice (1.4 log IFU reduction with Pval <0.05). The finding that the combination of TC0197+TC0261+TC0666 is able to protect mice against *C. muridarum* challenge (Figure 14) provides evidence that the combinations CT812+CT869+CT387 and CT812C+CT869+CT387 from *C. trachomatis* are protective against infection by *C. trachomatis*.

Protective activity of TC0197, TC0261 and TC0666 as single antigens against C. muridarum challenge

The protective activity of TC0197, TC0261 and TC0666 as single antigens against *C. muridarum* challenge was assessed. 3 Groups of BALB/c mice (16 mice per group) were immunized with the three recombinant antigens individually formulated with LTK63+CpG adjuvant (3 doses of 20 ug of each protein, at 2 week-interval, given intramuscularly). As a negative control, mice were immunized with the adjuvant only. Four weeks after the last immunization, animals were infected intranasally (I.N.) with 10³ IFU of infectious *C. muridarum*. After 12 days, the protective activity conferred by the three single antigens was measured by counting the infectious IFU in the lung of challenge animals. As shown in Figure 15, none of the 3 antigens individually were able to reduce significantly the number of IFU/lung in challenged mice as compared to adjuvant immunized mice.

Thus, the combination of TC0197+TC0261+TC0666 is able to protect mice against *C. muridarum* challenge (Figure 14). In particular, Figure 14 shows protection in terms of reduction in the mean number of IFUs recovered from lungs of immunized mice versus adjuvant immunized controls [p=0.0024]. In contrast, the three antigens are not protective when administered individually (Figure 15).

Example 14: Materials and methods

The experimental protocols used in Examples 1, 2, 7 (repeated experiments), 8, 10 and 11 are described in further detail in this Example.

Bacterial strains, cultures and reagents

5 *Chlamydia muridarum* Nigg and *Chlamydia trachomatis* serovar D strain D/UW-3/CX were grown on confluent monolayers of LLCMK2 (ATCC CCL7) or HeLa 229 cells (ATCC CCL 2.1) in Earle minimal essential medium (EMEM) as described (Caldwell *et al.* (1981) *Infect Immun* 31: 1161-1176). Purification of *C. trachomatis* and *C. muridarum* EBs was carried out by Renografin density gradient centrifugation as described (Montigiani *et al.* (2002) *Infect Immun* 70: 368-379.). Bacteria
10 were aliquoted and stored at -70°C in sucrose-phosphate-glutamine buffer (SPG) until use. When indicated, EBs were heat inactivated at 56°C for 3 hours.

E. coli DH5 α or BL21 (DE3) was grown aerobically in Luria Broth (LB) medium (Difco) at 37°C. When appropriate, ampicillin (100 μ g/ml) and isopropyl-beta-D-galactopyranoside (IPTG, 0.5 mM) were added to the medium.

15 Unless specified, all chemicals were purchased from Sigma. Restriction enzymes and DNA modification enzymes were from New England Biolabs. Unless differently stated, all reagents and antibody for intracellular cytokine staining were from BD Biosciences Pharmingen. Confocal microscopy reagents were from Molecular Probes.

Gene cloning, protein expression and preparation of antisera

20 To produce *C. trachomatis* recombinant proteins and their *C. muridarum* homologs, genes were PCR-amplified from *C. trachomatis* and *C. muridarum* chromosomal DNA using specific primers annealing at the 5' and 3' ends of either gene. The genes were cloned into plasmid pET21b⁺ (Invitrogen) or pGEXKG (Amersham) in order to express them both as a C-terminal His-tag fusion and as a double fusion protein with an N-terminal Glutathione transferase-encoding sequence and a
25 C-terminal His-tag.

Cloning and purification of His- and GST fusions were performed as already described (Montigiani *et al.*, 2002). CT0681 and TC0052, encoding for *C. trachomatis* and *C. muridarum* MOMP respectively (Ct MOMP and Cm MOMP, respectively) were expressed as His fusions and purified from the insoluble protein fraction. With the exception of TC0313 and TC0210, all the *C. muridarum*
30 proteins used in this work were purified only from the insoluble protein fraction in a denatured form.

For T cell in vitro stimulation assays, LPS-free proteins were prepared by washing of column-immobilized proteins with buffer Tris-HCl 10mM, pH 8, containing 1% Triton X114 (35 ml) at 4°C. The amount of residual endotoxin was determined using a *Limulus* Amebocyte Lysate Analysis Kit (QCL-100, BioWhittaker, Walkerville, MD).

Mouse antisera were generated and treated as described (Montigiani *et al.*, 2002). Where specified, sera from mice immunized with 20 µg of *E. coli* contaminant proteins (IMAC-purified proteins from *E. coli* bacteria containing pET21b+ empty vector) were used as negative control. Western blot, ELISA and Flow cytometry of *C. trachomatis* EBs were performed as described (Finco *et al.* (2005) 5 *Vaccine* 23: 1178-1188.).

Screening of antigen specific CD4-Th1 response in splenocytes from infected mice

Groups of 6 week-old female BALB/c mice purchased from Charles River Laboratories (3 mice/group) received a subcutaneous hormonal treatment with 2.5 mg of Depo-provera (Medroxyprogesterone acetate) and after five days mice were inoculated intravaginally with 15 µl of 10 SPG buffer containing 10^6 of *C. trachomatis* IFU. The level of infection was analyzed 7 days post-challenge, by collecting vaginal swabs and counting chlamydial inclusions 48h later stained with FITC-conjugated anti Chlamydia antibody (Merifluor) using a UV microscope.

The swabs were collected in 400 µl of SPG and were inoculated on LLCMK2 cell monolayers seeded on 96w flat bottom plates. After 48 hours incubation the number of infectious chlamydiae 15 was determined by counting chlamydial inclusions.

Ten days post challenge mice were sacrificed and their spleens were taken. Splenocytes were prepared by homogenization through a nylon filter (BD) and the erythrocytes were removed by hypotonic lysis in Ack lysis buffer (NH₄Cl 0,155 M, KHCO₃ 10 mM, Na₂EDTA 0,1 mM) for 3 minutes at RT, then the cells were plated in 96 wells plates at 2×10^6 cells per well and stimulated 20 with 20 µg/ml of endotoxin-free specific antigen or with 4 µg/ml of purified EBS in presence of 1 µg/ml anti-CD28 antibody (BD Biosciences Pharmingen) for 4 h at 37 °C. Brefeldin A (BFA; Sigma-Aldrich) was then added at a final concentration of 2.5 µg/ml and cells were incubated for an additional 16 h before intracellular cytokine staining. Cells were stained for viability with LIVE/DEAD® (Molecular Probes) dye according to the manufacturer's instructions. Cells were then 25 fixed and permeabilized using the Cytofix/Cytoperm kit (BD Biosciences Pharmingen) and stained with fluorochrome-labelled monoclonal antibodies for the detection of cells expressing CD3, CD4 on the surface and intracellular IFN γ and IL-4. Finally, cells were resuspended in PBS 1% BSA. All antibodies for intracellular cytokine staining were purchased from BD Pharmingen. Acquisition of the samples was performed using a BD Canto flow cytometer and data were analyzed using FlowJo 30 software (Tree Star Inc., Ashland, USA). The intracellular expression of IFN γ and IL-4 was analysed in CD4 expressing singlet cells, previously gated for, morphology, CD3 expression and viability. Cells were then harvested and stained for CD4 surface expression and IFN γ , or IL-4 intracellular production, to investigate whether the observed responses were of the Th1 (IFN γ) or Th2 (IL-4) type. As negative control, spleens from not infected mice were harvested and analyzed in parallel.

Preparation of CD4+ Th1 cell lines and of antigen presenting cells (APCs)

Splenocytes were prepared by homogenization from spleens from donor Balb/c mice that had previously been infected intranasally with 10^3 viable Elementary Bodies (EBs) of *Chlamydia muridarum* (*C. muridarum*) as described above. Following centrifugation at 1200 rpm and suspension in Macs Buffer (PBS PH 7,2 0,5% BSA and 2mM EDTA), the cells were incubated with CD4 (L3T4) microbeads (Milteny Biotec) for 15 minutes and then loaded on a LS columns. The CD4 cells bound to the magnet were recovered, washed and suspended in RPMI 1640 supplemented with 2,5% fetal bovine serum (Hyclone), antibiotics, L-Glutamine 2mM, Sodium Piruvate 1mM, MEM Not essential amino Acids, MEM Vitamins (Gibco) and Beta-mercaptoethanol 0.5 μ M. Then the cells were plated in 6 multiwell plates, 10^7 cells/wells. After the first stimulation, the purified CD4 were washed twice and then plated with APCs as described below.

Also a CD4+ cell line with *C. trachomatis* was obtained by spleens from donor Balb/c mice that had previously been infected intravaginally with 10^6 viable Elementary Bodies (EBs) of *Chlamydia trachomatis* and it was performed as described above for *Chlamydia muridarum*.

The CD4 cells were plated (6×10^6 /well) with APCs (2×10^7 /well) prepared by naive mice spleens. Splenocytes were prepared as described above, then were washed twice with the medium, gamma irradiated for 7 minutes washed again and suspended in medium.

Cultures were then incubated at 37°C in a humidified atmosphere containing 5%CO₂. After 24 h, Aldesleukin Proleukin (IL2) was added at a concentration of 20U/ml.

C. muridarum and C. trachomatis-mouse model of adoptive transfer

Groups of 6 week-old female BALB/c mice purchased from Charles River Laboratories (4 mice/group), were adoptively transferred by intravenous administration of 10^7 CD4+ T cells in 100 μ l of RPMI-1640 medium (Sigma). Mice were challenged intranasally 24 hours after with 10^3 IFUs of *C. muridarum* or 10^5 IFUs of *C. trachomatis*. The effect of adoptive immunization was evaluated by quantitating the number of IFUs recovered from lungs taken 10 days after *C. muridarum* challenge or 6 days after *C. trachomatis* challenge, as described above.

Characterization of the C. muridarum CD4+ T cell line

The same day of the adoptive transfer, an aliquot of purified CD4+ T cells were taken to assess the capability of *C. muridarum* antigens identified in the previous CD4+ Th1+ screening to stimulate them in vitro. 250000 cells/w were plated in 96 multiwell plates with 10^6 mouse splenocytes CD4 depleted as APC and stimulated with 20 μ g/ml of *C. muridarum* proteins, homologous to the *C. trachomatis* proteins identified as CD4+ Th1 inducers, in presence of 1 μ g/ml anti-CD28 antibody (BD Biosciences Pharmingen) for 3 h at 37 °C. Then BFA was added and intracellular staining was carried out as described for the splenocytes.

Mouse protection model

Groups of 6 week-old female BALB/c mice (10-15 mice/group), were immunized intramuscularly (i.m.) with 3 doses of the antigen combinations TC0551-TC890 (15 µg/dose) and TC0106-TC0431 (containing 10 µg of each protein/dose) at days 1, 15, and 28 formulated with 5µg of LTK63 (Ryan et al., 2000) + 10 µg of CpG (ODN 1826) adjuvant dissolved in 50 µl PBS. As negative control, groups of mice that received the adjuvant alone were included and treated in parallel.

Three weeks after the last immunization mice were inoculated intranasally (i.n.) with 40 µl of SPG buffer containing 10³ IFU of *C. muridarum*. The Chlamydia challenge dose given to each mouse was confirmed by culturing in triplicate serial dilutions of the inoculating dose on LLCMK2 cell monolayers seeded on 96 wells flat bottom plates. After 24 hours incubation the number of infectious chlamydiae was determined by counting chlamydial inclusions. In the time period between 10- and 12 days post challenge mice were sacrificed, lungs were isolated and their homogenates were used to assess chlamydia growth.

Analysis of antigen specific CD4-Th1 response in PBMC of mice

PBMC from mouse were isolated from up to 2 ml of heparinized blood, diluted 1/5 in HBSS (Hanks' Balanced Salt Solution) and separated by density gradient centrifugation over Lympholite-M (Cedarlane). 10⁶ PBMC were plated in duplicate in 96 multiwell plates with 10⁶ mouse splenocytes CD4 depleted as APC and stimulated and stained as described above for mouse splenocytes for 16 h. In this staining was analyzed the expression of IFNγ, TNFα and IL-2.

Confocal microscopy

To examine cellular localization of *C. trachomatis* proteins after infection, HeLa cells (20000) were plated on onto glass coverslides (Ø 13 mm) and after 24 hours were infected with CT EBs in 1:1 ratio as described above. At 6, 24, 48 and 72 hours post infection the cells were fixed in 2% paraformaldehyde in PBS buffer for 20 minutes at room temperature. After 2 washes with PBS the cells were permeabilized with a solution of 1%/saponin-0.1% Triton in PBS for 20 minutes.

After washing twice and blocking with PBS containing 1% BSA (PBS-BSA), the cell samples were subjected to antibody and chemical staining. The samples were incubated for 1h at RT (standard dilution 1:5000 in PBS-BSA) with polyclonal antisera obtained from mice immunized with TC601, TC279, TC733 and TC153, previously pre-adsorbed overnight at 4°C onto nitrocellulose strips containing *E. coli* BL21 cell total proteins. Goat anti-mouse Alexa Fluor (Molecular Probes) conjugated antibodies (excitation at 488) were used to visualize the localization of each antigen. Propidium Iodide and Phalloidin conjugated with Alexa Fluor dye A620 (Molecular Probes) were used to visualize respectively DNA and actin.

After extensive washes in PBS, cells were mounted with Anti-Fade reagent (Molecular Probes) and observed under a laser scanning confocal microscope (Bio-Rad) with 100X oil immersion objective lens.

It will be understood that the invention has been described by way of example only and modifications may be made whilst remaining within the scope and spirit of the invention.

TABLE 2

<i>C. pneumoniae</i> accession number & annotation	<i>C. trachomatis</i> accession number & annotation	CT No.
	Hypothetical protein (AAC67968)	CT372
	omcB (AAC68042)	CT443
	Hypothetical protein (AAC67634)	CT043
	Hypothetical protein (AAC67744)	CT153
	Nqr3 (AAC67872)	CT279
	papQ (AAC68203)	CT601
	Hypothetical protein (AAC68306)	CT711
	Hypothetical protein (AAC67705)	CT114
	oppA_4 (AAC68080)	CT480
	Hypothetical protein (AAC68056)	CT456
	ArtJ (AAC67977)	CT381
	IcrE (AAC67680)	CT089
	Hypothetical protein (AAC68329)	CT734
	Hypothetical protein (AAC67606)	CT016
gi 4376729 gb AAD18590.1 Polymorphic Outer Membrane Protein G Family	gi 3329346 gb AAC68469.1 Putative Outer Membrane Protein G	
gi 4376729 gb AAD18590.1 Polymorphic Outer Membrane Protein G Family	gi 3329346 gb AAC68469.1 Putative Outer Membrane Protein G	
gi 4376731 gb AAD18591.1 Polymorphic Outer Membrane Protein G/I Family	gi 3329346 gb AAC68469.1 Putative Outer Membrane Protein G	
gi 4376731 gb AAD18591.1 Polymorphic Outer Membrane Protein G/I Family	gi 3329350 gb AAC68472.1 Putative Outer Membrane Protein I	
gi 4376731 gb AAD18591.1 Polymorphic Outer Membrane Protein G/I Family	gi 3329346 gb AAC68469.1 Putative Outer Membrane Protein G	
gi 4376733 gb AAD18593.1 Polymorphic Outer Membrane Protein G Family	gi 3328840 gb AAC68009.1 Putative outer membrane protein A	
gi 4376731 gb AAD18591.1 Polymorphic Outer Membrane Protein G/I Family	gi 3329346 gb AAC68469.1 Putative Outer Membrane Protein G	
gi 4376754 gb AAD18611.1 Polymorphic Outer Membrane Protein (Frame-shift with C	gi 3329344 gb AAC68467.1 Putative Outer Membrane Protein E	
gi 4376260 gb AAD18163.1 Polymorphic Outer Membrane Protein G Family	gi 3329346 gb AAC68469.1 Putative Outer Membrane Protein G	
gi 4376262 gb AAD18165.1 hypothetical protein	gi 3328765 gb AAC67940.1 hypothetical protein	
gi 4376269 gb AAD18171.1 hypothetical protein	gi 3328825 gb AAC67995.1 hypothetical protein	
gi 4376270 gb AAD18172.1 Polymorphic Outer Membrane Protein G Family	gi 3329350 gb AAC68472.1 Putative Outer Membrane Protein I	
gi 4376272 gb AAD18173.1 Predicted OMP {leader peptide: outer membrane}	gi 3328772 gb AAC67946.1 hypothetical protein	CT351
gi 4376273 gb AAD18174.1 Predicted OMP {leader peptide}	gi 3328771 gb AAC67945.1 hypothetical protein	CT350
gi 4376296 gb AAD18195.1 hypothetical protein	gi 3328520 gb AAC67712.1 Ribulose-P Epimerase	
gi 4376362 gb AAD18254.1 YbbP family hypothetical protein	gi 3328401 gb AAC67602.1 hypothetical protein	
gi 4376372 gb AAD18263.1 Signal Peptidase I	gi 3328410 gb AAC67610.1 Signal Peptidase I	
gi 4376397 gb AAD18286.1 CHLPS hypothetical protein	gi 3328506 gb AAC67700.1 CHLPS hypothetical protein	
gi 4376402 gb AAD18290.1 ACR family	gi 3328505 gb AAC67699.1 ACR family	
gi 4376419 gb AAD18305.1 CT149 hypothetical protein	gi 3328551 gb AAC67740.1 possible hydrolase	
gi 4376446 gb AAD18330.1 hypothetical protein	gi 3329261 gb AAC68390.1 hypothetical protein	
gi 4376466 gb AAD18348.1 Oligopeptide Binding Protein	gi 3328604 gb AAC67790.1 Oligopeptide Binding Protein	CT198
gi 4376467 gb AAD18349.1 Oligopeptide Binding Protein	gi 3328604 gb AAC67790.1 Oligopeptide Binding Protein	
gi 4376468 gb AAD18350.1 Oligopeptide Binding Protein	gi 3328539 gb AAC67730.1 Oligopeptide Binding Protein	

gi 4376469 gb AAD18351.1 Oligopeptide Binding Protein	gi 3328579 gb AAC67766.1 Oligopeptide binding protein permease	
gi 4376520 gb AAD18398.1 Polysaccharide Hydrolase-Invasin Repeat Family	gi 3328526 gb AAC67718.1 predicted polysaccharide hydrolase-invasin repeat family	
gi 4376567 gb AAD18441.1 Inclusion Membrane Protein C	gi 3328642 gb AAC67825.1 Inclusion Membrane Protein C	
gi 4376576 gb AAD18449.1 Omp85 Analog	gi 3328651 gb AAC67834.1 Omp85 Analog	CT241
gi 4376577 gb AAD18450.1 (OmpH-Like Outer Membrane Protein)	gi 3328652 gb AAC67835.1 (OmpH-Like Outer Membrane Protein)	CT242
gi 4376601 gb AAD18472.1 Low Calcium Response D	gi 3328486 gb AAC67681.1 Low Calcium Response D	
gi 4376602 gb AAD18473.1 Low Calcium Response E	gi 3328485 gb AAC67680.1 Low Calcium Response E	CT089
gi 4376607 gb AAD18478.1 Phospholipase D Superfamily	gi 3328479 gb AAC67675.1 Phospholipase D Superfamily {leader (33) peptide}	
gi 4376615 gb AAD18485.1 YojL hypothetical protein	gi 3328472 gb AAC67668.1 hypothetical protein	CT077
gi 4376624 gb AAD18493.1 Solute Protein Binding Family	gi 3328461 gb AAC67658.1 Solute Protein Binding Family	
gi 4376639 gb AAD18507.1 Flagellar Secretion Protein	gi 3328453 gb AAC67651.1 Flagellar Secretion Protein	
gi 4376664 gb AAD18529.1 Leucyl Aminopeptidase A	gi 3328437 gb AAC67636.1 Leucyl Aminopeptidase A	CT045
gi 4376672 gb AAD18537.1 CBS Domain protein (Hemolysin Homolog)	gi 3328667 gb AAC67849.1 Hypothetical protein containing CBS domains	
gi 4376679 gb AAD18543.1 CT253 hypothetical protein	gi 3328664 gb AAC67846.1 hypothetical protein	
gi 4376696 gb AAD18559.1 CT266 hypothetical protein	gi 3328678 gb AAC67859.1 hypothetical protein	CT266
gi 4376717 gb AAD18579.1 Phospholipase D superfamily	gi 3328698 gb AAC67877.1 Phospholipase D superfamily	
gi 4376727 gb AAD18588.1 Polymorphic Outer Membrane Protein G/I Family	gi 3329346 gb AAC68469.1 Putative Outer Membrane Protein G	
gi 4376728 gb AAD18589.1 Polymorphic Outer Membrane Protein G Family	gi 3329346 gb AAC68469.1 Putative Outer Membrane Protein G	
gi 4376729 gb AAD18590.1 Polymorphic Outer Membrane Protein G Family	gi 3329350 gb AAC68472.1 Putative Outer Membrane Protein I	
gi 4376731 gb AAD18591.1 Polymorphic Outer Membrane Protein G/I Family	gi 3329350 gb AAC68472.1 Putative Outer Membrane Protein I	
gi 4376733 gb AAD18593.1 Polymorphic Outer Membrane Protein G Family	gi 3328840 gb AAC68009.1 Putative outer membrane protein A	
gi 4376735 gb AAD18594.1 Polymorphic Outer Membrane Protein (truncated) A/I Fam	gi 3328840 gb AAC68009.1 Putative outer membrane protein A	
gi 4376736 gb AAD18595.1 Polymorphic Outer Membrane Protein G Family	gi 3329346 gb AAC68469.1 Putative Outer Membrane Protein G	
gi 4376737 gb AAD18596.1 Polymorphic Outer Membrane Protein H Family	gi 3329347 gb AAC68470.1 Putative Outer Membrane Protein H	
gi 4376751 gb AAD18608.1 Polymorphic Outer Membrane Protein E Family	gi 3329344 gb AAC68467.1 Putative Outer Membrane Protein E	
gi 4376752 gb AAD18609.1 Polymorphic Outer Membrane Protein E Family	gi 3329344 gb AAC68467.1 Putative Outer Membrane Protein E	
gi 4376753 gb AAD18610.1 Polymorphic Outer Membrane Protein E/F Family	gi 3329344 gb AAC68467.1 Putative Outer Membrane Protein E	
gi 4376757 gb AAD18613.1 hypothetical protein	gi 3328701 gb AAC67880.1 PP-loop superfamily ATPase	
gi 4376767 gb AAD18622.1 Arginine Periplasmic Binding Protein	gi 3328806 gb AAC67977.1 Arginine Binding Protein	CT381
gi 4376790 gb AAD18643.1 Heat Shock Protein-70	gi 3328822 gb AAC67993.1 HSP-70	CT396
gi 4376802 gb AAD18654.1 CT427 hypothetical protein	gi 3328857 gb AAC68024.1 hypothetical protein	
gi 4376814 gb AAD18665.1 CT398 hypothetical protein	gi 3328825 gb AAC67995.1 hypothetical protein	CT398
gi 4376829 gb AAD18679.1 polymorphic membrane protein A Family	gi 3328840 gb AAC68009.1 Putative outer membrane protein A	
gi 4376830 gb AAD18680.1 polymorphic membrane protein B Family	gi 3328841 gb AAC68010.1 Putative outer membrane protein B	
gi 4376832 gb AAD18681.1 Solute binding protein	gi 3328844 gb AAC68012.1 Solute-binding protein	CT415
gi 4376834 gb AAD18683.1 (Metal Transport Protein)	gi 3328846 gb AAC68014.1 (Metal Transport Protein)	
gi 4376847 gb AAD18695.1 Tail-Specific Protease	gi 3328872 gb AAC68040.1 Tail-Specific Protease	
gi 4376848 gb AAD18696.1 15 kDa Cysteine-Rich Protein	gi 3328873 gb AAC68041.1 15kDa Cysteine-Rich Protein	
gi 4376849 gb AAD18697.1 60 kDa Cysteine-Rich OMP	gi 3328874 gb AAC68042.1 60kDa Cysteine-Rich OMP	CT443
gi 4376850 gb AAD18698.1 9 kDa-Cysteine-Rich Lipoprotein	gi 3328876 gb AAC68043.1 9kDa-Cysteine-Rich Lipoprotein	CT444

gi 4376878 gb AAD18723.1 2-Component Sensor	gi 3328901 gb AAC68067.1 2-component regulatory system-sensor histidine kinase	CT467
gi 4376879 gb AAD18724.1 similarity to CHLPS IncA	gi 3328451 gb AAC67649.1 hypothetical protein	
gi 4376884 gb AAD18729.1 CT471 hypothetical protein	gi 3328905 gb AAC68071.1 hypothetical protein	
gi 4376886 gb AAD18731.1 YidD family	gi 3328908 gb AAC68073.1 hypothetical protein	
gi 4376890 gb AAD18734.1 CT476 hypothetical protein	gi 3328911 gb AAC68076.1 hypothetical protein	
gi 4376892 gb AAD18736.1 Oligopeptide Permease	gi 3328913 gb AAC68078.1 Oligopeptide Permease	
gi 4376894 gb AAD18738.1 Oligopeptide Binding Lipoprotein	gi 3328915 gb AAC68080.1 oligopeptide Binding Lipoprotein	
gi 4376900 gb AAD18743.1 Glutamine Binding Protein	gi 3328922 gb AAC68086.1 Glutamine Binding Protein	
gi 4376909 gb AAD18752.1 Protease	gi 6578107 gb AAC68094.2 Protease	
gi 4376952 gb AAD18792.1 Apolipoprotein N-Acetyltransferase	gi 3328972 gb AAC68136.1 Apolipoprotein N-Acetyltransferase	
gi 4376960 gb AAD18800.1 FKBP-type peptidyl-prolyl cis-trans isomerise	gi 3328979 gb AAC68143.1 FKBP-type peptidyl-prolyl cis-trans isomerise	CT541
gi 4376968 gb AAD18807.1 CT547 hypothetical protein	gi 3328986 gb AAC68149.1 hypothetical protein	CT547
gi 4376969 gb AAD18808.1 CT548 hypothetical protein	gi 3328987 gb AAC68150.1 hypothetical protein	
gi 4376998 gb AAD18834.1 Major Outer Membrane Protein	gi 3329133 gb AAC68276.1 Major Outer Membrane Protein	CT681
gi 4377005 gb AAD18841.1 YopC/Gen Secretion Protein D	gi 3329125 gb AAC68269.1 probable Yop proteins translocation protein	
gi 4377015 gb AAD18851.1 FHA domain; (homology to adenylyate cyclase)	gi 3329115 gb AAC68259.1 (FHA domain; homology to adenylyate cyclase)	
gi 4377033 gb AAD18867.1 CHLPN 76 kDa Homolog_1 (CT622)	gi 3329069 gb AAC68226.1 CHLPN 76kDa Homolog	CT622
gi 4377034 gb AAD18868.1 CHLPN 76 kDa Homolog_2 (CT623)	gi 6578109 gb AAC68227.2 CHLPN 76kDa Homolog	CT623
gi 4377035 gb AAD18869.1 Integral Membrane Protein	gi 3329071 gb AAC68228.1 Integral Membrane Protein	
gi 4377072 gb AAD18902.1 CT648 hypothetical protein	gi 3329097 gb AAC68825.1 hypothetical protein	
gi 4377073 gb AAD18903.1 CT647 hypothetical protein	gi 3329096 gb AAC68824.1 hypothetical protein	CT647
gi 4377085 gb AAD18914.1 CT605 hypothetical protein	gi 3329050 gb AAC68208.1 hypothetical protein	
gi 4377090 gb AAD18919.1 Peptidoglycan-Associated Lipoprotein	gi 3329044 gb AAC68202.1 Peptidoglycan-Associated Lipoprotein	CT600
gi 4377091 gb AAD18920.1 macromolecule transporter	gi 3329043 gb AAC68201.1 component of a macromolecule transport system	
gi 4377092 gb AAD18921.1 CT598 hypothetical protein	gi 3329042 gb AAC68200.1 hypothetical protein	
gi 4377093 gb AAD18922.1 Biopolymer Transport Protein	gi 3329041 gb AAC68199.1 Biopolymer Transport Protein	CT597
gi 4377094 gb AAD18923.1 Macromolecule transporter	gi 3329040 gb AAC68198.1 polysaccharide transporter	
gi 4377101 gb AAD18929.1 CT590 hypothetical protein	gi 3329033 gb AAC68192.1 hypothetical protein	
gi 4377102 gb AAD18930.1 CT589 hypothetical protein	gi 3329032 gb AAC68191.1 hypothetical protein	CT589
gi 4377106 gb AAD18933.1 hypothetical protein	gi 3328796 gb AAC67968.1 hypothetical protein	
gi 4377111 gb AAD18938.1 Enolase	gi 3329030 gb AAC68189.1 Enolase	CT587
gi 4377127 gb AAD18953.1 General Secretion Protein D	gi 3329013 gb AAC68174.1 Gen. Secretion Protein D	
gi 4377130 gb AAD18956.1 predicted OMP {leader peptide}	gi 3329010 gb AAC68171.1 predicted OMP	CT569
gi 4377132 gb AAD18958.1 CT567 hypothetical protein	gi 3329008 gb AAC68169.1 hypothetical protein	CT567
gi 4377133 gb AAD18959.1 CT566 hypothetical protein	gi 3329007 gb AAC68168.1 hypothetical protein	
gi 4377140 gb AAD18965.1 Yop Translocation J	gi 3329000 gb AAC68161.1 Yop proteins translocation lipoprotein J	CT559
gi 4377170 gb AAD18992.1 Outer Membrane Protein B	gi 3329169 gb AAC68308.1 Outer Membrane Protein Analog	CT713
gi 4377177 gb AAD18998.1 Flagellar M-Ring Protein	gi 3329175 gb AAC68314.1 Flagellar M-Ring Protein	
gi 4377182 gb AAD19003.1 CT724 hypothetical protein	gi 3329181 gb AAC68319.1 hypothetical protein	
gi 4377184 gb AAD19005.1 Rod Shape Protein	gi 3329183 gb AAC68321.1 Rod Shape Protein	
gi 4377193 gb AAD19013.1 CT734 hypothetical protein	gi 3329192 gb AAC68329.1 hypothetical protein	
gi 4377206 gb AAD19025.1 CHLTR possible phosphoprotein	gi 3329204 gb AAC68339.1 CHLTR possible phosphoprotein	
gi 4377222 gb AAD19040.1 Muramidase (invasin repeat family)	gi 3329221 gb AAC68354.1 Muramidase (invasin repeat family)	CT759
gi 4377223 gb AAD19041.1 Cell Division Protein FtsW	gi 3329222 gb AAC68355.1 Cell Division Protein FtsW	
gi 4377224 gb AAD19042.1 Peptidoglycan Transferase	gi 3329223 gb AAC68356.1 Peptidoglycan Transferase	CT761
gi 4377225 gb AAD19043.1 Muramate-Ala Ligase & D-Ala-D-Ala Ligase	gi 3329224 gb AAC68357.1 UDP-N-acetylmuramate-alanine ligase	
gi 4377248 gb AAD19064.1 Thioredoxin Disulfide Isomerase	gi 3329244 gb AAC68375.1 Thioredoxin Disulfide Isomerase	

gi 4377261 gb AAD19076.1 CT788 hypothetical protein {leader peptide-periplasmi	gi 3329253 gb AAC68383.1 {leader (60) peptide-periplasmi	
gi 4377280 gb AAD19093.1 Insulinase family/Protease III	gi 3329273 gb AAC68402.1 Insulinase family/Protease III	
gi 4377287 gb AAD19099.1 Putative Outer Membrane Protein D Family	gi 3329279 gb AAC68408.1 Putative Outer Membrane Protein D	
gi 4377306 gb AAD19116.1 DO Serine Protease	gi 3329293 gb AAC68420.1 DO Serine Protease	CT823
gi 4377342 gb AAD19149.1 ABC transporter permease	gi 3329327 gb AAC68451.1 ABC transporter permease — pyrimidine biosynthesis protein	
gi 4377347 gb AAD19153.1 CT858 hypothetical protein	gi 6578118 gb AAC68456.2 predicted Protease containing IRBP and DHR domains	
gi 4377353 gb AAD19159.1 CT863 hypothetical protein	gi 3329337 gb AAC68461.1 hypothetical protein	
gi 4377367 gb AAD19171.1 Predicted OMP	gi 3328795 gb AAC67967.1 hypothetical protein	
gi 4377408 gb AAD19209.1 hypothetical protein	gi 3328795 gb AAC67967.1 hypothetical protein	
gi 4377409 gb AAD19210.1 Predicted Outer Membrane Protein (CT371)	gi 3328795 gb AAC67967.1 hypothetical protein	
gi 4376411 gb	gi 3328512 gb AAC67705.1 hypothetical protein	CT114
gi 4376508 gb	gi 3328585 gb AAC67772.1 hypothetical protein	CT181
gi 4376710 gb	gi 3328692 gb AAC67872.1 NADH (Ubiquinone) Oxidoreductase, Gamma	CT279
gi 4376777 gb	gi 3328815 gb AAC67986.1 hypothetical protein	CT389
gi 4376782 gb	gi 3328817 gb AAC67988.1 hypothetical protein	CT391
gi 4376863 gb	gi 3328887 gb AAC68054.1 Arginyl tRNA transferase	CT454
gi 4376866 gb	gi 3328889 gb AAC68056.1 hypothetical protein	CT456
gi 4376972 gb	gi 3328991 gb AAC68153.1 D-Ala-D-Ala Carboxypeptidase	CT551
gi 4377139 gb	gi 3329001 gb AAC68162.1 hypothetical protein	CT560
gi 4377154 gb	gi 3329154 gb AAC68295.1 hypothetical protein	CT700

SEQUENCE LISTING

SEQ ID NO: 1 - CT733 nucleotide sequence

ATGTTAATAAACTTTACCTTTTCGCAACTGTCTTTTGTTCCTTGTACACTGTCTAGTGTCCCTGTTTTCTCAGCACC
 TCAACCTCGCGGAACGCTTCCTAGCTCGACCACAAAAATTGGATCAGAAGTTTGGATTGAACAAAAAGTCCGCCAAT
 ATCCAGAGCTTTTATGGTTAGTAGAGCCGCTCTACGGGAGCCTCTTTAAAATCTCCTTCAGGAGCCATCTTTTCT
 CCAACATTATCCAAAAAAGGTCCTGCTTTGATATCGCAGTGCAGTTTGGATTCACTTACATTTATTAATCCA
 GGGTTCCCGCAAGCCTATGCTCAACTGATCCAACACAGACCAGCGAATCCCCTTAACATTTAAGCAATTCCCTG
 CATTGCATAAGCAATTAACCTCTATTTTTAAATTTCCCTAAGGAATTTTATGACTCTGTTAAAGTGTAGAGACAGCT
 ATCGTCTTACGTCACTTAGGCTGTTCAACTAAGGCTGTTGCTGCGTTTAAACCTTATTTCTCAGAAATGCAAAGAGA
 GGCTTTTTACACTAAGGCTCTGCATGTAACACACCTTCCAGAGCTAAGCCCATATTTGCTCGCTCTCTCCGG
 AGCAGAAAACTCTTCTTCTCCTTGAGAAAATTGGCGAATTACGATGAGTTACTCTCGCTGACGAACACCCCAAGT
 TTTGAGCTTTGTCTGCTGGGCGCTCGCAACGAGCTCTTTTAGCTCTGGACTTGTACCTCTATGCTTTGGATTCCCTG
 TGGAGAACAGGGGATGTCCTCTCAATTCCAACAACTTTCGCACCTCTACAGTCCATGTTGCAACAATACGCTACTG
 TAGAAGAGGCCCTTTCTCGTTATTTTACTTACCGAGCTAATCGATTAGGATTTGATGGCTCTTCTCGATCCGAGATG
 GCTTTAGTAAGAATGGCCACCTTGATGAACCTGCTCTCTCCGAAAGCTGCGATTTTAAACCACAAGCTTCAAAACCT
 TCCTACAGAAGAAGCGGATACTTTGATCAATAGTTTCTATACCAATAAGGGCGATTGCTTGGCTCTTTCTCTGCGAG
 GGTTGCTTACACTTGTATCCGAACCTGACGCGAAGCTCCCATGGCAATACCAATGCGAGAAGCTCGATCTCAGCAAAAT
 TATGCAACTACCCTATCGCTAGTAGTAAAGAGTCTGAAAGCGCACAAAGAAAATGCTAAACAAGCAAATTTCTTCTAA
 GGAAATGTTTTAGATTTCTCAGAACTGCAGCTTCTTGCACAGGATTGGATATCTTTCCGAGAATGTCGCTGTTT
 AAATTCACCTAAATGGAACCGTTAGTATCCATTTATAA

SEQ ID NO: 2 - CT733 protein sequence

MLINFTFRNCLLFLVTLSSVPVFSAPQPRGTLPSSTTKIGSEVWIEQKVRQYPELLWLVEPSSTGASLKSPSGAIF
 PTLFQKKVPAFDIAVRSLIHLHLIIQGSRQAYAQLIQLQTSSEPLTFKQFLALHKQLTLFLNSPKIFYDSVKVLETA
 IVLRHLGCKTKAVAAFKPYFSEMQRFAFYTKALHVLHTFPELSPSFLARLSPEQKTLFFSLRKLANYDELLSLTNTPS
 FQLLSAGRSQRALLALDLYLYALDSCGEQGMSSQFHTNFAPLQSMQRYATVEEAFSRYFTYRANRGLGFDGSSRSEM
 ALVRMATLMNLSPEAAAILTTSFKTLPTEEADTLINSFYTNKGDLSLALSLRGLPTLVSELTRTAHGNTNAEARSQRI
 YATTLVSLVVKSLKAHKEMLNKQILSKEIVLDFSETAASCQGLDIFSENVAVQIHLNGTVSIHL

SEQ ID NO:3 - CT153 nucleotide sequence

ATGACTAAGCCTTCTTTCTTATACGTTATTCAACCTTTTTCCGTATTTAATCCACGATTAGGACGTTTCTCTACAGA
 CTCAGATACTTATATCGAAGAAGAAAAACCGCTAGCATCGTTTCATTGAGAGTTTGGCCACTGGAGATCTTCGATATAC
 CTTCTTTTCATGGAAACCGGATTTCCAATAGCCCCATATTTTTATCTTTGGGAGACAACATAAAGACGGCGCTCTGTTT
 ACTATTTCTGAACCCAACTCTCAGCTTGGCGAGCCACTTGCCTGGTAGCCCCCTCTATACAAATGAAATCCGATGC
 GGAGCTCTTAGAAGAAATTAAGCAAGCGTTATTACGCACTCTCATGACGGTGTGAAATATCGCATCACAGAGAAT
 CTTCTCTCCAGAAAAAGAAAACCTCCTAAGGTTGCTAGTGCATGACGATATTGAAATGATTGATTGCGAATGCTGCT
 TTGGGTAGAGCTGTTGACATTGTCAAATTAGACCTTATTAATATTCTGAATACCGTAAGCGAAGAGAATATTCTAGA
 TTAATCTTTTTACAAGAAAAACGGCTCAGCTGAGCGCGGATGGTCTGTTTTGGTATTCCTCCAGGGACTAAGCTATTCC
 CTAACCTTCTTTGATGTAGAAATCAGTACCTCCATTTTTCGAAGAAACAACCTCAATTACTCGAAGTTTTCTGCA
 TCGGTTACTTTTGTGTAACAGACCTCGCGGCGACTATGCCTCTTCAAAGCCCTCCCATGGTAGAAAAATGGTCAAAA
 AGAAATTTGTGTCATTCAAAAACACTTATTCCCAAGCTACTCTCTAACTAGTCGATATTGTTAAACGATACAAAA
 GAGAGGCTAAGATCTTGATTAACAAGCTTTGCCCTTTGGAATGTTATGGCGACATCGGGCTAAAAGCCAAATCCTCAC
 GAGGGAAGCGTACGCTAGACTTACAAGGATTCACAGAATCGAAGTACAATTACCAGATTCAAGTAGGATCCCATAC
 GATTGCAGCTGTTAATCGATATGGATATTTCCAAGATTCAATCCAAATCAGAACAAAGCTTATGCAATTAGGAAAA
 TCAAATCAGGCTTTCAACGTAGCTTGGATGACTATCATATTTATCAAATGAAAGAAAAACAACTTTTCTTTTTCT
 CCGAAGCATCGCAGCCTCTCATCCACATCCCATTCCGAAGATTCTGATTTGGATCTTTCTGAAGCAGCCGCCCTTTT
 AGGAAGCTTACCTGCGAGTTTGTAAAAAAAAGCACTCAACATGCCAAGAATACCGTCAACATGTTCCACAGCCGCTC
 ATTCCTTATACACACTCAAAGAAGATGACAGCTCGAACCCCTCTGAAAAACGATTAGATAGTTGTTTCCGCAATTGG
 ATTGAAAAACAACTAAGCGCCAATTCTCCAGATTCCTGGTCAGCGTTTATTCAAAAATTCGGAACACACTATATTGC
 ATCAGCAACTTTTGGAGGGATAGGTTTCCAAGTGTCAAACCTATCTTTTGAACAGGTGGAGGATCTACATAGCAAAA
 AGCTCTCCTTAGAAAACCGCAGCAGCCAACCTCTATTAAGAGGTTCTGTATCCAGCAGCACAGAATCTGGATACTCC
 AGCTATAGCTCCACGCTCTTCTTCTCATAACGGTATTTTTAGGAGGAACGGTCTTACCTTCGGTTCATGATGAACGTTT
 AGACTTTAAAGATTGGTCGGAAAGTGTGCACCTGGAACCTGTTCTTATCCAGGTTTCTTTACAACCTATAACGAATT
 TACTAGTTCCCTCTCATTTTCTAATATCGGTGCTGCAGAGCTCTCTAATAAACGAGAATCTCTTCAACAAGCGATT
 CGAGTCTATCTCAAAGAACATAAAGTAGATGAGCAAGGAGAACGTAACATTTACATCAGGAATCGATAATCCTTC
 TTCCTGGTTTACCTTAGAAGCTGCCACTCTCTCTTATAGTACGTAACCTTACATTTGCTTCTGTTGCTACGCTTC
 CTTATTTGTTCCCAACATTAAGAGAACGTTCTTCGGCAACCCCTATTCGTTTTCTATTTTTGTGTAGATAAATAATGAA
 CATGCTTCGCAAAAAATATTAACAACATCGTATTGCTTCTCCTCGGTCCTTGGCTATTTCGACAAAAAATTTTTGGTAG
 CGAATTTGCTAGTTTCCCTATCTATCTTTCTATGGAAATGCAAAAAGAGGCGTACTTTGATAACACGTAACCCAA
 CGCGTTGTTGGGTTGATTGTTGAAAAGTAAATACTACACAAGATCAATTCCTCCGGGATGGAGACGAGGTGCGACTA
 AAACATGTTTCCAGCGGAAAGTATCTAGCAACAACCTCTTAAAGGATACCCATGGTACACTCACGCGTACAACGAA
 CTGTGAAGATGCTATCTTTATTTAAAAAATCTTCAGGTTATTGA

SEQ ID NO:4 - CT153 protein sequence

MTKPSFLYVIQPFVFNPRLGRFSTDSPTYIEENRLASFIESLPLEIFDIPSMETAISNSPYILSWETTKDGFALF
TILEPKLSACAATCLVAPSIQMKSDAELLEIEIKQALLRSSHDGVKYRITRESFSPEKKT PKVALVDDDIELIRNVDF
LGRAVDIVKLDPINILNVTVEENILDYSFTRETAQLSADGRFGIPPGTKLFPKPSFDVEISTSI FEETTSFTRSFSA
SVTFVSPDLAATMPLQSPPMVENGQKEICVIQKHLFPSYSPKLVDIVKRYKREAKILINKLAFGMLWRHRAKSQILT
EGSVRLDLQGFTESKYNYIQVGSHTIAAVLIDMDISKIQSKSEQAYAIRKIKSGFQRSLDDYHIYQIERKQTFSS
PKHRSLSSSTSHSESDLDLSEAAAFSSGLTCEVKKSTQHAKNVTVCSTAAHSLYTLKEDDSSNPSEKRLDSCFRNW
IENKLSANSPDSWSAFIQKFGTHYIASATFGGIGFQVLKLSFEQVEDLHSHKISLETAANSLKGSVSSSTESGYS
SYSSTSSSHTVFLGGTVLPSVHDERLDFKDWSESVHLEPVPIQVSLQPIITNLLVPLHFPNIGAAELSNKRESLQQA
RVYLKEHKVDEQGERTTFTSGIDNPSSWFTLEAAHSPDIVSTPYIASWSTLPYLFPTLRERSSATPIVYFVVDNNE
HASQKILNQSYCFGLSPIRQKIFGSEFASFPIYLFYGNAYEAYFDNTYPTRCGWIVEKLNNTTQDQFLRDGDEVRL
KHVSSGKYLATTPKLDTHGTLTRTTNCEDAIFIIKSSSY

SEQ ID NO:5 - CT601 nucleotide sequence

ATGCTCGCTAATCGCTTATTCTTAATAACCCCTTTAGGGTTAAGTTTCGTCTGTTTACGGCGCAGGTAAGCACCGTC
TTTGACAGGCTATTCTAGCCGAAGTCGAAGACACCTCCTCTCGTCTACACGCTCATCACAAATGAGCTTGCTATGATCT
CTGAACGCCTCGATGAGCAAGACACGAAACTACAGCAACTTTTCGTCAACACAAGATCATAACCTACCTCGACAAGTT
CAGCGACTAGAAAACGGACCAAAAAGCTTTGGCAAAAACACTGGCGATTCTTTTCGCAATCCGTCCAAGATATTCGGTC
TTCTGTACAAAATAAATTACAAGAAATCCAACAAGAACAAAAAATTAGCACAAAATTTGCGAGCGCTTCGTAACCT
CTTTACAAGCTCTCGTTGATGGCTCTTCTCCAGAAAATTATATTGATTTCTAAGTGGTGAACCCCGGAACATATT
CATATTGTTAAACAAGGAGAGACCTGAGCAAGATCGCGAGTAAATATAACATCCCGTCTGATAGAATTAATAAACT
TAATAAACTAAATTCGGATACTATTTTTACAGATCAAAGAATTCGCCTTCCGAAAAAGAAATAG

SEQ ID NO:6 - CT601 protein sequence

MLANRFLFITLLGLSSVYGAGKAPSLQAILAEVEDTSSRLHAHNLAMISERLDEQDITKLQQLSSTQDHNLPQV
QRLETDQKALAKTLAILSQSVQDIRSSVQNKLEIQEQEKKLAQNLRALRNSLQALVDGSSPENYIDFLTGETPEHI
HIVKQGETLSKIASKYNIPVVELKKNLNSDTIFTDQRIKPKK

SEQ ID NO:7 - CT279 nucleotide sequence

ATGGCATCCAAGTCTCGCCATTATCTTAATCAGCCTTGGTACATTATCTTATTCATCTTTGTTCTTAGTTAATTGC
TTGGTACCCTCTGTCTTCTGTGATTATGTCTTGCACCTATCCAACAGCAAGCTGCGGAATTCGATCGCAATCAAC
AAATGCTAATGGCTGCACAAGTAATTTCTCCGATAACACATTTCCAAGTCTATGAAAAGGGAGATTGGCACCCAGCC
CTATATAATACTAAAAAGCAGTTGCTAGAGATCTCCTCTACTCCTCCTAAAGTAACCGTGACAACCTTAAAGCTCATA
TTTTCAAACTTTGTTAGAGTCTTGCTTACAGATACACAAGGAAATCTTTCTTCATTGAAAGACCATAATCTCAATC
TAGAAGAATTTTTATCTCAACCAACTCCTGTAATACATGGTCTTGCCCTTTATGTGGTCTACGCTATCCTACACAAC
GATGCAGCTTCTCTAAATATCTGCTTCCCAAGTAGCGAAAAATCCAACAGCTATAGAATCTATAGTTCTTCTCAT
AGAAGTTTTGGTTTTGGGGACTATCTATGGATTCTTCTGCTATAGAAAAAGACGGGAATCCTGTTCTTGGTACTT
CTTGGTATCAACATGGCGAGACTCCTGGATTAGGAGCAATATCGCTAACCTCAATGGCAAAAAAATTTAGAGGC
AAAAAAGTATTTCTAGTCTCAGCTTCTGGAGAAACAGATTTGCTAAGACAACCTAGGACTGGAAGTTATAAAAGG
ATCTGTATCTGCAGCATTAGGAGACTCACCTAAAGCTGCTTCTTCCATCGACGGAAATTTAGGAGCTACTTTGACTT
GTAATGGTGTACCGAATCCTTCTCTCATTCTCTAGCTCCCTACCGCGCTTTGTTGACTTTCTTCCCAACTCTAAA
CCTAGTGGAGAGTCTCATGACCACTAA

SEQ ID NO:8 - CT279 protein sequence

MASKSRHYLNQPWYIILFIFVLSLIAGTLLSSVYVLAPIQDQAAEFDRNQMLMAAQVISSDNTFQVYEKGDWHPA
LYNTKKQLEISSPPKVTVTLLSSYFQNFVRVLLTDQGNLSSFEDHNLNLEEFLSQPTPVIHGLALVYVYAILHN
DAASSKLSASQVAKNPTAIESIVLPIEGFGLWGPYGFLEKDGNTVLGTSWYQHGETPGLGANIANPQWQKNFRG
KKVFLVSASGETDFAKTTLGLEVIKGSVSAALGDSPKAASSIDGISGATLTCNGVTESFSHSLAPYRALLTFFANSK
PSGESHDH

SEQ ID NO:9 - CT443 nucleotide sequence

ATGCGAATAGGAGATCCTATGAACAACTCATCAGACGAGCAGTGACGATCTTCGCGGTGACTAGTGTGGCGAGTTT
ATTTGCTAGCGGGGTGTTAGAGACCTCTATGGCAGAGTCTCTCTACAAACGTTATTAGCTTAGCTGACACCAAAG
CGAAAGACAACACTCTCATAAAAAGCAAAAAGCAAGAAAAACCACAGCAAAGAGACTCCCGTAGACCGTAAAGAG
GTTGCTCCGGTTCATGAGTCTAAAGCTACAGGACCTAAACAGGATTTCTGCTTTGGCAGAATGTATACAGTCAAAGT
TAATGATGATCGCAATGTTGAAATCACACAAGCTGTTCTTGAATATGCTACGGTAGGATCCTCCTATTGAAAA
TTACTGCTACAGGTAAGGGATTGTGTTGATGTTATCACTACTCAGCAATTACCATGTGAAGCAGAGTTCGTACGC
AGTGATCCAGCGACAACCTCCTACTGCTGATGGTAAGCTAGTTTGGAAAATTGACCGCTTAGGACAAGGCGAAAAGAG
TAAAATTAAGTATGGGTAACCTCTTAAAGAAGTTGCTGCTTTACAGCTGCAACAGTATGCGCTTGTCCAGAGA
TCCGTTCCGGTTACAAAATGTGGACAACCTGCTATCTGTGTTAAACAAGAAGGCCAGAGAATGCTTGTGTTGCGTTGC
CCAGTAGTTTACAAAATAATATAGTGAACCAAGGAACAGCAACAGCTCGTAACGTTGTTGTTGAAAATCCTGTTCC
AGATGGTTACGCTCATTCTTCTGGACAGCGTGTACTGACGTTTACTCTTGGAGATATGCAACCTGGAGAGCACAGAA
CAATTAAGTGTAGAGTTTTGTCGCTTAAACGTTGGTCTGCTACCAATATAGCAACGGTTTCTTACTGTGGAGGACAT
AAAAATACAGCAAGCGTAAACAAGTGTGATCAACGAGCCTTGGCTACAAGTAAGTATTGCAAGGAGCAGATTGGTCTTA
TGTTTTGTAAGCCTGTAGAATATGTGATCTCCGTTTCCAATCCTGGAGATCTTGTGTTGCGAGATGTCGTCGTTGAAG
ACACTCTTTCTCCCGGAGTACAGTCTTGAAGCTGCAGGAGCTCAAATTTCTGTAATAAAGTAGTTTGGACTGTG

AAAGAACTGAATCCTGGAGAGTCTCTACAGTATAAAGTTCTAGTAAGAGCACAAACTCCTGGACAATTCACAAATAA
TGTTGTTGTGAAGAGCTGCTCTGACTGTGGTACTTGTACTTCTTGGCGAGAAGCGACAACCTTACTGGAAAGGGAGTTG
CTGCTACTCATATGTCGCTAGTAGATACTTTGTGACCTGTTTGGTTTACACGGAGTTTCTGTAGGAGAAAATACTGTTTACCCTATTTGTGTC
ACCAACAGAGGTTCTGCAGAAAGATACAAATGTTTCTTTAATGCTTAAATTTCTTAAAGAACTGCAACCTGTATCCTT
CTCTGGACCAACTAAAGGAACGATTACAGGCAATACAGTAGTATTGATTTCGATTGTTACCTAGATTAGGTTCTAAAGAAA
CTGTAGAGTTTTCTGTAACATTGAAAGCAGTATCAGCTGGAGATGCTCGTGGGGAAGCGATTCTTTCTCCGATACA
TTGACTGTTCCAGTTTCTGATACAGAGAATACACACATCTATTA

SEQ ID NO:10 - CT443 protein sequence

MRIGDPMNKLIRRAVITFAVTSVASLFAAGVLETSMAESLSTNVISLADTKAKDNTSHKSKKARKNHSKETPVDRKE
VAPVHESKATGPKQDSCFGRMYTVKVNDDRNEITQAVPEYATVGSPIEITATGKRDCVDVVIITQQLPCEAEFVR
SDPATTPTADGKLVWKIDRLGQGEKSKITVWVKPLKEGCFATAATVCACPEIRSVTKGQPAICVKQEGPENACLRC
PVVYKINIVNQTATARNVVVENPVPDGYAHSSGQRVLTFTLGDMPGEHRTITVEFCPLKRGRATNIATVSYCGGH
KNTASVTTVINPCVQVSIAGADWSYVCKPVEYVISVSNPGLVLRDVVVEDTLPSPGVTVLEAAGAQISCNKVVWTV
KELNPGESLQYKVLVRAQTPGQFTNNVVVKSQSDCGTCTSCAEATTYWKGVAATHMCDVDTCDPVCVAGENTVYRICV
TNRGSAEDTNVSLMLKFSKELQPVSFSGPTKGTITGNTVVFDSLPRLGSKETVEFSVTLKAVSAGDARGEAILSSDT
LTVPVSDTENTHIY

SEQ ID NO:11 - CT372 nucleotide sequence

ATGCAGGCTGCACACCATCACTATCACCGCTACACAGATAAACTGCACAGACAAAACCATAAAAAAGATCTCATCTC
TCCCAAACCTACCGAACCAAGAGGCGTGAATACTTCTCCCTTAGTAAGGAATTAATCCCTCTATCAGAACAAAGAG
GCCTTTTATCCCCCATCTGTGACTTTATTTGGAACGCCCTTGGTTTACACGGAGTTTCTGTAGAAATCTCAAGCAA
GCGCTAAAAAATTCTGCAGGAACCCAAATTGCACTGGATTGGTCTATTCTCCCTCAATGGTTCAATCTCGGGTCTC
TCATGCCCCTAAGCTTCTATCCGAGACTTTGGGTATAGCGCACACCAAACCTGTTACCGAAGCCACTCCTCCTTGCT
GGCAAAACTGCTTTAAATCCATCTGCGGCCGTTACTATCTATGATTCTCATATGGGAAAGGGGCTTTTCAAATATCC
TATAACCTTGTCCGCTATTGGAGAGAGAATGCTGCGACTGCTGGCGATGCTATGATGCTCGCAGGGAGTATCAATGA
TTATCCCTCTCGTCAGAACATTTTCTCTCAGTTTACTTTCTCCCAAACCTCCCAAATGAACGGGTGAGTCTGACAA
TTGGTCAGTACTCACTCTATGCAATAGACGGAACATTATACAATAACGATCAACAACCTGGATTCAATAGTTACGCA
TTATCACAAAATCCAACGCAACTTATTTCTCTGGAAGTCTTGGAGCTTACCTACAAGTGCCTCCTACCGCAAGCAC
AAGTCTTCAAATAGGATTTCAAGACGCTTATAATATCTCCGGATCCTCTATCAAATGGAGTAACCTTACAAAAATA
GATACAATTTTCCGGTTTTGCTTCTGCGCTCCCGCTGTTGCTTAGGATCTGGCCAGTACTCCGTGCTTCTTTAT
GTGACTAGACAAGTTCCAGAACAGATGGAACAAACAATGGGATGGTCAGTCAATGCGAGTCAACACATATCTTCTAA
ACTGTATGTGTTTGAAGATACAGCGGTGTTACAGGACATGTGTTCCCGATTAAACCGCACGATTCATTCGGTATGG
CCTCTGCAAAATTTATTTAACCGTAACCCACAAGATTTATTTGGAATTGCTTGGCATTCAATAATGTACACCTCTCT
GCTTCTCCAAATACTAAAAGAAAATACGAAACTGTAATCGAAGGGTTTGAACATATCGGTTGCGGCCCTATCTTTC
TTTCGCTCCAGACTTCCAACCTACCTCTACCCAGCTCTTCGTCCAAACAAACAATCTGCCCGTGTATAGCGTGC
GAGCTAATTTAGCTATCTAA

SEQ ID NO:12 - CT372 protein sequence

MQAAHHHYHRYTDKLRHRNHHKDLISPKPTEQEAACNTSSLSKELIPLSEQRGLLSPICDFISERPCLHGVSVRNLKQ
ALKNSAGTQIALDWSILPQWFNPRVSHAPKLSIRDFGYSAHQTVTEATPPCWQNCFNPSAAVTIYDSSYKGVFQIS
YTLVRYWRENAATAGDAMMLAGSINDYPSRQNFISQFTFSQNFNERVSLTIGQYSLYIDGTLYNNQQLGFI SYA
LSQNPATYSSGSLGAYLQVAPTASTSLQIGFQDAYNISGSSIKWSNLTKNRYNFHGFASWAPRCCLGSGQYSVLLY
VTRQVPEQMEQTMGWSVNASQHISSKLYVFGRYSGVTGHVFPINRTYSFGMASANLFNRNPQDLFGIACAFNNVHLS
ASPNTKRYEYVIEGFATIGCGPYLSFAPDFQLYLYPALRPNKQ SARVYSVRANLAI

SEQ ID NO:13 - CT456 nucleotide sequence

ATGACGAATTCATATCAGGTTATCAACCTACTGTTACAACCTTCTACATCATCAACCCTTCCGGCATCAGGTGCTTC
CGGATCTCTGGGAGCTTCTTCTGTATCTACTACCGCAAACGCTACAGTTACACAAAACAGCAAACGCAACAAATTCAG
CGGCTACATCTTCTATCCAAACGACTGGAGAGACTGTAGTAAACTATACGAATTCAGCCTCCGCCCCCAATGTAAC
GTATCGACCTCCTTCTTCCACACAAGCCACAGCCACTTCCGAATAAACTTCCAAAGCCGTTGCTGGAAAAATCAC
TTCTCCAGATACTTCCAGAAAGCTCAGAAACTAGCTCTACTCTCAAGCGATCATATCCCTACAGCAATTTAGCAGT
TTGGTAGCAATAGTGGAGATATTAGCAACAACCTACGATGACGTAGGTAGTAACAACGGAGATATCAGTAGCAATTAT
GACGATGCTGCTGCTGATTACGAGCCGATAAGAACTACTGAAAATATTTATGAGAGTATTGGTGGCTCTAGAACAAG
TGGCCAGAAAAACAAGTGGTGGTGCAGCAGCAGCACTCAATTTCTTAAGAGGCTCCTCCTACAGCAATTTAGCAG
ATGCTGCTGCTGATTACGAGCCGATAAGAACTACTGAAAATATTTATGAGAGTATTGGTGGCTCTAGAACAAGTGGC
CCAGAAAATACGAGTGGTGGTGCAGCAGCAGCACTCAATTTCTTAAGAGGCTCCTCCTACAGCAATTTAGCAGTGC
TGCTGCTGATTACGAGCCGATAAGAACTACTGAAAATATTTATGAGAGTATTGGTGGCTCTAGAACAAGTGGCCAG
AAAATACGAGTGATGGTGCAGCAGCAGCAGCACTCAATTTCTTAAGAGGCTCCTCCTACACAACAGGGCCCTGTAAC
GAGGGTGTATTCGGCCCTGGACCGGAAGGACTACCAGACATGTCTCTTCTTATACGATCCTACAAATAAAACCTC
GTTATTTGACTTTCTCTCCAACCTCTATGTAAGTTCGAAAATGCTTGAACCTCGGGGCATTTCTGCTTCTATTGATA
CAGATAGAAGTAGTTTCAATTTCTTGTTCCTAACGAAAATTTGGACCAAGTCTGTTCAATTAAGTTCAAAATGGAAG
ACCAAAGAAGATCTCGACATCAAAGACTTGGAAAACATGTGTGCAAAATTTCTGTACAGGGTTTAGCAAAATTTCTGG
TGACTGGGACAGTCTTGTAGAACCCTATGGTGTGACGCAAAAGCTGGAGTGGCCAGCGGAGGCAATCTTCCCAATACAG
TGATTATCAATAATAAATTCAAAACCTTGGTGTGCTTATGGTCTTGGAAATAGCCAGGAAGCAAGTCTGGTTATACA
CCTTCTGCTTGGAGACGTGGTCACTCGAGTAGATTTTGGAGGAATTTTGGAGAAAGCCAAACGACTTTAATAAAATCAA
CTGGGGAACCTAAGCCGGGCTAGTAGCGAAGACGATGGCATTTCCTTCTCAATGAAACTCCTGGAGCTGGTCTCTG

CAGCTGCTCCATCACCAACGCCATCCTCTATTCTATCATCAATGTCAATGTCAATGTTGGCGGAACATAATGTGAAT
 ATTGGAGATACGAATGTCAACACGACTAACACCACACCAACAACCTCAATCTACAGACGCCCTACAGATACAAGCGA
 TATCGATGACATAAAATACCAACAACCAAACTGATGATATCAATACGACAGACAAAGACTCTGACGGAGCTGGTGGAG
 TCAATGGCGATATATCGGAAACAGAATCCTCTTCTGGAGATGATTAGGAAGTGTCTCTTCTCAGAATCAGACAAG
 AATGCCTCTGTGCGAAATGACGGACCTGCTATGAAAGATATCCTTTCTGCCGTGCGTAAACACCTAGACGTCGTTTA
 CCCTGGCGAAAATGGCGGTTCTACAGAAGGGCCTCTCCAGCTAACCAAACTCTCGGAGACGTAATCTCTGATGTAG
 AGAATAAAGGCTCCGCTCAGGATACAAAATTGTCAGGAAATACAGGAGCTGGGGATGACGATCCAACAACCACAGCT
 GCTGTAGGTAATGGAGCGGAAGAGATCACTCTTCCGACACAGATTCTGGTATCGGAGATGATGTATCCGATACAGC
 GTCTTCATCTGGGGATGAATCCGGAGGAGTCTCCTCTCCCTCTTCCAGAATCCAATAAAAACTGCCGTTGGAAATG
 ACGGACCTTCTGGACTAGATATCCTCGCTGCCGTACGTAACAATTTAGATAAGGTTTACCCTGGCGACAATGGTGGT
 TCTACAGAAGGGCCTCTCCAAGCTAACCAAACTCTTGGAGATATCGTCCAGGATATGGAAACAACAGGGACATCCCA
 AGAAACCGTTGTATCCCCATGGAAAGGAAGCACTTCTTCAACGGAATCAGCAGGAGGAAGTGGTAGCGTACAAACAC
 TACTGCCCTTACCACCTCCAACCCCGTCAACTACAACATTAAGAACGGGCACAGGAGCTACCACCACATCCTTGATG
 ATGGGAGGACCAATCAAAGCTGACATAATAACAACCTGGTGGCGGAGGACGAATTCCTGGAGGAGGAACGTTAGAAAA
 GCTGCTCCCTCGTATACGTGCGCACTTAGACATATCCTTTGATGCGCAAGGCGATCTCGTAAGTACTGAAGAGCCTC
 AGCTTGGCTCGATTGTAACAATTCGCAAGAACTGGTTCAAGAGGAATCTTAGCTTTGCTTGAGAGTGCCTCA
 GGCAAGCCGGGATCTGCACAGGCTTAAACGGGTACAGGGGGAGATAAAGGCAACCTATTCCAAGCAGCTGCCGCACT
 CACCCAAGCCTTAGGAAATGTTGACAGGAAAGTCAACCTTGGGATACAAGGCCAAAAACTATCATCCCTAGTCAATG
 ACGACGGGAAGGGTCTGTTGGAAGAGATTTATCCAAGCAGCAGCCAAACAACCTCAAGTGCTAAGCGCACTGATT
 GATACCGTAGGATAA

SEQ ID NO:14 - CT456 protein sequence

MTNSISGYQPTVTTSTSSSTTSASGASGLGASSVSTANATVTQTANATNSAATSSIQTTGETVVNYTNSASAPNV
 VSTSSSTQATATSNKTSQAVAGKITSPDTSSESSTSSSDHIPSDYDDVGSNSGDISNNYDDVGSNNGDISSNY
 DDAADYEPIRTTENIYESIGGSRSTSGPENTSGGAAAALNSLRGSSYSNYDDAAADYEPIRTTENIYESIGGSRSTSG
 PENTSGGAAAALNSLRGSSYSNYDDAAADYEPIRTTENIYESIGGSRSTSGPENTSDGAAAALNSLRGSSYTTGPRN
 EGVFPGPEGLPDMSLPSYDPTNKTSLLTFLSNPHVKSMLKLENSGHFVFIIDTRSSFILVPNGNWDQVCSIKVQNGK
 TKEDLDIKDLENMCAKFCSTGFSKFSGDWDSLVEPMVSAKAGVASGGNLPNTVIINNKFKTCVAYGPWNSQEAASSGYT
 PSAWRRGHRVDFGGIFEKANDFNKINWGTQAGPSSDDGISFSNETPGAGPAAAPSPPTPSSPIINNVNVGGTNNV
 IGDNTVNTTNTPTTQSTDASTDSTIDDDINTNNQTDINTTDKSDSDGAGGVNGDISETESSSGDSDSGVSSSES
 NNASVGNDDGPMKDIILSAVRKHLVVYPGENGGSTEGPLPANQTLGDVSDVENKGSQAQDTKLSGNTGAGDDPTTTA
 AVGNAGEEITLSDTDSGIGDDVSDTASSSGDES GGVSPPSESSENKNTAVGNDGPSGLDILA AVRKHL DKVYPGNDGG
 STEGPLQANQLGDIVQDMETTGTSQETVVSPPWKGSTSTESAGGSGSVQTLSPPTPTSTTLRTGTGATTTSLM
 MGGPIKADIITTGGRIPGGGTLEKLLPRIRAHLDISFDAQGDLVSTEEPQLGSI VNKLPFRQETSRGILAFVESA
 PKPQSAQVLTGTGGDKGNLFQAAA AVTQALGNVAGKVNLAIQGQKLSLVNDDGKGSVGRDLFQAAAQTTQVLSALI
 DTVG

SEQ ID NO:15: CT381 nucleotide sequence

ATGTGCATAAAAAGAAAAAACAATGGATAGCTTTTTTAGCAGTTGTCTGTAGTTTTTTGTTTGACGGGTTGTTTAAA
 AGAAGGGGGAGACTCCAATAGTAAAAAATTTATTGTAGGACTAATGCAACCTACCCTCTTTTGTAGTTTGTGATA
 AGCGAGGAGAGGTTGTAGGCTTCGATATAGACTTGGCTAGAGAGATTAGTAAACAAGCTGGGGAAAACGCTGGACGTT
 CGGGAGTTTTCTTTGATGCACTCATTCTAAACCTAAAAACAGCATCGGATTGATGCGGTTATAACAGGGATGTCCAT
 TACTCCTTCTAGATTGAAGGAAATTTTATGATCCCTATTATGGGGAGGAAATAAAACACTTGGTTTTAGTGTTTA
 AAGGAGAGAATAAGCATCCATTGCCACTCACTCAATATCGTTTCTGTAGCTGTTCAAACAGGAACCTATCAAGAGGCC
 TATTTACAGTCTCTTTCTGAAGTTTCAATTCGCTTTTTAGTAGACTCTAGAAGTACTCATGGAAGTCATGCATGG
 TAAATCTCCGTCGCTGTTTTAGAGCCATCTATCGCTCAAGTTGTCTTGAAGATTTCCCGGCTCTTTCTACAGCAA
 CCATAGATCTCCCTGAAGATCAGTGGGTTTTAGGATACGGGATTGGCGTTGCTTCAAGATCGCCCAGCTTTAGCCTTG
 AAAATCGAGGCAGCTGTGCAAGAGATCCGAAAAGAAGGAGTGCTAGCAGAGTTGGAACAGAAGTGGGGTTTGAACAA
 CTAA

SEQ ID NO:16: CT381 protein sequence

MCIKRKKTWIAFLAVVCSFCLTGCLKEGGDSNSEKFIGVTNATYPPFEFVDKRGEVVGFDIDLAREISNKLKGLD
 REFSFDALILNLKQHRIDAVITGMSITPSRLKEILMIPYYGEEIKHLVLFKGENKHPLPLTQYRSVAVQTGTQEA
 YLQSLSEVHIRSFDSTLEVLMEVMHGKSPVAVLEPSIAQVVLKDFPALSTATIDLPEQWVLYGIGVASDRPALAL
 KIEAAVQEIIRKEGVLAELEQKWGLNN

SEQ ID NO:17: CT043 nucleotide sequence

ATGTCAGGCGAGAATGCTGAGGAAAACTAAAAAATTTGCTAAAGAGCTTAAACTCCCCGACGTGGCCTTCGATCA
 GAATAATACGTGCATTTTGTGTTGATGGAGAGTTTCTCTTACCTGACCTACGAAGAACACTCTGATCGCCTTT
 ATGTTTACGCACCTCTTCTTACGGACTGCCAGACAATCCGCAAGAAGGTTAGCTCTATATGAGAAGTTGTTAGAA
 GGCTCTATGCTCGGAGGCCAAATGGCTGGTGGAGGGGTAGGAGTCTACTAAGGAACAGTTGATCTTAATGCACTG
 CGTGTTAGACATGAAGTATGACAGAGACCAACCTACTCAAAGCTTTTGCACAGCTTTTTATTGAAACCGTTGTAAAT
 GGCGAACTGTTTGTCTGATATCAGCGCTGGACGAGAACCCTACTGTTGATACCATGCCACAAATGCCTCAAGGGGGT
 GGCGGAGGAATCAACCTCCTCCAGCAGGAATCCGTGCATAA

SEQ ID NO:18: CT043 protein sequence

MSRQNAEENLNKNAKELKLPDVAFDQNNTCILFVDGEFSLHLTYEEHSDRLVYAPLLDGLPDPNRRLALYEKLL
 GSMLGGQMGAGGGVGVATKEQLILMHCVLDMKYAETNLLKAFALFIETVVKWRTVCSDISAGREPTVDTMPQMPQGG
 GGGIQPPPAGIRA

SEQ ID NO:19: CT711 / hypothetical protein (AAC68306)

MSIQPTISISLTKNITAALAGEQVDAAAVYMPQAVFFFQQLDEKSKGLKQALGLLEEVDLEKFIPLSLEKSPTPITTTGT
 TSKISADGIEIVGELSSETILADPNKAAAQVFGGLADSFDDWLRLENGGIQDPTAIEEEIVTKYQTELNLRNKL
 KQQLSLTDDDEYTKLYAIPQNFVKEIESLKNENNVRILPKSKVTNFQWQNIMLTYNSVTSLEPVTDAAMNTTMAEYSLYI
 ERATEAAKLIREITNTIKDIFNPVWDVREQTGIFGLKGAEYNALEGNMIQSLLSFAGLFRQLMSRTATVDEIGALYP
 KNDKNEDEVIIHTAIDDYVNSLADLKANEQVKLNGLLSLVYAYASTLGFACKDVFNNAQASFTDYTNFLNQEIQYWTP
 RETSSFNISNQLQTFKNKPSADYNGVYLFDNKGLETNLFNPTFFFQVSLMTADPTKMSRQDYNKVITASESSIQ
 KINQAITAWELAIACGTTKAKLEPSSLNYNFAMVEAKKTFVETSPIQMVYSSMLDKYLPNQYILETLGSQMTFS
 NKAARYLNDIAYAVSFQADVYVYSLGMYLRQMNQEFPEVISRANDTVKKEIDRSRADLFHCCKAIEKIKELVTSV
 NADTELTSSQRAELLETLASYAFEFENLYHNL SNVYVMVSKVQISGVSKPDEVDEAF TAKIGSKEFDTWIQQLTTFE
 SAVIEGGRNGVMPGGEQVLQSLQSKQDYTSFNQNLALQMESAAIQEWTMVAAAALMNNQIFAKLIRRFK

SEQ ID NO:20: CT114 / hypothetical protein (AAC67705)

MCFIGIGSLLLPTALRATERMRKEPIPLLDKQSFQWVNDPYCLECICACFVAHRDPLSAKQLMYLFPQLSEEDVSVF
 ARCILSSKRPEYLFKSEELFAKLILPRVSLGVHRDDDLARVLAEPSAEEQKARYYSLYLDVLAALRAYVERERL
 ASAAHGDPERIDLATIEAINTILFQEEGWRYPSKQEMFENRSELAAVTDSKFGVCLGTVVLYQAVQRLDLSLDPV
 TPPGHIYLRKDKVNIETTSGGRHLPTERYCECIKESQLKVRSMELIGLTFMNRGAFFLQKGEFLQASLAYEQADS
 YLSDEQISDLLGITYVLLGKKAAGEALLKKSAEKTRRGSIIYDYFQGYISPEILGLVFAADSGVTYQETLEYRKKLVM
 LSKKYPKSGSLRRLATTALELGLVKEGVQLLEESVKDAPEDLSLRLQFCILCNRHDYVRAKYHFDQAQALLIKEG
 LFSEKTSYTLTKTIGKLSLAFPS

SEQ ID NO:21: CT480 / oppA_4 (AAC68080)

MIDKIIRTILVLSLFLLYWSSDLLEKDVKSIRELKAHEVDVLELVRISHQKKNVQSTDFSVSPEISVLKDCGPA
 FPNLLCEDPYVEKVVPSLLKEGFVVKGILRTAQVGRPDNLSPFNGFVNIYRFYELCVPNLAVEHVKGKYEAFAPSLAL
 KIEEHYVEDGSGDKEFHILRPNMFWEPIDPTLFPKNITLADSFRLRPHVTAHDVKFYDDVVMNPPYVAEMRAVAMRS
 YFEDMVSVRVENDLKLIVRWRRAHTVRNEQGEEEKVLYSAFANTLALQPLPCFYQHFANGEKIVPESDSDPTYRKD
 SVWAQNFSSHWAYNYIVSCGAFRFAGMDDEKITLVRNPNYHNPFAALVEKRYIYMKDSTDSLQDFKAGKVDIAYFP
 PNHVNDLASFMQTSAYKEQAARGEAILEKNSSDRSYIYGNCLSLFFNRSVRQAMNMLIDRDRIEEQCLDGRGVS
 VSGPFSLCSPSYNRDVEGWQYSPEEAARKLEEEGWIDADGDGIREKVIDGVVVPFRFLCYVVKSVTARTIAEYVAT
 YCKEVGIECCLLGLDMADYSQALEEKNFDAILSGWCLGTPPEDPRALWHSEGALEKGSANAVGFCNEEADRIIEQLS
 YEYDSNKRQALYHRFHEVIHEESPYAFLYSRQYSLVYKEFVKNIFVPTHEQDLIPGAQDET VNL SMLWVDKEEGRC
 AIS

SEQ ID NO:22: CT089 / lcrE (AAC67680)

MTASGGAGGLGSTQTVDVARAQAATAQDAQEVIGSQEASEASMLKGCEDLINPAAATRIKKKGEKFESELEARRKPT
 ADKAEEKSESTEKGDTPLEDRFTEDLSEVSGEDFRGLKNSFDSSPDEILDALTSKFSQDPTIKDLALDYLIQTAP
 SDGKMLSTLIQAKHQLMSQNPQAIIVGGRNVLLASETFASRANTSPSSLRSLYFQVTSPPSNCANLHMLASLYPSEK
 TAVMVLVNGMVDLQSEGPSIPPAKLQVYMTLSNLQALHSVNSFFDRNIGNLENSLKHGHAPIPSLTGGLTKT
 FLQLVEDKFPSSSKAQKALNELVGPDTGPTQTEVNLNFFRALNGCSPRIFSGAEKKQQLASVITNTLDAINADNEDYP
 KPGDFPRSSFSSTPPHAPVPQSEIPTSTSTQPPSP

SEQ ID NO:23: CT734 / hypothetical protein (AAC68329)

MKKFIYKYSFGALLLSGLSGLSSCCANSYGSTLAKNTAEIKEESVTLREKPDAGCKKKSSCYLRKFFSRKPKPEKT
 EPVLPNFKSYADPMTDSERKDLDFVVSAAADKSSIALAMAQGEIKGALSRIEIHPLALLQALAEDPALIAGMKKMQ
 GRDWVWNIFITELSKVFSQAASLGAFSVADVAFASTLGLDSGTVTSIVDGERWAELIDVVIQNPAI

SEQ ID NO:24: CT016 / hypothetical protein (AAC67606)

MKVKINDQFICISPYISARWNQIAFIESCDGGTEGGITLKLHLIDGETVSIPLNGQAIIVDEVFQEHLLYLESTAPQK
 NKEEEKISSLLGAVQMAKAGCEVQVFSQKGLVSMLLGGAGSINVLLQHSPEHKDHPDLPTDLLERIAQMMRSLSIGP
 TSILAKPEPHCNCLHCQIGRATVEEEDAGVSDDELTFRSWDISQSGEKMYTVTDPLNPEEQFNVYLGTPIGCTCGQP
 YCEHVKAVLYT

SEQ ID NO:25: CM homolog of CT279 = TC_0551

ATGGCATCCAAGTCTCGTCATTATCTTAACCCAGCCTTGGTACATTATCTTATTCATCTTTGTTCTTAGTCTGGTTGC
 TGGTACCCCTTCTTTCTCAGTTTCCTATGTTCTATCTCCAATCCAAAAACAAGCTGCAGAATTTGATCGTAATCAGC
 AAATGTTGATGGCCGACAAATTTTCTATGACAAATAAATTCCAAATATATGCTGAAGGGGATTGGCAACCTGCT
 GTCTATAATACAAAAAACAAGATACTAGAAAAAAGCTCTTCCACTCCACCACAAGTGACTGTGGCGACTCTATGCTC
 TTATTTTCAAAATTTGTTAGAGTTTTGCTTACAGACTCCCAAGGGAATCTTTCTTTCTTTGAAGATCACAACTTA
 ACCTAGAAGAGTTCTTATCCACCCACATCTTACGTACAAGATCACTCTCTGCATGTAATTTATGCTATTCTAGCA
 AACGATGAATCCTCTAAAAAGTTATCATCTCCCAAGTAGCAAAAAATCCGGTATCCATAGAGTCTATTATCTTCC

TATAAAAGGATTTGGTTTATGGGGACCAATCTATGGATTTCTTGCTTTAGAAAAGGACGGTAATACGGTTCTAGGGA
CATGCTGGTATCAACATGGTGAGACTCCAGGATTAGGAGCAAATAAATACTAATCCCCAATGGCAACAAAATTTTCAGA
GGAATAAAAGTATTTCTCCTCTCCGGAGAACCGATTTTGCTAAAACAACCTAGGACTAGAAGTTATAAAA
AGGATCTGTTTCTGCATTATTAGGGGACTCTCCCAAAGCTAATTCGCTGTTGATGGAATTTTCAGGAGCTACACTGA
CCTGTAATGGAGTTACTGAAGCTTTTGTAAATTCGCTAGCTCCTTACCGCCCCCTTATTGACTTTCTTCGCCAATCTT
AACTCTAGTGGAGAATCTCATGACAACCAATAA

SEQ ID NO:26: CM homologue of CT279 protein sequence = TC_0551 protein sequence

MASKSRHYLNQWPYIILFIFVLSLVAGTLLSSVSIVLSPIQKQAAEFDRNQMLMAAQIISYDNKFIYAEGDWQPA
VYNTKKQILEKSSSTPPQVTVALCSYFQNFVRVLLTDSQGNLSSFEDHNLNLEEFLSHPTSSVQDHSLHVIYAILA
NDESSKLLSSQVAKNPVSIIESIILPIKGFGLWGPYIGFLALEKDGNTVLGTCWYQHGTEPLGANITNPQWQRFNR
GKKVFLASSSGETDFAKTTLGLEVIKGSVSALLGDSPKANSVDGISGATLTCNGVTEAFANSLAPYRPLLTFANL
NSSGESHDNQ

SEQ ID NO:27: CM homologue of CT372 = TC_0651 nucleotide sequence

ATGAATGGAAAAGTCTGTGTGAGGTTTCTGTGTCCTTCCGTTTCGATTCTGCTGACGGCTCTGCTTTCACCTTTCTTT
TACAAACACTATGCAGGCTGCACACCATCATTATCACCGTTATGATGATAAACTACGCAGACAATACCATAAAAAAGG
ACTTGCCCACTCAAGAGAAATGTTCCGAAAGAGTTTTGTAATCCCTACTCTCATAGTAGTGATCCTATCCCTTTGTCA
CAACAACGAGGAGTCCATCTCCTATCTGTGATTTAGTCTCAGAGTGCTCGTTTTTGAACGGGATTTCCGTTAGGAG
TCTTAAACAAAACACTGAAAAATTTCTGCTGGGACTCAAGTTGCTTTAGACTGGTCTATCCTTCCCTCAATGGTTCAATC
CTAGATCCTCTTGGGCTCCTAAGCTCTCTATTGAGATCTTGGATATGGTAAACCCAGTCCCTTATTGAAGCAGAT
TCCCCTTGTTCAAAACCTGCTTCAACCCATCTGCTGCTATTACGATTTACGATTCTTCATGTGGGAAGGTTGTTGT
CCAAGTGTACATACCCCTTGTTCGTTATTGGAGAGAAACGGCTGCACCTTGCAGGGCAAACATGATGCTTGCAGGAA
GTATTAATGATTATCCTGCTCGCCAAAACATATTTCTCAACTTACATTTTTCCCAAACCTTCCCTAATGAGAGAGTA
AATCTAACTGTTGGTCAATACTCTCTTTACTCGATAGACGGAACGCTGTACAACAATGATCAGCAGCTAGGATTTAT
TAGTTATGCGTTGTGCGAAAATCCAACAGCGACTTATTCCTCTGGAAGCCTTGGCGCCTATCTACAAGTGCCTCCAA
CAGAAAGCACCTGTCTCAAGTTGGGTTCCAAGATGCCTATAATATTTTCAGGTTCCCTCGATCAAATGGAATAATCTT
ACAAAAAATAAGTATAAACCCTCCATGGCTATGCATCTTGGGCTCCACACTGTTGCTTAGGACCTGGACAATACTCTGT
TCTTCTTTATGTAACCAGAAAGGTTTCTGAGCAATGATGCAGACAATGGGCTGGTCTGTGAATGCAAGTCAATACA
TCTCTTCTAAACTTTATGATTTGGAAGATACAGCGGAGTACAGGCCAATTGTCTCTATTAACCGAACCTATTCA
TTTGGCTTAGTCTCTCCTAATTTATTGAACCGTAACCCACAAGACTTATTTGGAGTAGCTTGCGCATTCAATAATAT
ACACGCCTCCGCTTTCAAATGCTCAAAGAAAATATGAAACTGTGATCGAGGGATTTGCAACTATTGGTTGCGGAC
CTTACATCTCCTTTGCTCCAGATTTCCAACCTTACTCTATCCTGCTCTGCTCCAAATAAACAAAAGCGCCGAGTC
TATAGCGTTCGCGCAAACCTAGCTATTTAG

SEQ ID NO:28: CM homologue of CT372 = TC_0651 protein sequence

MNGKVLCEVSVSFRSILLTALLSLSFTNTMQAHHHHYHRYDDKLRQYHKKDLPTQENVRKEFCNPYSHSSDPIPLS
QQRGVLSPICDLVSECSFLNGISVRSLSKQTLKNSAGTQVALDWSILPQWFNPRSSWAPKLSIRDLGYGKQPQSLIEAD
SPCCQTCFNPSAAITIYDSSCGKGVVQVSYTLVRYWRETAALAGQTMMLAGSINDYPARQNIQSRLTFSQTFPNERV
NLTVGQYSLSIDGTYLNNDDQLGFISYALSQNPATYSSGSLGAYLQVAPTESTCLQVGFQDAYNISGSSIKUNNL
TKNKYNFHGYASWAPHCLGPGQYSVLLYVTRKVPQMMQTMGWSVNASQYISSKLYVFGRYSGVTGQLSPINRTYS
FGLVSPNLLNRNPQDLFGVACAFNNIHASAFQNAQRKYETVIEGFATIGCGPYISFAPDFQLYLYPALRPNKQ SARV
YSVRANLAI

SEQ ID NO:29: CM homologue of CT443 = TC_0727

ATGCGAATAGGAGATCCTATGAACAACTCATCAGACGAGCTGTGACGATCTTCGCGGTGACTAGTGTGGCGAGTTT
ATTTGCTAGCGGGGTGTTAGAGACCTCTATGGCAGAGTCTCTCTTACCAACGTTATTAGCTTAGCTGACACCAAAG
CGAAAGAGACCACCTCTCATCAAAAAGACAGAAAAAGCAAGAAAAAATCATCAAAAATAGGACTTCCGTAGTCCGTA
GAGGTTACTGCAGTTCGTGATACTAAAGCTGTAGAGCCTAGACAGGATTCCTGCTTTGGCAAAAATGTATACAGTCAA
AGTTAATGATGATCGTAATGTAGAAATCGTGCAGTCCGTTCCCTGAATATGCTACGGTAGGATCTCCATATCCTATTG
AGATTACTGCTATAGGGAAAAGAGACTGTGTTGATGTAATCATTACACAGCAATTACCATCGGAAGCAGAGTTTGT
AGCAGTACTCCAGTACTCCTACTCTGCTGATGGTAAGCTAGTTTGGAAAAATTGATCGGTTAGGACAGGGCGAAAA
GAGTAAAATTACTGTATGGGTAAAACCTCTTAAAGAAGGTTGCTGCTTTACAGCTGCAACGGTTTTGTGCTTGTCCAG
AGATCCGTTCCGTTACGAAATGTGGCCAGCCTGCTATCTGTGTTAAACAGGAAGGTCAGAAAAGCGCATGTTTGGCT
TGCCACAGTAACTTATAGAATTAATGTAGTCAACCAAGGAACAGCAACAGCAGTAATGTTGTTGTGGAAAAATCCTGT
TCCAGATGGCTATGCTCATGCATCCGGACAGCGTGTATTGACATATACTCTTGGGGATATGCAACCTGGAGAACAGA
GAACAATCACCGTGGAGTTTTGTCCGCTTAAACGTGGTCCGAGTCAACAATATTGCTACAGTTTTCTACTGTGGTGGGA
CACAAAAATACTGCTAGCGTAACAACAGTGTCAATGAGCCTTGCCTGCAAGTTAACATCGAGGGAGCAGATTGGTC
TTATGTTTGTAAAGCCTGTAGAATATGTTATCTCTGTTTCTAACCTGGTGACTTAGTTTTACGAGACGTTGTAATTG
AAGATAAGCCTTCTCCTGGAATAACTGTTGTTGAAAGCAGCTGGAGCTCAGATTTCTTGTAAATAAATTGGTTTGGACT
TTAAGAAGAACTCAATCCTGGAGAGTCTTTACAATAAAGGTTCTAGTAAGAGCTCAAACCTCAGGGCAATTCACAAA
CAACGTTGTTGTGAAAAGTTGCTCTGATTGCGGTATTTGACTTCTTGGCAGAAAGCAACAACCTTACTGGAAAAGGAG
TTGCTGCTACTCATATGTGCGTAGTAGATACTTGTGATCCTATTTGCGTAGGAGAGAACTGTTTATCGTATCTGT
GTGACAAACAGAGGTTCTGCTGAAGATACAAATGTGTCCTTAATTTTGAATTTCTTAAAGAATTACAACCTATATC
TTTCTCTGGACCAACTAAAGGAACATTACAGGAAACACGGTAGTGTGTTGATTGCTTACCTAGATTAGGTTCTAAAG
AACTGTAGAGTTTTCTGTAACGTTGAAAGCAGTATCCGCTGGAGATGCTCGTGGGGAAGCTATTTCTTCCGAT
ACATTTGACAGTTCCTGTATCTGATACGGAGAATACACATATCTATTA

SEQ ID NO:30: CM homologue of CT443 = TC_0727

MRIGDPMNKLIRRAVTFITFAVTSVASLFAASVLETSMAESLSTNVISLADTKAKETTSHQKDRKARKNHQNRTSVVRK
EVTAVRDTKAVEPRQDSCFGKMYTVKVNDDRNVIEIVQSVPEYATVGSPPYIEITAIGKRDVVDVITQQLPCEAEFV
SSDPATTPADGKLVWKIDRLGQGEKSKITVWVKPLKEGCCFTAATVCACPEIRSVTKCGQPAICVKQEGPESACL
CPVTYRINVVNQGTARNVVVENPVPDGYAHASGRVLYTLGDMQPGEQRTITVEFCPLKRGRVTNIATVSYCGG
HKNTASVTTVINPCVQVNIIEGADWSYVCKPVEYVISVSNPGLVLRDVVIEDTLSPGITVVEAAGAQISCNKLVWT
LKELNPGESLQYKVLVRAQTPGQFTNNVVVKSCSDCGICTSCAEATTYWKGVAATHMCCVVDTCDPICVGENTVYRIC
VTNRGSAEDTNVSLILKFSKELQPIFSFGPTKGTITGNTVVFDSLPRLGSKETVEFSVTLKAVSAGDARGEAILSSD
TLTVPVSDTENTHIY

SEQ ID NO:31: CM homologue of CT043 = TC_0313 nucleotide sequence

ATGTCCAGACAGAATGCTGAGGAAAATCTAAAAAATTTTGTCTAAAGAGCTCAAGCTCCCCGACGTGGCCTTCGATCA
GAATAATACGTGCATTTTGTGTTGATGGAGAGTTTTCTCTTACCTGACCTACGAAGAGCACTCTGATCGCCTTT
ATGTTTACGCACCTCTCCTTGACGGACTCCAGATAATCCGCAAAGAAAGTTGGCTCTGTATGAGAAATTTGTTGGAA
GGCTCTATGCTCGGAGGCCAAATGGCTGGTGGAGGAGTAGGAGTTGCTACTAAAGAACAGTTGATCCTAATGCATTG
CGTGTTAGATATGAAATATGCAGAGACTAATCTATTGAAAGCTTTTGCACAGCTTTTCATTGAAACTGTTGTGAAAT
GGCGAACGGTCTGTTCTGATATCAGCGCTGGACGAGAACCTCCGTTGACACTATGCCCTCAAATGCCCTCAAGGAGGC
AGCGGAGGAATCAACCTCCTCCAACAGGAATTCGTGCGTAG

SEQ ID NO:32: CM homologue of CT043 = TC_0313 protein sequence

MSRQNAEENLKNFAKELKLPDVAFDQNNTCILFVDGEFSLHLTYEEHSDRLVYVAPLLDGLPDPNQRKLLALYEKLL
GSMLGGQMGAGGGVGVATKEQLILMHCVLDMKYAETNLLKAFQFLFIETVVKWRTVCSDISAGREPSVDTMPQMPQGG
SGGIQPPPTGIRA

SEQ ID NO:33: CM homologue of CT601 = TC_0890 nucleotide sequence

ATGCTCGCTAATCGGTTATTTCTAATCACCCCTATAGGTTTTGGCTATTCTGCTTACGGTGCCAGCACAGGGAAATC
ACCTTCTTTACAGGTTATTTAGCTGAAGTCGAGGATACATCTTCGCGTTACAAGCTCATCAGAATGAGCTTGTTA
TGCTCTCGGAACGTTTAGATGAGCAAGACACAAAACCTCAACAACCTCTCGTCAACTCAGGCCCGTAATCTTCCCTCAA
CAAGTTCAACGGCTTGAGATTGATCTGAGAGCTCTGGCTAAACAGCTGCTGTGCTCTCGCAATCTGTTTCAGGATAT
CCGATCATCCGTGCAAAAATAAATTACAAGAAATCCAACAAGAACAACAAAAAATTTAGCTCAAAATTTACGAGCGCTTC
GCAACTCCTTACAAGCACTAGTTGATGGCTCTTCCCGAGAAATATATTGATTTTTGGCCGGGAGACACCTGAA
CATATTCACGTTGTTAAACAAGGAGAAACCTGAGTAAATCGCTAGTAAAGTACAATATCCCTGTGCGAGAATTGAA
AAAACCTTAATAAATTAATTCGGATACTATTTTTACTGATCAAAGAATCCGACTTCCAAAAAAGAAATAA

SEQ ID NO:34: CM homologue of CT601 = TC_0890 protein sequence

MLANRFLITLIGFGYSAYGASTGKSPSLQVILAEVEDTSSRLQAHQNELVMLSERLDEQDTKLQQLSSTQARNLPQ
QVQRLEIDLRLAKTAAVLSQSVQDIRSSVQNKLEIQEQKNAQNLRALRNSLQALVDGSSPENYIDFLAGETPE
HIHVVKQGETLSKIASKYNIIPVAELKLNKLNLSDTIFTDQRIRLPKPK

SEQ ID NO:35: CM homologue of CT456 = TC_0741

ATGACGACTCCAATAAGTAATCTCCATCTTCTATTTCCAACCTGTTACAGTATCAACTACTACAGCATCTTCTGGATC
TCTCGGAACCTTCTACTGTATCATCAACGACTACAAGTACTTCAGTCGCACAAACAGCAACAACAACATCTTCTGCTT
CTACATCTATAAATCAGTCTAGTGGAGAAAACATCCAATCCAACACTACAGGTACCCCTTCTCCTATTACGCTTAGTGTT
TCAACATCCGCTCCATCTCTAAAGCCTCCGCACTGCAAAAACCTTCAAGCGCTGTTTCTGGGAAAATACCTC
ACAAGAACTTCTGAGGAATCCGAAACCCAAGCCACTACATCTGATGGAGAAGTTAGTAGTAATTAACGATGATGTTG
ATACCCCGACCAATTCGTCGATTGACAGTTGATAGTATTACCAAGATGTTGAGACTCAGTACAAAACAATTAGC
AACAAATGGTGAACACTTATGAAACAATCGGAAGTCAATGGTGGAGAAAACACACACGTCAGGAAAGCCATGCATC
CGGAACAGGAAATCCATAAATAATCAGCAAGAAGCTATTAGACAGCTCCGATCATCTACCTATAACAACAGCCCTC
GTAATGAGAATATATTAGTCCAGGACCGGAAGGCTACCTAATATGTCTCTTCTTAGTTACAGCCCTACAGATAAA
AGTTCTCTACTAGCTTCTATCTAATCCCAATACAAAAGCAAAAATGCTCGAACACTCCGGGCATTTAGTCTTTAT
AGACACAACCTAGAAGTAGCTTTATCTTTGTTCCGAATGGAAATGGGATCAAGTCTGTTCCATGAAGGTTGAGAATG
GGAAAACCTAAAGAAGACCTTGGCTTAAAGGACTTAGAAGATATGTGTGCAAAGTTTTGCACAGGATACAATAAATTC
TCCTCTGATTGGGGAAATCGAGTTGACCCCTTGGTCTCTTCAAGGCCGGGATAGAAAGTGGGGGGCACCTCCCAAG
CTCAGTTATCATCAACAACAATTTAGAACCTGTGTTGCCTATGGGCCGTGGAAACCCAAAGAAAACGGCCCCAATT
ATACTCCTTACGCCCTGGAGACGTGGGCATCGAGTAGATTTTGGAAAGATCTTTGATGGAAACAGCGCCGTTAATAAAA
ATCAACTGGGGCTCTTCCCTACCCCTGGTGTGACGGCATCTCCTTCTCTAATGAAACTATTGGGTCTGAACCATT
CGCGACACCTCCCTCATCCCATCGCAAACCCCGTTATCAACGTCAATGTTAATGTCGGTGGAAACCAATGTTAATA
TTGGGGATACAAACGATCTAAAGGATCCGGCACACCAACATCTTCTCAATCTGTGGACATGTCTACAGATACTAGC
GATTTAGATAACAGTGATATTGATACAAAACAACCAAACTAACCGCGATATCAACACGAATGACAACCTCAATAATGT
CGATGGAAAGTTTATCTGACGTTGATTCAAGGGTGGAAAGACGATGACGGTGTATCGGATACAGAGTCCACTAATGGCA
ATGACTCTGGTAAAACCTTCCACAGAAGAAAATGGTGAACCAAGCGGACAGACATCTGGCTGTCTGTAGTAAA
CACCTAGACACTGTCTATCCAGGAGAAAATGGCGGATCTACAGAAGGACCTCTCCCTGCTAATCAAAATCTGGGGAA
CGTTATCCATGATGTGGAGCAGAATGGATCTGCTAAAGAAACTATTATCACTCCAGGAGATACAGGGCCTACAGACT
CAAGCTCCTCTGTAGATGCTGATGCAGACGTTGAAGATACTTCTGATACTGACTCTGGAAATCGGAGACGACGACGGT
GTATCGGATACAGAGTCCACTAATGGTAATAACTCTGGTAAAACCTACTTCCACAGAAGAAAATGGTGAACCAAGCGG
ACCAGACATCCTGGCTGCTGTACGTAACACCTAGACACTGTCTATCCAGGAGAAAATGGCGGATCTACAGAAGGAC

CTCTCCCTGCTAATCAAAATCTGGGGAACGTTATCCATGATGTAGAACAAAACGGAGCCGCTCAAGAACTATTATC
 ACTCCAGGAGATACGGAATCTACAGACACAAGCTTAGTGTAAATGCTAATGCAGACTTAGAAGATGTTTCTGATGC
 TGATTCAGGATTCGGGGATGATGACGGTATATCGGTATACAGAGTCCACTAATGGTAAACGACTCTGGAAAAAATACTC
 CTGTAGGGGATGGTGGTACACCAAGCGGACAGATATCTTAGCTGTACGCAAAACATCTAGACACTGTCTATCCA
 GGAGAAAATGGTGGATCTACAGAGAGACCTTTACCCGCTAATCAAAATTTAGGAGATATCATTATGATGTAGAACA
 AAACGGAAGCGCTAAAGAACTGTAGTATCGCCTTATCGAGGAGGAGGAGGAAATACATCTTCCCAATTGGATTAG
 CCTCCCTGCTTCCAGCAACACCATCCACACCTTTGATGACAACACCTAGAACAAATGGGAAAGCTGCAGCTTCTTCT
 TTGATGATAAAAAGGAGGAGAACTCAAGCCAAGCTAGTTAAGAATGGCGGCAATATCCCTGGAGAAACCACATTAGC
 AGAATTACTCCCTCGTTTAAAGAGGACACCTTGACAAAGCTTTACTTTCAGACGGGAAGTTTACAAATCTTAATGGAC
 CTCAACTTGGAGCCATCATAGACCAATTCCGCAAAGAAACGGGTTCCGGAGGAATCATAGCTCATAAGATAGTGT
 CCAGGAGAGAACGGAACAGCCTCTCTCTCACAGGAAGTTCAGGGGAAAAAGTCTCTCTATGATGCAGCGAAAAA
 CGTCACTCAAGCTTTAAACAAGTGTACGAACAAAGTAACCCCTAGCAATGCAAGGACAAAAACTGGAAAGGAATTATAA
 ACAACAACAATAACCCCTCTTCTATTGGACAAAATCTTTTCGACGACGCGAGGGCAACGACACAATCCCTCAGTTCA
 TTAATTTGGAACCGTACAATAA

SEQ ID NO:36: CM homologue of CT456 = TC_0741 protein sequence

MTPPISNSPSSIIPTVTVSTTTASSGSLGTSTVSSSTTSTVSAQTATTTSSASTSIIQSSGENIQSTTGTSPITSSV
 STSAPSPKASATANKTSSAVSGKITSQETSEESETQATTS DGEVSSNYDDVDTPNTSSDSTVDSYQDVETQYKTIS
 NNGENTYETIGSHGEKNTHVQESHASGTGNPINNQEAIRQLRSSSTYTTSPRNENIFSPGPEGLPNMSLPSYSPTDK
 SLLAFLSNPNTKAKMLEHSGHLVFIIDTRSSSIFVPNGNWQVCSMKVQNGKTKEDLGLKDLMDCAKFCGTGYNKF
 SSDWGNRVDPLVSSKAGIESGGHLPSSVIINNKFRTCVAYGPWNPKEGPNYTPSAWRRGHRVDFGKIFDGTAPFNK
 INWGSSTPQDDGISFNETIGSEPFATPPSSPSQTPVINVNPNVGGTNNVIGDITVSKSGSPTSSQSVDMSTDT
 DLDTSDIDTNNQTNQDINTNDNSNNVDGSLSDVDSRVEDDDGVSDTESTNGNDSGKTTSTEENGDPSPDILAAVRK
 HLDTVYPGENGGSTEGPLPANQNLGNVIHDVEQNGSAKETIITPGDTGPTDSSSSVDADADVEDTSDTDSGIGDDG
 VSDTESTNGNNSGKTTSTEENGDPSPDILAAVRKHLDTVYPGENGGSTEGPLPANQNLGNVIHDVEQNGAAQETII
 TPGDTESTDTSSSVNANADLEDVSDADSGFGDDGISDTESTNGNDSGKNTPVGDGGTPSGPDILAAVRKHLDTVYP
 GENGGSTERPLPANQNLGDIHDVEQNGSAKETVVSYPYRGGGNTSSPIGLASLLPATPSTPLMTTPRTNGKAAASS
 LMIKGGETQAKLVKNGGNIPEGTTLAELLPRLRGHLKVFVTSQDGKFTNLNGPQLGAIIDQFRKETGSGGIIAHTDSV
 PGENGASPLTSSSGEKVSLYDAAKNVTQALTSVTKVTLAMQGRKLEGIINNNTPSSIGQNLFAAARATTQSLSS
 LIGTVQ

SEQ ID NO:37: CM homologue of CT381 = TC_0660

GTGAGTATGTATATAAAAAAGAAAGAAAGCTTGGATGACTTTCTTAGCAATTGTCTGTAGTTTCTGTTTGGCGGGCTG
 TTCAAAAAGAGAGCAAAGACTCTGTTAGTGAAAAATTTATTGTAGGAACTAACGCAACGTATCCTCCTTTTGAGTTTG
 TTGATGAAAAGAGGTGAGACGGTTGGCTTTGATATTGATTTAGCTAGGGAGATTAGTAAAAAGCTAGGGAAAAAATTA
 GAAGTCCGAGAATTTGCTTTTGGATGCACTCGTTCTCAATTTAAAACAGCATCGTATTGATGCAATTATGGCAGGGGT
 GTCCATTACGCTTCTCGATTGAAAGAAATTTGATGATTTCCCTACTATGGCGAAGAAATAAAGAGTTTGGTTTTAG
 TGTTTAAGGATGGAGACTCAAAGTCTTTACCACCTAGATCAGTATAAATTTCTGTTGCTGTTCAAACCTGGCACGTACCA
 GAGGAATATTTACAGTCTCTTCCAGGGGTGCGTATTGCTCTTTTGTAGTACTTTAGAAGTGCTTATGGAAGTTT
 GCATAGCAAGTCTCCTATAGCTGTTTTAGAACCCTCTATTGCGCAGGTGCTTTTAAAAGATTTTCCGACGCTCACTA
 CTGAAACGATAGATCTTCTGAAAGATAAATGGGTTTTAGGGTATGGAATTGGAGTTGCTTCTGATCGACCATCTCTA
 GCTTCTGATATAGAAGCTGCTGTACAAGAGATCAAGAAAGAAGGAGTGTAGCAGAGTTAGAGCAAAAATGGGGTTT
 GAACGGCTAA

SEQ ID NO:38: CM homologue of CT381 = TC_0660

MSMYIKRKKAWMTFLAIVCSFCLAGCSKESKDSVSEKFIIVGTNATYPPFEFVDERGETVGFIDDLAREISKKLKGLK
 EVREFAFDALVNLKQHRIDAIMAGVSISSRLKEILMIPYGGEEIKSLVLFKDGDSKSLPLDQYNSVAVQTGTYYQ
 EEYLQSLPGVIRISFDSTLEVLMEVLHKSPIAVLEPSIAQVVLKDFPTLTTEITDLPEDKQVLYGIGVASDRPSL
 ASDIEAAVQEIKKQVLALELQKWLNG

SEQ ID NO:39 – CT255 nucleotide sequence

ATGGAAGAAAAAGGCATCTTACAATTGGTTGAAATTTGCGGAGCAATGGCTTTACAGGGAGTTTGTCTTGGACTAA
 TTTACAGAGTGTGGAGTCTATGTTGCAGTATATAGCAGGGGAGTGTGAGGAGTTGGCTGATGCTGTACAAGAAAAATA
 AAGCTTTCGTTGGAAATCGCTTCGGAAGCCGGAGACGTACTTACTTTAGTATTGACCTTGTGTTTCTTGCTAGAAAAGA
 GAAGGAAAGCTTAAAGCTGAAGAAGTATTTGTAGAAGCTTTGGCTAAGTTGCGTCGATCTCCTCATGTTTTTGA
 TCCTCATAATCAAATTTCTTTAGAACAGGCTGAAGAACTGCGCTCGTATGAAACAGCAAGAAAAAATTTCTTAA

SEQ ID NO:40 – CT255 protein sequence

MEEKGILQLVEISRAMLQGVCPWNLQSVESMLQYIAGECQELADAVQENKASLEIASEAGDVLTLVLTLCFLLER
 EGKLAEEVFVEALAKLRRRSPHVFDPHNQISLEQAEYWARMKQKEKIS

SEQ ID NO:41 – CT341 nucleotide sequence

ATGGATTACTACAGTATTGGGTGTAGCGAAGACTGCTACTCCTGAAGAAATAAAGAAAGCTTACCGTAAGCTCGC
 TGTAAGTACCATCCAGATAAGAATCCTGGGGATGCTGAAGCGGAGCGACGCTTAAAGAAAGTTTCTGAAGCCTATG
 AAGTATTAGGTGATGCGCAGAAGCGGGAGTCATATGATCGTTACGGCAAAGACGGTCCATTTGCTGGTGTGGAGGA
 TTCGGTGGCGCTGGCATGGGGAATATGGAAGACGCTTTGCGAACATTTATGGGAGCTTTTGGCGCGCATTTCTGGTGG

TAATGGAGGCGGTTTTCTTTGAAGGGCTTTTTGGAGGACTTGGAGAAGCTTTGGAATGCGTGGAGGCTCAGAAAGTT
 CTCGACAAGGAGCTAGTAAGAAGGTGCATATTACGCTGTCTTCGAGGAGGCGGCAAAAGGTGTTGAAAAAGAAGTT
 CTTGTTTTAGGCTATAAATCTTGTGATGCTTGTCTGGTAGTGGAGCCAATACTGCTAAAGGTGTAAGGTTTGTGA
 TCGATGCAAGGGCTCTGGTCAGGTAGTGCAAAGCCGAGGCTTTTTCTCCATGGCTTCTACTTGCCTGATTGTAGTG
 GTGAAGGTCGGGTTATCACAGATCCTTGTTCAGTTTGTCTGGGACAGGACGTATCAAGGATAAACGTAGCGTCCAT
 GTTAATATCCCAGCTGGAGTCGATTCTGGGATGAGATTAAGATGGAAGGCTATGGAGATGCTGGCCAAAATGGAGC
 GCCTGCAGGGGATCTGTATGTTTTTATTGATGTAGAGCCTCATCTGTTTTCGAGCGCCATGGGGATGATTTAGTTT
 TAGAGCTTCTATTGGATTTGTTGATGCGGCTTTAGGGATCAAGAAGGAAATCCCTACACTCTTAAAAGAAGGTACT
 TGCCGTTTTGAGTATCCCAGAAGGGATTACAGAGCGGAACAGTTCTTAAAGTTAGAGGGCAGGGATTCCCTAATGTGCA
 TGGGAAATCCAGAGGAGATCTTTTAGTAAGAGTATCTGTGGAGACTCCCAGCACCTATCTAATGAACAAAAGATT
 TATTGAGACAGTTTGTCTGCTACGGGAAGGCTGAAAATTTCCCTAAGAAACGGAGTTTCTTAGACAAAATCAAAGGT
 TTTTTTTCTGACTTTGCTGTATAG

SEQ ID NO:42 – CT341 protein sequence

MDYYTILGVAKTATPEEIKKAYRKLAVKYHPDKNPGDAEAERRFKEVSEAYEVLGDAQKRESYDRYKDGPFAGAGG
 FGGAGMGNMEDALRTFMGAFGGDFGGNGGGFFGLFGLGEAFGMRGGSESSRQAGASKKVHITLSFEEAAKGVKEL
 LVSGYKSCDACS GSGANTAKGVKVC DRCKGSGQVVQSRGFFSMASCTPDCS GEGRVITDPCSVCRGQGRKDKRSVH
 VNIPAGVDSGMRLKMEGYGDAGQNGAPAGDLYVFDVPHVFERHGDDLVLLEPIGFVDAALGIKKEIPTLLKEGT
 CRLSIPEGIQSGTVLKVRRGQGFPNVHGKSRGDLVVRVSVETPQHL SNEQKDLLRQFAATEKAENFPKRSFLDKIKG
 FFSDFAV

SEQ ID NO:43 – CT716 nucleotide sequence

ATGAATAAAAAAAGCTCCAAGATCTGTCTAAACTGCTCACTATTGAGCTTTTCAAGAAACGTACACGGTTGGAAACAGT
 AAAAAAAGCGCTCTCCACAATAGAACATCGCTTACAACAAAATACAGGAGCACATCGCGAAAATTTCTTAAACAAGGC
 ACAAACAATTCCTATGTCGGTCATATACCCATGAATATGACCAACATTTAGAACAATTTACAAGAGAGCAAACTTCT
 CTATATAAACAGCATCAGACCCTGAAAACGTCTTTGAAAGATGCTTATGGCGACATACAAAAACAAGTACCAAAAG
 AAAAAATTATCGAAAAGATCCATGACAGTAAATATCTATAAAGAGCGCGAATAACTAA

SEQ ID NO:44 – CT716 protein sequence

MNKKLQDLSKLLTIELFKKRTRLETVKKALSTIEHRLQIQEHI AKISL TRHKQFLCRSYTHEYDQHLHLQREQTS
 LYKQHQTLKTSKDAYGDIQKQLDQRKIIEKIHDSKYPIKSANN

SEQ ID NO:45 – CT745 nucleotide sequence

ATGAAACATGCTCTCATTGTTGGCTCAGGTATTGCCGGCTTTCTGCCGCGTGGTGGCTACACAAACGATTCCCTCA
 TGTGCAGCTGTCTATTCTAGAAAAAGAGTCTCGATCTGGAGGGCTAATTGTACACAGAGAAACAACAAGGGTTTTCC
 TCAATATGGGCCCTAAAGGTTTTGTTTTAGCTCATGATGGGCAACACACCCTTACCTCATTAGCTTTAGGCCTA
 GCAGACGAGCTATTATATAGCTCTCCAGAGGCTAAAAACCGCTTTATCCACTATAATAATAAAACCCGAAAAGTCTC
 GCCTTGGACTATTTTCAAACAATACTCCCTCTCTTTTGTCTAAGGATTTCTTTGCGCGTCTTACAACAACAAGACA
 GCTCCGTGGAAAGCCTTCTTTAAAAGACACAGTCTTCCAAGCTTAGAAGAAATCTTTTAAATCCCATTAGCATTGCT
 ATTCGTGCAGGACATAGTCATATATTGTCTGCACAGATGGCTTACCCAGAATTAACACGAAGAGAAGCTCAAACAGG
 ATCGTTGTTACGTAGTTATCTCAAAGATTTTCTTAAAGAGAAACGCACAGGCCCTTATTTAGCTACCTTGGGCTCTG
 GGATGGGAATGCTAACCCAGGCTTTGCATGATAAATTGCCTGCTACCTGGTATTTTTCTGCACCCGTCAGCAAAATC
 GCTCAGTTGGCGAATGGGAAAATTTCTTTTCTCTTCTCAAGGAGAAATAACGGGAGATATGCTATTTATGCTGG
 CTCCGTGCACGATCTCCCTTCTGTCTAGAAGGGATCCCTGAAACCAAGCTTATCAAGCAACGACTTTCATCTTGGG
 ATCTCTCTTGTGTATCTTTAGGATGGCATGCATCCTTCCCTATCCCTCATGGATATGGCATGCTTTTCTGCTGATACG
 CCTCCCTTATTAGGGATCGTGTTTAAACGGAAGTGTCCCTCAACCCGAGCGGCTAATACAATAGTCTCTCTTCT
 TTTAGAAGGTGATGGCACCAAGAAGAAGCGTATGCTTTCTCACTAGCAGCTATTTCTGAGTACCTGCAAATTTACA
 CTCTCCCCAAGCTTTCTCACTATTCTCTCCTCGAGAGGGACTTCCCCAACACCATGTTGGATTTATCCAATCCCGC
 CAACGCCTTCTATCTAAACTTCTCACAATATAAAAAATTTGATGGGCAGAATTTTGCAGGTCAGGCTCTCAACCGCGC
 TACAGCGTCTGCTTATAAAGCTATAGCTTCTTTACTATCATGA

SEQ ID NO:46 – CT745 protein sequence

MKHALIVGSGIAGLSAAWHLKRFPHVQLSILEKESRSGGLIVTEKQQGFSLNMGPKGFVL AHDGQHTLHLIQSLGL
 ADELLYSSPEAKNRFIHYNNKTRKVSPWTFIKQNLPLSFAKDF FARPYKQDSSVEAFFKRHSSSKLRRNLLNPIA
 IRAGHSHILSAQMAYPELTRREAQTGSLRLSYLKD FPKKERTGPYLATLRSGMGLTQALHDKLPATWYF SAPVSKI
 RQLANGKISLSSPQGEITGDM LIYAGSVHDLPSCLEGIPETKLIKQTTSSWDLSCVSLGWHSFPIPHGYGMLFADT
 PPLLGI VFNTEVFPQPERPNTIVSLLLEGRWHQEEAYFSLAAISEYLQIYTPPQAFSLFSPREGLPQHHVGF IQSR
 QRLLSKLP HNIKIVGQNFAGPGLNRATASAYKAIASLLS

SEQ ID NO: 47 – CT387 nucleotide sequence

ATGACGCTCTTTTCATTCTCATCATGATGCCGCTCTCCAGACAGCTACCTATGTTCTTCCCTTTCAGTTAGTTGGTAC
 TGGCGTATACGAAGGAGAAATCGAGATTCAAAATATCCCTCTTATTTCCCTGGATTCCAATTACCTCTCATTGCA
 TACACCTTAATTTAAAGAGCTCTCTAGCTCAATTAAGGAATAGATGCTCCCTTCTCACTGCGAATTGAGCAAAAAT
 CAACATCGAGCACATATACATGCTCAATTTACCGGT CATGGCCCCATTGCTGAATCTATGCTAGCCCTTCTCCAACC
 AGGAGATCGTGTAGCAAAACTATTTGCTGCAGACGATCGCAGACTGGTCCGATCTCCAGATTACCTCGAAAGCATGC
 TGA AAAATACAGATAAAGCTGGCCATCCTTTGCTCTGTTTTGGGAAAAAATTAGAACA CTGATTTCTTTTGTATGTG

GTAGATGATCGCCTTGTCTGCTCCCTTCTACCTGCCGGGAGTTGTTGTTATGATTCGGATATTTATGGACTCCT
 TCCTCTTATTCAAAAATCACTCAGTAATCCCAAACCTCAGCATTTCGTCACCTTTTAGCTCTGTACCAACAGATTGTGG
 AAGGGCAACATGTCTCTTGGCGAAACCATATTTCTTCTGATCAAAAACAGAACCGCTGCACATCCGCACTGTATTTGCT
 CGCGTGGTAAATCAACTCCTCCCTCAAGGTCTCTCCACACTTCTGCCAATATTTTGGAAACCAACCACTCGAGAATC
 CGGGGATATCTTTGAATTTTTTGGGAACCCCTTCTGCACAGATAGAAAAGAATTCCTTTAGAATTTTTCACTATCGAAC
 CCTATAAAGAACATTCTTACTTCTGTAATCGGGATTTATTACAAAACATCTTACAATCAGAAAGCGAAATCAAAAA
 ATATTCGAAACAGCGCCAAAGAACCTGTCAAAGCTGCCACCTATTTATCAAAGGCGAGTGAATCTCTTCCCTGCA
 CACAGACTCTTGGCTCACAGGATCCGCGAGCTGCCTATCAATATAGTGAGCAAGCAGATAAAAAACGAGTACACTCATG
 CTCAACCTTGTCTCTTTCTTAGAAGCAATGGAAATGGGCCTGATCAATAGCGAAGGAGCCTTACTCACTCGTTAT
 TTCCTTTCAGCTAGCTTAAAAGGAATGTTGATTTCCATCCATGTGCGCCACTATCTCAAACAAATCTACTTTCAAGT
 TCCCTCTTATACACATGGAAACTATTTCTCTCATAATGACAGAGGTTTGCTATTAGATCTGCAGCAAGCAGATATTG
 ATGTTTTCTGGGCAGATGAAGAAAGCGGCCGTGTGTTGCAATATACAAAACGACGCGATAAGAATAGCGGTATGTTT
 GTGATCAAAAATCGTGTGGAAGAGTTTCGATCAGCTTATTTTATTGCTATTTATGGCTCTCGTCTCCTTGAGAATAA
 TTTCTCTGCTCAGCTCCATACCCTCCTAGCGGGCTTACAGCAAGCAGCACATACTCTCGGCATTCTGGATTCTCAA
 AGCCTACCCCACTTGCAGTCAACCCGAGGCGGCACTGGAGTTATGGCCACAGGAAATCGTGTAGCTAAAGAACTA
 GGAATCCTATCTTGTGGAACCGTTCTTGTATTAGAAAGCTTCTCCAGCACAAATCGACCAACTACCAATGAATCTT
 AGATGCTAAAATGACATACCGCCTACCTCAACTTATAGAAAAGCAAGAACACTTTTTATGCAGACCTTCTATCCTTG
 TAGTTGGCGGTGTAGGAACCGATTTTGAACCTTACCTAGAACTTGTCTATCTCAAAAACAGGAGCTAAACCACCGACT
 CCCATTTTCTAATTGGAACCTATTGAATACTGGAAAGAAAAGTGGCCACGCTACGAGATCAACCTCAAAGCAGG
 AACCATCCGTGGATCCGAATGGATCAGCAACTGCCATATTGTATCACTTCTCCGGAAGCTGGAATTTGCCGTATTCG
 AACAAATCCTAGCTGGAGAATCCCTATAGGATACGACTATCCTCCAGCTCCAGATGGATTAGTGATCGTCTAA

SEQ ID NO:48 – CT387 protein sequence

MTLFHSHHDAVSPDSYLCSSLQLVGTGVYEGEIEIQNIPSYFLGFQLPSHCIIHLNLKSSLAQLGIDASLLHCELSKN
 QHRAHIHAQFTGHGPIAESMLALLQPGDRVAKLFAADRRRLVLRSPDYLESMLKNTDKAGHPLLFCFGKLEHLISFDV
 VDDRLVVSPLPLPGVVRYDSDIYGLLPLIQKLSNPKLSIRHFLALYQQIVEGQHVSCGNHILLIKTEPLHIRTVFA
 RVVNQLLPQGLSHTSANILEPTTRESGDIFEFFGNPSAQIERIPLEFFTIEPYKEHSYFCNRDLLQILQSESEIKK
 IFETAPKEPVKAATYLSKGEISSLHTDSWLTGSAAYQYSEQADKNEYTHAQPYPFLEAMEMGLINSEGALLTRY
 FPSASLKGMLISYHVRHYLKQIYFQVPSYTHGNYFSHNDRGLLLDLQADIDVFWADEESGRVLQYTKRRDKNSGMF
 VIKNRVEEFRSAYFIAIYGSRLLENFSAQLHTLLAGLQQAHTLGIPGFSKPTPLAVITGGGTGVMATGNRVAKEL
 GILSCGTVLDEASPAQIDQPTNEFLDAKMTYRLPQLIERQEHFYADLPILVVGVGVDFFELYLELVYKTKGAKPPT
 PIFLIGPIEYWKQVAHAYEINLKAGTIRGSEWISNCLYCITSPEAGIAVFEQFLAGELPIGYDYPPAPDGLVIV

SEQ ID NO:49 – CT812 nucleotide sequence

ATGAGTCCGAGAAAGATATAAAAAGCACCTGTTCTAAGTTTTCTTTGCTGTAGTAGCAGCTATCCTTGCCCTCTGT
 TAGCGGGTTAGCTAGTTGCGTAGATCTTCATGCTGGAGGACAGTCTGTAAATGAGCTGGTATATGTAGGCCCTCAAG
 CGGTTTTATTGTTAGACCAAATTCGAGATCTATTCGTTGGGTCTAAAGATAGTCAGGCTGAAGGACAGTATAGGTTA
 ATTGTAGGAGATCCAAGTTCTTTCCAAGAGAAAGATGCGGATACTCTTCCCGGGAAGGTAGAGCAAAGTACTTTGTT
 CTCAGTAACCAATCCCGTGGTTTTCCAAGGTGTGGACCAACAGGATCAAGTCTCTTCCAAGGGTTAATTTGTAGTT
 TTACGAGCAGCAACCTTGATTCTCCTCGTGACGGAGAATCTTTTTAGGTTATTGCTTTTTGTTGGGGATAGTAGTAA
 GCTGGAATCACATTAAGTGAAGCTTCTTTGCTGAGCGGCTTTTATTTCTACAGAAGATCTTATCTTTGA
 AAAGATTAAGGGTGGATTGGAATTTGCATCATGTTCTTCTCTAGAACAGGGGGAGCTTGTGCAGCTCAAAGTATTT
 TGATTCATGATTGTCAAGGATTGCAGGTTAAACACTGTACTACAGCCGTGAATGCTGAGGGGTCTAGTGCGAATGAT
 CATCTTGGATTTGGAGGAGGCGCTTTCTTTGTTACGGGTTCTTTTCTGGAGAGAAAAGTCTCTATATGCCCTGCAGG
 AGATATGGTAGTTGCGAATTTGTATGGGGCTATATCTTTTGAAGGAAACAGCGCGAAGCTTGTAAATGGAGGAGCGA
 TTGCTGCCTCTGGGAAAGTGTCTTTTGTGCTAATGATAAAAAGACTTCTTTTATAGAGAACCGAGCTTTGTCTGGA
 GGAGCGATTGCAGCCTCTTCTGATATTGCCTTTCAAACCTGCGCAGAAGTATTTTCAAAGGCAATTTGTGCAATTTGG
 AACAGAGGATAAAGGTTCTTTAGGTGGAGGGGCTATATCTTCTCTAGGCACCGTCTTTTTGCAAGGGAATCACGGGA
 TAACTTTGTGATAAAGATGAGTCTGCTTCCGAAGGAGGCGCCATTTTGGCAAAAATTTGTAGATTTCTGACAACGAG
 GGGCCAGTGGTTTTCAAGATAGTACAGCTTGTCTTAGGAGGAGGCGCTATTGCAAGCTCAAGAAATTTGTTCTATTCA
 GAACAATCAGGCTGGGATTTCTTTCGAGGGAGGTAAGGCTAGTTTTCGGAGGAGGTAATGCGTGTGGATCTTTTTCTT
 CCGCAGGTGGTGCCTTCTGTTTTAGGGACCATGATATTTGAAAGATTTAGGCGGATTTCTGTTCTCTCGTACTTTA
 TGTAAGCCTCAGATTTAGGACAAATGGAGTACCAGGGAGGAGGAGCTCTATTTGGTGAATAATTTCTTTCTGTA
 GAATGCTGGTGTGCTCACCTTAAAGACAACATTTGTGAAGACTTTTGTCTCGAATGGGAAAATTTCTGGGAGGAGGAG
 CGATTTTAGCTACTGGTAAGGTGGAATTAATAAATTTCCGAAGGAATTTCTTTTACAGGAAATGCGAGAGCTCCA
 CAAGCTCTTCAAACCAAGAGGAGTTTCTTTTATTGCAAAAAGAAGGGCGACCACTCTCTTCAAGGATTTCTGG
 GGGAGGAGCGATTTTAGGAAGAGAAGTAGCTATTTCTCAACAACGCTGCAGTAGTATTTGAGCAAAAATCGTTTGCAGT
 GCAGCGAAGAAGAAGCGACATTATTAGGTTGTTGTGGAGGAGGCGCTGTTTCATGGGATGGATAGCACTTTCGATTGTT
 GGCAACTCTTCAAGTAAAGATTTGGTAATAATTACGCAATGGGACAAGGAGTCTCAGGAGGAGCTTTTTATCTAAAAC
 AGTGCAGTTAGCTGGGAATGGAAGCGTCGATTTTTCTGAAATATTGCTAGTTTGGGAGGAGGAGCTTTCAAGCTT
 TGAAGGAAATTTGAGACTAGTTGATAACGGCTATGCTGATTCAGAGATAAATCGAGGGAGGTTCTTTGGGGTGCCT
 ATTTCTTGTCTACGTGGAGATGTAGTCAATTTCTGGAAAACAAGGGTAGAGTTGAATTTAAAGACAACATAGCAACACG
 TCTTTATGTGGAAGAACTGTAGAAAAGGTTGAAGAGGTAGAGCCAGCTCCTGAGCAAAAAGACAATAATGAGCTTT
 CTTTCTTAGGGAGAGCAGAACAGAGTTTTATTACTGCAGCTAATCAAGCTCTTTTCTGCATCTGAAGATGGGGATTTA
 TCACCTGAGTCACTCAATTTCTTCTGAAGAAGTGGCAAAAAGAGAGTGTGCTGGAGGAGCTATTTTTGCCAAAACG
 GGTTCTGATTTGTAGATAAAGAGGCGTTGTTGTTCTGAAATCTTCTGATATTTATGGCGGCGCCATTTTAA
 CAGGTTCTTCTGAGAAAGAGGATAAGTTAGATGGGCAAAATCCTGAAAGTCTTGATCTCAGGCAATGCAGGGGATGTT
 GTTTTTTCCGGAATTTCTCGAAGCGTGATGAGCATCTTCTCATAACAGGTGGGGGAGCCATTTGTACTCAAAAATTT

GACGATTTCTCAGAATACAGGGAATGTTCTGTTTTATAACAACGTGGCCTGTTCTGGGAGGAGCTGTTCTGATAGAGG
 ATCATGGTAATGTTCTTTTGAAGCTTTTGGAGGAGATATTGTTTTAAAGGAAATCTTCTTTTCAGAGCACAAAGGA
 TCCGATGCTATCTATTTTGCAGGTAAGAATCGCATATTACAGCCCTGAATGCTACGGAAAGGACATGCTATTGTTTT
 CCACGACGCATTAGTTTTGAAAATCTAGAAGAAAGGAAATCTGCTGAAGTATTGTTAATCAATAGTCGAGAAAATC
 CAGGTTACACTGGATCTATTTCGATTTTTAGAAGCAGAAAGTAAAGTTCTCAATGTATTTCATGTACAACAAGGAAGC
 CTTGAGTTGCTAAATGGAGCCACATTATGTAGTTATGGTTTTAAACAAGATGCTGGAGCTAAGTTGGTATTGGCTGC
 TGGAGCTAAACTGAAGATTTAGATTTCAGGAACCTCCTGTACAACAAGGGCATGCTATCAGTAAACCTGAAGCAGAAA
 TCGAGTCATCTTCTGAACCAGAGGGTGCACATTCTCTTTGGATTGCGAAGAATGCTCAAAACAACAGTTCTATGGTT
 GATATCCATACTATTTCTGTAGATTTAGCCTCCTTCTCTTCTAGTCAACAGGAGGGGACAGTAGAAGCTCCTCAGGT
 TATTGTTCTCTGGAGGAAGTTATGTTTCGATCTGGAGAGCTTAATTTGGAGTTAGTTAAACAACAAGGTAAGTTATG
 AAAATCATGCTTTTATTGAAGAATGAGGCTAAAGTTCCATTGATGTCTTTCTGTTGCTTCTGGTGTGAAGCTTCAGCC
 GAAATCAGTAACTTGTGGTTTTCTGATTTACAGATTCATGTAGTAACTCCAGAGATTGAAGAAGACACATACGGCCA
 TATGGGAGATTGGTCTGAGGCTAAAATTCAGATGGAACCTCTTGTCAATTAGTTGGAATCCTACTGGATATCGATTAG
 ATCCTCAAAAAGCAGGGGCTTTAGTATTTAATGCATTATGGGAAGAAGGGGCTGTCTTGTCTGCTCTGAAAAATGCA
 CGCTTTGCTCATAATCTCACTGCTCAGCGTATGGAATTCGATTATTCTACAATGTGTGGGGATTCCCTTTGGTGG
 TTTCCGAACCTCTATCTGCAGAGAATCTGGTTGCTATTGATGGATACAAGGAGCTTATGGTGGTCTTCTGCTGGAG
 TCGATATTCAATTGATGGAAAGATTTTGTCTAGGAGTTAGTGGAGCTGCTTTCTAGGTAAAATGGATAGTCAGAAG
 TTTGATGCGGAGGTTTCTCGGAAGGGAGTTGTTGGTTCTGTATATACAGGATTTTTAGCTGGATCCTGGTTCTTCAA
 AGGACAATATAGCCTTGGAGAAACACAGAACGATATGAAAACGCGTTATGGAGTACTAGGAGAGTTCGAGTGCCTTCT
 GGACATCTCGAGGAGTACTGGCAGATGCTTTAGTTGAATACCGAAGTTTGTGGTCTGTGAGACCACTTTTAT
 GATTTGCTTTCAATCTTATGTCTGAAGTATCTTATGCTTCTGATGAAATTCCTGGCTTTACAGAACCAAGGAGAGA
 AGCGCGTTCTTTTGAAGACGCTTCCCTTACCAATATCACCATTCTTTAGGGATGAAGTTTGAATTGGCGTTCTATAA
 AAGGACAGTTTTTCAGAGGTGAACCTCTTTGGGAATAAGTTATGCATGGGAAGCTTATCGAAAAGTAGAAGGAGGCGCG
 GTGCAGCTTTTGAAGCTGGGTTTATTGGGAGGGAGCTCAATGGATCTTCTAGACAGGAGCTGCGTGTGCTCT
 GGAAAATAATACGGAATGGAGTTCTTACTTTCAGCACAGTCTTAGGATTAACAGCTTTTTGTGGAGGATTTACTTCTA
 CAGATAGTAAACTAGGATATGAGGCGAATACTGGATTGCGATTGATCTTTTAA

SEQ ID NO:50 – CT812 protein sequence

MSEKDIKSTCSKFSLSVVAAILASVSVGLASCVDLHAGGQSVNELVYVGPQAVLLLQIRDLFVGSKDSQAEQYRL
 IVGDPSSFQEKDADTLPGKVEQSTLFSVTNPVVFQVVDQDDQVSSQGLICSFTSSNLDSPRDGESFLGIAFVGDSSK
 AGITLTDVKASLSGAALYSTEDLIFEKIKGGLEFASCSLLEQGGACAAQSSILIHDCQGLQVKHCTTAVNAEGSSAND
 HLGFGGGAFFVTGSLSGEKSLYMPAGDMVVANCDGAISEFEGNSANFANGGAIASGKLVFVANDKKTFSFIENRALS
 GAIAASSDIAFQNCALVFKGNCAIGTEDKGSLLGGAISSLLGTVLLQGNHGITCDKNESASQGGAIQKNCQISDNE
 GPVVFRDSTACLGGGAIAAQEIVSIQNNQAGISFEGGKASFGGIACGSFSSAGGASVLTIDISKNLGAISFRTL
 CTTSDLGQMEYQGGGALFGENISLSENAGVLTFKDNIVKTFASNGKILGGGAILATGKVEITNNSEGISFTGNARAP
 QALPTQEEFPLFSKKEGRPLSSGYSGGGAILGREVALHNAAVVFEQNRLLQCEEEATLLGCCGGGAVHGMDSSTIV
 GNSSVRFGNYYAMQGVSGGALLSKTVQLAGNGSVDFSRNIASLGGGALQASEGNCELVDNGYVLFDRNDRGRVYGG
 ISCLRGDVVISGNKGRVFEKDNIAATRLYEETVEKVEEVEPAPEKQDNNELSFLGRAEQSFITAANQALFASDGL
 SPESSISSEELAKRRECAAGAIQAKRVRIVDNQEAUVFSSNNFSDIYGGAIFTGSLREEDKLDGQIPEVLISGNAGDV
 VFSGNSSKRDEHLPHTTGGGAICTQNLTISQNTGNVLFYNNVACSGGAVRIEDHGNVLEAFGGDIVFKGNSSFRAG
 SDAIYFAGKESHITALNATEGHAIVFHDALVFENLEERKSAEVLLINSRENPGYTGSRIFLEAESKVPQCIHVQGG
 LLELNGATLCSYGFQKQDAGAKLVLAAGAKLKILDSGTPVQGGHAIKPEAEIESSEPEGAHSLWIAKNAQTTPMV
 DIHTISVDLSTGCGATTGGTTATGCGAGTCCCAATCTCCTACCVNTGGAGATTCTGTGATACAATAGGTCTGTAATCT
 TTGAAAATAACTTGTGTCAGACTATTTACATGGAGAAATCCTTATGCTGCTGATAAAAATAAGAGAAGGCGGAGCC
 ATTCATGCTCAAAATCTTACATAAATCATAATCATGATGTGGTGGGATTTATGAAGAACTTTTCTTATGTCCAAGG
 AGGAGCCATTAGTACCCTAATACCTTTGTTGTGAGCGAGAATCAGTCTTGTCTTCTTTATGGACAACATCTGTA
 TTCAAACATAACAGCAGGAAAAGGTGGCGCTATCTATGCTGGAACGAGCAATCTTTTGGAGAGTAATAACTGCGAT
 CTCTTCTTCAATAACGCCTGTTGTGCAGGAGGAGCGATCTTCTCCCTATCTGTTCTTAACAGGAAATCGTGG
 TAACATCGTTTTCTATAACAATCGCTGCTTTAAAAATGTAGAAAACAGCTTCTTCAAGAACTCTGATGGAGGACAA
 TTAAGTAACACTACTCGCCTAGATGTTACAGGCAATCGTGGTAGGATCTTTTTTAGTGACAATATCAAAAAAATTAT
 GCGGGAGCTATTTACGCTCCTGTAGTTACCCTAGTGGATAATGGCCCTACCTACTTTATAACAATATCGCCAATAA
 TAAGGGGGGGCGCTATCTATATAGACGGAACAGTAACTCCAAAATTTCTGCCGACCGCCATGCTATTTATTTTAAATG
 AAAATATTGTGACTAATGTAACATAATGCAATGGTACCAGTACGTACGTAACTCCTCCTAGAAGAAATGCAATAACA
 TAGCAAGCTCCTCTGTAAGAAATCTATTAGGAGCAGGAGTAGCCAAAATTTAATTTTTTATGATCCTATTGAAGT
 TAGCAATGCAGGGTCTCTGTGCTTCAATAAGGAAGCTGATCAACAGGCTCTGTAGTATTTTTCAGGAGCTACTG
 TTAATTTCTGCAGATTTTCATCAACGCAATTTACAAAACAAAACACCTGCACCCCTACTCTCAGTAATGGTTTTCTA

SEQ ID NO:51 – CT869 nucleotide sequence

ATGAAAAAAGCGTTTTTCTTTTCTTATCGGAAACTCCCTATCAGGACTAGCTAGAGAGGTTCTTCTAGAATCTT
 TCTTATGCCCCAAGTCCAGATCCTACGAAAGAGTGCCTATCAAATAAAATAGTTTTGACAGGAGACACTCACA
 ATCTCACTAACTGCTATCTCGATAACCTACGCTACATACTGGCTATTCTACAAAAAATCCCAATGAAGGAGCTGCT
 GTCAACAATAACAGATTACCTAAGCTTTTTGATACACAAAAAGAAGGTATTTATTTTGCAAAAAATCTCACCCCTGA
 AAGTGGTGGTGGATTGGTTATGCGAGTCCCAATCTCCTACCVNTGGAGATTCTGTGATACAATAGGTCTGTAATCT
 TTGAAAATAACTTGTGTCAGACTATTTACATGGAGAAATCCTTATGCTGCTGATAAAAATAAGAGAAGGCGGAGCC
 ATTCATGCTCAAAATCTTACATAAATCATAATCATGATGTGGTGGGATTTATGAAGAACTTTTCTTATGTCCAAGG
 AGGAGCCATTAGTACCCTAATACCTTTGTTGTGAGCGAGAATCAGTCTTGTCTTCTTTATGGACAACATCTGTA
 TTCAAACATAACAGCAGGAAAAGGTGGCGCTATCTATGCTGGAACGAGCAATCTTTTGGAGAGTAATAACTGCGAT
 CTCTTCTTCAATAACGCCTGTTGTGCAGGAGGAGCGATCTTCTCCCTATCTGTTCTTAACAGGAAATCGTGG
 TAACATCGTTTTCTATAACAATCGCTGCTTTAAAAATGTAGAAAACAGCTTCTTCAAGAACTCTGATGGAGGACAA
 TTAAGTAACACTACTCGCCTAGATGTTACAGGCAATCGTGGTAGGATCTTTTTTAGTGACAATATCAAAAAAATTAT
 GCGGGAGCTATTTACGCTCCTGTAGTTACCCTAGTGGATAATGGCCCTACCTACTTTATAACAATATCGCCAATAA
 TAAGGGGGGGCGCTATCTATATAGACGGAACAGTAACTCCAAAATTTCTGCCGACCGCCATGCTATTTATTTTAAATG
 AAAATATTGTGACTAATGTAACATAATGCAATGGTACCAGTACGTACGTAACTCCTCCTAGAAGAAATGCAATAACA
 TAGCAAGCTCCTCTGTAAGAAATCTATTAGGAGCAGGAGTAGCCAAAATTTAATTTTTTATGATCCTATTGAAGT
 TAGCAATGCAGGGTCTCTGTGCTTCAATAAGGAAGCTGATCAACAGGCTCTGTAGTATTTTTCAGGAGCTACTG
 TTAATTTCTGCAGATTTTCATCAACGCAATTTACAAAACAAAACACCTGCACCCCTACTCTCAGTAATGGTTTTCTA

TGTATCGAAGATCATGCTCAGCTTACAGTGAATCGATTACACACAACTGGGGGTGTTGTTTCTCTTGGGAATGGAGC
 AGTTCTGAGTTGCTATAAAAAATGGTACAGGAGATTCTGCTAGCAATGCCTCTATAACACTGAAGCATATTGGATTGA
 ATCTTTCTTCCATTCTGAAAAGTGGTGTGAGATTCTTTTATTGTGGGTAGAGCCTACAAATAACAGCAATAACTAT
 ACAGCAGATACTGCAGCTACCTTTTATTAAGTGTGTAATACTCTACTCATTGATGACTACGGGAACCTCTCCTTA
 TGAATCCACAGATCTGACCCATGCTCTGTATCAGCAGCCTATGCTATCTATTTCTGAAGCTAGCGATAACCCAGCTAC
 AATCAGAAAATATAGATTTTTTCGGGACTAAATGTCCCTCATTATGGATGGCAAGGACTTTGGACTTGGGGCTGGGCA
 AAAACTCAAGATCCAGAACAGCATCTTCAGCAACAATCACTGATCCACAAAAAGCCAATAGATTTTCATAGAACCTT
 ACTACTAACATGGCTTCTGCGGGTATGTTCCCTAGCCCAAAACACAGAAGTCCCCCATAGCTAACACCTTATGGG
 GGAATATGCTGCTTGCACAGAAAGCTTAAAAAATAGTGCAGAGCTGACACCTAGTGGTCACTCTTCTGGGGAATT
 ACAGGAGGAGGACTAGGCATGATGGTTTACCAAGATCCTCGAGAAAATCATCCTGGATTCCATATGCGCTCTTCCGG
 ATACTCTGCGGGGATGATAGCAGGGCAGACACACACCTTCTCATTGAAATTCAGTCAGACCTACACCAAATCAATG
 AGCGTTACGCAAAAAACAACGTATCTTCTAAAAATTACTCATGCCAAGGAGAAATGCTCTTCTCATTGCAAGAAGGT
 TTCTTGCTGACTAAATAGTTGGGCTTTACAGCTATGGAGACCATAACTGTCCACATTTCTATACTCAAGGAGAAAA
 TCTAACATCTCAAGGGACGTTCCGCGAGTCAAACGATGGGAGGTGCTGTCTTTTTTGATCTCCCTATGAAACCTTTG
 GATCAACGCATATACTGACAGCTCCCTTTTTAGGTGCTCTTGGTATTTATTCTAGCCTGTCTCACTTTACTGAGGTG
 GGAGCCTATCCGCGAAGCTTTTCTACAAAGACTCCTTTGATCAATGCTAGTCCCTATTGGAGTTAAAGGTAGCTT
 TATGAATGCTACCCACAGACCTCAAGCCTGGACTGTAGAATTGGCATAACCAACCGTTCTGTATAGACAAGAACCAG
 GGATCGCAGCCAGCTCCTAGCCAGTAAGGGTATTTGGTTCCGTAGTGGAAAGCCCCCATCGCGTCAATGCCATGTCC
 TATAAAATCTCACAGCAAAACACAACCTTTGAGTTGGTTAACTCTCCATTTCCAGTATCATGGATTCTACTCCTCTT
 AACCTTCTGTAATTCATCAATGGGGAATTTGCTCTGCGATTCTAG

SEQ ID NO:52 – CT869 protein sequence

MKKAFFFFLIGNSLSGLAREVPSRIFLMPNSVDPDKESLSNKISLTDGTHNLNLCYLDNLRYLAILQKTPNEGAA
 VTITDYLFFDTQKEGIYFAKNTLPESGGAIGYASPNSTVEIRDITGPVIFENNTCCRLFTWRNPYAADKIREGGA
 IHAQNLINHNHDVVGFMKNFSYVQGGAIISTANTFVSENQSCFLFMDNICIQNTAGKGGAIYAGTSNSFESNND
 LFFINNACCAGGAFSPICSLTGNRGNIVFYNNRCKFNVETASSEASDGGAIKVTTRLDVTGNRGRIFSDNITKNY
 GGAIYAPVVTLDVNGPTYFINNIANNKGGAIYIDGTSNSKISADRHAIIFNENIVTNVTNANGTSTSNPPRRNAIT
 VASSSGEILLCGGSSQNLIFYDPIEVSNAGVSVFNKEADQTSVVFSGATVNSADFHQRNLRQTKTAPPLTSLNGFL
 CIEDHAQLTVNRFQTGGVVSLGNGAVLSCYKNGTGDASNASITLKHIGLNLSSILKSGAEIPLLVPEPTNNSNNY
 TADTAATFSLSDVKLSLIDYDGNPSESTDLTHALSSQPMLSISEASDNQLQSENIQDFSLNVPHYGWQGLWTWGW
 KTQDPEPASSATITDPQKANRFHRTLLLTLWLPAGYVPSPKHRSPLIANTLWGNMLLATESLKNSEALTPSGHPFWGI
 TGGGLGMMVYQDPRENHPGFHMRSSGYSAGMIAGQTHFSLKFSQTYTKLNERYAKNNVSSKNYSQGMFLSLQEG
 FLLTKLVGLYSYGDNHCHHFFYTQGENLTSQGTFRQDLPMPKPFGSTHILTAFLGALGIYSSLSHFTEV
 GAYPRSFSTKTPLINVLVPIGVKGSFMNATHRPQAWTVELAYQPVLYRQEPGIAAQLLASKGIWFGSGSPSSRHAMS
 YKISQQTQPLSWLTLHFQYHGFYSSSTFCNYLNGEIALRF

SEQ ID NO:53 – CT166 nucleotide sequence

GTGAACGTTTCGTACTCTGTTTCAGAGGGGGGGGTA AAAACGATTTCTGCTAGTGCAGTTCCTCCTACAGCAGC
 TGTTTTATCGAGAAAAAGCGTGTCTATAGAAGAGAAGAAGGAGGAAGCTTCTTCTGGAAAGATAGAAAATCTTGATG
 CTAGCAAAATACGATCTTACTCCCAAGAACATAGAAGAAAAACTAGGAATTACTCCTGAACAGAAAATCTACTGTTAAA
 GACCTATTAATAAACTGAAAAAGGTCATTAGTGTCTACAACCTCTATGCCAGATAAAAAATTCGGAAGCGGGACAGAA
 TTCCTTGATTCAACAAGGAAAATACGTCGATGCCATTGAGAAGAAGCTTCCAGCATCATCGCAGGCTCAGCCTAAAC
 AGGCAAAAGCTAAGGAACAGAAAAGCCGAAGAAAAACCTAAGACGACTCCGATTGAAGGTGTTCTTGA AACCATCAA
 ACAGAAATTAAGGCCATCGTGTACCTGTTGAGAAAAATCATCCATGGAATATGGATCGCAGGAGCGCTCCGGATGG
 TATCGAAGATTATATGCGAGTCTTTTTAGATACTTATGAAGGTTTTGACTTCTACTTCTGGGTAGATGAGAATGCTT
 ATGCAGCAGCTAAATTTTCTAGCATTTTGAAGAAGGTCGCTTTCGATGCGGCTATTCAAGATCTACGATCTGCCACA
 GATGAGTCTACGAAGGCTTTTGTAAAGACTACGATGAATTA AAAACAGAAAATGAAAAGAAAAGTTGCGGAGACGAC
 TTCTCAAGCAGAAAAAGACCAATATCTCAAAGATCTAAAGGATCTTTTAGAGAAAATTTACAAAAATCAGTGATGAGA
 TTCGTGGAAAATTTGATCGGCTGTTTCTTAAGAATGTGATTGTTGCTCAGAACGGATTCTTTAATTTCTGCTTGTG
 AAAGGCTCTGGCAATATCAATGACGAAAACCGCTGCAGAGATTATAGAGAAAAGAACTCAAACCTTCTACTGAGGAGAT
 CGAACAGTATAAAAAAGCTTAAAGAGACGAAACAAAAGAGAGATAGCCGCTATTGTA AAAACAACTAAACAGAAACTTG
 GATCGGATCGGGTAAAAATCAAAGACATTAAGAGCTGCAATCTATGAAGCAAGCTCGAAATGTCTACAATTAAGAA
 CAGGAAAATGTTTCTGCGCTGGAACATATGCAGCCGCAACAGATCAGATTGCTATGATATGTTGGAGGAACTTGGAGG
 TCTTTATACTGATCTGGATATGATGCCTTATACTCTCAGGAAGTATTGGAGCTTATCAAAAAGCACAGTGATGGAA
 ACCGAATGTTTGGAGATATGAGCTCTAGACGGGCGATTTCTGATGCGGTTTTAAAGATGGCTGTAGGTAAGGCGACA
 ACAGTTTTCCATGGAAGAGGTAGCAAAGGATATCGATGTTTCTCGCTTAACAGAAGAGGATAAGACAAAATTAATGCT
 TCTATTTAAGGATCTAGAGCCATTTGCAAAACCGGATTCTAAAGGAGCTGAAGCAGAAGGGGGTGAAGGAGCAAAAAG
 GTATGAAAAAGAGCTTTTTCCAGCCCATAGATCTGAATATTGTGAGAAAATACCATGCCTATCTTGAGACGCTATCAT
 CACTATCCTGAGTTAGGATGGTTTTATTGAGGATTGAACGGATTGATGGTCTCTATAAGGGAAGCACTGCGGTTTC
 TGCTGTCAATTGAGGGCAACAGGCTGCCTACCAGGAATCAGCAGCACTTAGACAAGATGTCCTTTACAGGGAGTTTT
 TCCATCTTTAGAAAATTTGACACATAGAAAACCAATAAGGACGATTTGGAAAATCATCTCGCTCAGTTATTTGGCT
 AAAAGTCTCTTTTTGATTACTGCCAAGATTCAGTGATGCCGAGGCTGTAAGTACCTTAGGTATTAGATGA

SEQ ID NO: 54 – CT166 protein sequence

MNVRTYSVQRRGVKTIASAVPPTAAVL SRKKRAIEEKKEEASSGKIENLDASKYDLTPKNIEEKLGITPEQKSTVK
 DLLNKLKKVISAYNSMPDKNSEAGQNSLIQDQKGYVDAIQKKLPASSQAQPKQAKAKEQKAEKPKTPIEGVLETIK
 TEFKGRHVPVEKIIHGIWIAGAPPDGIEDYMRVFLDTYEGDFYFVWVDENAYAAKFSSILKKVAFDAAIQDLRSAT
 DESTKAFVKDYDELKQKYEKVAETTSQAEDQYLDKDLLEKFTKISDEIRGKFDRLFLKNVIVAQNGFFNFCLL

KGLGNINDETRAELYLEKELKLPTEEIEQYKCLKETNKEKIAAIVKQLNEKLGSDRVKIKDIKELQSMKQARNVYNYE
QEMFLRWNYAAATDQIRMYMLEELGGLYTDLDMMPSYSQEVLELIKHSDDGNRMFEDMSSRRAISDAVLKMAVGKAT
TVSMEEVAKDIDVSRLEEDKTKLNLALFKDLEPFKPKDQSKGAEAEQGGEGAKGMKKSFFQPIDLNIIVRNTMPLRRYH
HYPELGFIRGLNGLMVSHKGSTAVSAVIVGQQAAYQELAAALRQDVLSEFFHSLLENLTHRNHKERIGNHLVANYLA
KSLFFDYCQDSVMPEAVSTLGIR

SEQ ID NO:55 – CT175 nucleotide sequence

ATGCATCACAGGAAGTTTTAGCAGTTTTCCATTGCTTTCGTAAGTTTTAGCTTTTGGGCTAACATCTTGTATCATAA
AAAAGAAGAACCAAAAGATGTTTTGCGGATTGCGATCTGTATGATCCAATGTCTTTAGATCCGCGTCAGGTTTTTT
TAAGCAAAGATGTTTTCTATTGTAAGCTCTCTATGAAGGGTTAGTCCGGGAAAAAGAAAGCTGCGTTCCAGCTAGCT
TTGGCAGAAAGATATCATCAATCTGATGATGTTGTGTTTTATACTTTTTTTTCAAAAAATACATTCTGGAGCAACGG
AGATGTTGTAACAGCATATGATTTTTGAAGAGTCTATTAACAAATTTATTTCCGAGAAATTGATAACCCCTCGTTAC
GCTCTCTTGCATTAATTAATAATTCTCATGCTGTTTTAACAGGAGCTCTCCCTGTTGAAGATTTAGGTGTTAGAGCT
TTGAATGCGAAAACCTAGAAAATTGTTTTAGAAAACCCGTTTTCTTATTTTCTAGAGATATTGGCGCACCCGGTTTT
TTATCCGGTGCACACCTCTTTACGAGAATATTACAAAGATAAGCGTAACAAACGCGTTTTTCCCGATAATTTCTAATG
GTCTTTTTGCGATTCAATGTTATGAGCCGCAAGATATTTACTAATCAACAAAAACCTCTGTATCATGCCAAGCAC
GATGTTCTGTTAAATTCGGTATGTTTTGCAGATAGTTCCTGATATCCATACAGCTATGCAAGTTATTTCAAAAAATCA
TATCGATTTAGTTGGGTTACCTGGAGCTCCTCCTTTTTCTTTAGAAGAACAAGAAATCTCCCTAGAGAAAAATTA
TTGATTATCCTGTATTGAGTTGCTCTGTTTTATTCTGTAACATTCATCAAACACCTTTAAATAATCCCTCGCTGAGA
ACAGCCCTCTCTTTAGCAATCAATCGAGAACTTTATTAAGACTAGCAGGTAAGGCTGTAGCGCTACGAGCTTTGT
TCACCCACAATTAATCTCAGATACTTACTACTTTGCTCAAGATGAGCGGATTGCTTTAGCAAAAGGCTACTTGA
CCGAAGCTTTAAAGACTTTATCTCAAGAAGATTTAGAAAAAATACATTAATTTATCCTATAGAATCTGTTTGTCTTA
CGAGCCGTTGTTCAAGAAATTCGCAACAATTAATTTGATGTAAGTGGGATTTAAAAATTTCTACATTAGGATTAGAATA
TCATTGTTTTTTAGACAAACGTTCCAGAGGAGAATTTCTCCTTAGCAACTGGTAATTGGATTGCAGACTATCATCAAG
CTAGTGCTTTCTGCTGCTCCTAGGTAATGGGACAAGATATAAAGACTTTCAATTGATTAAGTGGCAGAACCAAAAG
TACACAAATATAGTTGCTCAACTTCTGATTCAAGAATCAAGCGACCTACAGCTTATGGCAGAGCAGTTGTTGCTTAA
AGAAAGTCTCTTATCTCTATACCACCTCGATTAATGTGTATGCGAAACAGCCCTCGGGTGTCTGATCTCCAAACCT
CTTCTCGTGGAGAAATGATTTAAAAAGAGTTTCATTAGCTGAAGGATAG

SEQ ID NO:56 – CT175 protein sequence

MHRKFLAVSIAFVSLAFGLTSCYHKKEEPKDLVLRIAICHDPMSLDPRQVFLSKDVSIVKALYEGLVREKEAAFQLA
LAERYHQSDDGCVYFFLKNFTWSNGDVVTAAYDFEESIKQIYFREIDNPSLRSLALIKNSHAVLTYGALPVEDLGVRA
LNAKTLEIVLENPFYFLEILAHVPVFPVHTSLREYKDKRNRVFPFIISNGPFAIQCYEPQRYLLINKNPLYHAKH
DVLNLSVCLQIVPDIHTAMQLFQKNHIDLVLGWPWSSSFLEEQRNLPREKLFYDYPVLSCSVLFQNIHQTPLNPSLR
TALSLAINRETLKLAGKGSATSFVHPQLSIPATTLSDERIALAKGYL TEALKTLSQEDLEKITLIYPIESVCL
RAVVQEIQRQLFDVLGFKISTLGLLEYHCFLDKRSRGEFSLATGNWIADYHQASAFLSVLGNGTRYKDFQLINWQNZK
YTNIVAQLLIQESSDLQLMAEQLLLKESPLIPLYHLDYVYAKQPRVSDLQTSRGEIDLKRVSLAEG

SEQ ID NO: 57 – TC0666 nucleotide sequence (homologue of CT387)

ATGAGGATCCAAATGACACTCTTTCACACTCATCAGATGCCGTCTCTCCGGACGGCTACTTATGTTCTTCCCTTCA
GTTAGTTGGCTCTGGCACATATGAAGGAGAAATCGAAATCCAAAATATTCCTTCTTATTTCTTGGATTCCGATTAC
CCACCCATTGCGTTTCACTTAATTTGAAGAGTTCTCTAGCCCAGTTAGGAGTAGATGCATCTCTTCTTCACTGCGAA
CTAAGCAAAAATCAACAACGTGCACATATGCACGTGCAGTTCACCCGGCTATGGCCCTATCGCTGAGTCCATGCTATC
TCTTCTCAAACCCGGAGATCGAGTAGCCAAACTGTTTGTGTCAGATGATCGTAGACTAGTCCGCTCCCTGATTATC
TTGAAAGCATGCTAAAAAATCTGATAAGACAGGACATCTCTGCTCCGATTTGGAAAAAATCTCGAGCATTTATC
TCTTTTGTAGTGGTGGACGATCGCTCGTTGTACTCCCCACTTGGCAGGCATAGTCAATTAAGACCCAGACAT
CTATGGACTTCTTCCCTTAATTCAAAAATCACTAAGCAATCCTAAATTTAGTATTTCGCCACTTCTTGTCTCTCTATC
AGAAGATCGTAGAAGGACACACATCCCTTATGAAGGAAACATTTTGTAAATCAAAAACAGAGCCTCTTCATATCCGC
ACAGTATTTGCTCGCGTGGTTCGATCAAATGCTCCCTCAAGGTCTATTTTACACTTCTGCCAACATTTTAGAACCAC
AACCGGAGAGTCTGGAGATATTTTTGAATTTTTTGGAAATCCCTCCACTCTGTAGAAAAGAAATCCCTCTAGAATTCT
TCACTATCGAACCCCTACAAAAGAACACTCTTACTTCTGTAATCGAGATCTATTGCAAACTACCTTGAATCGGAAAGT
GAAATCAAAAAAATATTGATACAGCTCCTCAAGAGCCTGTAAGAGCCGCCACTTATTTATCAAAAAGGAAGTGAAT
TTCTTCTCTTGATGCAGATTCTTGGCTTACGGGATCCGCAGCTGCATACCAATGTAGCGAAAAACAGGCAGCTAAAG
ACGAATACATCCACGCTCAACCCCTGTTATCCATTTTGGAAAGCAATGGAAACGGGACTCATCAATAGCGAAGGAGCT
TTACTCACTCGGTTTTTCCCTCTTCCAGCTTAAAAAGGGATGTTGATCTCTCATCATGTACGCCACTATCTTAAGCA
AATTTACTTTCAAGTCTCTTATACATATGGAGACTCTTCTCATAATGACCGAGGATTAATCTGTTAGATCTAT
ATCAGGCGAACATTGATGTGTTCTGGGCTGATGAAGAGAGCGGCCGTGATTGCAATATACAAAAACGGCGCGACAAA
AATAGTGGAAATGTTCTGCTGTTAAAAATCGAGTAGAAGAGTTCGAATCAGCATATTTCTGAGCGATTTATGGATCACG
TCTCCTGGAAAAATAATTTCTCGGCCCACTAAACAGCTTCTTGCAGGGTTACAAAAAGCTGCACACACTCTAGGCA
TTCCAGGCTTCTCAAAAACCACTCCTCTTGCCTGAATCACAGGAGGAGGGACTGGCGTTATGGCTACAGGAAATCGT
GTTGCAAAAAGATTTGGGAATTTCTTGGGGACCGTTCTCGATTTGGAAAGCTTACCTGCACAAAATAGATCAGCC
TGCACAAAGAAATTTTAGATGCCAAAATGACATACCGTCTACCCGCACTTATAGAAAAGACAAGAACATTTTTTACG
ACCTTGCCATTTTGTGTTGGTGGTGTGGAAACAGATTTGCAACTTTACCTAGAACTCGTCTACTTGGAAACAGGC
GCCAAACCTCCTACTCCAATTTTCTTATTTGGGCTGTTGAATACTGGAAAGAGAAAGTTGCTCATGCCTATGAGAT
TAATCTTAAAGCAGGAACTATTCGTGGTTCTGAGTGGATCAGCAACTGCTTATTTCTGCATTACATCTCCTGAAGCAG
GAATTGCTGTATTGCAACAGTTCCTCGCTGGAGAACTCCCATAGGATATGATTATCTCCAGCTCCAGACGGATTA
GTTATCGTCTAA

SEQ ID NO: 58 - TC0666 protein sequence (homologue of CT387)

MRIPMTLFHTHHDVAVSPDGYLCSSSLQLVGSGETYEGEIEIQNIPSYFLGFRLPTHCVHLNLKSSLAQLGVDASLLHCE
 LSKNQQRRAHMHVQFTGYGPIAESMLSLLKPGDRVAKLFAADDRRLVLRSPDYLESMLKNTDKTGHPLLRFGKLEHLI
 SFDVVDDRLVLSPLPLPGIVNYDDPIYGLPLIQLKSLSNPKLSIRHFLSLYQKIVEGPHIPYEGNILLIKTEPLHIR
 TVFARVVDQMLPQGLFHTSANILEPTTRESGDIFEFFGNPSTLVERIPLEFFFTIEPYKEHSYFCNRDLLQTTLQSES
 EIKKIFDTAPQEPVKAATYLSKGSEISSLDADSWLTGSAAYQCEKQAAKDEYIHAQPCYPFLEAMETGLINSEGA
 LLTRFFPSSSLKGM LISYVHRHYLKQIYFQVPSYTYG DYF SHNDRGLLLDLYQANIDVFWADEESGRVLQYTKRRDK
 NSGMFVVKNRVEEFQSAFYVAIYGSRLLENFSAQLNLLAGLQKAAHTLGIPGFSKPTPLAVITGGGTGVMATGNR
 VAKELGILSCGTVDLEASPAQIDQPANEFLDAKMTRYLPQLIERQEHFYSDLAILVGGVGTDFELYLELVYLKTG
 AKPPTPIFLIGPVEYWKQVAHAYEINLKAGTIRGSEWISNCLFCITSPEAGIAVFEQFLAGELPIGYDYPPAPDGL
 VIIV

SEQ ID NO: 59 - TC0197 nucleotide sequence

ATGAGTTCGGAGAAAGATAAAAAAAAAACTCCTGTTCTAAGTTTTCTTATCGGTAGTAGCAGCTATTCTCGCTTCTAT
 GAGTGGTTTTATCGAATTGTTCCGATCTTTATGCCGTAGGAAGTTCTGCAGACCATCCTGCCTACTTGATTCTCAAG
 CGGGTTATTATGGATCATATTAAGGATATATTCATTGGCCCTAAAGATAGTCAGGATAAGGGGCAGTATAAGTTG
 ATTATTGGTGAGGCTGGCTCTTTCCAAGATAGTAATGCAGAGACTCTTCTCAAAAAGGTAGAGCACAGCACTTTGTT
 TTCAGTTACAACACCTATAATTGTGCAAGGAATAGATCAACAAGATCAGGTCTCTTCGCAGGGATTGGTCTGTAATT
 TTTCAGGAGATCATTAGAGGAGATTTTTGAGAGAGAATCCTTTTTAGGGATCGCTTTCTAGGGGAATGGTAGCAAG
 GATGGAATCACGTTAACAGATATAAAATCTTCGTTATCTGGTGCTGCCTTGTATTCTCAGATGATCTTATTTTTGA
 AAGAATTAAGGGAGATATAGAGCTTTCTCTTGTTCATCTTTAGAAAGAGGAGGAGCTTGTTCAGCTCAAAGTATTT
 TAATTCATGATTGTCAAGGATTAACGGTAAAACATTGTGCCGCGAGGGGTGAATGTTGAAGGAGTTAGTGTAGCGAC
 CATCTCGGATTTGGGGCGGGGCTTCTCTACTACAAGTTCTCTTTCTGGAGAGAAGAGTTTGTATATGCCTGCAGG
 CGATATTGTGGTGGCTACCTGCGATGGTCTGTGTGTTTGAAGGAAATAGTGCTCAGTTAGCAAATGGTGGCGCTA
 TTGCCGCTTCTGGTAAAGTTCTTTTTGTAGCTAACGAAAAAAGATTTCTTTACAGACAACCAAGCTTTGTCTGGA
 GGAGCTATTTCTGCATCTTCTAGTATTTCTTTCAAAAATGTGCTGAGCTTGTGTTCAAGAGTAATCTTGCAAAAGG
 GTTAAGATAAAATGTTCTTTGGGAGGAGGTGCTTTAGCCTCTTTAGAATCCGTAGTTTTGAAGGAGTATCTCGGTA
 TTACTTATGAAAAAATCAGTCTTATTCCGGAAGGAGGGGCTATTTTTGGGAAGGATTGTGAGATTTTTGAAAAAGG
 GGGCTGTTGTATTAGAGATAATACAGCTGCTTTAGGAGGCGGAGCTATTTTTGGCGCAACAACTGTGGCGATTTG
 TGGTAATAAGTCTGGAATATCTTTTGAAGGAAGTAAGTCTAGTTTTGGAGGGGCCATTGCTTGTGGAAATTTCTCTT
 CTGAGATAAATTTCTCAGCTTTGGGATCAATTGATATCTTAACAATCTAGGAGATATCTTTTTCTTCGGACTCTG
 TGTACTACTTCGGATTAGGGCAAAACGGATTACCAAGGGGAGGGCCCTTATTTCGCTGAAAAATTTCTCTTTGTA
 GAATGCTGGTGCAATTAATTCGCTCAAAGACAATATTGTGAAGACATTTGCCTCAAATGGAATAATGTTGGGTGGAGGGG
 CAATTTTAGCTTCAGGAAATGTTTTGATTAGCAAAAACCTCTGGAGAGATTTCTTTTGTAGGGAATGCTCGAGCTCCT
 CAGGCTATTCCGACTCGTTCATCTGACGAATTGCTTTTTGGCGCACAATTAACCTCAAACCTACTTCAGGATGTTCTGG
 AGGAGGAGCTCTTTTTGGTAAAGAGGTTGCCATTGTTCAAAAATGCCACTGTTGATTTCGAGCAAAAATCGCTTACAGT
 GTGGCGAGCAGGAAACACATGGTGGAGGCGGTGCTGTTTATGGTATGGAGAGTGCCTCTATTATTGAAACTCTTTT
 TGGAGATTCGGAAATAATTCGCTGTAGGGAATCAGATTTCTGGAGGAGCTCTTTTATCCAAGAAGGTCCTTTAGC
 TGAATAACAAGGGTAGATTTTTCTCGAAATATCGCTACTTTCTGCGGCGGGGCTGTTCAAGTTTCTGATGGAAGTT
 GCGAATTGATCAACAAATGGGTATGTGCTATTAGAGATAACCGAGGGCAGACATTTGGTGGGGCTATTTCTTGCTTG
 AAAGGAGATGTGATCATTTCCGGAATAAAGATAGGGTTGAGTTTAGAGATAACATTGTGACCGCGGCTTATTTTGA
 AGAAAATGAAGAAAAAGTTGAGACAGCAGATATTAATTCAGATAAGCAAGAAGCAGAAGAGCGCTCTTTATTAGAGA
 ACATTTGAGCAGAGCTTTATTACTGCAACTAATCAGACCTTTTTCTTAGAGGAAGAGAAAACCTCCATCAGAAGCTTTT
 ATCTCTGCTGAAGAACTTTCAAAGAGAAGAGAATGTGCTGGTGGGGCGATTTTTGCAAAAACGGGTCTACATTACGGA
 TAATAAAGAACCCTATCTGTTTTCTGCATAATTTTTCTGATGTTTATGGGGGAGCTATTTTTACGGGTTCTCTACAGG
 AAACCTGATAACAAGATGTTGTAACCTCTGAAGTTGTGATATCAGGCAACGATGGGGATGTCATTTTTCTGGAAT
 GCAGCTAAACATGATAAGCATTACCTGATACAGGTGGTGGAGCCATTTGTACACAGAATTTGACGATTTCCCAAAA
 ACATGGGAATGTCTGTTCTTGAACAATTTGCTGTTCTGGTGGAGCAGTTTCGCTATAGAGGATTCAGGAAAGTTCT
 TTTTAGAGGCTTTTGGGGGAGATATTAATTTCAATGGAACCTCTTCTTTAGAGCTCAAGGATCGGATGCGATCTAT
 TTTGCTGGTAAGGACTCTAGAATTAAGCTTTAAATGCTACTGAAGGACATGCGATTGTGTTCCAAGATGCATTGGT
 GTTTGAAAAATAGAAAGAAAGAAAGTCTTCGGGACTATTGGTGATTAACCTCTCAGGAAAAATGAGGGTTATACGGGAT
 CCGTCCGATTTTTAGGATCTGAAAGTAAGGTTCCCTCAATGGATTATGTGCAACAGGGAGGTCTTGAGTTGCTACAT
 GGAGCTATTTTTATGTAGTTATGGGGTTAAACAAGATCCTAGAGCTAAAATAGTATTATCTGCTGGATCTAAAATTGAA
 GATTCTAGATTCAGAGCAAGAAAAATAACGCAGAAAATTGGAGATCTTGAAGATTCTGTTAATTCAGAAAAAACCAAT
 CTCTTTGGATTGGGAAGAACGCTCAAGCAAAAGTCCCTCTGGTTGATATCCATACTATTTCTATTGATTTAGCATCA
 TTTTCTTCTAAAGCTCAGGAAACCCCTGAGGAAGCTCCACAAGTCATCGTCCCTAAGGGAAGTTGTGTCCACTCGGG
 AGAGTTAAGTTGGAGTTGGTTAATAACAACAGGAAAAAGTTAGATGAGAATCATGCGTTGTTAAAAAATGATACTCAGG
 TTTCTCTATGCTCTTCAAAGAGGAAAAATGATGGATCTTTAGAAGATTTGAGTAAGTTGTCTGTTTCGGAATTTACGC
 ATTAAGGTTCTACTCCAGATATTGTAGAAGAAACTTATGGCCATATGGGGGATTTGGTCTGAAGTCAAAATCAAGA
 TGGGGCTCTTGTCAATTAATTGGCATCCTACTGGATATAAATTAGATCCGCAAAAAGCTGGTTCTTTGGTATTCAATG
 CATTATGGGAGGAAGAGGCTGTATTGTCTACTCTAAAAAATGCTCGGATTGCCATAACCTTACCATTACAGAGAATG
 GAATTTGATTATCTACAAATGCTTGGGGATTAGCTTTTAGTAGCTTTAGAGAGCTATCTTCAGAGAAGCTTGTTC
 TGTTGATGGAATAGAGGCTCTTATATAGGGGCTTCTGCAGGCAATTTGATACTCAGTTGATGGAAGATTTTGTGTTGG
 GAATCAGCAGGCTTCTTCTTCCGGGAAAAATGCAATAGTCAGAATTTTGTGATGCAGAGATTTCTCGACATGGTTTTGT
 GTTTCGGTCTATACAGGCTTCTAGCTGGGGCTGGTTCTTCAAGGGGCGTACAGTCTTGGCGAAACACATAACGA
 TATGACAACCTCGTTACGGGGTTTTGGGAGAATCTAATGCTACTTTGGAAGTCTCGAGGAGTACTAGCAGATGCTTTAG
 TTGAATATCGTAGTTTAGTCGGTCCAGCACGACCTAAATTTTATGCTTTGCATTTAATCCTTATGTGAGGATCT

TATGCATCTGCGAAGTTCCCTAGTTTTGTAGAACAAGGAGGAGAAGCTCGTGCTTTTGAAGAAACCTCTTTAACAAA
CATTACCGTTCCCTTTGGTATGAAATTTGAACTATCTTTTACAAAAGGACAGTTTTTCAGAGACTAATTCTCTGGAA
TAGGTTGTGCATGGGAAATGTATCGGAAAGTCGAAAGGAGATCTGTAGAGCTACTAGAAGCTGGTTTTGATTGGGAA
GGATCTCCTATAGATCTCCCTAAACAAGAGCTGAGAGTGGCTTTAGAAAACAATACGGAATGGAGTTTCTATTTTAG
TACAGCTCTAGGATAACAGCATTGTTGTGGAGGATTTCTTCTATGGATAATAAACTAGGATACGAAGCGAATGCTG
GAATGCGTTTATTCTAG

SEQ ID NO: 60 - TC0197 protein sequence

MSSEKDKKNSCSKFSLSVVAAILASMSGLSNCSLDYAVGSSADHPAYLIPQAGLLLDHIKDFIFGPKDSQDKGQYKL
IIGEAGSFQDSNAETLPQKVEHSTLFSVTPPIIVQGDQDDQVSSQGLVCNFSGDHSEEIFERESFLGIAFLNGSK
DGITLTDIKSSLSGAALYSSDDLIFERIKGDIELSSCSSLERGGACSAQSILIHDCQGLTVKHCAAGVNVEGVSASD
HLGFGGGAFSTTSSLSGKSLYMPAGDIVVATCDGPVCFEGNSAQLANGGAIASGKVLVFNKISFTDNQALS
GAISASSSISFQNCALVFKSNLAKGVKDKCSLGGGALASLESVVLKDNLGITYEKNQSYSEGGAIKDKCEIFENR
GPVVFRDNTAALGGGAILAQQTVAICGNKSGISFEFGSKSSFGGAIACGNFSSENNSALGSDIDISNNGDISFLRTL
CTTSDLGQTDYQGGGALFAENISLSENAGAITFKDNIVKTFASNGKMLGGGAILASGNVLISKNSGEISFVGNARAP
QAIPTRSSDELQFQAQLTQTTSGCSGGGALFGKEVAIVQNAIVVFEQNRQLQCGEQETHGGGGAVYGMESASIIIGNSF
VRFGNYYAVGNQISGALLSKKVRLEAENTRVDFSRNIATFCGGAVQVSDGSCLELNNGYVLFDRDNRGQTFGGASL
KGDVVIISGNKDRVEFRDNIVTRPYFEENEKVEADINSQKQEAEEERSLLENIEQSFITATNQTFLEEEKLPSEAF
ISAEELSKRRECAGGAIFAKRVYITDNKEPILFSHNFSQVYGGAIFTGSLQETDKQDVVTPPEVVISGNDGDIVFSGN
AAKHDKHLPDTGGGAICTQNLTISQNNGNVLFNNFACSGGAVRIEDHGEVLEAFGGDIIFNGNSSFRAQGSDAIY
FAGKDSRIKALNATEGHAIQVFDALVFENIEERKSSGLLVINSQENEGYTGSRVFLGSESKVPQWIHVQGGLELLH
GAILCSYGVKQDPRAKIVLSAGSKLKILDSEQENNAEIGDLEDSVNSEKTPSLWIGKNAQAKVPLVDIHTISIDLAS
FSSKAQETPEEAPQVIVPKGSCVHSGELSLELVNNTTGKGYENHALLKNDTQVSLMSFKEENDGSLDLSKLSVSLR
IKVSTPDIVEETYGHMGDWSEATIQDQALVINWHPGTGYKLDQKAGSLVFNALWEEAVLSTLKNARIAHNLTIQRM
EFDYSTNAWGLAFSSFRELSSEKLVSDGYRGSYIGASAGIDTQLMEDFVLGISTASFFGKMHSQNFDAEISRHFV
GSVYTGFLAGAWFFKQYSLGETHNDMTRYGVLGESNATWKSRLVADALVEYRSLVGPAPKPFYALHFNPHYEVS
YASAKFSPFVEQGGEARAFEETSLTNITVPFGMKFELSFTKQFSETNSLGIGCAWEMYRKVEGRSVELLEAGFDWE
GSPIDLPKQELRVALENTEWSSYFSTALGVTAFCGGFSSMDNKLGYEANAGMRLIF

SEQ ID NO: 61 - TC0261 nucleotide sequence

ATGAAAAAAGCTGTTCTTTTTTGTCTTATTGGAAGCTCTATACTGGGATTTACTCGAGAAGTCCCTCCTTCGATTCT
TTTAAAGCCTATACTAAATCCATACCATATGACCGGGTATTTTTTCCCAAGGTAATTTGCTTGGAGACACACATA
ATCTCACGATTACCATTTGGATAATCTAAATGCATTCTGGCTTGCCTACAAAGAACTCCTTATGAAGGAGCTGCT
TTCACAGTAACCGATTACTTAGGTTTTTCAGATACACAAAAGGATGGTATTTTTTGTAAAAATCTTACTCCAGA
TGCAGTGGAGGGTTATTGGTTCCCAACTCAAAACACTCTACTATAAAAAATTCATAATAACAATCGGACCTCCTTT
TCGAAAAATAATACCTGTACATAGACTGTGGACACAGACCGATCCCGAAAATGAAGGAAAACAAGCACGCGAAGGCGGG
GCAATTCATGCTGGGACGTTTACATAAGCAATAACAGAACCTTGTGCGATTCAATAAGAACTTTGCTTATGTTCA
AGGTGGAGCTATTAGTGTAACTTTTTGCTATAAAAGAAAATAAATCGAGCTTTCTTTGCTAAATAACTCTTGTA
TACAAAATAAGACGGGAGGAAAAGGTGGTGTATTTACGTTAGTACGAGCTGCTCTTTCGAGAACAATAACAAGGAT
CTGCTTTTCATCCAAAACCTCCGGCTGTGCGAGGAGGACTATCTTCTCCTCAACCTGTTCTCTAATAGGAAACCAAG
AGATATTTGTTTTTACAGCAACCACGGTTTTAAAAATGTTGATAATGCAACTAACGAATCTGGGGATGGAGGAGCTA
TTAAAGTAACCTACCGCTTGGACATCACCAATAATGGTAGTCAAACTTTTTTTCTGATAATATCTCAAGAAATTTT
GGAGGAGCTATTGATGCTCCTTGTCTTATCTTGTGGTAATGGGCCAACCTATTTTACAAACAATATAGCTAATCA
CACAGGTGGGCTATTTATATAACAGGAACAGAAAACCTCAAAGATTTCTGCAGATCACCATGCTATTTTGTGATA
ATAAATTTCTGCAAAACGCCACCAATGCGGACGGATCTAGCAGCAACAATACTCTCCTCACAGAAATGCGATCACT
ATGGACAATTCGCTGGAGGAATAGAACTTGGTGCAGGGAAGAGCCAGAATCTTATTTCTATGATCCTATTTCAAGT
GACGAATGCTGGAGTTACCGTAGACTTCAATAAGGATGCCTCCCAAACCGGATGTGTAGTTTTCTCTGGAGCGACTG
TCCTTTCTGCAGATATTTCTCAGGCTAATTTGCAAACTAAAACACCTGCAACGCTTACTCTCAGTCAACGGTCTTCTG
TGTATCGAAGATCGTGCTCAGCTCACAGTGAACAAATTTACACAAAACAGGAGGGATTGTAGCCTTAGGAAATGGAGC
AGTTTTAAGCAGCTACCAACACAGCACTACAGACGCCACTCAAACTCCCTTACAAACCACTACAGATGCTTCCG
TAACTCTTAATCACATTGGATTAATCTCCCTCTATTCTTAAGGATGGAGCAGAGATGCCTCTATTATGGGTAGAA
CCTATAAGCACAACTCAAGGTAACACTACAACATATACGTGAGATACCGCGGCTTCTTCTCATTAAATGGAGCCAC
ACTCTCTCATTGATGAAGATGGAAATTTCTCCCTATGAAAACACGGACCTCTCTGCTGATTGTACGCTCAACCTA
TGCTAGCAATTTCTGAGGCCAGTGATAACCAATTTGCAATCCGAAAGCATGGACTTTTCTAAAGTTAATGTTCCCTCAC
TATGGATGGCAAGGACTTTGGACCTGGGGGTGGGCAAAAACCTGAAAATCCAAACAACAACCTCCTCCAGCAACAATTAC
TGATCCGAAAAAAGCTAATCAGTTTATAGAACTTTATTATTAACGTTGGCTCCCTGCTGGTTATATCCCAGCCCTA
AACATAAAAGCCCTTAAATAGCTAATACTTGTGGGGGAATATACTTTTTGCAACGGAAAACCTTAAAAAATAGCTCA
GGGCAAGAATCTTGTATGCTCTTTCTGGGAAATACAGGAGGGGGCTGGGGATGATGGTCTATCAAGAACCTAG
AAAAGACCATCTGGATTCCACATGCATACCTCCGGATATTCAGCAGGAATGATTACAGGAAACACACATACCTTCT
CATTACGATTCAGCCAGTCTTACAAAACCTCAATGAACGTTATGCCAAGAATATGTGTCTTCAAAAATTTACTCT
TGCCAAGGGGAAATGCTTTTGTCTTACAGAAGGACTATGCTGACTAACTAAATTTGGTCTCTATAGTTATGGGAA
TCACAACAGCCACCATTTCTATAACCAAGGAGAAGACCTATCGTCTCAAGGGGAGTTCCATAGTCAGACTTTTGGAG
GGGCTGTCTTTTTTGTATCTACCTCTGAAACCTTTTGAAGAACACACATACTTACAGCTCCTTTCTTAGGTGCCATT
GGTATGTATTCTAAGCTGTCTAGCTTTACAGAAGTAGGAGCCTATCCAAGAACCTTTATTACAGAAAACGCTTTAAT
CAATGCTCTGATTTCTATCGGAGTAAAAGGTAGCTTATGAATGCCACCCATAGACCTCAGGCTGGACTGTAGAGC
TTGTTTACCAACCTGTTCTTTACAGACAAGAACCTAGTATCTTCAACCAATTTACTCGCTGGCTAAAGGATGTGGTTT
GGGCATGGAAGTCTGCATCTCGCCACGCTCTAGCTTATAAAATTTACAGAAAACACAGCTTTTGGGATTTGCAAC

ACTTCAACTCCAGTATCACGGATACTATTTCGTCTTCCACTTTCTGTAATTATCTGAATGGAGAGGTATCTTTACGTT
TCTAA

SEQ ID NO: 62 - TC0261 protein sequence

MKKLFFFVLIGSSILGFTREVPSSILLKPIILNPYHMTGLFFPKVNLGLDTHNLTDYHLNLIKILACLQRTPEGAA
FTVTDYLGFSDTQKDGIFCFKNLTPESGGVIGSPTQNTPTIKIHNTIGPVLFENNTCHRLWTQTPENEGNKAREGG
AIHAGDVYISNNQNLVGFINKFAYVQGGAISANTFAYKENKSSFLCLNNSCIQTKTGGKGGAIYVSTSCSFENNKD
LLFIQNSGCAGGAIFSPTCSLIGNQGDIVFYSNHGFKVNDNATNESGDDGGAIKVTRLDITNNGSQIFFSDNISRN
GGAIHAPCLHLVGNPTYFTNNIANHTGGAIYITGTEYSKISADHHAIIIFDNNISANATNADGSSSNTNPPHRNAIT
MDNSAGGIELGAGKSQNLIFYDPIQVVTNAGVTVDFNKQASQTGCVVFSGATVLSADISQANLQTKTPATLTLSHGLL
CIEDRAQLTVNNFTQTTGGIVALGNLSSYQHSSTDATQTPPTTTTTDASVTLNHIGLNLPSILKDGAEMLLWVE
PISTTQGNNTTYSDTAASFSLNGATLSLIDEDGNSPYENTDLRSLALYAQPLAISEASDNQLQSESMDFSKVNVPH
YQWQGLWTGWAKTENPTTTPPATITDPKANQFHRLLTQLPAGYIPSPKHKSPLIANTLWGNILFATENLKNSS
GQELLDRPFWGITGGGLGMMVYQEPKRDHPGFHMHTSGYSAGMITGNHTFSLRFSQSYTKLNERYAKNYVSSKNYS
CQGEMLLSLQEGMLTKLIGLSYGNHNSHHFYTQGEDLSSQGEFHSQTFGGAVFFDLPLKPFGRTHILTAPFLGAI
GMYSKLSSTFEVGAYPRTFITETPLINVLIPIGVKGSFMNATHRPQAWTVELAYQPVLYRQEPSISTQLLAGKGMWF
GHGSPASRHALAYKISQKTQLLRFATLQLQYHGYSSSTFCNYLNGEVSLRF

SEQ ID NO: 63 - CT733 fragment nucleotide sequence

GCACCTCAACCTCGCGGAACGCTTCCCTAGCTCGACCACAAAAATTGGATCAGAAGTTTGGATTGAACAAAAAGTCCG
CCAATATCCAGAGCTTTTATGGTTAGTAGAGCCGCTCTACGGGAGCCTCTTAAATCTCCTTCAGGAGCCATCT
TTTCTCCAACATTATTCCAAAAAAGGTCCCTGCTTTCGATATCGCAGTGCAGTGGCAGTTTGATTCACTTACATTTATTA
ATCCAGGGTTTCCCGCAAGCCTATGCTCAACTGATCAACTACAGACCAGCGAATCCCTCTAACATTTAAGCAATT
CCTTGCAATTGCATAAGCAATTAAGTCTATTTTTAAATCCCTAAGGAATTTTATGACTCTGTTAAAGTGTAGAGA
CAGCTATCGTCTTACGTCACTTAGGCTGTTCAACTAAGGCTGTTGCTGCGTTTTAAACCTTATTTCTCAGAAATGCAA
AGAGAGGCTTTTTACACTAAGGCTCTGCATGACTACACACCTTCCAGAGCTAAGCCCATCATTTGCTCGCCTCTC
TCCGGAGCAGAAAACTCTCTTCTCTCTTGGAGAAAAATTGGCGAATTACGATGAGTTACTCTCGCTGACGAACACCC
CAAGTTTTCAGCTTCTGTCTGCTGGGCGCTCGCAACGAGCTCTTTTAGCTCTGGACTGTACTCTATGCTTTGGAT
TCTCTGTGGAGAACAGGGGATGCTCTCTCAATCCACACAAACTCTCGACCTTACAGTCCATGTTGCAACAATACGC
TACTGTAGAAGAGGCTTTTTCTCGTTATTTTACTTACCGAGCTAATCGATTAGGATTTGATGGCTCTTCTCGATCCG
AGATGGCTTTAGTAAAGATGGCCACCTTGATGAACCTGTCTCTTCCGAAGCTGCGATTTTAAACCACAAGCTTCAA
ACCTTCTCAGAGAAGAGCGGATACTTTGATCAATAGTTTCTATAACCAATAAGGGCGATTGCTTGGCTCTTTCTCT
CGGAGGTTGCTTACACTGTATCCGAAGTACGCGCAACTGCCCATGGCAATACCAATGCAGAAGCTCGATCTCAGC
AAATTTATGCAACTACCTTATCGTATAGTAAAGAGTCTGAAAGCGCACAAAGAAATGCTAAACAAGCAAACTCTT
TCTAAGGAAATGTTTTAGATTTCTCAGAACTGCAGCTTCTTCCAAAGGATTGGATATCTTTTCCGAGAAATGTCG
TGTTCAAATTCACTTAAATGGAACCGTTAGTATCCATTTA

SEQ ID NO: 64 - CT733 fragment protein sequence

APQPRGTLPSSTTKIGSEVWIEQKVRQYPELLWLVEPSSSTGASLKSPSGAFISPTLQKKVPAFDIAVRSLIHLHL
IQGSRQAYAQLIQLTSESPITFKQFLALHKQLTLFLNSPKEFYDSVKVLETAIVLRHLGCSTKAVAAPKPYFSEMQ
REAFYTKALHVLHTFPELSPS FARLSPEQKTLFFSLRKLANYDELLSLTNTPSFQLLSAGRSQRALLALDLYLALD
SCGEQGMSSQFHTNFAPLQSMQLQYATVEEAFSRYFTYRANRLGFDGSSRSEMALVRMATLMNLSPEAAIILTTSFK
TLPTEEADTLINSFYTNKGDLSLALSLRGLPTLVSELTRTAHGNTNAEARSQIYATLTLVVKSLKAHKEMLNKQIL
SKEIVLDFSETAASCQGLDIFSENVAVQIHLNGTVSIIHL

SEQ ID NO:65 – CT153 fragment nucleotide sequence

ACTAAGCCTTCTTTCTTATACGTTATTCAACCTTTTTCCGTATTTAATCCACGATTAGGACGTTTCTCTACAGACTC
AGATACTTATATCGAAGAAGAAAAACCGCCTAGCATCGTTCATTGAGAGTTTGCCACTGGAGATCTTCGATATACCTT
CTTTCATGGAAACCGCGATTTCCAATAGCCCTATATTTTATCTTGGGAGACAACATAAGACGGCGCTCTGTTCACT
ATTCCTGAACCCAACTCTCAGCTTGGCGAGCCACTTGCCTGGTAGCCCCCTTCTATACAAATGAAATCCGATGCGGA
GCTCCTAGAAGAAATTAAGCAAGCGTTATTACGCGACTCTCATGACGGTGTGAAATATCGCATACCCAGAGAAATCCT
TCTCTCCAGAAAAAGAAACTCCTAAGGTTGCTCTAGTCTGATGACGATATTGAATGATTGCGCAATGTCGATTTTTG
GGTAGAGCTGTTGACATTTGTCAAATTAAGACCCTATTAATTTCTGAATACCGTAAGCGAAGAGAATATTCTAGATTA
CTCTTTTACAAGAGAAACGGCTCAGCTGAGCGGGATGGTGGTTTTGGTATTCCTCCAGGGACTAAGCTATTCCTTA
AACCTTCTTTTGTAGTAAAGATCAGTACCTCCATTTTTCGAAGAAACAACCTTCAATTTACTCGAAGTTTTTCTGCATCG
GTTACTTTTGTAGTACCAGACCTCGCGGCGACTATGCCTCTTCAAAGCCCTCCCATGGTAGAAAAATGGTCAAAAAAGA
AATTTGTGTCATTCAAAAAACCTTATCCCAAGCTACTCTCTAACTAGTTCGATATTGTTAAACGATACAAAAAGAG
AGGCTAAGATCTTGATTAACAAGCTTGGCTTTGGAAATGTTATGGCGACATCGGGCTAAAAGCCAAATCTCACCGAG
GGAAGCGTACGTCTAGACTTACAAGGATTACAGAATCGAAGTACAATTAACAGATTCAAGTAGGATCCCATACGAT
TGCAGCTGTATTAATCGATATGGATATTTCCAAGATTCAATCCAAATCAGAACAAGCTTATGCAATTAGGAAAAATCA
AATCAGGCTTTCAACGTAGCTTGGATGACTATCATATTTATCAAATTTGAAAGAAAAACAACCTTTTTCTTTTTCTCCG
AAGCATCGCAGCCTCTCATCCACATCCCATTCCGAAGATTCTGATTTGGATCTTTCTGAAGCAGCCGCTTTTTCAGG
AAGCTTACCTCGAGTTGTAATAAAAAAGCACTCAACATGCGAAGAATACCGTCAATGTTCCACAGCCGCTCAT
CCTTATACACTCAAAAAGAGATGACAGCTCGAACCCCTCTGAAAAACGATTAGATAGTTGTTTTCCGCAATTGGATT
GAAAAACAATAAGCGCAATTTCTCCAGATTCCTGGTTCAGCGTTTTATTCAAAAAATTCGGAACACACTATATTGCATC
AGCAACTTTTGGAGGGATAGGTTTTCAAGTGCTCAAACATCTTTTTGAACAGGTGGAGGATCTACATAGCAAAAAAGA

TCTCCTTAGAAAACCGCAGCAGCCAACCTCTCTATTAATAAAGGTTCTGTATCCAGCAGCACAGAATCTGGATACTCCAGC
 TATAGCTCCACGTCTTCTCTCATAACGGTATTTTTAGGAGGAACGGTCTTACCTTCGGTTCATGATGAACGTTTAGA
 CTTTAAAGATTGGTCGAAAAGTGTGCACCTGGAACCTGTTCTTATCCAGGTTTCTTTACAACCTATAACGAATTTAC
 TAGTTCCTCTCCATTTTCTAATATCGGTGCTGCAGAGCTCTCTAATAAACGAGAATCTCTTCAACAAGCGATTGCGA
 GTCTATCTCAAAGAACATAAAGTAGATGAGCAAGGAGAACGTAACATTTACATCAGGAATCGATAATCCTTCTC
 CTGGTTTACCTTAGAAGCTGCCCACTCTCCTCTTATAGTCAGTACTCTTACATTGCTTCGTGGTCTACGCTTCTCT
 ATTTGTTCCCAACATTAAGAGAAGCTTCTTCGGCAACCCCTATCGTTTTCTATTTTTGTGTAGATAATAATGAACAT
 GCTTCGCAAAAAATATTAACCAATCGTATTGCTTCTCGGGTCTTGCCTATTGACAAAAATTTTTGGTAGCGA
 ATTTGCTAGTTTCCCCTATCTATCTTTCTATGGAAATGCAAAAGAGGCGTACTTTGATAACACGTAACCCAACGC
 GTTGTGGGTGGATTGTTGAAAAGTTAAATACTACCAAGATCAATTCCTCCGGGATGGAGACGAGGTGCGACTAAAA
 CATGTTTCCAGCGAAAAGTATCTAGCAACAACCTCTTAAAGGATACCCATGGTACACTACGCGGTACAACGAACCTG
 TGAAGATGCTATCTTTATTATTAATAAATCTTCAGGTTAT

SEQ ID NO:66 - CT153 fragment protein sequence

TKPSFLYVIQPFVFNPRLRGRFSTDSPTYIEEENRLASFIESLPLEIFDIPSPMETAISNSPYILSWETTKDGALFT
 ILEPKLSACAATCLVAPSIQMKSDAELLEEEKQALLRSSHQGVKYRITRESFSPEKKTPKVALVDDDIELIRNVDFL
 GRAVDIVKLDPINILNTVSEENILDYSFTRETAQLSADGRFGIPPGLKLPKPSFQVEISTSIFFEETTSFTRFSAS
 VTFSVPDLAATMPLQSPPMVENGRQKEICVIAQKHLFPSYSPKLVDIVKRYKREAKILNKLAFGLWRHRAKSLTE
 GSVRLDLQGFTESKYNYQIQVGSHTIAAVLIDMDISKIQSKSEQAYAIRKIKSGFQRLDDYHIYQIERKQTFSP
 KHRSLSSSTSHSESDLDLSEAAAFSGSLTCEFVKKSTQAKNTVTCTAAHSLYTLKEDDSSNPSEKRLDSCFRNWJ
 ENKLSANSPDSWSAFIQKFGTHYIASATFGGIGFQVLKLSFEQVEDLHKKISLETAANSLLKGSVSSSTESGYSS
 YSSTSSSHTVFLGGTVLPSVHDERLDFKDWSESVHLEPVPIQVSLQPIITNLLVPLHFPNIGAAELSNKRESLQAIR
 VYLKEHKVDEQGERTTFTSGIDNPSSWFTLEAAHSPLIVSTPYIASWSTLPYLPFLRERSSATPIVYFVVDNNEH
 ASQKILNQSYCFLGSLPIRQKIFGSEFASFPYLSFYGNAKEAYFDNTYYPTRCGWIVEKLNNTQDQFLRDGDEVRLK
 HVSSGKYLATTPKLDTHGTLTRTTNCEDAIFIKKSSGY

SEQ ID NO:67 - CT601 fragment nucleotide sequence

GGTAAAGCACCGTCTTTGCAGGCTATTCTAGCCGAAGTCGAAGACACCTCCTCTCGTCTACACGCTCATCACAATGA
 TCTTGCTATGATCTCTGAACGCCTCGATGAGCAAGACACGAAACTACAGCAACTTTCTGCAACACAAGATCATAACC
 GACCTCGACAAGTTCAGCGACTAGAAACGGACCAAAAAAGCTTTGGCAAAAACACTGGCGATTCTTTGCAATCCGTC
 CAAGATATTCGGTCTTCTGTACAAAATAAATTAACAAGAAATCCAACAAGAACAACAAAAAATTAGCACAAAATTTGCG
 AGCGCTTCGTAACCTTTACAAGCTCTCGTTGATGGCTCTTCTCCAGAAAATTATATTGATTTTCTAAGTGGTGAAA
 CCCCAGAACATATTCAATTTGTTAAACAAGGAGAGACCTGAGCAAGATCGCGAGTAAATATAACATCCCCGTCGTA
 GAATTAAAAAAATTAATAAACTAAATTCGGATACTATTTTTACAGATCAAAGAATTCGCCTTCGAAAAAGAAA

SEQ ID NO:68 - CT601 fragment protein sequence

GKAPSLQAILAEVEDTSSRLHAHNELAMISERLDEQDTKLQQLSSTQDHNLPQVQRLETQKALAKTLAILSQSV
 QDIRSSVQNKLRQEQEKKLAQNLRALRNSLQALVDGSSPENYIDFLTGETPEHIHIVKQGETLSKIASKYNIPVV
 ELKKNLNSDTIFDQRIKPKK

SEQ ID NO:69 - CT279 fragment nucleotide sequence

GCACAAGTAATTTCTCCGATAACACATTCCAAGTCTATGAAAAGGGAGATTGGCACCCAGCCCTATATAACTAA
 AAAGCAGTTGCTAGAGATCTCCTCTACTCCTCTAAAGTAACCGTGACAACCTTAAAGCTCATATTTTCAAACTTTG
 TTAGAGTCTTGCTTACAGATACACAAGGAAATCTTTCTTATTGAAAGCATAATCTCAATCTAGAAGAATTTTAA
 TCTCAACCAACTCCTGTAATACATGGTCTTGCCCTTTATGTGGTCTACGCTATCCTACACAACGATGCAGCTTCTC
 TAAATTATCTGCTTCCCAAGTAGCGAAAAATCCAACAGCTATAGAATCTATAGTTCTTCTATAGAAGTTTTGGTT
 TGTGGGGACCTATCTATGGATTCTTGTCTAGAAAAAGACGGGAATACTGTTCTTGGTACTTCTTGGTATCAACAT
 GCGGAGACTCCTGGATTAGGAGCAAATATCGCTAACCTCAATGGCAAAAAAATTTAGAGGCAAAAAAGTATTTCT
 AGTCTCAGCTTCTGGAGAAAACAGATTTTGGTAAGACAACCTTAGGACTGGAAGTTATAAAAGGATCTGTATTCGAC
 CATTAGGAGACTCACCTAAAGCTGCTTCTTCCATCGACGGAATTTAGGAGCTACTTTGACTTGTAAATGGTGTACC
 GAATCCTTCTCTATTCTCTAGCTCCCTACCGCGCTTGTGACTTTCTTCGCAACTCTAAACCTAGTGGAGAGTC
 TCATGACCAC

SEQ ID NO:70 - CT279 fragment protein sequence

AQVISSDNTFQVYEKGDWHPALYNTKKQLLEISSTPPKVTVTTLSSYFQNFVRVLLTDQGNLSSFEHDHNLNLEEF
 SQPTPIHGLALYVYVAILHNDAAASSKLSASQVAKNPTAIESIVLPIEGFLWGPIYGFLEALEKDGNTVLGTSWYQH
 GETPGLGANIANPQWQKNFRGKKVFLVSASGETDFAKTTLGLEVIKGSVSAALGDSPKAASSIDGISGATLTCNGVT
 ESFSLAPYRALLTFFANSKPSGESHDH

SEQ ID NO:71 - CT443 fragment nucleotide sequence

GGGGTGTAGAGACCTCTATGGCAGAGTCTCTCTCAACAAAGTTATTAGCTTAGCTGACACCAAGCGAAAGACAA
 CACTTCTCATAAAAGCAAAAAAGCAAAAAAACCACAGCAAAAGAGACTCCCGTAGACCGTAAAGAGGTTGCTCCGG
 TTCATGAGTCTAAAGCTACAGGACCTAAACAGGATTTCTGCTTTGGCAGAATGTATACAGTCAAAGTTAATGATGAT
 CGCAATGTTGAAATCACACAAGCTGTTCTGAATATGCTACGGTAGGATCTCCCTATCCTATTGAAATTAAGTCTAC
 AGGTAAAAGGGATTGTGTTGATGTTATCATTACTCAGCAATTACCATGTGAAGCAGAGTTCGTACGCAGTGATCCAG
 CGCAACTCCTACTGCTGATGGTAAGCTAGTTTGGAAAATTGACCGCTTAGGACAAGGGCAAAAGAGTAAATTAAT

GTATGGGTAAAACCTCTTAAAGAAGGTTGCTGCTTTACAGCTGCAACAGTATGCGCTTGTCCAGAGATCCGTTCCGGT
TACAAAATGTGGACAACCTGCTATCTGTGTTAAACAAGAAGGCCAGAGAATGCTTGTGCTTGCCTGCCCAGTAGTTT
ACAAAATTAATAGTGAACCAAGGAACAGCAACAGCTCGTAACGTTGTTGTTGAAAATCCTGTTCCAGATGGTTAC
GCTATTCTTCTGGACAGCGTGTACTGACGTTTACTCTTGGAGATATGCAACCTGGAGAGCACAGAACAATTACTGT
AGAGTTTTGTCCGCTTAAACGTTGGTCTGCTACCAATATAGCAACGGTTTTCTTACTGTGGAGGACATAAAAAATACAG
CAAGCGTAACAACCTGTGATCAACGAGCCTTGCCTACAAGTAAGTATTGCAGGAGCAGATTGGTCTTATGTTTGTAAAG
CCTGTAGAATATGTGATCTCCGTTTCCAATCCTGGAGATCTTGTGTTGCGAGATGTCGTCGTTGAAGACACTCTTTC
TCCCGGAGTACAGTCTTGAAGCTGCAGGAGCTCAAATTTCTTGAATAAAGTAGTTTGGACTGTGAAAGAACCTGA
ATCCTGGAGAGTCTCTACAGTATAAAGTTCTAGTAAGAGCACAAACTCCTGGACAATTACAAAATAATGTTGTTGTG
AAGAGCTGCTCTGACTGTGGTACTTGTACTTCTTGCAGAGAACGACAACTTACTGGAAAGGAGTTGCTGCTACTCA
TATGTGCGTAGTAGATACTTGTGACCTGTTTGTGTAGGAGAAAATACTGTTTACCGTATTTGTGTCAACCAACAGAG
GTTCTGCAGAAGATACAAATGTTTCTTTAATGCTTAAATTTCTTAAAGAACTGCAACCTGTATCCTTCTCTGGACCA
ACTAAAGGAACGATTACAGGCAATACAGTAGTATTCGATTTCGTTACCTAGATTAGGTTCTAAAGAACTGTAGAGTT
TTCTGTAACATTGAAAGCAGTATCAGCTGGAGATGCTCGTGGGGAAGCGATTCTTCTCCGATACATTGACTGTTCC
AGTTTTCTGATACAGAGAATACACACATCTAT

SEQ ID NO:72 - CT443 fragment protein sequence

GVLETSMAESLSTNVISLADTKAKDNTSHKSKKARKNHSKETPVDRKEVAPVHESKATGPKQDSCFGRMYTVKVNDD
RNVEITQAVPEYATVGSPPYIEITATGKRDCVDVITQQLPCEAEFVRS DPATTPADGKLVWKIDRLGQGEKSKIT
VWVKPLKEGCCFTAAATVCACPEIRSVTKCGQPAICVKQEGPENALRCPVVYKINIVNQGTATARNVVENPVPDGY
AHSSGQRVLTFTLGDMPGEHRTITVEFCPLKRGRATNIATVSYCGGHKNTASVTTVINEPCVQVSIAGADWSYVCK
PVEYVISVSNPGLVLRDVVVEDTLSPGVTVLEAAGAQISCNKVVWTVKELNPGESLQYKVLVRAQTPGQFTNNVVV
KSCSDCGTCTSCAEATYWKVAATHMVCVVDTCDPVCGENTVYRICVTNRGSAEDTNVSLMLKFSKELQPVFSFSGP
TKGTITGNTVVFDSLPRLLGSKETVEFSVTLKAVSAGDARGEAILSSDTLTVPVSDTENTHIY

SEQ ID NO:73 - CT372 fragment nucleotide sequence

CAGGCTGCACACCATCACTATCACCGCTACACAGATAAACTGCACAGACAAAACCATAAAAAAGATCTCATCTCTCC
CAAACCTACCGAACAGAGGCGTGCAATACTTCTTCCCTTAGTAAGGAATTAATCCCTCATCAGAACAAAGAGGCC
TTTTATCCCCCATCTGTGACTTTATTTCCGGAACGCCCTTGTCTACACGGAGTTTCTGTTAGAAATCTCAAGCAAGCG
CTAAAAAATTTCTGCAGGAACCCAAATTCGACTGGATTGGTCTATTCTCCCTCAATGGTTCAATCCTCGGGTCTCTCA
TGCCCCAAGCTTTCTATCCGAGACTTTGGGTATAGCGCACACCAAACTGTTACCGAAGCCACTCCTCCTTGTGCGC
AAAACTGCTTTAATCCATCTGCGGCCGTTACTATCTATGATTCTCATATGGGAAAGGGGTCTTTCAAATATCCTAT
ACCTTGTCCGCTATTGGAGAGAGAATGCTGCGACTGCTGGCGATGCTATGATGCTCGCAGGGAGTATCAATGATTA
TCCCTCTCGTCAGAACATTTTCTCTCAGTTTACTTTCTCCCAAACTTCCCAAATGAACGGGTGAGTCTGACAATTG
GTCAGTACTCACTCTATGCAATAGACGGAAACATTATACAATAACGATCAACAACCTGGATTCAATAGTTACGCATTA
TCACAAAATCCAACAGCAACTTATTCCTCTGGAAGCTTTGGAGCTTACCTACAAGTCGCTCCTACCGCAAGCACAAAG
TCTTCAAATAGGATTTCAAGACGCTTATAATATCTCCGGATCCTCTATCAAATGGAGTAACCTTACAAAAAATAGAT
ACAATTTTACGGTTTTTGTCTTCTGCGCTCCCGCTGTTGTTGTTAGGATCTGGCCAGTACTCCGTGCTTTTATGTG
ACTAGACAAGTTCCAGAACAGATGGAAACAAACAATGGGATGGTCAGTCAATGCGAGTCAACACATATCTTCTAAACT
GTATGTGTTTGGAAAGATACAGCGGTGTTACAGGACATGTGTTCCCGATTAACCGCACGATTCATTCCGGTATGGCCT
CTGCAAAATTTATTTAACCGTAACCCACAAGATTTATTTGGAATTGCTTGGCGATTCAATAATGTACACCTCTCTGCT
TCTCCAAATACTAAAGAAAATAACGAAACTGTAATCGAAGGGTTTTGCAACTATCGGTTGCGGCCCTATCTTTCTTT
CGCTCCAGACTTCCAACCTTACCTCTACCCAGCTCTTCGTCCAAACAAACAATCTGCCCGTGTATATAGCGTGGCAG
CTAATTTAGCTATC

SEQ ID NO:74 - CT372 fragment protein sequence

QAAHHYHRYTDKLRQNHKKDLISPKPTEQEAACNTSSLKELIPLSEQRGLLSPICDFISERPCLHGVSVRNLKQA
LKNSAGTQIALDWSILPQWFNPRVSHAPKLSIRDFGYSAHQTVTEATPPCWQNCFNPSAAVTIYDSSYGKGVFQISY
TLVRYWRENAATAGDAMMLAGSINDYPSRQNFISQF TFSQNFNERNVSLTIGQYSLY AIDGTLYNNDDQLGFISYAL
SQNPATYSSGSLGAYLQVAPTASTLSLIGFQDAYNISGSSIKWSNLTKNRYNFHGFASWAPRCDLGSGBQYSVLLYV
TRQVPEQMEQTMGWSVNASQHISSKLYVFGRYSGVTVGHVFPINRTYSFGMASANLNFNRNPQDLFGIACAFNNVHLSA
SPNTRKRYETVIEGFATIGCGPYLSFAPDFQLYLYPALRPNKQ SARVYSVRANLAI

SEQ ID NO:75 - CT456 fragment nucleotide sequence

ACAAATTCAGCGGCTACATCTTCTATCCAAACGACTGGAGAGACTGTAGTAAACTATACGAATTCAGCCTCCGCCCC
CAATGTAACGTATCGACCTCCTCTTCCACACAAGCCACAGCCACTTCGAATAAAACTTCCCAAGCCGTTGCTG
GAAAAATCACTTCCAGATACTTCAGAAAGCTCAGAAACTAGCTCTACCTCATCAAGCGATCATATCCCTAGCGAT
TACGATGACGTTGGTAGCAATAGTGGAGATATTAGCAACAACACTACGATGACGTAGGTAGTAACAACGGAGATATCAG
TAGCAATTTAGCAGTGTCTGCTGCTGATTACGAGCCGATAAGAATACTGAAAAATTTTATGAGAGTATTGGTGGCT
CTAGAACAAGTGGCCCAGAAAATACAAGTGGTGGTGCAGCAGCAGCACTCAATTCCTAAGAGGCTCCTCCTACAGC
AATTAATGACGATGCTGCTGCTGATTACGAGCCGATAAGAATACTGAAAAATTTTATGAGAGTATTGGTGGCTCTAG
AACAAAGTGGCCCAGAAAATACGAGTGGTGGTGCAGCAGCAGCACTCAATTCCTAAGAGGCTCCTCCTACAGCAATT
ATGACGATGCTGCTGCTGATTACGAGCCGATAAGAATACTGAAAAATTTTATGAGAGTATTGGTGGCTCTAGAACA
AGTGGCCCAGAAAATACGAGTGTGGTGCAGCAGCAGCACTCAATTCCTAAGAGGCTCCTCCTACACAACAGG
GCCTCGTAACGAGGGTGTATTCCGGCCCTGGACCGAAGGACTACCAGACATGTCTTCTTCCATACGATCCTACAA
ATAAAACCTCGTTATTGACTTCTCTCCAACCTCATGTAAAGTCGAAAATGCTTGAAGAACTCGGGGCATTTGCTC
TTCATTGATACAGATAGAAGTAGTTTCAATCTTGTTCCTAACGGAAATTTGGGACCAAGTCTGTTCAATTAAGTTCA

AAATGGAAAGACCAAAGAAGATCTCGACATCAAAGACTTGGAAAAATGTGTGCAAAATTTCTGTACAGGGTTTAGCA
AATTCTCTGGTGACTGGGACAGTCTTGTAGAACCTATGGTGTGAGCCAAAGCTGGAGTGGCCAGCGGAGGCAATCTT
CCCAATACAGTGATTATCAATAATAAATTCAAAACCTTGCCTTGGCTTATGGTCTTGGAAATAGCCAGGAAGCAAGTTC
TGGTTATACACCTTCTGCTTGGAGACGTGGTTCATCGAGTAGATTTTGGAGGAATTTTTGAGAAAGCCAACGACTTTA
ATAAAATCAACTGGGGAACCTCAAGCCGGGCTTAGTAGCAAGACGATGGCATTTCCTTCCAATGAAACTCCTGGA
GCTGGTCTGCAGCTGCTCCATCACCAACGCCATCCTCTATTCTATCATCAATGTCAATGTCAATGTTGGCGGAAC
TAATGTGAATATTGGAGATACGAATGTCAACACGACTAACACCACACCAACAACCTCAATCTACAGACGCCTCTACAG
ATACAAGCGATATCGATGACATAAATACCAACAACCAAACTGATGATATCAATACGACAGACAAAGACTCTGACGGA
GCTGGTGGAGTCAATGGCGATATATCCGAAACAGAATCCTCTTCTGGAGATGATTCAGGAAGTGTCTCTTCTCAGA
ATCAGACAAGAATGCCCTCTGTGCGAAATGACGGACCTGCTATGAAAGATATCCTTTCTGCCGTGCGTAAACACCTAG
ACGTCGTTTTACCCTGGCGAAAATGGCGGTTCTACAGAAGGGCCTCTCCAGCTAACCAAACTCTCGGAGACGTAATC
TCTGATGTAGAGAATAAAGGCTCCGCTCAGGATACAAAATTTGTGAGGAAATACAGGAGCTGGGGATGACGATCCAAC
AACCACAGCTGCTGTAGGTAATGGAGCGGAAGAGATCACTCTTTCCGACACAGATTCTGGTATCGGAGATGATGTAT
CCGATACAGCGTCTTCACTCTGGGGATGAATCCGGAGGAGTCTCCTCTCCCTCTTCCAGAAATCCAATAAAAACTGCC
GTTGGAATGACGGACCTTCTGGACTAGATATCCTCGCTGCCGTACGTAACATTTAGATAAGGTTTACCCTGGCGA
CAATGGTGGTTCTACAGAAGGGCCTCTCCAAGCTAACCAAACTCTTGGAGATATCGTCCAGGATATGGAAACAACAG
GGACATCCCAAGAAACCGTTGTATCCCATGGAAAGGAAGCACTTCTTCAACGGAATCAGCAGGAGGAAGTGGTAGC
GTACAAAACACTACTGCCTTACCACCTCCAACCCCGTCAACTACAACATTAAGAACGGGCACAGGAGCTACCACCAC
ATCCTTGATGATGGGAGGACCAATCAAAGCTGACATAATAACAACCTGGTGGCGGAGGACGAATTCCTGGAGGAGGAA
CGTTAGAAAAGCTGCTCCCTCGTATACGTGCGCACTTAGACATATCCTTTGATGCGCAAGGCGATCTCGTAAGTACT
GAAGAGCCTCAGCTTGGCTCGATTGTAACAATAATCCGCAAGAACTGGTTCAAGAGGAATCTTAGCTTTCTGTTGA
GAGTGTCTCAGGCAAGCCGGGATCTGCACAGGTCTTAAACGGGTACAGGGGGAGATAAAGGCAACCTATTCCAAGCAG
CTGCCGCAGTACCCCAAGCCTTAGGAAATGTTGCAGGAAAAGTCAACCTTGGGATACAAGGCCAAAAACTATCATCC
CTAGTCAATGACGACGGGAAGGGGTCTGTTGGAAGAGATTTATTCCAAGCAGCAGCCAAACAACCTCAAGTGCTAAG
CGCACTGATTGATACCGTAGGA

SEQ ID NO:76 - CT456 fragment protein sequence

TNSAATSSIQTGTGETVVNYTNSASAPNVTVSTSSSSSTQATATS NKTSQAVAGKITSPTDTSSESSETSSSTSSSDHIPSD
YDDVGSNSGDISNNYDDVGSNNGDISSNYDDAAADYEPIRTTENIYESIGGSR TSGPENTSGGAAAALNSLRGSSYS
NYDDAAADYEPIRTTENIYESIGGSR TSGPENTSGGAAAALNSLRGSSYSNYDDAAADYEPIRTTENIYESIGGSR
SGPENTSGGAAAALNSLRGSSYTTGPRNEGVF GPGPEGLPDMSLPSYDPTNK TSLTFLSNPHVKS KMLENSGHFV
FIDTDRSSFILVPNGNWDDQVCSIKVQNGKTKEDLDIKDLENMCAKFC TGF SKFSGDWDSLVEPMVSAKAGVASGGNL
PNTVIINNKFKTCVAYGPWNSQEAASSGYTPSAWRRGHRVDFGGIFEKANDFNKINWGTQAGPSEDDGISFSNETPG
AGPAAAPSPTPSSIPIINNVNVVGGT NVNIGD TNVNTTNTPTTQSTDASTD TSDIDDINTNNQTD DINTTDKDS DG
AGGVNGDISE TESSSGDDSGSVSSSES DKNASVGN DGPAMKDILSAVRKHL DVVYPGENGGSTEGPLPANQTLGDVI
SDVENKGSAQDTKLSGNTGAGDDPTTTAAVNGAEETLSDTDSGIGDDVSDTASSSGDES GGVSSPSES NKNTA
VGNDGPSGLDILAAVRKHL DKVYPGDNGGSTEGPLQANQTLGDIVQDMETTGT SQTETVVSPWKGSTSSTESAGGSGS
VQTL LPSPPPTSTTLRTGTGATTTSLMMGGPIKADIITGGGGRI PGGGTLEKLLPRIRAHLDISFDAQDGLVST
EERQLGSI VNKFRQETGSRGILAFVESAPGKPGSAQVLTGTGGDKGNLFQAAA AVTQALGNVAGKVNLAIQGQRLSS
LVNDDGKGSVGRDLFQAAAQTTQVLSALIDTVG

SEQ ID NO:77: CT381 fragment nucleotide sequence

TGTTTTAAAAGAAGGGGGAGACTCCAATAGTGAAAAATTTATTGTAGGGACTAATGCAACCTACCCCTCCTTTTGAGTT
TGTTGATAAGCGAGGAGAGGTTGTAGGCTTCGATATAGACTTGGCTAGAGAGATTAGTAACAAGCTGGGGAAAACCG
TGGACGTTCCGGGAGTTTTCTTTGATGCACTCATTCTAAACCTAAACACAGCATCGGATTGATGCGGTTATAACAGGG
ATGTCCATTACTCCTTCTAGATTGAAGGAAATCTTATGATTCCCTATTATGGGGAGGAAATAAAAACACTTGGTTTT
AGTGTTTAAAGGAGAGAATAAGCATCCATTGCCACTCAATATCGTTCTGTAGCTGTTCAAACAGGAACCTATC
AAGAGGCCTATTTACAGTCTCTTTCTGAAGTTCATATTCGCTCTTTTGTAGCACTCTAGAAGTACTCATGGAAGTC
ATGCATGGTAAATCTCCCGTCTGCTGTTTTAGAGCCATCTATCGCTCAAGTTGTCTTGAAAAGATTTCCCGGCTCTTTC
TACAGCAACCATAGATCTCCCTGAAGATCAGTGGGTTTTAGGATAACGGGATTGGCGTTGCTTCAGATCGCCAGCTT
TAGCCTTGAAAATCGAGGCAGCTGTGCAAGAGATCCGAAAAGAAGGAGTGCTAGCAGAGTTGGAACAGAAAGTGGGGT
TTGAACAAC

SEQ ID NO:78: CT381 fragment protein sequence

CLKEGGDSNSEKFI VGTNATYPPEFV DDKRGEVVGFDIDLAREISNKL GKTL DVREFSFDALILNLKQHRIDAVITG
MSITPSRLKEILMIPYYGEEIKHLV LVFKGENKHLPLTQYRSVAVQTGT YQAYLQSLSEVHIRSFDSTLEVLMEV
MHGKSPVAVLEPSIAQVV LKDFPALSTATIDLPEQWV LGGYIGVASDRPALALKIEAAVQEI RKEGVLAELEQKWG
LNN

SEQ ID NO:79: CT043 fragment nucleotide sequence

TCCAGGCAGAATGCTGAGGAAAATCTAAAAAATTTTGCTAAAGAGCTTAAACTCCCGACGTGGCCTTCGATCAGAA
TAATACGTGCATTTTGTGTTGATGGAGAGTTTTCTTTCACCTGACCTACGAAGAACACTCTGATCGCCTTTATG
TTTACGCACCTCTTCTTGACGGACTGCCAGACAATCCGCAAAAGAGTTAGCTCTATATGAGAAGTTGTTAGAAGGC
TCTATGCTCGGAGGCCAAATGGCTGGTGGAGGGTAGGAGCTACTAAGGAACAGTTGATCTTAATGCACCTCCGT
GTTAGACATGAAGTATGCAGAGACCAACTACTCAAAGCTTTTGCACAGCTTTTTATTGAAAACCGTTGTGAAATGGC
GAACTGTTTGTCTGATATCAGCGCTGGACGAGAACCCTACTGTTGATACCATGCCACAAAATGCCTCAAGGGGGTGGC
GGAGGAATTC AACCTCTCCAGCAGGAATCCGTGCA

SEQ ID NO:80: CT043 fragment protein sequence

SRQNAEENLKNFAKELKLPDVAFDQNNCTILFVDGEFSLHLTYEEHSRLYVYAPLLDGLPDNPQRRLALYEKLLLEG
SMLGGQMAGGGVGVATKEQLILMHCVLDMKYAETNLLKFAQFLFIETVVKWRTVCSDISAGREPTVDTMPQMPQGGG
GGIQPPPAIRA

SEQ ID NO:81: CT711 fragment nucleotide.seq Length: 2298

TCAATACAACCTACATCCATTTCTTTAACTAAGAATATAACGGCAGCTTTAGCCGGAGAGCAGGTCGATGCTGCTGC
AGTGTATATGCCCGCAGGCTGTTTTTTCTTTCAGCAACTGGATGAAAAAAGCAAGGGGCTGAAACAGGCTTTAGGAT
TGCTCGAAGAGGTTGATCTAGAAAAATTTATACCGTCTTTAGAAAAATCACCTACACCTATCACTACGGGAACAACG
AGTAAAAATTTCCGCTGATGGGATTTGAGATTGTTGGAGAGCTTTCTTCAGAAACAATTTTGGCAGATCCTAATAAAGC
TGCAGCTCAGGTTTTGGAGAGGGGCTTGCAGATAGTTTTGATGATTGGCTCAGATTATCTGAAAAATGGGGGGATT
AAGATCCTACAGCAATAGAAGAAGAGATTGTTACTAAGTATCAAAACAGAACTCAATACTCTGCGCAATAAACTCAAG
CAACAATCTTTAACAGACGATGAGTATACGAAGCTTTATGCTATTCTCAAACTTTGTTAAAGAGATAGAAAGCTT
AAAGAATGAAAAATAATGTGAGGTTAATTTCCAAAAGTAAAGTCACTAACTTTTGGCAGAATATCATGCTCACTTACA
ACTCGGTAACCTCGTTATCAGAACCTGTTACCGATGCGATGAATACGACTATGGCGGAGTACTCTCTTTATATTGAG
AGAGCTACAGAGGCTGCCAAGTTGATACGGGAGATAACCAACACGATCAAAGACATTTTCAATCCAGTTTGGGATGT
GCGTGAACAAAACAGGAATTTTTGGGTTAAAAGGAGCTGAGTATAACGCTTTAGAAGGCAATATGATTTCAAAGCTTGC
TTAGCTTTGCGGGTCTATTCCGGCAGTTAATGAGTCGTAAGTGAACAGTTGATGAGATAGGCGCACTTTATCCTAAA
AATGATAAAAACGAAGACGTCATTCATACTGCTATTGATGATTATGTGAATTTCTTAGCTGATTTGAAAGCCAATGA
ACAGGTCAAACCTCAACGGTCTGTTGAGTTTAGTATATGCTTATTATGCTAGTACTTTAGGTTTTGCTAAGAAGGATG
TATTCAATAATGCACAAGCTTTCTTTACAGATTACTAATTTTCTAAACCAAGAGATCCAATATTGGACGCCCTAGA
GAGACTTCAAGTTTTAATATCTCCAATCAAGCATTGCAAACTTTAAAAATAAGCCTTCCGGCTGATTATAACGGCGT
ATATCTTTTTGATAATAAAGGATTAGAGACTAATCTCTTAACTCTACGTTCTTCTTTGATGTTGTGAGTCTCATGA
CAGCTGATCCTACGAAGACTATGTCTCGACAGGATTACAATAAGGTGATTACAGCCTCGGAATCCAGTATTCAGAAG
ATTAATCAGGCTATTACCGCTTGGGAACTAGCTATTGCGAGAATGTGGGACTAAAAAAGCGAAGCTCGAACCATCCAG
TTTTAAATTTAATGCTATGGTTCGAAGCGAAGAAGACCTTCGTAGAGACCTTCCAATACAGATGGTCTATTTCAT
CTTTGATGTTGGATAAGTATCTTCCGAATCAGCAGTACATATTAGAGACATTAGGAAGTCAGATGACTTTCTCTAAC
AAGGCTGCTCGGTATTTAAATGATATCATTGCGTATGCGAGTTAGCTTCCAAACAGCTGACGCTCTATTATCTTTAGG
GATGTATCTTCGACAAATGAACCAGCAGGAATTTCTGAGGTGATTTCTCGTGCTAACGATACTGTGAAAAAAGAGA
TAGATCGGAGTCGTGCGGATCTCTTTCACTGTAAAAAAGCTATCGAAAAGATTAAGAATTAGTGACTTCTGTAAT
GCGGATACTGAATTGACCTCATCTCAGCGTGCAGAGTTATTAGAGACGTTAGCTAGTTATGCTTTTGAATTTGAGAA
TCTCTATCACAACCTCTAATGTTTACGTCATGGTTCTAAGGTACAGATTTCTGGCGTAAGCAAGCCTGATGAAG
TGGATGAGGCTTTTACTGCTAAGATTGGATCGAAGGAATTCGATACTTGGATTGAGCAGCTTACAACATTTGAAAGT
GCTGTGATTGAAGGTGGGCGTAATGGTGTGATGCCGTTGGGGAGAGCAGCAGGTTTTACAGAGTTTAGAGAGCAAGCA
GCAAGATTACACGTCGTTCAACCAGAATCAGCAATTAGCTCTACAAATGGAGTCCGACGCGATTCAACAAGAGTGG
CTATGGTAGCAGCAGCCTTAGCATTAAATGAATCAGATTTTTGCTAAGTTGATCCGTAGATTTAA

SEQ ID NO:82: CT711 fragment protein sequence (AAC68306)

SIQPTSISLTKNITAALAGEQVDAAVYMPQAVVFFQQLDEKSKGLKQALGLLEEVDLEKFIKPSLEKSPITTTGTT
SKISADGIEIVGELSSSETILADPNKAAAQVFGGLADSFDDWLRLENGGIQDPTAIEEEIVTKYQTELNLRNKLK
QQLSLTDDDEYTKLYAIPQNFVKEIESLKNENNVRLIPKSKVTNFQNIIMLTYNVTSLSLSEPVTDAMNTTMAEYSLYIE
RATEAAKLIREITNTIKDIFNPVWDVREQTGIFGLKGAEYNALEGNMIQSLLSFAGLFRQLMSRTATVDEIGALYPK
NDKNEDVIHTAIDDDYVNSLADLKANEQVKLNLGLLSLVYAYASTLGF AKKDVFNNAQASFTDYTNFLNQEIQYWT
PR ETSFNISNQLQTFKNKPSADYNGVYLFDNKGLETNLFNPTFFDVVSLMTADPTKMSRQDYNKVITASESSIQK
INQAITAWELAIACGTTKAKLEPSSLNYNFAMVEAKKTFVETSPIQMVYSSMLMDKYLPNQYIILETLGSQMTFSN
KAARYLNDIIAYAVSFQADVYYSLSGMYLRQMNQEFPEVISRANDTVKKEIDRSRADLFHCKKAIEKIKELVTSVN
ADTELTSQRAELLETLASYAFEFENLYHNL SNVYVMVSKVQISGVSKPDEVDEAFTAKIGSKEFDTWIQLTTFES
AVIEGGRNGVMPGGEQVQLQSLQSLQSDYTSFNQNLQALQMESAAIQEWTMVAALALMNQIFAKLIRRFK

SEQ ID NO:83: CT114 fragment nucleotide sequence - Length: 1296

GATCCTTTGAGTGCAAAACAGTTAATGTATCTGTTTCTCAGCTCTCAGAAGAGGATGTATCTGTTTTTGGCTCGATG
CATTTTTGCTTCAAAGCGTCCAGAATACCTTTTTCAAATCGGAGGAAGAGCTCTTTGCAAAATTTGATTTTGCCAA
GGGTTTTCTTAGGTGTTTCATCGGGACGATGATTTAGCGAGAGTGTGGTGTAGCGGAGCCTTCTGCAGAAGAGCAG
AAGGCTCGATACTATTCAATTGTATCTGGATGTTTTAGCTTTGCGTGCATACGTTGAAAGAGAGCGTTTGGCGAGTGC
TGCAACCGGAGATCCTGAGCGGATAGATTTGGCAACCATAGAAGCTATTAATACCATCCTTTTTCAGGAAGAAGGAT
GGAGGTATCCTTCAAACAAGAGATGTTTTGAAAAACAGGTTTTCTGAGTTAGCTGCTGTTACAGATAGTAAGTTTTGA
TTTTGCTTGGGAACCTGTAGTGCTTTATCAAGCTGTCGCCACGCGGCTTGATTTGCTCTGACCCTGTACCCCTCC
TGGACATATTTACTTACGCTATAAGGACAAGGTGAATATTGAAACCACTTCTGGAGGAAGGCATCTTCTACTGAAA
GGTATTGTGAATGCATAAAAGAGTCGCAGTTAAAGGTGCGTTCGCAGATGGAGCTTATAGGGTTAACTTTTATGAAT
AGAGGAGCTTTCTTTTTGCAAAAAGGAGAGTTTTCTCAGGCGTCTTAGCTTATGAGCAAGCTCAATCATATTTATC
AGACGAGCAGATTTCTGATTTGTTAGGGATTACTTATGTTCTTTTAGGAAAGAAGGCGGCGGAGAGGCTCTTTTAA
AGAAATCTGCAGAAAAGACTCGGCGAGGTTCACTATGACTATTTTCAAAGGATATATTTCCCCCGAAATCCTA
GGGGTGTGTTTTGCCGATTGAGGGGTGACCTATCAAGAACTTTGGAGTATCGAAAAAACTAGTGATGCTTTCCAA
GAAGTATCCAAAAGTGGATCTCTTAGGTTGAGGTTGGCGACAACAGCATTGGAGCTAGGGCTGGTCAAGGAGGGGG
TGCAGTTGTTAGAAGAGAGTGTAAAGGATGCCCCAGAGGACCTCTTTTACGTCGTCAGTTTTGTAATTTCTTTGC

AATCGACATGATTATGTCCGAGCAAAATATCATTTTGATCAAGCGCAAGCTCTTCTCATTAAAGAAGGGTTGTTTT
CGAAAAAACTTCTATACTCTCTTAAAACTATCGGGAAAAAGCTATCTTTTTGCTCCGAGT

SEQ ID NO:84: CT114 fragment protein sequence (AAC67705)

DPLSAKQLMYLFPQLSEEDVSVFARCI LSSKRPEYLF SKSEELFAKLILPRVSLGVHRDDDLARVVLVAEPSAEEQ
KARYYSLYLDVLA LRAYVERERLASAAHGDPERIDLATIEAINTILFQEEGWRYPSKQEMFENRFSELAAVTDSKFG
VCLGTVVLYQAVARLDLSLDPVTPPGHIYLRKDKVNIETTSGGRHLPTERYCECIKESQLKVVRSQMEIIGLTFMN
RGAFFLQKGEFLQASLAYEQAAQSYLSDEQISDLLGITVYLLGKKAAGEALLKKS AEKTRRGSSIYDYFQGYISPEIL
GVLFADSGVTTYQETLEYRKKLVMLSKKYPKSGSLRLRLATTALELGLVKEGVQLLEESVKDAPEDLSLRLQFCILC
NRHDYVRAKYHFDQAQALLIKEGLFSEKTSYLLKTI GKKLSLFAFS

SEQ ID NO:85: CT480 fragment nucleotide sequence

TCTTCAGATCTACTTGAAAAAGATGTGAAATCGATCAAAAAGAGAACTCAAGGCTTTACATGAAGATGTTCTTGAGTT
AGTCCGGATCTCGCATCAGCAAAAAAATGGGTCCAGTCTACAGATTTTTCTGTTTCTCCAGAGATCAGTGTATTGA
AGGATTGCGGAGATCCTGCTCCCTAATTTATTATGCGAAGACCCTTATGTTGAAAAAGGGTCCCTTCTGTTGTTA
AAGGAAGGTTTTGTTCCGAAAGGTATTTTGCCTACAGCTCAAGTAGGAAGGCCGTGATAACCTAAGTCCGTTAATGG
CTTTGTTAATATCGTTGATTTTATGAATTGTGCGTTCCCTAATTTGGCTGTTGAGCATGTTGGTAAATACGAGGAGT
TTGCGCCTAGTTTAGCCTTAAAGATAGAAGAGCATTATGTAGAGGATGGGTCTGGGGATAAAGAATTTTCATATTTAT
TTGCGTCTAATATGTTTTGGGAGCCGATAGATCTACGCTGTTCCCTAAAAATATAACTTTAGCAGACAGCTTCTT
AAGACCACATCCTGTACCCTCATGATGTGAAGTTCTAATACGATGTAGTCATGAATCCCTATGTTGCAGAAAATGC
GTGCAGTGGCTATGAGATCTTATTTTGAAGATATGGTTTTCGGTTCCGGTAGAAAACGATTTGAAATTAATCGTTCGT
TGGAGAGCTCATACTGTACGTAATGAACAGGGAGAGGAAGAGAAAAAAGTGCTCTATCTGCTTTCCGCAATACATT
GGCACTCCAACCGTTACCTTGTTCGTGTATCAGCATTTCGCAATGGAGAGAAGATCGTTCAGAAAGATTCTGATC
CCGATACGTATCGCAAAAGATTCGGTATGGGCGCAAACTTTTCTTACATTTGGGCGTATAATTACATAGTGAGCTGT
GGAGCATTCCGATTTGCAGGGATGGATGATGAGAAAAATTAAGTTAGTTGTAATCCCTAATTTATCATAATCCGTTTGC
GGCTCTTGTGGAGAAGCGCTATATCTATATGAAAGATAGTACAGATTCTCTCTTCCAAGATTTCAAAGCTGGGAAGG
TGGATATTGCGTATTTCCCTCCTAACCATGTCGATAATCTAGCGAGCTTCATGCAAACTCTGCTTATAAGGAACAA
GCTGCTAGAGGAGAGGCAATTTTAGAAAAAATTCATCAGACCGGTCCTATTTTACATCGGATGGAATTTGCTTTT
TCTTTTCTTAAACAATCGTTCGGTACGACAAGCCATGAATATGTTGATCGATCGGGATCGCATTATTGAGCAGTGCT
TGGATGGTCTGGAGTCTCTGTGAGTGGGCTTTTTCTCTCTGCTCTCCATCATAACAACAGAGATGTAGAGGGATGG
CAATACTCTCGGAAGAGGCGCACGTAAATTAGAGGAAGAGGGCTGGATCGATGCTGATGGAGATGGTATTCTGTA
GAAAGTAATCGATGGAGTTGTAGTGCCCTTTCCGTTTCCGGTTATGCTACTATGTGAAAAGTGTAACAGCACGAACGA
TTGCCGAATATGTAGCTACGGTATGTAAGAGGGTGGGTATCGAGTGTGCTTACTCGGGTTAGATATGGCGGATTA
TCACAAGCCCTCGAGGAGAAAAATTTGATGCTATTTTCCGGATGGTGTTTAGGAACCCCTCCAGAAGATCCTCG
TGCTCTATGGCATTCCGAAAGGAGCTTTGGAGAAAGGATTCGCAATGCTGTTGGATTTTGTAAATGAGGAAGCAGACC
GTATCTCGAACAGCTCAGTTACGAGTATGATTCTAATAGCGCCAAAGCCTTGTATCACCGTTTACGAGGTTGATT
CATGAGGAATCTCCTTACGCGTTTCTCTATTCAAGACAGTACTCCCTTGTCTATAAGGAGTTTGTAAAAAATATTTT
TGTGCCAACAGAACATCAGGATTTGATTCCTGGAGCTCAAGATGAGACAGTGAATTTATCCATGTTGTGGGTAGATA
AAGAGGAGGGTCTGATGCTCCGCTATATCT

SEQ ID NO:86: CT480 / oppA_4 fragment protein sequence (AAC68080)

SSDLLEKDVKSIKRELKALHEDVLELVLRISHQKKNWVQSTDFSVSPEISVLKDCGDPAPFNLLCEDPYVEKVVPSLL
KEGFVPKGLIRTAQVGRPDNLSPFNQVNIYRFLYELCVPNLAVEHVKGKYEAFAPSLALKIEEHYVEDGSGDKEFH
LRPNMFWEPIIDPTLFPKNITLADSFLRPHVTAHDVKFYDDVVMNPPYVAEMRAVAMRSYFEDMVSVRVENDLKLIVR
WRAHTVRNEQGEEEKVLYSAFANTLALQPLPCFYVQHFANGEKIVPEDSDPDYRKDSVWAQNFSSHWAYNYIVSC
GAFRFAGMDDEKITLVRPNYHNPFAALVEKRYIYMKDSTDSLQDFKAGKVDIAYFPNHVNDLASFMTSAYKEQ
AARGEAIKKNSSDRYSYIGWNCLSLFFNNRSVRQAMNMLIDRRIEQLDGRGVSVSGPFTTLCACGAGGTVEGW
QYSPEEAARKLEEEGWIDADGDGIREKVIDGVVVPFRFLCYVKSVTARTIAEYVATVCKEVGIECCLLGLDMADY
SQALEEKNFADAILSGWCLGTPPEDPRALWHSEGALEKGSANAVGFCNEEADRIEQLSYEYDSNKRQALYHRFHEVI
HEESPYAFLYSRQYSLVYKEFVKNIFVPTHEQDLIPGAQDETVNLSMLWVDKEEGRCSAIS

SEQ ID NO:87: CT089 fragment nucleotide sequence - Length: 1194

GCTGCAGCTACTCAAGATGCACAAGAGGTTATCGGCTCTCAGGAAGCTTCTGAGGCAAGTATGCTCAAAGGATGTGA
GGATCTCATAAATCCTGCAGCTGCAACCCGAATCAAAAAAAGGAGAGAAGTTGAATCATTAGAAGCTCGTCGCA
AACCAACAGCGGATAAAGCAGAAAAGAAATCCGAGAGCACAGAGGAAAAAGGCGATACTCCTCTTGAAGATCGTTT
ACAGAAGATCTTCCGAAGTCTCCGGAGAAGATTTTTCGAGGATTGAAAAATTCGTTGATGATGATTCTTCTCCTGA
CGAAATCTCGATGCGCTCACAAGTAAATTTCTGATCCCAATAAAGGATCTAGCTCTTGATTATCTAATCAA
CAGCTCCCTCTGATGGGAAACTTAAAGTCCACTCTCATTTCAGGCAAAGCATCAACTGATGAGCCAGAATCTCAGGCG
ATTGTTGAGGAGCAGCAATGTTCTGTTAGCTTCAGAAAACCTTTGCTTCCAGAGCAAAATACATCTCCTTACGCTTCG
CTCCTTATATTTTCAAGTAACCTCATCCCCCTCTAATTTGCGCTAATTTACATCAAAATGCTTGTCTTACTTCCAT
CAGAGAAAACCGCTGTTATGGAGTTTCTAGTAAATGGCATGGTAGCAGATTTAAAAATCGGAGGGCCCTTCCATTCCT
CCTGCAAAATTTGCAAGTATATATGACGGAACCTAAGCAATCTCCAAGCCTTACACTCTGTAATAGCTTTTTGATAG
AAATATTGGGAACCTTGGAAAAATAGCTTAAAGCATGAAGGACATGCCCTTATCCATCCTTAAACGACAGGAAATTTAA
CTAAACCTTCTTACAATAGTAGAAGATAAATCCCTTCTTCCAAAGCTCAAAAGGCAATTAATGAACATGGTA
GGCCACAGTACGGTCTCAAACTGAAGTTTTAACTTACTTCCGCGCTCTTAAATGGCTTTCGCTAGAAATATT
CTCTGGAGCTGAAAAAAGCAGCAGCTGGCATCGGTTATCAAAATACGCTAGATGCGATAAATGCGGATAATGAGG

ATTATCCTAAACCAGGTGACTTCCCACGATCTTCTTCTAGTACGCCTCCTCATGCTCCAGTACCTCAATCTGAG
 ATTCCAACGTCACCTACCTCAACACAGCCTCCATCACCC

SEQ ID NO:88: CT089 / *lcrE* fragment protein sequence (AAC67680)

AAATQDAQEVIGSQEASEASMLKGCEDLINPAAATRIKKKGEKFESLEARRKPTADKAEKKSESTEKGDTPLEDRF
 TEDLSEVSGEDFRGLKNSFDDSSPDEILDALTSKFSPTIKDLALDYLIQTAPSDGKLGKSTLIQAKHQLMSQNPQA
 IVGGRNVLLASETFASRANTSPSSLRSLYFQVTVSSPNCANLHQMLASYLPSEKTAVMEFLVNGMVADLKSEGPSIP
 PAKLQVYMTLSNLQALHSVNSFFDRNIGNLENSLKHEGHAPIPSLTTGNLTKTFLQLVEDKFPSSSKAQKALNELV
 GPDTGPQTEVLNLFRRALNGCSPRIFSGAEKKQQLASVITNTLDAINADNEDYKPKGDFPRSSFSSTPPHAPVQSE
 IPTSPTSTQPPSP

SEQ ID NO:89: CT734 fragment nucleotide sequence - Length: 591

TGTTGCGCCAACCTCTTATGGATCGACTCTTGCAAAAAATACAGCCGAGATAAAAAGAAGAATCTGTTACACTTCGCGA
 GAAGCCGGATGCCGGCTGTAAAAAGAAATCTTCTTGTACTTGAGAAAATTTTTCTCGCGCAAGAAACCTAAAGAGA
 AGACAGAGCCTGTGTTGCCGAACCTTAAAGTCTTACGCAGATCCAATGACAGATCCGAAAAGAAAGACCTTCTTTC
 GTAGTATCTGCTGCTGCTGATAAGTCTTCTATTGCTTTGGCTATGGCTCAGGGGGAAATTAAGGCGCATTATCGCG
 TATTAGAGAGATCCATCCTTGCATTGTTACAAGCTTGCAGAAGATCCTGCTTTAATTGCTGGAATGAAAAAGA
 TGCAAGGACGGGATGGGTCTGGAATATCTTTATCACAGAATTAAGCAAAGTTTTTCTCAAGCAGCATCTTAAAGG
 GCTTTCAGCGTTGCGAGACGTTGCCGCGTTGCGCTCGACCTTAGGATTAGACTCGGGGACCGTTACCTCAATTGTTGA
 TGGGGAAAGGTGGGCTGAGCTGATCGATGTCGTGATTAGAACCTGCTATA

SEQ ID NO:90: CT734 fragment protein sequence (AAC68329)

CCANSYGSTLAKNTAEIKESVTLREKPDAGCKKKSSCYLRKFFSRKKPKKEKTEPVLPNFKSYADPMTDSERKDLSE
 VVSAAADKSSIALAMAQGEIKGALSRIREIHLALLQALAE DPALIA GMKMMQGRDWWVWNIFITELSKVFSQAASLG
 AFSVADVAFASTLGLDSGTVTSIVDGERWAEIDVVIQNP AI

SEQ ID NO:91:CT016 fragment nucleotide sequence

AAAGTTAAAATTAATGATCAGTTTCATTTGTATTTCCCATACATTTCTGCTCGATGGAATCAGATAGCTTTCATAGA
 GTCTTGTGATGGAGGGACGGAAGGGGGTATTACTTTGAACTCCATTTAATTGATGGAGAGACAGTCTCTATACCTA
 ATCTAGGACAAGCGATTGTTGATGAGGTGTTCCAAGGACACTTGCTATATTTAGAGTCCACAGCTCCTCAGAAAAAC
 AAGGAAGAGGAAAAAATTAGCTCTTTGTTAGGAGCTGTTCAACAAATGGCTAAAGGATGCGAAGTACAGGTTTTTTC
 TCAAAAGGGCTTGGTTCTATGTTACTAGGAGGAGCTGGTTCGATTAATGTGTTGTTGCAACATTCTCCAGAACATA
 AGGATCATCCTGATCTTCTACCGATTTACTGGAGAGGATAGCGCAAATGATGCGTTCATTATCTATAGGACCAACT
 TCTATTTTAGCTAAGCCAGAGCCTCATTGCAACTGTTGCAATTGTCAAATGGACGAGCTACAGTGGAGAAGAGGA
 TGCCGGAGTATCGGATGAGGATCTTACTTTTCTGTTTCAATGGGATATCTCTCAAAGTGGAGAAAAGATGTACTGTTA
 CAGATCCTTTGATCCAGAAGAGCAGTTTAATGTGATTTAGGAACCGCGATTGGATGCACATGTGGGCAGCCATAC
 TGTGAACACGTAAGCTGTTCTTTATACT

SEQ ID NO:92: CT016 fragment protein sequence (AAC67606)

KVKINDQFICISPYISARWNQIAFIESCDGGTEGGITLKLHLIDGETVSIPLNGQAIVDEVFQEHLLYLESTAPQKN
 KEEEKISSLLGAVQMAKGEVQVFSQKGLVSMLLGGAGSINVLLQHSPEHKDHPDLPTDLLERIAQMMRSLSIGPT
 SILAKPEPHCNCLHCQIGRATVEEEDAGVSDDELTFRSWDISQSGEKMYTVTDLNPEEQFNVYLGTPIGCTCGQPY
 CEHVKAVALYT

SEQ ID NO:93: CM homolog of CT279 = TC_0551 fragment nucleotide sequence

GCATCCAAGTCTCGTCATTATCTTAACCAGCCTTGGTACATTATCTTATTTCATCTTTGTTCTTAGTCTGGTTGCTGG
 TACCCTTCTTTCTCAGTTTCTATGTTCTATCTCCAATCCAAAAACAAGCTGCAGAATTTGATCGTAATCAGCAAA
 TGTTGATGGCCGCACAAATTTCTATGACAATAAATCCAAATATATGCTGAAGGGGATTGGCAACCTGCTGTC
 TATAATACAAAAAACAGATACTAGAAAAAAGCTTCCACTCCACCACAAGTACTGTGGCGACTCTATGCTCTTA
 TTTTCAAAATTTGTTAGAGTTTTGCTTACAGACTCCCAAGGGAATCTTTCTTTTGAAGATACAATCTTAACC
 TAGAAGAGTTCTTATCCACCCACATCTTCAGTACAAGATCACTCTCTGCATGTAATTTATGCTATTCTAGCAAAC
 GATGAATCCTTAAAAAGTTATCATCCTCCCAAGTAGCAAAAAATCCGGTATCCATAGAGTCTATTATTTCTTCTAT
 AAAAGGATTTGGTTTATGGGGACCAATCTATGGATTTCTTGCTTTAGAAAAGGACGGTAATACGGTTCTAGGGACAT
 GCTGGTATCAACATGGTGAGACTCCAGGATTAGGAGCAAATATAACTAATCCCCAATGGCAACAAAAATTTAGAGGA
 AAAAAAGTATTTCTGCTTCTCTTCCGGAGAAACCGATTTTGTCTAAAACAACCTTAGGACTAGAAGTTATAAAGG
 ATCTGTTTCTGCATTATTAGGGGACTCTCCAAAGCTAATTCCGCTGTTGATGGAATTTAGGAGCTACACTGACCT
 GTAATGGAGTTACTGAAGCTTTTGCTAATTCGCTAGCTCCTTACCGCCCCCTTATTGACTTTCTTCGCCAATCTTAAC
 TCTAGTGGAGAATCTCATGACAACCAA

SEQ ID NO:94: CM homologue of CT279 = TC_0551 fragment protein sequence

ASKSRHYLNQWPYIILFIFVLSLVAGTLLSSVSYVLSPIQKQAAEFDRNQMLMAAQIISYDNKFQIYAEGDWQPAV
 YNTKKQILEKSSSTPPQVTVATLCSYFQNFVRVLLTDSQGNLSSFEDHNLNLEEFLSHPTSSVQDHSLSHVIYAILAN
 DESSKKLSSSQVAKNPVSIESIILPIKGFGLWGPYIGFLALEKDGNTVLGTCWYQHGTEPGLGANITNPQWQNFQ
 KKVFLASSSGETDFAKTTLGLEVIKGSVSALLGDSPKANSVAVDGISGATLTCNGVTEAFANSLAPYRPLLTFANLN
 SSGESHNDQ

SEQ ID NO:95: CM homologue of CT372 = TC_0651 fragment nucleotide sequence

AATGGAAAAGTTCTGTGTGAGGTTTCTGTGTCCTCCGTTTCGATTCTGCTGACGGCTCTGCTTTCACTTTCTTTTAC
AAACACTATGCAGGCTGCACACCATCATTATCACCGTTATGATGATAAACTACGCAGACAATACCATAAAAAGGACT
TGCCCACTCAAGAGAATGTTTCGAAAAGAGTTTTGTAATCCCTACTCTCATAGTAGTATCCTATCCCTTTGTACAA
CAACGAGGAGTCCATCTCCTATCTGTGATTTAGTCTCAGAGTGCTCGTTTTTGAACGGGATTTCCGTTAGGAGTCT
TAAACAAACACTGAAAAATTCTGCTGGGACTCAAGTTGCTTTAGACTGGTCTATCCCTTCTCAATGGTTCAATCCTA
GATCCTCTTTGGGCTCCTAAGCTCTCTATTCCGAGATCTGGATATGGTAAACCCAGTCCCTTATTGAAGCAGATTCC
CCTTGTGTCAAACCTGCTTCAACCCATCTGCTGCTATTACGATTTACGATTCTTCATGTGGGAAGGGTGTGTCCA
AGTGTACATACACCTTGTTCGTTATTGGAGAGAAACGGCTGCACCTTGACGGGCAAACTATGATGCTTGCAGGAAGTA
TTAATGATTATCCTGCTCGCCAAAACATATTCTCTCAACTTACATTTTCCAAAACCTTCCCTAATGAGAGAGTAAAT
CTAACTGTTGGTCAATACTCTCTTTACTCGATAGACGGAAACGCTGTACAACAATGATCAGCAGCTAGGATTTATTAG
TTATGCGTTGTGCGAAAAATCCAACAGCGACTTATTCCTCTGGAAGCCTTGGCGCCTATCTACAAGTCGCTCCAACAG
AAAGCACCTGTCTTCAAGTTGGGTTCCAAGATGCCTATAATATTTTCAGGTTCTCGATCAAATGGAATAATCTTACA
AAAAATAAGTATAACTTCCATGGCTATGCATCTTGGGCTCCACACTGTTGCTTAGGACCTGGACAATACTCTGTTCT
TCTTTATGTAACCCAGAAAGGTTCTGAGCAATGATCGAGACAATGGGCTGGTCTGTGAATGCAAGTCAATACATCT
CTTCTAAACTTTATGATTTTGGAAAGATACAGCGGAGTCAAGGCAATTGTCTCTATTAACCGAACCTATTCATTT
GGCTTAGTCTCTCCTAATTTATTGAACCGTAACCCACAAGACTTATTTGGAGTAGCTTGCAGCTTCAATAATATACA
CGCCTCCGCTTTCAAATGCTCAAAGAAAATATGAACTGTGATCGAGGGATTGCAACTATTGGTTGCGGACCTT
ACATCTCCTTTGCTCCAGATTTCAAACCTTACCTCTATCTGCTCTGCGTCAAATAAAACAAAGCGCCGAGTCTAT
AGCGTTCGCGCAAACCTAGCTATT

SEQ ID NO:96: CM homologue of CT372 = TC_0651 fragment protein sequence

NGKVLCEVSVSFRSILLTALLSLSFTNTM Ω AAHHHYHRYDDKLRRQYHKKDLPTQENVRKEFCNPYSHSSDPIPLSQ
QRGVLSPICDLVSECSFLNGISVRS Ω TLKNSAGTQVALDWSILPQWFNPRSSWAPKLSIRD Ω LYGKPKQSLIEADS
PCCQTCFNPSAAIT Ω YDSSCGKGVVQVS Ω YTLVRYWRETAALAGQTMMLAGSINDYPARQNI Ω IFSQ Ω LTFSQ Ω TFPNERVN
LTVGQYSLYSIDGTYLNN Ω Q Ω LG Ω FISYALS Ω NPATYSSGSLGAYLQVAPTESTCLQV Ω FDAYNISGSSIKWNNLT
KNYNFHGYASWAPHCLGPG Ω YSVLLYVTRKVP Ω MM Ω TMGWSVNAS Ω YISSKLYVFG Ω RYSGVTG Ω LSPINR Ω YTSF
GLVSPNLLNRNP Ω DLFGVACFN Ω IHASAF Ω NA Ω RKYETVIEGFATIGCGPYISFAPDF Ω LYLYPALRPNK Ω SARVY
SVRANLAI

SEQ ID NO:97: CM homologue of CT443 = TC_0727 fragment nucleotide sequence

AGCGGGGTGTTAGAGACCTCTATGGCAGAGTCTCTCTACCAACGTTATTAGCTTAGCTGACACCAAAGCGAAAGA
GACCACTTCTCATCAAAAAGCAGAAAAGCAAGAAAAATCATCAAAATAGGACTTCCGTAGTCCGTAAGAGGTTA
CTGCAGTTCTGTGATACTAAAGCTGTAGAGCCTAGACAGGATTTCTGCTTTGGCAAAATGTATACAGTCAAAGTTAAT
GATGATCGTAATGTAGAAATCGTGCACTCCGTTCTGAAATATGCTACGGTAGGATCTCCATATCCTATTGAGATTAC
TGCTATAGGGAAAAGAGACTGTGTTGATGTAATCATTACACAGCAATTACCATGCGAAGCAGAGTTTGTAGCAGTG
ATCCAGCTACTACTCTACTGCTGATGGTAAGCTAGTTTGGAAAATTGATCGGTTAGGACAGGGCGAAAAGAGTAAA
ATTACTGTATGGGTAAAACTCTTAAAGAAGGTTGCTGCTTTACAGCTGCAACGGTTTGTGCTTGTCCAGAGATCCG
TTCGGTTACGAAATGTAGCAGCCTGCTATCTGTGTTAAACAGGAAGGTCAGAAAAGCGCATGTTTGGCTTGGCCAG
TAACTTATAGAATTAATGTAGTCAACCAAGGAACAGCAACAGCACGTAATGTTGTTGTGGAAAATCCTGTTCCAGAT
GGCTATGCTCATGCATCCGGACAGCGTGTATTGACATATACTCTTGGGGATATGCAACCTGGAGAACAGAGAACAAAT
CACCGTGGAGTTTTGTCCGCTTAAACGTTGGTCCGAGTCACAAATATTGCTACAGTTTCTTACTGTGGTGGACACAAAA
ATACTGCTAGCGTAACAACAGTGATCAATGAGCCTTGGCTGCAAGTTAACAATCGAGGGAGCAGATTGGTCTTATGTT
TGTAAGCCTGTAGAAATGTTATCTCTGTTTCAACCTGGTGACTTAGTTTTACGAGACGTTGTAATTGAAGATAC
GCTTTCTCCTGGAATAACTGTTGTTGTAAGCAGCTGGAGCTCAGATTTCTTGTAAATAAATTGGTTTGGACTTTGAAAGG
AACTCAATCCTGGAGAGTCTTTACAATATAAGGTTCTAGTAAGAGCTCAAACCTCAGGGCAATTCACAAACAACGTT
GTTGTGAAAAGTTGCTCTGATTGCGGTATTTGACTTCTTGGCAGAAAGCAACAACCTTACTGGAAAGGAGTTGCTGC
TACTCATATGTGCGTAGTAGATACTTGTGATCCTATTTGCGTAGGAGAGAACAACCTGTTTATCGTATCTGTGTGACAA
ACAGAGGTTCTGCTGAAGATACAAATGTGTCCTTAAATTTGAAATTTCTAAAGAATTACAACCTATATCTTTCTCT
GGACCAACTAAAGGAACCAATTACAGGAAACACGGTAGTGTGTTGATTCGTTACCTAGATTAGGTTTAAAGAAACTGT
AGAGTTTTCTGTAACGTTGAAAGCAGTATCCGCTGGAGATGCTCGTGGGGAAGCTATTCTTTCTTCCGATACATTGA
CAGTTCCTGTATCTGATACGGAGAATACACATATCTAT

SEQ ID NO:98: CM homologue of CT443 = TC_0727 fragment protein sequence

SGVLETSMAESLSTNVISLADTKAKETTSHQKDRKARKNHQNR Ω TSVVRKEVTAVRD Ω TKAVEPRQDS Ω CFGKMYTVKVN
DDRNEIV Ω SVPEYATV Ω SPYIEITAIGKRD Ω VDVIT Ω QLPCEAEFVSSDPATTP Ω ADGKLVWKIDRL Ω GQGEKSK
ITVWVKLKEGCCF Ω TAATV Ω CACPEIRSVTKCG Ω PAICVK Ω QEGPESACLRC Ω PVYRINVVN Ω GTATARNVV Ω ENPVP Ω
GYAHASGQRVLT Ω Y Ω LD Ω M Ω Q Ω PE Ω QRTITVEFC Ω PLKRGRV Ω TNIATV Ω SYCGGHKNTASV Ω TTVIN Ω PCV Ω VNIEGAD Ω WSYV
CKPVEYVISVSNP Ω DLVLR Ω VDVIED Ω TLSPGITV Ω VEAGA Ω QISCNKLV Ω WTLKELN Ω PGESL Ω QYKVLVRA Ω TPG Ω Q Ω TNNV
VVKSCSD Ω CGICTSCAEAT Ω Y Ω WK Ω VAA Ω THM Ω CV Ω DTCD Ω PICVGENTVYR Ω ICV Ω TNR Ω GS Ω AED Ω TNVSL Ω LK Ω FSKEL Ω Q Ω ISFS
GPTKGTITGNTV Ω DFSLPRL Ω GSKETVEFSV Ω TLKAVSAGDARGE Ω AILSSD Ω TLV Ω VPVSD Ω TENTHIY

SEQ ID NO:99: CM homologue of CT043 = TC_0313 fragment nucleotide sequence

TCCAGACAGAATGCTGAGGAAAATCTAAAAATTTTGGCTAAAGAGCTCAAGCTCCCCGACGTGGCCTTCGATCAGAA
TAATACGTGCATTTTGTGTTGATGGAGAGTTTTCTCTCACCTGACCTACGAAGAGCACTCTGATCGCCTTTATG
TTTACGCACCTCTCCTTGACGGACTCCAGATAATCCGCAAAGAAAGTTGGCTCTGTATGAGAAATTGTTGGAAGGC

TCTATGCTCGGAGGCCAAATGGCTGGTGGAGGAGTAGGAGTTGCTACTAAAGAACAGTTGATCCTAATGCATTGCGT
GTTAGATATGAAATATGCAGAGACTAATCTATTGAAAGCTTTTGCACAGCTTTTCATTGAAACTGTTGTGAAATGGC
GAACGGTCTGTTCTGATATCAGCGCTGGACGAGAACCTCCGTTGACACTATGCCTCAAAATGCCTCAAGGAGGCAGC
GGAGGAATTC AACCTCCTCCAACAGGAATTCGTGCG

SEQ ID NO:100: CM homologue of CT043 = TC_0313 fragment protein sequence

SRQNAEENLKNFAKELKLPDVAFDQNNTCILFVDGEFSLHLYEEHSDRLYVYAPLLDGLPDNPQRKLALYEKLLLEG
SMLGGQ MAGGGVGVATKEQLILMHCVLDMKYAETNLLKAF AQLFIETVVKWRTVCS DISAGREPSVDTMPQMPQGGG
GGIQQPPPTGIRA

SEQ ID NO:101: CM homologue of CT601 = TC_0890 fragment nucleotide sequence

CTCGTAATCGGTTATTTCTAATCACCCCTTATAGGTTTTGGCTATTCTGCTTACGGTGCCAGCACAGGGAAATCAC
TTCTTTACAGGTTATTTAGCTGAAGTCGAGGATACATCTTCGCGCTTACAAGCTCATCAGAATGAGCTTGTATGC
TCTCGAACGTTTAGATGAGCAAGACACAAAACCTCAACAACCTCTCGTCAACTCAGGCCCGTAATCTTCTCAACAA
GTTCAACGGCTTGAGATTGATCTGAGAGCTCTGGCTAAAAACAGCTGCTGTGCTCTCGCAATCTGTTCCAGGATATCCG
ATCATCCGTGCAAAAATAAATTACAAGAAATCCAACAAGAACAAAAAATTTAGCTCAAAATTTACGAGCGCTTCGCA
ACTCCTTACAAGCACTAGTTGATGGCTCTTCCCAGAAAAATTATATTGATTTTTGGCCGGGAGACACCTGAACAT
ATTCAGTTGTTAAACAAGGAGAAACCTTGAGTAAATCGTAGTAAGTACAATATCCCTGTGCGAGAATTGAAAAA
ACTTAATAAATAAATTCGATACTATTTTTACTGATCAAGAATCCGACTTCAAAAAAGAAA

SEQ ID NO:102: CM homologue of CT601 = TC_0890 fragment protein sequence

LANRFLITLIGFGYSAYGASTGKSPSLQVILAEVEDTSSRLQAHQNELVMLSERLDEQDTKLQQLSSTQARNLPQ
VQRLEIDLRLAKTAAVLSQSVQDIRSSVQNKLRQEIQRQKNLRAQNLRLRNSLQALVDGSSPENYIDFLAGETPEH
IHVVKQGETLSKIASKYNIPVAELKKNLNSDTIFDQRIRLPK

SEQ ID NO:103: CM homologue of CT456 = TC_0741 fragment nucleotide sequence

ACGACTCCAATAAGTAATCTCCATCTTCTATTCCAACCTGTTACAGTATCAACTACTACAGCATCTTCTGGATCTCT
CGGAACCTTACTGTATCATCAACGACTACAAGTACTTCAGTCGCACAAACAGCAACAACAACATCTTCTGCTTCTA
CATCTATAATTCAGTCTAGTGGAGAAAACATCCAATCCTACAGGTACCCCTTCTCTATTACGTCTAGTGTTC
ACATCCGCTCCATCTCCTAAAGCCTCCGCCACTGCAACAAAACCTCAAGCGCTGTTTCTGGGAAAATACCTCAC
AGAACTTCTGAGGAATCCGAAACCCAAAGCCACTACATCTGATGGAGAAGTTAGTAGTAATTACGATGATGTTGATA
CCCCGACCAATTCGTCGATTTCGACAGTTGATAGTGATTAACCAAGATGTTGAGACTCAGTACAAAACAATTAGCAAC
AATGGTGAAAACACTTATGAAAACAATCGGAAGTCAATGGTGAGAAAAACACACACGTCCAGGAAAGCCATGCATCCGG
AACAGGAAATCCCATAAATAATCAGCAAGAAGCTATTAGACAGCTCCGATCATCTACCTATAACAACCAGCCCTCGTA
ATGAGAATATATTTAGTCCAGGACCGGAAGGTCTACCTAATATGTCTCTTCTAGTTACAGCCCTACAGATAAAAGT
TCTCTACTAGCTTTCTATCTAATCCCAATACAAAAGCAAAAATGCTCGAACACTCCGGGCATTTAGTCTTTATAGA
CACACTAGAAAGTAGCTTTATCTTTGTTCCGAATGGAAATTTGGGATCAAGTCTGTTCCATGAAGGTTCCAGATGGGA
AACTAAAGAAGACCTTGGCTTAAAGGACTTAGAAGATATGTGTGCAAGTTTTGCACAGGATACAATAAATCTCC
TCTGATTGGGGAAATCGAGTTGACCCCTTGGTCTCTTCTAAGGCCGGGATAGAAAGTGGGGGGCACCTCCCAAGCTC
AGTTATCATCAACAACAATTTAGAACCTGTGTTGCCATGAGGCGTGGAAACCCAAAGAAAACGGCCCAATTTATA
CTCCTTACGCTGGAGACGTGGGCATCGAGTAGATTTTGGAAAGATCTTTGATGGAACAGCGCCGTTTAAATAAAATC
AATCGGGCTCTTCCCTTACCCCTGGTGATGACGGCATCTCCTTCTAATGAAACTATTGGGTCTGAACCATTCGC
GACACCTCCCTCATCCCATCGCAACCCCGTTATCAACGTCAATGTTAATGTCGGTGGAAACCAATGTTAATATTG
GGGATACAAACGTATCTAAAGGATCCGGCACACCAACATCTTCTCAATCTGTGGACATGTCTACAGATACTAGCGAT
TTAGATACCAGTGATATTGATACAAACAACCAACCTAACGGCGATATCAACACGAATGACAACCTCCAATAATGTCGA
TGGAAGTTTATCTGACGTTGATTCAAGGGTGGAAAGACGATGACGGTGTATCGGATACAGAGTCCACTAATGGCAATG
ACTCTGGTAAAACACTTCCACAGAAGAAAATGGTGACCCAAAGCGGACAGACATCCTGGCTGCTGTACGTAAACAC
CTAGACACTGTCTATCCAGGAGAAAATGGCGGATCTACAGAAGGACCTCTCCCTGCTAATCAAAATCTGGGGAACGT
TATCCATGATGTGGAGCAGAATGGATCTGCTAAAGAAAATATTATCACTCCAGGAGATACAGGGCCTACAGACTCAA
GCTCCTCTGTAGATGCTGATGCAGACGTTGAAGATACTTCTGATACTGACTCTGGAATCGGAGACGACGACGGTGT
TCGGATACAGAGTCCACTAATGGTAATAACTCTGGTAAAACACTTCCACAGAAGAAAATGGTGACCCAAAGCGGACC
AGACATCCTGGCTGCTGTACGTAACACCTAGACACTGTCTATCCAGGAGAAAATGGCGGATCTACAGAAGGACCTC
TCCCTGCTAATCAAAATCTGGGGAACGTTATCCATGATGTAGAAACAAAACGGAGCCGCTCAAGAAAATATTACTACT
CCAGGAGATACGGAATCTACAGACACAAGCTCTAGTGTAATGCTAATGCAGACTTAGAAGATGTTTCTGATGCTGA
TTCAGGATTCGGGGATGATGACGGTATATCGGATACAGAGTCCACTAATGGTAAACGACTCTGGAAAAAATACTCCTG
TAGGGGATGGTGGTACACCAAGCGGACAGATATCCTAGCTGCTGTACGCAACATCTAGACACTGTCTATCCAGGA
GAAAATGGTGGATCTACAGAGAGACCTTTACCCGCTAATCAAAAATTTAGGAGATATCATTACATGATGTAGAACAAA
CGGAAGCGCTAAAGAACTGTAGTATCGCCTTATCGAGGAGGAGGAGAAAATACATCTTCCCAATTTGGATTAGCTC
CCCTGCTTCCAGCAACACCATCCACACCTTTGATGACAACACCTAGAACAAAATGGGAAAGCTGCAGCTTCTTCTTG
ATGATAAAAGGAGGAGAACTCAAGCCAAGCTAGTTAAGAATGGCGGCAATATCCCTGGAGAAAACACATTAGCAGA
ATTACTCCCTCGTTTAAAGAGGACACCTTGACAAAGTCTTTACTTCAGACGGGAAAGTTTACAAATCTTAATGGACCTC
AACTTGGAGCCATCATAGACCAATTCGCAAAAGAAAACGGGTTCCGGAGGAATCATAGCTCATAACAGATAGTGTCCA
GGAGAGAACGGAAACAGCCTCTCCTCTCACAGGAAGTTCAGGGGAAAAAGTCTCTCTATGATGCAGCGAAAAACGT
CACTCAAGCTTTAAACAAGTGTACGAACAAAGTAACTTAGCAATGCAAGGACAAAAACTGGAAGGAATTTAAACA
ACAACAATACCCCTCTTCTATTGGACAAAATCTTTTCGACGACGAGGGGCAACGACACAATCCCTCAGTTCATTA
ATTGGAACCGTACAA

SEQ ID NO:104: CM homologue of CT456 = TC_0741 fragment protein sequence

TTPISNSPSSIIPTVTVSTTTASSGSLGTSTVSSSTTTSTVSVAQTATTTSSASTSIIQSSGENIQSTTGTPSPITSSVS
TSAPSPKASATANKTSSAVSNGKITSQETSESEETQATTSDEGEVSSNYDDVDPTNSSDSTVDSYQDVETQYKTISN
NGENTYETIGSHGEKNTHVQESHASGTGNPINNQQEAI RQLRSSTYTTSPRNENIFSPGPEGLPNMSLPSYSPTDKS
SLLAFLSNPNTKAKMLEHSGHLVFDITTRSSFIFVPNGNWQVCSMKVQNGKTKEDLGLKLEDMCAKFCCTGYNKFS
SDWGNRVDPLVSSKAGIESGGHLPSSVIINNKFRTCVAYGPWNPKENGPNYTPSAWRRGHRVDFGKIFDGTAPFNKI
NWGSSPTPGDDGISFSNETIGSEPFATPPSSPSQTPVINVNPNVGGTNNVIGDTNVS KGS GTP TSSQSVDMSTDTSD
LDTSDIDTNNQTNQDINTNDNSNNVDGSLSDVDSRVEDDDGVSDTESTNGNDSGKTTSTEENGDPSPGDILA AVRKH
LDTVYPGENGGSTEGPLPANQNLGNVIHDVEQNGSAKETIITPGDTGPTDSSSSVDADADVEDTSDTDSGIGDDDG
SDTESTNGNNSGKTTSTEENGDPSPGDILA AVRKHLDTVYPGENGGSTEGPLPANQNLGNVIHDVEQNGAAQETIIT
PGDTESTDTSSSVNANADLEDVSDADSGFGDDDGISDTESTNGNDSGKNTPVGDGGTSPGPDILA AVRKHLDTVYPG
ENGGSTERPLPANQNLGDIHDVEQNGSAKETVVSYPYRGGGNTSSPIGLASLLPATPSTPLMTTPRTNGKAAASSL
MIKGGETQAKLVKNGGNIPGETTLAELLPRLRGHLKVFVTS DGKFTNLNGPQLGAIIDQFRKETGSGGIIAHTDSSVP
GENGTASPLTGSSGEKVS LYDAAKNVTQALTSVTNKVTLAMQGQKLEGIINNNTPSSIGQNLFAAARATTQSLSSL
IGTVQ

SEQ ID NO:105: CM homologue of CT381 = TC_0660 fragment nucleotide sequence

TGTTCAAAAGAGAGCAAAGACTCTGTTAGTGAAAAATTTATTGTAGGAACTAACGCAACGTATCCTCCTTTTGAGTT
TGTTGATGAAAGAGGTGAGACGGTTGGCTTTGATATTGATTTAGCTAGGGAGATTAGTAAAAAGCTAGGGAAAAAAT
TAGAAGTCCGAGAATTTGCTTTTGTGACTCGTTCTCAATTTAAAACAGCATCGTATTGATGCAATTTAGCCAGGG
GTGTCCATTACGTCTTCGATTGAAAGAAAATTTGATGATTTCCCTACTATGGCGAAGAAATAAAGAGTTTTGTTTT
AGTGTTTAAGGATGGAGACTCAAAGTCTTTACCACTAGATCAGTATAAATCTGTTGCTGTTCAAACCTGGCACGTACC
AAGAGGAATATTTACAGTCTCTCCAGGGGTGCGTATTCGCTCTTTTGTAGTACTTTAGAAGTGCTTATGGAAGTT
TTGCATAGCAAGTCTCCTATAGCTGTTTTAGAACCGTCTATTGCGCAGGTGCTTTTAAAAGATTTTCCGACGCTCAC
TACTGAAACGATAGATCTTCTGAAGATAAATGGGTTTTAGGGTATGGAATTGGAGTTGCTTCTGATCGACCATCTC
TAGCTTCTGATATAGAAGCTGCTGTACAAGAGATCAAGAAAGAAGGAGTGTAGCAGAGTTAGAGCAAAAAATGGGGT
TTGAACGGC

SEQ ID NO:106: CM homologue of CT381 = TC_0660 fragment protein sequence

CSKESKDSVSEKFI VGTNATYPPFEFVDERGETVGF DIDLAREISKKLGKKLEVREFAFDALVNLKQHRIDAIMAG
VSITSSRLKEILMIPYGGEEIKSLVLFKDGDSKSLPLDQYNSVAVQGTGYQEEYLQSLPGVIRISFDSTLEVLMEV
LHKSPIAVLEPSIAQVVLKDFPTLTETIDLPEDKWLGYGIGVASDRPSLASDIEAAVQEI KKEGVLAELEQKWG
LNG

SEQ ID NO:107 – CT255 fragment nucleotide sequence

GAAGAAAAAGGCATCTTACAATTTGGTTGAAATTTGCGGAGCAATGGCTTTACAGGGAGTTTGTCTTGGACTAATTT
ACAGAGTGTGGAGTCTATGTTGCAGTATATAGCAGGGAGTGTGAGGAGTTGGCTGATGCTGTACAAGAAAAATAAG
CTTCGTTGGAAATCGCTTCGGAAGCCGGAGACGTACTTACTTTAGTATTGACCTTGTGTTTTCTTGCTAGAAAGAGAA
GAAAGCTTTAAAGCTGAAAGATATTTGTAGAAGCTTTGGCTAAGTTGCGTCGTCGATCTCCTCATGTTTTGATCC
TCATAATCAAATTTCTTTAGAACAGGCTGAAGAATACTGGGCTCGTATGAAACAGCAAGAAAAAATTTCT

SEQ ID NO:108 – CT255 fragment protein sequence

E EK G I L Q L V E I S R M A L Q G V C P W T N L Q S V E S M L Q Y I A G E C Q E L A D A V Q E N K A S L E I A S E A G D V L T L V L T L C F L L E R E
G K L K A E E V F V E A L A K L R R R S P H V F D P H N Q I S L E Q A E E Y W A R M K Q Q E K I S

SEQ ID NO:109 – CT341 fragment nucleotide sequence

GATTACTACACGATATTGGGTGTAGCGAAGACTGCTACTCCTGAAGAAATAAAGAAAGCTTACCCTAAGCTCGCTGT
AAAGTACCATCCAGATAAGAATCCTGGGGATGCTGAAGCGGAGCGACGCTTTAAAGAAAGTTTCTGAAGCCTATGAAG
TATTAGGTGATGCGCAGAAGCGGGAGTCATATGATCGTTACGGCAAAGACGGTCCATTTGCTGGTGCTGGAGGATTC
GGTGGCGCTGGCATGGGAATATGGAAGACGCTTTGCGAACATTTATGGGAGCTTTTGGCGGCGATTTCCGGTGGTAA
TGGAGGGCGTTTTCTTTGAAGGGCTTTTTGGAGGACTTTGGAGAAGCTTTCCGAAATGCGTGGAGGCTCAGAAAGTTCTC
GACAAGGAGCTAGTAAGAAGGTGCATATTACGCTGTCCTTCGAGGAGGCGGCAAAAAGGTGTTGAAAAAGAACTTCTT
GTTTCAGGCTATAAATCTTGTGATGCTTGTCTGGTAGTGGAGCCAATACTGCTAAAGGTGTAAAAGTTTTGTGATCG
ATGCAAGGGCTCTGGTCAGGTAGTGCAAAGCCGAGGCTTTTTCTCCATGGCTTCTACTTGCCTGATTGTAGTGGTG
AAGGTCCGGTTATCACAGATCCTTGTTCAGTTTGTGCTGGGCGAGGACGTATCAAGGATAAACGTAGCGTCCATGTT
AATATCCCAGCTGGAGTCGATTCTGGGATGAGATTAAGATGGAAGGCTATGGAGATGCTGGCCAAAATGGAGCGCC
TGCAGGGGATCTGTATGTTTTTATTGATGTAGAGCCTCATCCTGTTTTCGAGCGCCATGGGGATGATTTAGTTTTAG
CGTTTCTTATTGGATTTGTTGATGCGGCTTTAGGGATCAAGAAGGAAATCCCTACACTTTAAAAGAAGGTACTTGC
AGTTTTGAGTATCCCAAGAGGATTCAGAGCGGAACAGTTCTTAAAGTTAGAGGGCAGGGATTCCTAATGTGCATGG
GAAATCCAGAGGAGATCTTTTAGTAAGAGTATCTGTGGAGACTCCCAGCACCTATCTAATGAACAAAAAGATTTAT
TGAGACAGTTTGTCTACGGAGAAGGCTGAAAAATTTCCCTAAGAAACGGAGTTTCTTAGACAAAATCAAAGTTTTT
TTTTCTGACTTTGCTGTA

SEQ ID NO:110 – CT341 fragment protein sequence

DYYTILGVAKTATPEEIKKAYRKLAVKYHPDKNPGDAEAERRFKEVSEAYEVLGDAQKRESYDRYKDGPFAGAGGF
GGAGMGNMEDALRTFMGAFGGDFGGNGGGFFEGFLGGLGAEAFGMRGGSESSRQGASKKVHITLSFEEAAKGVKELL
VSGYKSCDACSGSGANTAKGVKVCDRCKGSGQVVQSRGFFSMASCTPDCSSEGRVITDPCSVCRGQGRICKDKRSVHV
NIPAGVDSGMRKMEGYGDAGQNGAPAGDLVYFIDVPEHPVFERHGDLDVLELPIGFVDAALGIKKEIPTLLKEGTC
RLSIPEGIQSGTVLKVRRQGFQPNVHGKSRGDLVVRVSVETPQHLSENEQKDLLRQFAATEKAENFPKKRSFLDKIKGF
FSDFAV

SEQ ID NO:111 – CT716 fragment nucleotide sequence

AATAAAAACTCCAAGATCTGTCTAAACTGCTCACTATTGAGCTTTTCAAGAAACGTACACGGTTGGAACAGTAAA
AAAAGCGCTCTCCACAATAGAACATCGCTTACAACAAATACAGGAGCACATCGCGAAAATTTCTTAACAAGGCACA
AACAAATTCCTATGTCGGTCATATACCCATGAATATGACCAACATTTAGAACATTTACAAAGAGAGCAAACCTTCTCTA
TATAAACAGCATCAGACCTGAAAACGTCTTTGAAAGATGCTTATGGCGACATACAAAAACAACCTAGACCAAAGAAA
AATTATCGAAAAGATCCATGACAGTAAATATCCTATAAAGAGCGCGAATAAC

SEQ ID NO:112 – CT716 fragment protein sequence

NKKLQDLSKLLTIELFKRTRLETVKKALSTIEHRLQIQIEHIAKISLTRHKQFLCRSYTHEYDQHLLEHLQREQTSL
YKQHQTLKTSKDAYGDIQKQLDQRKIIEKIHDSKYPIKSANN

SEQ ID NO:113 – CT745 fragment nucleotide sequence

CGGTGGTGGCTACACAAAACGATTCCCTCATGTGCAGCTGTCTATTCTAGAAAAAGAGTCTCGATCTGGAGGGCTAAT
TGTCACAGAGAAAACAACAAAGGGTTTTCCCTCAATATGGGCCCTAAAGGTTTTGTTTTAGCTCATGATGGGCAACACA
CCCTTACCTCATTAGCTTTTAGGCCTAGCAGACGAGCTATTATATAGCTCTCCAGAGGCTAAAAACCGCTTTATC
CACTATAATAATAAAAACCGAAAAGTCTCGCCTTGACTATTTTCAAACAAAATCTCCCTCTCTTTTTGCTAAGGA
TTTTCTTTGCGCGTCTTACAACAACAGACAGCTCCGTGGAAGCCTTCTTTAAAAGACACAGTTCTTCCAAGCTTAGAA
GAAATCTTTTAAATCCCATTAGCATTGCTATTCGTGCAGGACATAGTCATATATTGCTGCACAGATGGCTTACCCA
GAATTAACACGAAGAGAAGCTCAAACAGGATCGTTGTTACGTAGTTATCTCAAAGATTTTCTAAAGAGAAAACGCAC
AGGCCCTTATTTAGCTACCTTGCAGTCTGGGATGGGAATGCTAACCCAGGCTTTGCATGATAAATTGCCTGCTACCT
GTAATTTTTCTGACCCGTCAGCAAATCCGTCAGTTGGCGAATGGGAAAATTTCTTTTCACTCTCTCAAGGAGAA
ATAACGGGAGATATGCTCATTATGCTGGGTCCGTGCACGATCTCCCTTCTGTCTAGAAGGGATCCCTGAAAACCAA
GCTTATCAAGCAAACGACTTCATCTTGGGATCTCTCTTGTGTATCTTTAGGATGGCATGCATCCTTCCCTATCCCTC
ATGGATATGGCATGCTTTTTCGCTGATACGCCTCCCTTATTAGGGATCGTGTAAATACGGAAGTGTCCCTCAACCC
GAGCGGCCTAATACAATAGTCTCTCTTTTAGAAGGTCGATGGCACAAGAAGAAGCGTATGCTTTCTCACTAGC
AGCTATTTCTGAGTACCTGCAAATTTACACTCCCTCCCAAGCTTTCTCACTATTCTCTCTCTCAGAGAGGGACTTCCCC
AACACCATTGTGGATTTATCCAATCCCGCCAACGCTTCTATCTAAACTTCTCACAAATATAAAAATTTGATGGGCGAG
AATTTTGCAGGTCCAGGTCTCAACCGCGCTACAGCGTCTGCTTATAAAGCTATAGCTTCTTTACTATCA

SEQ ID NO:114 – CT745 fragment protein sequence

AWWLHKRFPVHQLSILEKESRSGGLIVTEKQQGFSLNMGPKGFVLAHDGQHTLHLIQLSLGLADELLYSSPEAKNRFI
HYNNKTRKQVSPWTFKQNLPLSFAKDF FARPYKQDSSVEAFFKRHSSSKLRRNLLNPISIAIRAGHSHILSAQMAYP
ELTRREAGTGLSLLRSYLKDFPKEKRTGYPYLATLRSGMGLTQALHDKLPATWYFAPVSKIRQLANGKISLSSPQGE
ITGDMLIYAGSVHDLPSCLEGIPETKLIKQTTSSWDLSCVSLGWHASFPIPHGYGMLFADTPPLLGIIVFNTVEVFPQ
ERPNTIVSLLLEGRWHQEEAYAFSLAAISEYLQIYTPPQAFSLFSPREGLPQHHVGFQSRQRLLSKLPNHIKIVGQ
NFAGPGLNRATASAYKAIASLLS

SEQ ID NO: 115 – CT387 fragment nucleotide sequence

ACGCTCTTTTCACTCATCATGATGCCGTCTCTCCAGACAGCTACCTATGTTCTTCCCTTCAGTTAGTTGGTACTGG
CGTATACGAAGGAGAAATCGAGATTCAAATATCCCTCTTATTTCTTGGATTCCAATTACCCTCTCATTGCATAC
ACCTTAATTTAAAGAGCTCTCTAGCTCAATTAGGAATAGATGCCTCCCTTCTCACTGCGAATTGAGCAAAAAACA
CATCGAGCACATATACATGCTCAATTTACCGGTCATGGCCCCATTGCTGAATCTATGCTAGCCCTTCTCCAACAGG
AGATCGTGTAGCAAAACTATTTGCTGCAGACGATCGCAGACTGGTCCGATCTCCAGATTACCTCGAAAGCATGCTGA
AAAATACAGATAAAGCTGGCCATCCTTTGCTCTGTTTTGGGAAAAAATTAGAACAACCTTGATTTCTTTTGATGTGGTA
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AGACTCTTGGCTCACAGGATCCGCAGCTGCCTATCAATATAGTGAGCAAGCAGATAAAAAAGAGTACACTCATGCTC
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CCTTCAGCTAGCTTAAAAGGAATGTTGATTTCTACCATGTGCGCCACTATCTCAAACAAATCTACTTTCAAGTTCC
CTCTTATACACATGGAAACTATTTCTCTATAATGACAGAGTTTGTCTATTAGATCTGCAGCAAGCAGATATTGATG
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CTACCCCACTTGCAGTCATCACCGGAGGCGGCACTGGAGTTATGGCCACAGGAAATCGTGTAGCTAAAGAACTAGGA
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TGCTAAAATGACATACCGCTACCTCAACTTATAGAAAAGGCAAGAACACTTTTATGACAGCTTCTATCCTTGTAG
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CATCCGTGGATCCGAATGGATCAGCAACTGCCTATATTGTATCACTTCTCCGGAAGCTGGAATTGCCATTGAAAC
AATTCCTAGCTGGAGAATCCCTATAGGATACGACTATCTCCAGCTCCAGATGGATTAGTGATCGTC

SEQ ID NO:116 – CT387 fragment protein sequence

TLFHSHHDAVSPDSYLCSLQLVGTGVYEGEIEIQNIPSYFLGFQLPSHCIHLNLKSSLAQLGIDASLLHCELSKNQ
HRAHIIHAQFTGHGPIAESMLALLQPGDRVAKLFAADRRLLVRSPTYLESMLKNTDKAGHPLLFCGKLEHLISFDVV
DDRLVLSLPTLPVVRYSYDIYGLLPLIQKLSLNPKLSIRHFLALYQIIVEGQHVSCGNHILLIKTEPLHIRTVIFAR
VVNQLLPQGLSHTSANILEPTTRESGDIFFGNPKAQIERIPLEFFTIEPYKEHSYFCNRDLLQITLQSESEIKKI
FETAPKEPVKAATYLSKGSEISSLHTDSWL TGSAAAYQYSEQADKNEYTHAQPYPFLEAMEMGLINSEGALLTRYF
PSASLKGMLISYHVRHYLKQIYFQVPSYTHGNYFSHNDRGLLLDLQADIDVFWADEESGRVLQYTKRRDKNSGMFV
IKNRVEEFRSAYFIAIYGSRLLENFSAQLHTLLAGLQAAHTLGIPGFSKPTPLAVITGGGTGVMATGNRVAKELG
ILSCGTVLDLEASPAQIDQPTNEFLDAKMTYRPLQIERQEHFYADLPILVGGVGTDFELYLELVYLKTGAKPPTP
IFLIGPIEYWKEKVAHAYEINLKAGTIRGSEWISNCLYCITSP EAGIAVFEQFLAGELPIGYDYPPAPDGLVIV

SEQ ID NO:117 – CT812 fragment nucleotide sequence

TGCGTAGATCTTCATGCTGGAGGACAGTCTGTAATGAGCTGGTATATGTAGGCCCTCAAGCGGTTTTATTGTTAGA
CCAAATTCGAGATCTATTCGTTGGGTCTAAAGATAGTCAGGCTGAAGGACAGTATAGGTTAATTGTAGGAGATCCAA
GTTCTTTCCAAGGAAAGATGCGGATACTCTCCCGGGAAGGTAGAGCAAAGTACTTTGTTCTCAGTAACCAATCCC
GTGGTTTTCCAAGGTGTGGAACCAACAGGATCAAGTCTTTCCCAAGGTTAATTTGTAGTTTTACGAGCAGCAACT
TGATTTCTCTCGTACGGGAGAAATCTTTTTAGGATTGCTTTTTGTTGGGATAGTAGTAAGGCTGGAATCACATTAA
CTGACGTGAAAGCTTCTTTGCTGGAGCGGCTTTATATTCTACAGAAGATCTTATCTTTGAAAAGATTAAGGGTGGAA
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AATTTGTGATGGGCTATATCTTTTGAAGGAAACAGCGCGAACTTTGCTAATGGAGGAGCGATTGCTCCTCGGAAA
AGTGCTTTTTGTGCTAATGATAAAAAGACTTCTTTTATAGAGAACCAGCTTTGTCTGGAGGAGCGATTGCAGCCT
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ACAGGGAAATGTTCTGTTTTATAACAACGTTGGCTGTTCCGGGAGGAGCTGTTCTGATAGAGGATCATGGTAATGTTCT
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CGGTTTTCTGATTTACAGATTGATGTAAGTCCAGAGATTGAAGAAGACACATACGGCCATATGGGAGATTGGTCT
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GCAGAGAATCTGGTTGCTATTGATGGATACAAAGGAGCTTATGGTGGTGCTTCTGCTGGAGTCGATATTCAATTGAT
GGAAGATTTTGTCTAGGAGTTAGTGGAGCTGCTTCTAGGTAATAATGGATAGTCAGAAGTTTGTATGCGGAGGTTT
CTCGGAAGGGAGTTGGTTCTGTATATACAGGATTTTATAGCTGGATCCTGGTCTTCAAAGGACAATATAGCCTT
GGAGAAACACAGAACGATATGAAAACGCGTTATGGAGTACTAGGAGAGTCGAGTGCCTTCTGGACATCTCGAGGAGT
NCDGAISFEGNSANFANGGAIASAASGKLVFVANDKKTFSFIENRALS SGGAI AASSDIAFQNC AELVFKGNCAIGTE DKG
ACTGGCAGATGCTTTAGTTGAATACCGAAGTTTAGTTGGTCTGTGAGACCTACTTTTTATGCTTTGCAATTTCAATC
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GACGCTTCCCTTACCAATATCACCATTCTTTAGGGATGAAGTTTGAATTGGCGTTCAAAAAGGACAGTTTTTCAGA
GGTGAACCTTTGGGAATAAGTTATGCATGGGAAGCTTATCGAAAAGTAGAAGGAGGCGCGGTGCAGCTTTTAGAAG
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SEQ ID NO:118 – CT812 fragment protein sequence

CVDLHAGGQSVNELVYVGPQAVLLLDQIRDLFVGSKDSQAEQRYRLIVGDPSSFQEKDADTLP GKVEQSTLFSVTNP
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LEFASCSLSLEGGACAQAASILIHDCQGLQVKHCTTAVNAEGSSANDHLGFGGGAFFVTGSLSGEKSLYMPAGDMVVA
NCDGAISFEGNSANFANGGAIASAASGKLVFVANDKKTFSFIENRALS SGGAI AASSDIAFQNC AELVFKGNCAIGTE DKG
SLGGGAISSLGTVLLQGNHGITCDKNESASQGGAI FGNKQISDNEGPVVFRDSTACLGGGAI AQAQEI VSIQNNQAG
ISFEGGKASFGGGIACGSFSSAGGASVLGTIDISKNLGAISFSRTLCTTSDLGQMEYQGGGALFGENISLSENAGVL
TFKDNIVKTFASNGKILGGGAILATGKVEITNNSGISFTGNARAPQALPTQEEFPLFSKKEGRPLSSGYSGGGAIL
GREVAIHLHNAAVFEQNR LQCEEEATLLGCCGGGAVHGM DSTSIVGNSSVRF GNNYAMGQGVSGGALLSKTVQLAG
NGSVD FSRNIASLGGGALQASEGNCELVDNGYVLF RDNRRGRVYGGGAI SCLRGD VVISGNKGRVEFKDNIATRL YVEE
TVEKVEEVEPEAPEQKDNNELSFLGRAEQSFITAAANQALFASDGDLSPESSISSEELAKRRECAGGAIFAKRVRIVD
NQEAVVFSNNFSDIYGGAI FTGSLREEDKLDGQIPEVLISGNAGDVVFSGNSSKRDEHLPH TGGGAICTQNL TISQ N
TGNVLFYNNVACSGGAVRIEDHGNVLL EAFGGDIVFKGNSSFRAGSDAIYFAGKESHITALNATEGHAI VFHDLV
FENLEERKSAEVLINSRENPGYTG SIRFLEAESKVPQCIHVQGSLELLNGATLCSYGFQDAGAKLVLAAGAKLK
ILD SGT PVQ QGHAI SKPEAEIESSSEPEGAHSLWIAKNAQTTPMVVDIHTISVDLASFSSSQEGTVEAPQVIVP GG
SYVRS GELNLELVNTTGTGYENHALLKNEAKVPLMSFVASGDEASAEISNLSVSDLQIHVVTP EIEEDTYGHMGDWS
EAKIQDGTLVISWNPTGYRLDPQKAGALVFNALWEEGAVLSALKNARFAHNLTAQRMEFDYSTNVWGF AFGGFRTLS
AENLVAIDGYK GAYGGASAGVDIQLMEDFVLGVSGAAFLGKMDSQKFD AEVSRKGVVGSVYTGFLAGSWFFKQYSL
GETQNDMKTTRYGLGESSASWTSRGLADALVEYRSLVGPVPRPTFYALHFNPYVEVSYASMKFPGFTEQGREARSFE
DASLTNITIVLMKFFELAFIKGQFSEVNSLGISYAWEAYR KVEGGAVQLLEAGFDWEGAPMDLPRQELRVALENNTE
WSSYFSTVLGLTAFCGGFTSTDSKLGYEANTGLRLIF

SEQ ID NO:119 - CT812N nucleotide sequence

TGCGTAGATCTTCATGCTGGAGGACAGTCTGTA AATGAGCTGGTATATGTAGGCCCTCAAGCGGTTTTATTGTTAGA
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GTTCTTTCCAAGAGAAAAGATGCGGATACTCTTCCCGGGAAGGTAGAGCAAAGTACTTTGTTCTCAGTAACCAATCCC
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SEQ ID NO:120: CT812N protein sequence

CVDLHAGGQSVNELVYVGPQAVLLLDQIRDLFVGSKDSQAEQYRLIVGDPSSFQEKDADTLPGKVEQSTLFSVTNP
 VVFQGVDDQDQVSSQGLICSFTSSNLDSPRDGESFLGIAFVGDSSKAGITLTDVKASLSGAALYSTEDLIFEKIKGG
 LEFASCSLSLEQGGACAAQSIILHDCQGLQVKHCTTAVNAEGSSANDHLGFGGGAFFVTGSLSGEKSLYMPAGDMVVA
 NCDGAISEFEGNSANFANGGAIASGKVLVANDKKTFSIENRALSGGAIASSDIAFQNCALVFKGNCAIGTEDKG
 SLGGGAISSLGTVLLQGNHGITCDKNESASQGGAIQKNCQISDNEGPVVRDSTACLGGGAIQAQEIIVSIQNNQAG
 ISFEGGKASFGGGIACGSFSSAGGASVLGTIDISKNLGAIQSFRTLCTTSDLGQMEYQGGALFGENISLSENAGVL
 TFKDNIVKTFASNGKILGGGAILATGKVEITNNSGISFTGNARAPQALPTQEEFPLFSKKEGRPLSSGYSGGGAIL
 GREVAIHLHNAAVVFEQNRQCSEEEATLLGCCGGGAVHGMDSISIVGNSSVRFGNMYAMGQVSGGALLSKTVQLAG
 NGSVDFSRNIASLGGGALQASEGNCELVDNGYVLFNRDNRGRVYGGAIISCLRQDVVISGNKGRVEFKDNIATRLYVEE
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SEQ ID NO:121: CT812C nucleotide sequence

GAAGAAGCTTGCAGAAAAGAGAGAGTGTGCTGGAGGAGCTATTTTTGCAAAACGGGTTCGTATTGTAGATAACCAAGA
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 AGTTAGATGGGCAAACTCCCTGAAGTCTTGATCTCAGGCAATGCAGGGGATGTTGTTTTTCCGGAATTCCTCGAAG
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 TGTTCTGTTTTATAACAACGTGGCCTGTTCCGGGAGGAGCTGTTCTGATAGAGGATCATGGTAATGTTCTTTTAGAAG
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 AATTCAGATGGAAGCTTTGTCATTAGTTGGAATCCTACTGGATATCGATTAGATCCTCAAAAAGCAGGGGCTTTAG
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 GATTGGGAGGGAGCTCCAATGGATCTTCTAGACAGGAGCTGCGTGTGCTCTGGAAAAATAATACGGAATGGAGTTC
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 CGAATACTGGATTGCGATTGATCTTT

SEQ ID NO:122: CT812C protein sequence

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 KESHITALNATEGHAIIVFDALVFENLEERKSAEVLINSRENPGYTGSIKIRFLAESAQKVPQCIHVQGGSLLELNGAT
 LCSIYGFKQDAGAKLVLAAGAKLKILDSGTPVQGHAIKPEAIESSSEPEGHSLWIAKNAQTTPVPMVDIHTISVD
 LASFSSSQEGTVEAPQIVPQGSYVRSSELNLELVNTTGTGYNHALLKNEAKVPLMSFVASGDEASAEISNLVSVS
 DLQIHVVTPTEIEEDTYGHMGDWSEAKIQDGTLVISWNPTGYRLDPQKAGALVFNALWEEGAVLSALKNARFAHNLTA
 QRMEFDYSTNVWGFAGGFRTLAENLVAIDGYKGYGGASAGVDIQLMEDFVLGVSGAFLGKMDSQKFDAEVSRK
 GVVGSVYTGFLAGSWFFKQYSLGETQNDMKTRYGVLGESSASWTSRGLVADALVEYRSLVGPVPRPTFYALHFNYPV
 EVSYASMFKPFGTEQGREARSFEDASLTNITPLGMKFEAFIKGQFSEVNSLGISYAWEAAYRKYVEGGAVQLLEAGF
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SEQ ID NO:123: CT869 fragment nucleotide sequence

AGAGAGGTTCTTCTAGAACTTTCTTATGCCCAACTCAGTTCAGATCCTACGAAAAGAGTGCCTATCAAATAAAAT
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 AAATCCCAATGAAGGAGCTGCTGTACAATAACAGATTACCTAAGCTTTTTGATACACAAAAAGAAGGTATTTAT
 TTTGCAAAAAATCTCACCCCTGAAAGTGGTGGTGGATTGGTTATGCGAGTCCCAATTCCTACCGTGGAGATTCTG
 TGATACAATAGGTCCTGTAATCTTTGAAAATAAATCTTGTTCAGACTATTTACATGGAGAAATCCTTATGCTGCTG
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 AGAAGCTTCTGATGGAGGAGCAATTAAGTAACACTACTCGCTAGATGTTACAGGCAATCGTGGTAGGATCTTTTTTA
 GTGACAATATCACAAAAAATATGGCGGAGCTATTTACGCTCCTGTAGTTACCTAGTGGATAATGGCCCTACCTAC
 TTTATAAAACAATACGCCAATAATAAGGGGGGGCGCTATCTATATAGACGGAACCAGTAACTCCAAAATTTCTGCCGA
 CCGCCATGCTATTATTTTAAATGAAAATATTGTGACTAAATGTAACATAATGCAAATGGTACAGTACGTACGTAATC
 CTCCTAGAAGAAATGCAATAACAGTAGCAAGCTCCTCTGGTGAATTTCTATTAGGAGCAGGGAGTAGCCAAAATTTA
 ATTTTTTATGATCCTATTGAAGTTAGCAATGCAGGGGTCTCTGTGTCTTCAATAAGGAAGCTGATCAAAACAGGCTC
 TGTAGTATTTTTCAGGAGCTACTGTTAATTCTGCAGATTTTCATCAACGCAATTTACAAAACAAAAACACCTGCACCCC
 TTAATCTCAGTAATGGTTTTCTATGTATCGAAGATCATGCTCAGCTTACAGTGAATCGATTACACAAAACCTGGGGGT
 GTTGTCTCTTGGGAATGGAGCAGTTCTGAGTTGCTATAAAAAATGGTACAGGAGATTCTGCTAGCAATGCCTCTAT
 AACACTGAAGCATATTGGATTGAATCTTTCTCCATTCTGAAAAGTGGTGTGAGATTCTTTATTGTGGGTAGAGC
 CTACAAAATAACAGCAATAACTATAACAGCAGATACTGCAGCTACCTTTTCATTAAGTGTGTA AAACTCTCACTCATT
 GATGACTACGGGAACCTCCTTATGAATCCACAGATCTGACCCATGCTCTGTCATCACAGCCTATGCTATCTATTTT
 TGAAGCTAGCGATAACAGCTACAATCAGAAAATATAGATTTTTCGGGACTAAATGTCCTCATTATGGATGGCAAG
 GACTTTGGACTTGGGGCTGGGCAAAAACCTCAAGATCCAGAACAGCATCTTCAGCAACAATCACTGATCCACAAAAA
 GCCAATAGATTTTATAGAACCTTACTACTAACATGGCTTCTGCGGGTATGTTCTTAGCCAAAACACAGAAGTCC
 CCTCATAGCTAACCTTATGGGGGAATATGCTGCTTGCACAGAAAAGCTTAAAAAATAGTGCAGAGCTGACACCTA
 GTGGTTCATCTTTCTGGGAATTAACAGGAGGAGGACTAGGCATGATGGTTTACCAAGATCCTCGAGAAAATCATCCT
 GGATTCATATGCGCTCTTCCGGATACTCTGCGGGGATGATAGCAGGGCAGACACACACCTTCTCATTGAAATTCAG
 TCAGACCTACACAAAACCTCAATGAGCGTTACGCAAAAAACAACGTATCTTCTAAAAATTAATCATGCCAAGGAGAAA
 TGCTCTTCTCATTGCAAGAAAGTTTTCTGCTGACTAAATTAAGTTGGGCTTTACAGCTATGGAGACCATAAAGTGCAC
 CTTTTCTATACTCAAGGAGAAAATCTAACATCTCAAGGGACGTTCCGCAGTCAAACGATGGGAGGTGCTGTCTTTTT
 TGATCTCCCTATGAAACCCTTTGGATCAACGCATATACTGACAGCTCCCTTTTTAGGTGCTCTTGGTATTTATTCTA
 GCCTGTCTCACTTTACTGAGGTGGGAGCCTATCCGCGAAGCTTTTTCTACAAAGACTCCTTTGATCAATGTCTTAGTC
 CCTATTGGAGTTAAAGGTAGCTTTATGAATGCTACCCACAGCTCAAGCCTGGACTGTAGAATGGCATAACCAACC
 CGTCTGTATAGACAAGAACCAGGGATCGCAGCCAGCTCCTAGCCAGTAAGGGTATTTGGTTCCGTTAGGTAAGCC
 CCTCATCGCGTCATGCCATGTCTTATAAAAATCTCACAGCAAAACACAACCTTTGAGTTGGTTAACTCTCATTCCAG
 TATCATGGATTCTACTCTTCAACCTTCTGTAATTATCTCAATGGGGAATTTGCTCTGCGATT

SEQ ID NO: 124: CT869 fragment protein sequence

REVPSRIFLMPNSVDPDKESLSNKISLTDGTHNLNLCYLNLRYILAILQKTPNEGAAVITDYLSSFDTQKEGIY
 FAKNLTPESSGGAIGYASPNSTVEIRDITIGPVIFENNTCCRLFTWRNPYAADKIREGGAIIHAQNLNHNHDDVVGF
 KNFSYVQGGAIQSTANTFVVSENQSCFLFMDNICIQNTAGKGGAIYAGTSNSFESNNCDLFFINNACCAGGAI
 CSLTGNRGNIVFYNNRCFKNVETASSEASDGGAIKVTTRLVDVTGNRGRIFSDNITKNYGGAIYAPVVTLVDNGPT
 FINNIANNKGGAIYIDGTSNSKISADRHAIIFNENIVTNTNANGTSTSANPPRRNAITVASSSSGEILLGAGSSQNL
 IFYDPIEVSNAGVSVFNKEADQTSVVFSGATVNSADFHRNLQTKTAPLTLNGLFCIEDHAQLTVNRFQTGG
 VVSLGNGAVLSYKNGTGDASNASITLKHIGLNLSSILKSGAEIPLLVVEPTNNSNNTADTAATFSLSDVKLSLI
 DDYGNFSPEYESTDLTHALSSQPMLSISEASDNQLSENIDFSGLVNPHYGWQGLWTWGWAKTQDDPEPASSATIDP
 ANRFHRTLLLWLPAGYVPSPKHRSPLIANTLWGNMLLATESLKNSAELTPSGHPFWGTTGGGLGPMVYQDIPRENH
 GFHMRSSGYSAGMIAGQTHTFSLKFSQTYTKLNERYAKNNVSSKNYSQGEMLFSLQEGFLTKLVGLYSYGDHNC
 HFYTDGENLTSQGFERSQTMGGAVFFDLPMKPFGSTHILTAFLGALGIYSSLSHFTEVGAYPRFSFKTPLINLV
 PIGVKGSFMNATHRPQAWTVELAYQPVLYRQEPGIAAQLLASKGIWFGSGSPSSRHAMSYSKISQRTQPLSWLTLHFQ
 YHGFYSSSTFCNYLNGEIALRF

SEQ ID NO:125: CT166 fragment nucleotide sequence

AACGTTTCGTACGTACTCTGTTTCAGAGGGGGGGGTA AAAACGATTTCTGCTAGTGCAGTTCCTCCTACAGCAGCTGT
 TTTATCGAGAAAAAAGCGTGCTATAGAAGAGAAGAAGGAGGAAGCTTCTTCTGGAAAAGATAGAAAATCTTGATGCTA
 GCAAAATACGATCTTACTCCCAAGAACATAGAAGAAAACTAGGAATTAATCCTGAACAGAAAATCTACTGTTAAAGAC
 CTATTAATAAACTGAAAAAGGTCATTAGTGCTTACAACCTATGCCAGATAAAAAATTCGGAAGCGGGACAGAATTC
 CTTGATTTCAACAAGGAAAAATACGTCGATGCCATTCAGAAGAAGCTTCCAGCATCATCGCAGGCTCAGCCTAACAGG
 CAAAAGCTAAGGAACAGAAAAGCCGAAGAAAAACCTAACAGCAGCTCCGATTGAAGGTGTTCTTGAAAACCATCAAAA
 GAATTTAAAGGCCATCGTGTACCTGTTGAGAAAATCATCCATGGAATATGGATCGCAGGAGCGCCTCCGGATGGTAT
 CGAAGATTATATGCGAGTCTTTTTAGATACTTATGAAGGTTTTGACTTCTACTTCTGGGTAGATGAGAATGCTTATG
 CAGCAGCTAAATTTCTAGCATTGTTGAAGAAGGTGCTTTTCGATGCGGCTATTCAGATCTACGATCTGCCACAGAT
 GAGTCTACGAAGGCCTTTGTAAAGACTACGATGAATTA AAAACAGAAATATGAAAAGAAAAGTTGCGGAGACGACTTC
 TCAAGCAGAAAAAGCCAATATCTCAAAGATCTAAAAGGATCTTTTAGAGAAAATTTACAAAATCAGTGATGAGATTC
 GTGGAAAATTTGATCGGCTGTTTCTTAAGAATGTGATTGTTGCTCAGAACGGATTCTTTAATTTCTGCTTGTGAAA
 GGCCTCGGCAATATCAATGACGAAACGCGTGCAGAGTATTTAGAGAAAAGA ACTCAAACCTTCTACTGAGGAGATCGA
 ACAGTATAAAAAGCTTAAAGAGACGAACAAAGAGAAGATAGCCGCTATTGTA AAAACA ACTAAACGAGAAAACCTGGAT
 CGGATCGGTTAAAAATCAAAGACATTAAGAGCTGCAATCATGAAGCAAGCTCGAAAATGTCTCAATTAATGAACAG
 GAAATGTTTTCTGCGCTGGAACCTATGCAGCCGCAACAGATCAGATTCGATGTATATGTTGGAGCAACTGGAGGCTCT
 TTATACTGATCTGGATATGATGCCTTCTACTCTCAGGAAGTATTGGAGCTTATCAAAAAGCACAGTGTGGA AACCC
 GAATGTTTGGAGATATGAGCTCTAGACGGGCGATTTCTGATGCGGTTTTAAAGATGGCTGTAGGTAAGGCGACAACA
 GTTCCATGGAAGAGGTAGCAAAGGATATCGATGTTTCTCGCTAACAGAAAGGATAAGACAAAATTAATGCTCT
 ATTTAAAGGACTAGAGCCATTTGCAAAAACCGGATTTCTAAAGGAGCTGAAGCAGAAAGGGGTGAAGGAGCAAAGGTA
 TGAAAAGAGCTTTTTCCAGCCCATAGATCTGAATATTGTCAGAAATACCATGCCTATCTTGAGACGCTCATCATC
 TACTCTGAGTTAGGTTGTTTATTTCGAGGATTGAACCGGATGATGGTCTCTCATAAAGGGAAGCAGCTCGGTTTTCTG
 TGTCAATTGTAGGGCAACAGGCTGCCTACCAGGAAC TAGCAGCACTTAGACAAGATGTCCTTTACGGGGAGTTTTTCC

ATTCTTTAGAAAATTTGACACATAGAAAACCATAAGGAGCGTATTGGAAATCATCTCGTCTGCTAATTATTTGGCTAAA
AGTCTCTTTTTTGGATTACTGCCAAGATTGATGATGCCGGAGGCTGTAAGTACCTTAGGTATTAGA

SEQ ID NO: 126 – CT166 fragment protein sequence

NVRTYSVQRGGVKTISASAVPPTAAVLSRKKRAIEEKKEEASSGKIENLDASKYDLTPKNIEEKLGITPEQKSTVKD
LLNKLKKVISAYNSMPDKNSEAGQNSLIQGGKYVDAIQKKLPASSQAQPKQAKAKEQKAEEKPKTTPIEGVLETIKT
EFKGRVPVEKIIHGIIWAGAPPDGIEDYMRVFLDTYEGFDFYFVVDENAYAAAFSSILKKVAFDAAIQDLRSATD
ESTKAFVKDYDELKQKYEKKVAETTSQAEKDQYLLKDLKDLLEKFTKISDEIRGKFDRLFLKNVIVAQNGFFNFCLLK
GLGNINDETRAEYLEKELKLPTEEIEQYKLLKETNKEKIAAIVKQNLNEKLGSDRVKIKDIKELQSMKQARNVYNYE
EMFLRWNYAAATDQIRMYMLEELGGLYTDLDMMPYSYQEVLELICKHSDGNRMFEDMSSRRAISDAVLKMAVGKATT
VSMEEVAKDIDVSRLEEDKTKLNALFKDLEPFAKPDSSKGAEEGGEGAKGMKKSFFQPIDLNIVRNTPILRRYHH
YPELGFIRGLNGLMVSHKGTAVSAVIVGQAAAYQELAAALRQDVLSGEFFHLENLTHRNHKERIGNHLVANYLAK
SLFFDYCQDSVMPEAVSTLGR

SEQ ID NO:127 – CT175 fragment nucleotide sequence

TGTTATCATAAAAAAGAAGAACCAAAAGATGTTTTGCGGATTGCGATCTGTCTATGATCCAATGTCTTTAGATCCGGC
TCAGGTTTTTTAAGCAAAGATGTTTCTATTGTAAGGCTCTCTATGAAGGGTTAGTCCGGGAAAAAGAAGCTGCGT
TCCAGCTAGCTTTGGCAGAAAGATATCATCAATCTGATGATGGTTGTGTTTATACTTTTTTCTAAAAATACATTC
TGGAGCAACGGAGATGTTGTAACAGCATATGATTTGAAGAGTCTATTAACAAAATTTATTTCCGAGAAATTTGATAA
CCCTTCGTTACGCTCTCTTGCATTAATTAATAAATTCATGCTGTTTTAACAGGAGCTCTCCCTGTTGAAGATTTAG
GTGTTAGAGCTTTGAATGCGAAAACCTCTAGAAATGTTTTAGAAAACCCGTTTCTTATTTTCTAGAGATATTGGCG
CACCCGGTTTTTATCCGGTGCACACCTCTTACGAGAATATTACAAAGATAAGCGTAACAAACGCGTTTTCCCGAT
AATTTCTAATGGTCTTTTGCAGTCAATGTTATGAGCCGCAAAGATATTTACTAATCAACAAAAACCTCTGTATC
ATGCCAAGCAGATGTTCTGTTAAATTCGGTATGTTTGCAGATAGTTTCTGATATCCATACAGCTATGCGATTTATC
CAAAAAATCATATCGATTTAGTTGGGTTACCTCGGAGCTCTCTCTTTTCTTAGAAGAAAGAAATCTCCCTAG
AGAAAAATTTGATTATCCTGTATTGAGTTGCTCTGTTTTATTCTGTAACATTCATCAACACCTTTAAATAATC
CCTCGCTGAGAACAGCCCTCTCTTTAGCAATCAATCGAGAACTTTATTAATAACTAGCAGGTAAAGGCTGTAGCGCT
ACGAGCTTTGTTACCCACAATTATCTCAGATACCTGCTACTACTTTGTCTCAAGATGAGCGGATTGCTTTAGCAA
AGGCTACTTTGACCGAAGCTTTAAAGACTTTATCTCAAGAAGATTTAGAAAAAATTACATTAATTTATCCTATAGAA
CTGTTTGTCTACGAGCCGTTGTTCAAGAAATTCGCCAACAAATTTATTTGATGTACTGGGATTTAAATTTCTACATTA
GGATTAGAATATCATTGTTTTTTAGACAAACGTTCCAGAGGAGAATTTCTCCTTAGCAACTGGTAATTGGATTGCAGA
CTATCATCAAGCTAGTCTTTCTGTCTGTCTTAGGTAATGGGACAAGATATAAAGACTTTCAATTGATTAAGTGGC
AGAACCAAAAGTACACAAATATAGTTGCTCAACTTCTGATTCAAGAATCAAGCGACCTACAGCTTATGGCAGAGCAG
TTGTTGCTTAAAGAAAGTCTCTTATTTCTCTATACCACCTCGATTATGTGTATGCGAAACAGCCTCGGGTGTCTGA
TCTCCAAACCTCTCTCGTGGAGAAATTTGATTTAAAAAGAGTTTCATTAGCTGAAGGATAG

SEQ ID NO:128 – CT175 fragment protein sequence

CYHKKEEPKDVLRIAICHDPMSLDPRQVFLSKDVSIVKALYEGLVREKEAAFQLALAEYHQSDDGCVYTFFLKNTF
WSNGDVVTAYDFEESIKQIYFREIDNPSLRSLALIKNSHAVLTGALPVEDLGVRALNAKTLEIVLENPFYPFLEILA
HPVYFPVHTSLREYYKDKRNKRVPFIISNGPFAIQCEYEPQRYLLINKNPLYHAKHDVLLNSVCLQIVPDIHTAMQLF
QKNHIDLVLGWPSSSFLSEEQRNLPREKLFDYPVLSVSVLFCNIHQTPLNPSLRALSLAINRETLKLAGKGC
TSFVHPQLSIPATTLSDERIALAKGYLTELKTLGSDLEKITLIYPIESVCLRAVVQEIQRQLFDVLFKISTL
GLEHYHFLDKRSRGEFSLATGNWIADYHQASAFLSVLGNGTRYKDFQLINWQNKYTNIVAQLLIQESSDLQMAEQ
LLLKESPLIPLYHLDYVYAKQPRVSDLQTSRGEIDLKRVSLAEG

SEQ ID NO: 129 – TC0666 fragment nucleotide sequence (homologue of CT387)

ATGCACTCTTTCACACTCATCAGATGCCGCTCTCCGGACGGCTACTTATGTTCTTCCCTTCAGTTAGTTGGCTC
TGGCACATATGAAGGAGAAATCGAAATCCAAAATATCCTTCTTATTTCTTGGATTCCGATTACCCACCCATTGCG
TTCATCTTAATTTGAAGAGTTCTCTAGCCCAGTTAGGAGTAGATGCATCTCTTCTTCACTGCGAACTAAGCAAAAA
CAACAACGTGCACATATGCACGTGCAGTTACCCGGCTATGGCCCTATCGCTGAGTCCATGCTATCTCTTCTCAAACC
CGGAGATCGAGTAGCCAAACTGTTTGTGCTGCAGATGATCGTAGACTAGTCCGCTCCCTGATTATCTTGAAGCATGC
TAAAAAATACTGATAAGACAGGACATCCTCTGCTCCGATTTGGAAAAAACTCGAGCATCTTATCTCTTTGATGTG
GTGGACGATCGCCTCGTTGTATCACTCCCCACCTTGCAGGCATAGTCAATTATGACCCAGACATCTATGGACTTCT
TCCCTTAATTTCAAAAACTACTAAGCAATCCTAAATTTGAGATTTCCGCACTTCTTGTCTCTATCAGAAGATCGTAG
AAGGACCACACATCCCTTATGAAGGAAACATTTTGTAAATCAAAAACAGAGCCTCTTCAATCCGCACAGTATTTGCT
CGCGTGGTGCATCAAAATGCTCCCTCAAGGTCTATTTCACTTCTGCCAACATTTTAGAACCCACAACGCGAGAGTC
TGGAGATATTTTGAATTTTGGAAATCCCTCCACTCTTGTAGAAAGAATCCCTCTAGAATTTCTACTATCGAAC
CCTACAAAGAACACTCTTACTTCTGTAATCGAGATCTATTTGCAAACTACCTTGCAATCGGAAAGTGAATCAAAAA
ATATTCGATACAGCTCCTCAAGAGCCTGTAAGAGCCGCACTATTTATCAAAAAGGAAAGTGAATTTCTCTTGA
TGCAGATTCTTGGCTTACGGGATCCGCAGCTGCATACCAATGTAGCGAAAAACAGGACGCTAAAGACGAATACATCC
ACGCTCAACCTGTTATCCATTTTTGGAAGCAATGGAACCGGACTCATCAATAGCGAAGGAGCTTTACTCACTCGG
TTTTTCCCTCTTCCAGCTTAAAGGGATGTTGATCTCTATCATGTACGCCACTATCTTAAGCAAATTTACTTTCA
AGTTCTTCTTATACATATGGAGACTACTTCTCTATAATGACCGAGGATTACTGTTAGACTATATCAGGCGAAC
TTGATGTGTTCTGGCTGATGAAGAGAGCGGCGGTGATTGCAATATACAAAAACGGCGCAAAAACTAGTGGAAATG
TTCGTCTTTAAAAATCGAGTAGAAGAGTTCCAATCGATATTTCTGATAGCGATTTATGGATCAGCTTCCGAAAA
TAATTTCTCGGCCCAACTAAACACGCTTCTTGCAGGGTTACAAAAAGCTGCACACACTCTAGGCATTTCCAGGCTTCT
CAAAACCCACTCTCTTCCGCTAATCACAGGAGGAGGACTGGCGTTATGGCTACAGGAAATCGTGTGCAAAAGAG

TTGGGAATTCCTTCTTGC GGGACCGTTCTCGATTTGGAAGCTTACCTGCACAAATAGATCAGCCTGCAAACGAATT
 TTTAGATGCCAAATGACATACCGTCTACCGCAACTTATAGAAAAGACAAGAACATTTTTTATTAGACCTTGCCATTT
 TAGTTGTTGGTGGTGTGGAAACAGATTTGCAACTTTACCTAGAACTCGTCTACTTGAAAAACAGGCGCCAAACCTCTT
 ACTCCAATTTCTTATTGGGCTGTTGAATACTGGAAGAGAAAAGTTGCTCATGCCATGAGATTAATCTTAAAGC
 AGGAACTATTCTGTTCTGAGTGGATCAGCAACTGCTTATTCTGCATTACATCTCCTGAAGCAGGAATTGCTGTAT
 TCGAACAGTTCTCTCGTGGAGAAGTTCCCATAGGATATGATTATCCTCCAGCTCCAGACGGATTAGTTATCGTC

SEQ ID NO: 130 - TC0666 fragment protein sequence (homologue of CT387)

MTLFHTHDAVSPDGYL CSSLQLVGS GTYEGEIEIQNIPSYFLGFR LPTHCVHLNLKSSLAQLGV DASLLHCELSKN
 QRRAHMHVQFTGYGPIAESMLSLLKPGDRVAKLFAADDRRLVRS PDYLESMLKNTDKTGHPLLRFGKLEHLISFDV
 VDDRLVVS LPTLPGLVNYDPDIYGLLPLIQKSLSNPKLSIRHFLSLYQKIVEGPHIPYEGNILLIKTEPLHIRTVFA
 RVVDQMLPQGLFH TSANILEPTTRESGDIFEFFGNPSTLVERIPLEFFTIEPYKEHSYFCNRDLLQTTLQSESEIKK
 IFDTAPQEPVKAATYLSKGSEISSLDADSWLTGSAAYQCEKQAAKDEYIHAQPCYPFLEAMETGLINSEGALLTR
 FFPSSSLKGM LISYHVRHYLKQIYFQVPSYTYGDYF SHNDRGLLLDLYQANIDVFWADEESGRVLQYTKRRDKNSGM
 FVVKNRVEEFQSA YFVAIYGSRLLENFSAQLNLLAGLQKAAHTLGIPGFSKPTPLAVITGGGTGMATGNRVAKE
 LGILSCGTVLDLEASPAQIDQ PANEF L DAKM TYR L PQLIERQEHFYSDLAILVGGVGTDFELYLELVYKTKGAKPP
 TPIFLIGPVEYWK EKV AHAYEINLKAGTIRGSEWISNCLFCITSPEAGIAVFEQFLAGELPIGYDYPPAPDGLVIV

SEQ ID NO: 131 - TC0197 fragment nucleotide sequence

AATTGTTCCGATCTTTATGCCGTAGGAAGTTCTGCAGACCATCCTGCCTACTTGATTCTCAAGCGGGGTATTATT
 GGATCATATTAAGGATATATTCAATTGGCCCTAAAGATAGTCAGGATAAGGGGCAGTATAAGTTGATTATTGGTGAGG
 CTGGCTCTTTCCAAGATAGTAATGCAGAGACTCTTCTCAAAGGTAGAGCACAGCACTTTGTTTTTCAGTTACAACA
 CCTATAAATTTGTCAAGGAATAGATCAACAAGATCAGGTCCTCTTCGAGGGATTGGTCTGTAATTTTTTCAGGAGATCA
 TTCAGAGGAGATTTTTGAGAGAGAATCCTTTTTAGGGATCGCTTCTCAGGGAAATGGTAGCAAGGATGGAATCACGT
 TAACAGATATAAAATCTTCGTTATCTGGTGCTGCCCTGTATTCTTCAGATGATCTTATTTTTGAAAGAATTAAGGGA
 GATATAGAGCTTTCTTCTGTTTCATCTTTAGAAAGAGGAGGAGCTTGTTCAGCTCAAAGTATTTTAATTCATGATTG
 TCAAGGATTAACGGTAAAAACATTGTGCCG CAGGGGTGAATGTTGAAGGAGTTAGTGCTAGCGACCATCTCGGATTTG
 GGGCGGGGCTTCTCTACTACAAGTTCTCTTCTGGAGAGAAGAGTTGTATATGCTTCAGGCGATATTGTGGTG
 GCTACCTGCGATGGTCTGTGTGTTTGAAGGAAATAGTGCTCAGTTAGCAAAATGGTGGCGCTATTGCGCTTCTGG
 TAAAGTTCTTTTTGTAGCTAACGAAAAAAGATTTCTTTACAGACAACCAAGCTTTGTCTGGAGGAGCTATTTCTG
 CATCTTCTAGTATTTCTTTCCAAAATTTGTGCTGAGCTTGTGTTCAAGAGTAATCTTGCAAAAAGGAGTTAAAGATAAA
 TGTTCTTTGGGAGGAGGTGCTTTAGCCTCTTTAGAAATCCGTAAGTTTGAAGATAATCTCGGTATTAATGAAAA
 AAATCAGTCTTCTCGGAAGGAGGGGCTATTTTTGGGAGGATTGTGAGATTTTTGAAAAACAGGGGGCTGTGGTAT
 TCAGAGATAATACAGCTGCTTTAGGAGGCGGAGCTATTTTTGGCGCAACAAACTGTGGCGATTTGTGGTAATAAGTCT
 GGAATATCTTTGAAGGAAGTAAGTCTAGTTTTGGAGGGCCATTGCTTGTGGAAATTTCTCTTCTGAGAATAATTC
 TTCAGCTTTGGGATCAATGATATCTCTAACAACTAGGAGATATCTTTTTCTTCGGACTCTGTGTACTACTTCGG
 ATTTAGGGCAAACGGATTACCAAGGGGGAGGGGCTTATTCTGCTGAAAATATTTCTTTCTGAGAATGCTGGTGCA
 ATTAATTTCAAAGACAATATTGTGAAGACATTTGCCTCAAATGGAAAAATGTTGGTGGAGGGGCAATTTAGCTTC
 AAGAAATGTTTTGATTAGCAAAAACCTCTGGAGAGATTTCTTTTTGTAGGGAATGCTCGAGCTCTCAGGCTATTCGA
 CTCGTTTCATCTGACGAATTGTCTTTTTGGCGCACAAATTACTCAAACACTCTTCAGGATGTTCTGGAGGAGGAGCTCTT
 TTTGGTAAAGAGGTTGCCATTGTTCAAATGCCACTGTTGTATTTCGAGCAAAAATCGCTTACAGTGTGGCGAGCAGGA
 AACACATGGTGGAGGCGGTGCTGTTTATGGTATGGAGAGTGCCTCTATTTATGGAACTCTTTTTGTGAGATTCGGAA
 ATAATTACGCTGTAGGGAATCAGATTTCTGGAGGAGCTCTTTTTATCCAAGAAGGTCGTTTAGCTGAAAATACAAGG
 GTAGATTTTTCTCGAAATATCGCTACTTTCTGCCGCGGGGCTGTTTCAAGTTTCTGATGGAAGTTGCGAATTTGATCA
 CAATGGGTATGTGCTATTAGAGATAACCGAGGGCAGACATTTGGTGGGGCTATTTCTTGCTTGAAGGAGATGTGA
 TCATTTCCGGAAATAAAGATAGGGTTGAGTTTAGAGATAACATTGTGACGCGGCCTTATTTTGAAGAAAATGAAGAA
 AAAGTTGAGACAGCAGATATTAATTCAGATAAGCAAGAAGCAGAAGAGCGCTCTTTATTAGAGAACATTGAGCAGAG
 CTTTATTACTGCAACTAATCAGACCTTTTTCTTAGAGGAAGAGAAAACCCATCAGAAGCTTTTATCTCTGCTGAAG
 AACTTTCAAAGAGAAAGAAATGTGCTGGTGGGGCTATTTTGAAGAAAGGGTCTACATTCAGGATAAAGAAACCT
 ATCTTTGTTTTCGCATAATTTTTCTGATGTTTATGGGGGAGCTATTTTTACGGGTTCTCTACAGGAAACTGATAAACA
 AGATGTTGTAACCTCTGAAGTTGTGATATCAGGCAACGATGGGGATGTCATTTTTCTGGAAATGCAGCTAAACATG
 ATAAGCATTTACCTGATACAGGTGGTGGAGCCATTTGTACACAGAATTTGACGATTTCCCAAAAACAATGGGAATGTC
 TTGTTCTTGAACAAATTTGCTTGTCTGGTGGAGCAGTTCCATAGAGGATCATGGAGAAGTTCTTTTAGAGGCTTT
 TGGGGGAGATATTATTTTCAATGGAAACTCTCTTTTCAGAGCTCAAGGATCGGATGCGATCTATTTTTGCTGGTAAAG
 ACTCTAGAATTAAGCTTTAAATGCTACTGAAGGACATGCGATTGTGTTCCAAGATGCATTGGTGTGTTGAAAAATA
 GAAGAAAGAAAGCTTCGGGACTATTGGTGATTAACCTCTCAGGAAAATGAGGGTTATACGGGATCCGTCGGATTTTT
 AGGATCTGAAAGTAAGGTTCTCAATGGATTCATGTGCAACAGGGAGGCTTGAGTTGCTACATGGAGCTATTTTAT
 GTAGTTATGGGGTTAAACAAGATCCTAGAGCTAAAAATAGTATTAATCTGCTGGATCTAAATTAAGGATTTAGATTCA
 GAGCAAGAAAAAACAAGCAGAAATTTGAGATCTTGAAGATTTGTTAATTCAGAAAAAACCCATCTCTTTGGATTGG
 GAAGAACGCTCAAGCAAAAAGTCCCTCTGGTTGATATCCATACTATTTCTATTGATTTAGCATATTTTCTTAAAG
 CTCAGGAAACCCCTGAGGAAGCTCCACAAGTCACTGCTCCCTAAGGGAAAGTTGTGTCCTCAGGAGAGTTAAGTTTG
 GAGTTGGTTAATAACAACAGGAAAAGGTTATGAGAATCATGCGTTGTTAAAAAATGATACTCAGGTTTCTCTCATGTC
 TTTCAAAGAGGAAAATGATGGATCTTTAGAAAGATTTGAGTAAGTTGTCTGTTTCGGATTTACGCATTAAGTTTCTA
 CTCCAGATATTGTAGAAGAACTTATGGCCATATGGGGGATTTGGTCTGAAGCTACAATTCAGATGGGGCTCTTGTCT
 ATTAATTTGGCTACTGATATAAATTAGATCCGCAAAAAGCTGGTTCTTTGGTATTCAATGCAATTTAGGGAGGA
 AGAGGCTGTATTGTCTACTCTAAAAAATGCTCGGATTGCCATAACCTTACCATTAGAGAAATGGAATTTGATTATT
 CTACAAATGCTTGGGGATTAGCTTTTAGTAGCTTTAGAGAGCTATCTTCAGAGAAGCTTGTCTTCTGTTGATGGATAT
 AGAGGCTCTTATATAGGGGCTTCTGCAGGCAATTGATACTCAGTTGATGGAAGATTTTGTGTTTGGGAATCAGCACGGC

TTCCTTCTTCGGGAAATGCATAGTCAGAATTTTGATGCAGAGATTTCTCGACATGGTTTTGTTGGTTCGGTCTATA
 CAGGCTTCTAGCTGGGGCTGGTTCTTCAAGGGGCAGTACAGTCTTGGCGAAACACATAACGATATGACAACCTCGT
 TACGGGGTTTTGGGAGAATCTAATGCTACTTGGAAAGTCTCGAGGAGTACTAGCAGATGCTTTAGTTGAATATCGTAG
 TTTAGTCCGGTCCAGCAGACCTAAATTTTATGCTTTGCATTTTAATCCTTATGTGCGAGGTATCTTATGCATCTGCCA
 AGTTCCCTAGTTTTGTAGAACAAGGAGGAGAAGCTCGTGCTTTTGAAGAAACCTTTTAAACAAACATTACCGTTCCC
 TTTGGTATGAAATTTGAACTATCTTTTACAAAAGGACAGTTCAGAGACTAATTCTCTTGGAAATAGGTTGTGCATG
 GGAATGTATCGGAAAGTGAAGGAAGATCTGTAGAGCTACTAGAAGCTGGTTTTGATTGGGAAGGATCTCCTATAG
 ATCTCCCTAAACAAGAGCTGAGAGTGGCTTTAGAAAACAATACGGAATGGAGTTCTGATTTTTAGTACAGCTCTAGGA
 GTAACAGCATTTTGTGGAGGATTTCTTCTATGGATAATAAACTAGGATACGAAGCGAATGCTGGAATGCGTTTGAT
 TTTCTAG

SEQ ID NO: 132 - TC0197 fragment protein sequence

NCSDLAVGSSADHPAYLIPQAGLLLDHIKIDIFIGPKDSQDKGQYKLIIGEAGSFQDSNAETLPQKVEHSTLFSVTT
 PIIVQGDIDQDDQVSSQGLVCFNSGDHSEEIFERESFLGIAFLNGSKDGITLTDIKSSLSGAALYSSDDLIFERIKG
 DIELSSCSSLERGGACSAQSILIHDCQGLTVKHCAAGVNVGVSASDHLGFGGGAFSTSSLSGKSLYMPAGDIVV
 ATCDGPVCFEGNSAQLANGGAIAASGKVLVFAVEKKISFTDNQALSGGAISASSSISFQNCALVFKSNLAKGVKDK
 CSLGGGALASLESVVLKDNLGITYEKNQSYSEGGAIKDFGKCEIFENRGPVVFRDNTAALGGGAILAQQTVAICGNKS
 GISFEGSKSSFYGGAIKDFGSLQETDKQDVVTEPVVISGNDGDVIFSGNAAKHDKHLPTDGGGAICTQNLTISQNNGNV
 ITFKDNIVKTFASNGKMLGGGAILASGNVLISKNSGEISFVGNARAPQAIPTRSSDELSTFGAQLTQTTSGCSGGGAL
 FGKEVAIVQNAVTVVFEQNRQLQCGEQETHGGGGAVYGMESASIIIGNSFVRFVFNQYAVGNQISGGALLSKKVRLAENTR
 VDFSRNIATFCGGAVQVSDGSCLEINNGYVLFDRNRGQTFGGAISCLKGDVVISGNKDRVEFRDNIVTRPYFEENE
 KVETADINSQKQEAERSLLENIEQSFITATNQFFLEEEKLPSEAFISAELSKRRECGGAIFAKRVYITDNKEP
 ILFSHNFSDVYGGAIKDFGSLQETDKQDVVTEPVVISGNDGDVIFSGNAAKHDKHLPTDGGGAICTQNLTISQNNGNV
 LFLNNAFCSGGAVRIEDHGEVLLFAFGGDIIFNGNSSFRAQGSDAIYFAGKDSRIKALNATEGHAIVFQDALVFENI
 EERKSSGLLVINSQENEGYTGSRVFLGSESKVPQWIVHQQGGLELLHGAILCSYGVKQDPRAKIVLSAGSKLKILDS
 EQENNAEIGDLEDVNSEKTPSLWIGKNAQAKVPLVDIHTISIDLASFSSKAQETPEEAPQVIVPKGSCVHSGELSL
 ELVNTTGKGYENHALLKNDTQVSLMSFKEENDGSLEDLSKLSVSDLRIVKSTPDIVEETYGHMGDWSEATIQDQALV
 INWHPTGYKLDQPKAGSLVFNALWEEEAFLSTLKNARIAHNLTIQRMEFDYSTNAWGLAFSSFRELSSEKLVSDGY
 RGSYIGASAGIDTQLMEDFVLGISTASFFGKMHSQNFDAEISRHFVGSVYTGFLAGAWFFKGQYSLGETHNDMTR
 YGVLGESNATWKSRLVADALVEYRSLVGPAPKPYALHFNPHYVEVSYASAKFPSFVEQGGEARAFEETSLENTITVP
 FGMKFELSFTKGQFSETNSLGIGCAWEMYRKVEGRSVELLEAGFDWEGSPIDLKQELRVALENNTWSSYFSTALG
 VTAFCGGFSSMDNKLGYEANAGMRLIF

SEQ ID NO: 133 - TC0261 fragment nucleotide sequence

ACTCGAGAAGTCCCTCCTTCGATTCTTTTAAAGCCTATACTAAATCCATACCATATGACCGGGTTATTTTTTCCCAA
 GGTTAATTTGCTTGGAGACACATAATCTCACTGATTACCATTTGGATAATCTAAAAATGCATTCTGGCTTGCCTAC
 AAAGAAGTCCCTTATGAAGGAGCTGCTTTTACAGTAACCGATTACTTAGGTTTTTTCAGATACACAAAAGGATGGTATT
 TTTTGTTTTTAAAAATCTTACTCCAGAGAGTGGAGGGGTATTGGTTCCCCAACTCAAAACACTCCTACTATAAAAAAT
 TCATAATAACAATCGGCCCCGTTCTTTTCGAAAAATAACTGTGTCATAGACTGTGGACACAGACCGGATCCCGAAAAATG
 AAGGAAACAAAGCAGCGAAGGCGGGGCAATTCATGCTGGGGACGTTTACATAAGCAATAACCAAGCAACTTGTCCGGA
 TTCATAAAGAAGCTTTGCTTATGTTCAAGGTGGAGCTATTAGTGCTAATACTTTTGCCTATAAAGAAAATAAATCGAG
 CTTTCTTTGCCTAAATAACTCTTGTATACAAACTAAGACGGGAGGGAAAAGGTGGTGTATTTACGTTAGTACGAGCT
 GCTCTTTTCGAGAACAATAACAAGGATCTGCTTTTCACTCAAAAACCTCCGGCTGTGCAGGAGGAGCTATCTTCTCTCCA
 AACTGTTCTCTAATAGGAAACCAAGGAGATATTGTTTTTACAGCAACCAACGGTTTTAAAAATGTTGATAATGCAAC
 TAACGAACTCTGGGGATGGAGGACTATTAAGTAACCTACCCGCTTGGACATCACCAATAATGGTAGTCAAATCTTTT
 TTTCTGATAATATCTCAAGAAAATTTGGAGGAGCTATTCATGCTCCTTGTCTTCTTGTGGTAATGGGCCAACCC
 TATTTTACAAACAATATAGCTAATCACACAGGTGGGGCTATTTATATAACAGGAACAGAAACCTCAAAGATTTCTGC
 AGATCACCATGCTATTTATTTGATAATAACATTTCTGCAAAACGCCACCAATGCGGACGGATCTAGCAGCAACACTA
 ATCCTCCTCACAGAAATGCGATCACTATGGACAATTCGCTGGAGGAATAGAACTTGGTGCAGGGAAAGAGCCAGAAT
 CTTATTTTCTATGATCCTATTCAAGTGACGAATGCTGGAGTTACCGTAGACTTCAATAAGGATGCCTCCCAAACTTTT
 ATGTGTAGTTTTCTCGGAGCGACTGTCTTTCTGCAAGATATTTCTCAGGCTAATTTGCAAACTAAAAACACTGCAA
 CGCTTACTCTCAGTACGGTCTTCTGTGTATCGAAGATCGTGCTCAGCTCACAGTGAACAATTTTACACAAACAGGA
 GGGATTGTAGCCTTAGGAAATGGAGCAGTTTTAAAGCAGCTACCAACACAGCACTACAGACGCCACTCAAACCTCCCC
 TACAACCACCACTACAGATGCTTCCGTAACCTTAAATCACATTGGATTAAATCTCCCCCTATTCTTAAGGATGGAG
 CAGAGATGCCTCTATTATGGGTAGAACCCTATAAGCACAACTCAAGGTAACACTACAACATATACGTACAGATACCGGG
 GCTTCTCTCTCATTAAATGGAGCCACACTCTCTCATTGATGAAGATGGAATTTCCCTATGAAAAACACGGACCT
 CTCTCGTGCATTGTACGCTCAACCTATGCTAGCAATTTCTGAGGCCAGTGATAACCAATTGCAATCCGAAAGCATGG
 ACTTTTCTAAAGTTAATGTTCTCACTATGGATGGCAAGGACTTTGGACCTGGGGTGGGCAAAAACCTGAAAATCCA
 ACAACAACCTCCTCAGCAACAATTAAGTATCCGAAAAAAGCTTAATCAGTTTCTATAGAACTTTATTATTAACGTGGCT
 CCTGCTGGTTATATCCCAAGCCCTAAACATAAAAGCCCTTAAATAGCTAATAACCTTGTGGGGGAATATACTTTTTG
 CAACGGAAAACTTAAAAAATAGCTCAGGGCAAGAACTTTGATGCTCTTTCTGGGGAATTAACAGGAGGGGGCTTG
 GGGATGATGGTCTATCAAGAACCTAGAAAAAGACCCTGGATTCCACATGCATACCTCCGGATATTACAGCAGGAAT
 GATTACAGGAAACACACATACTTCTCATTACGATTACAGCCAGTCTATACAAAACCTCAATGAACGTTATGCCAAGA
 ACTATGTGTCTTCTAAAAAATTAAGTCTTGGCAAGGGGAAATGCTTTTGTCTTACAAGAAGGACTCATGCTGACTAAA
 CTAATTTGGTCTATAGTTATGGGAATCACACAGCCACCATTTCTATACCCAAGGAGAAGACCTATCGTCTCAAGG
 GGAGTTCCATAGTACAGCTTTTGGAGGGGCTGTTTTTGTGATCTACCTCTGAAACCTTTGGAAGAACAACACATAAC
 TTACAGCTCTTTCTTAGGTGCCATTGGTATGATTCTAAGCTGTCTAGCTTTACAGAAGTAGGAGCTATCCAAGA
 ACCTTTATTACAGAAACGCCCTTAAATCAATGCTCTGATCTCTATCGGAGTAAAAGGTAGCTTCTGAAATGCCACCCA

TAGACCTCAGGCCTGGACTGTAGAGCTTGCTTACCAACCTGTTCTTTACAGACAAGAACCTAGTATCTCTACCCAAT
TACTCGCTGGTAAAGGTATGTGGTTTGGGCATGGAAGTCTGCATCTCGCCACGCTCTAGCTTATAAAATTTACAG
AAAACACAGCTTTTGGCATTGCAACACTTCAACTCCAGTATCACGGATACTATTCTGCTTCCACTTTCTGTAATTA
TCTGAATGGAGAGGTATCTTTACGTTTC

SEQ ID NO: 134 - TC0261 fragment protein sequence

TREVPPIILLKPIINPYHMTGLFFPKVNLGDTNHLTDYHLNKLKILACLQRTPYEGAAFTVTDYLGFSDTQKDG
FCFKNLTPESSGGVIGSPTQNTPTIKIHNTIGPVLFENNTCHRLWTQTDPENEGNKAREGGAIHAGDVYISNNQNLV
FIKNFAYVQGGGAISSANTFAYKENKSSFLCLNNSCIQTKTGKGGAIYVSTSCSFENNNDLLFIQNSGCAGGAI
TCSLIGNQGDIVFYSNHGFKNVDNATNESGDDGAIKVTTRLDITNNGSQIFFSDNISRNFGGAIHAPCLHLVGN
YFTNNIANHTGGAIYITGETSKISADHHAIIIFDNNISANATNADGSSSNTNPPHRNAITMNSAGGIELGAGKS
LIFYDPIQVTVNAGVTVDFNKDSQTGCVVFSGATVLSADISQANLQTKTPATLTLSHGLLCIEDRAQLTVNNTQ
GIVALGNLAVLSSYQHSSTDATQTPPTTTTTDASVTLNHLNLSILKDGAEMLLWVEPISTTQGNNTTYSDTA
ASFSLNGATLSLIDEDGNSPYENTDLRSLALYAQPLAISEASDNQLQSESMDFSKVNVPHYGWQGLWTGWAKTEN
TTTTPPATITDPKKAQFHRLLLLTWPAGYIPSPKHKSPLIANTLWGNILFATENLKNSSGQELLDLRFWGITGG
GMMVYQEPKRDHPGFHMHTSGYSAGMITGNHTFSLRFSQSYTKLNERYAKNYVSSKNYSQGEMLLSLQEGLMLTK
LIGLYSYGNHNSHHFYTGEDLSSQGEFHSQTFGGAVFFDLPLKPFGRTHILTAPFLGAIQMYSKLSSFEVGYPR
TFITETPLINVLIPIGVKGSMNATHRPQAWTVELAYQPVLYRQEPSISTQLLAGKGMWFGHGSPASRHALAYKISQ
KTQLLRFATLQLQYHGYSSSTFCNYLNGEVSRLF

SEQ ID NO: 135 - CT600 nucleotide sequence

ATGAGAAAGACTATTTTTAAAGCGTTTAATTTATTATCTCCCTTCTTTTTCTTCTTCATGCTCTTATCCTTGCGAG
AGATTTGGGAATGCCATGGTTGCGACTCCGCAAGACCTCGTAAATCCTCTTTTGGATTTCGTACCTTTCTACTCCGATG
AAGAAATTCACAAGCTTTTGTGGAAGATTTGATTCCAAAAGAGAGCAGCTGTACAAAACGAGCGCACAGAGTACC
TCTTTCCGAAATATCACTTTGCTACAGATAGTTATTCTATTAAGGAGAGGATAACCTCACGATTCTTGCAAGCTT
AGTTTCGTCAATTTGCATAAATCTCCTAAAGCTACGCTATATATAGAGGGCCATACAGATGAACGTGGAGCTGCAGCTT
ATAACCTAGCTTTAGGAGCTCGTCTGCGAATGCTGTAAAAACAATACCTCATCAAAACAGGGAATCGCTGCAGACCGC
TTATTCACTATTTCTACGGAAAAGAACATCCTGTTTCATCCAGGCCATAATGAATTAGCTTGGAACAAAATCGTCG
TACTGAATTTAAGATCCATGCTCGCTAA

SEQ ID NO: 136 - CT600 protein sequence

MRKTIKAFNLLFSLFLSSCSYPCRDWECHGCD SARPRKSSFGFVFPFYSDEEIQQAFVEFDSDSKEEQLYKTSARST
SFRNITFATDSYSIKGEDNLTILASLVRHLHKSPKATLYIEGHTDERGAAAYNLALGARRANAVKQYLIKQGIADR
LFTISYKQHPVHPGHNELAWQQRNRTEFKIHAR

SEQ ID NO: 137 - CT600 fragment nucleotide sequence

TGCTCTTATCCTTGCGAGAGATTGGGAATGCCATGGTTGCGACTCCGCAAGACCTCGTAAATCCTCTTTTGGATTTCGT
ACCTTTCTACTCCGATGAAGAAATTCACAAGCTTTTGTGGAAGATTTGATTCCAAAAGAGAGCAGCTGTACAAA
CGAGCGCACAGAGTACCTCTTTCCGAAATATCACTTTGCTACAGATAGTTATTCTATTAAGGAGAGGATAACCTC
ACGATTCTTGCAAGCTTAGTTTCGTCAATTTGCATAAATCTCCTAAAGCTACGCTATATATAGAGGGCCATACAGATGA
ACGTGGAGCTGCAGCTTATAACCTAGCTTTAGGAGCTCGTCTGCGAATGCTGTAAAAACAATACCTCATCAAAACAGG
GAATCGCTGCAGACCGCTTATTCACTATTTCTACGGAAAAGAACATCCTGTTTCATCCAGGCCATAATGAATTAGCT
TGGCAACAAAATCGTCTACTGAATTTAAGATCCATGCTCGC

SEQ ID NO: 138 - CT600 fragment protein sequence

CSYPCRDWECHGCD SARPRKSSFGFVFPFYSDEEIQQAFVEFDSDSKEEQLYKTSARST SFRNITFATDSYSIKGEDN
LTILASLVRHLHKSPKATLYIEGHTDERGAAAYNLALGARRANAVKQYLIKQGIADR LFTISYKQHPVHPGHNELA
WQQRNRTEFKIHAR

SEQ ID NO: 139 - CT823 nucleotide sequence

ATGATGAAAAGATTATTATGTGTGTTGCTATCGACATCAGTTTTCTCTTCGCCAATGCTAGGCTATAGTGCCTCAA
GAAAGATTCTAAGGCTGATATTTGCTTGCAGTATCCTCAGGAGATCAAGAGGTTTCAACAAGAAGATCTGCTCAAAG
AAGTATCCCAGGATTTTCTCGGGTCGCTGCTAAGGCAACGCTGGAGTTGTATATATAGAAAATTTTCTAAAACA
GGGAACCAGGCTATTGCTTCTCCAGGAAACAAAAGAGGCTTTCAAGAGAACCCTTTTGATTATTTTAATGACGAATT
TTTTAATCGATTTTTTGGATTGCCTTCGCATAGAGAGCAGCAGCGTCCGCAGCAGCGTGATGCTGTAAGAGGAACTG
GGTTCATTGTTTTGAAAGATGGTTATGTTGTTACTAACCATCATGTAGTCGAGGATGCAGGAAAATTCATGTTACT
TCCACAGCAGGACAAAATAACACAGATCAAGATCGTGGGGTTAGATCCAAAACAGATCTTGTCTGTATCAAAATTC
AGCGGAGAAATTAACATTTTTGACTTTTTGGGAATTTGATCAGCTGCAGATAGGTGACTGGGCTATTGCTATTGGAA
ATCCTTTTGGATTGCAAGCAACGGTCACTGTGGGGTCAATTAGTGCTAAAGGAAGAAAATCAGCTACATATTGTAGAT
TTCGAAGACTTTATTCAAAACAGATGCTGCCATTAATCCTGGGAATTCAGGCGGTCCATTGTTAAACATCAATGGTCA
AGTTATCGGGGTTAATACTGCCATTGTCAAGTGGTAGCGGGGATATATTGGAATAGGGTTTGTATTCTAGCTTGA
TGGCTAAACAGGATCATTGATCAATTGATTAGTGTGGCAGGTAACAAGAGGCTTTTGGGAGTTACCTTGCAACCG
ATAGATTCTGAATGGCTACTTGTACAAAATGGAAAAAGTGACGGAGCTTTGGTGACGGATGTTGTTAAAGGTTT
TCCAGCAGAAAAGCAGGCTGCGCCAAGAAGATGTCATTGTGGCTTACAATGGAAAAGAAGTAGAGTCTTTGAGTG
CGTTGCGTAATGCCATTTCCCTAATGATGCCAGGGACTCGTGTGTTTTAAAAATCGTTCGTGAAGGGAAAACAATC
GAGATACCTGTGACGGTTACACAGATCCCAACAGAGGATGGCGTTTACGCGTTGCAGAAGATGGGAGTCCGTGTTCA

GAACATTACTCCAGAAATTTGTAAGAAACTCGGATTGGCAGCAGATACCCGAGGGATTCTGGTAGTTGCTGTGGAGG
CAGGCTCGCCTGCAGCTTCTGCAGGCGTCTGCTCCTGGACAGCTTATCTTAGCGGTGAATAGGCAGCGAGTCGCTTCC
GTTGAAGAGTTAAATCAGGTTTTGAAAAACTCGAAAGGAGAGAATGTTCTCCTTATGGTTTCTCAAGGAGATGTGGT
GCGATTATCGTCTTGAATCAGACGAGTAG

SEQ ID NO: 140 - CT823 protein sequence

MMKRLLCVLLSTSVFSSPMLGYSASKKDSKADICLAVSSGDQEVSDLLKEVSRGFSRVAAKATPGVVYIENFPKT
GNQAIASPGNKRGFQENPFDFYNDEFNRFFGLPSHREQRRPQRDAVRGTGFIVSEDDGYVVTNHHVVEDAGKIHVT
LHDGQKYTAKIVGLDPKTDLAVIKIQAEKLPFLTFGNSDQLQIGDWAIAIGNPFLQATVTVGVISAKGRNQLHIVD
FEDFIQTDAAINPNSGGPLLNINGQVIGVNTAIVSGSGGYIGIGFAIPSLMAKRVIDQLISDQVTRGFLGVTLQPI
DSELATCYKLEKVVYALVTDVVKGSPAEEKAGLRQEDVIVAYNGKEVESLSALRNAISLMMPGTRVVLKIVREGKTI
EIPVTVTQIPTEDEVSALQKMGVVRVQNIPEICKLGLAADTRGILVVAVEAGSPAASAGVAPGQLILAVNRQVRAS
VEELNQLVKNKSGENVLLMVSQGDVVRFIVLKSDE

SEQ ID NO:141 - CT823 fragment nucleotide sequence

TCGCCAATGCTAGGCTATAGTGGCTCAAAGAAAGATTCTAAGGCTGATATTTGCTTGCAGTATCCTCAGGAGATCA
AGAGGTTTCCACAAGAAGATCTGCTCAAAGAAGTATCCCGAGGATTTTCTCGGGTGCCTGCTAAGGCAACGCCTGGAG
TTGTATATATAGAAAAATTTTCTAAAAACAGGGAACAGGCTATTGCTTCTCCAGGAAAAAAAAGAGGCTTTCAAGAG
AACCTTTTGGATTATTTAATGACGAATTTTTAATCGATTTTTTGGATTGCCCTTCGCATAGAGAGCAGCAGCGTCC
GCAGCAGCGTGTACTGCTAAGAGGAACCTGGGTTTATTGTTTCTGAAGATGGTTATGTTGTTACTAACCATCATGTAG
TCGAGGATGCAGGAAAAATTCATGTTACTCTCCACGACGACAAAAATACACAGCTAAGATCGTGGGGTTAGATCCA
AAAAACAGATCTTGTGTGATCAAAATTCAGCGGAGAAATACCATTTTTGACTTTTGGGAATTCGTATCAGCTGCA
GATAGGTGACTGGGCTATTGCTATTGGAAATCCTTTTGGATTGCAAGCAACGGTCACTGTGCGGGTCAATAGTGCTA
AAGGAAGAAATCAGCTACATATTGTAGATTTTGAAGACTTTTATCAACAGATGCTGCCATTAATCCTGGGAATTC
GGCGGTCCATTGTTAAACATCAATGGTCAAGTTATCGGGGTTAATACTGCCATTGTCAGTGGTAGCGGGGATATAT
TGGAATAGGGTTTGTCTATTCTAGCTTGTAGGCTAAACGAGTCATTGATCAATTGATTAGTGATGGGCAGGTAACAA
GAGGCTTTTTGGGAGTTACCTTGGCAACCGATAGATTCTGAATTTGGCTACTTGTACAAATGGAAAAAGTGTACGGA
GCTTTGGTGACGGATGTTGTTAAAGGTTCTCCAGCAGAAAAAGCAGGGCTGCGCCAAGAAGATGTCATTGTGGCTTA
CAATGGAAAAAGAAGTAGAGTCTTTGAGTGGCTTGGTAATGCCATTTCCCTAATGATGCCAGGGACTCGTGTGTTT
TAAAAATCGTTCTGTAAGGGAAAAACAATCGAGATACCTGTGACGGTTACACAGATCCCAACAGAGGATGGCGTTTCA
GCGTTGCAGAAGATGGGAGTCCGTGTTGAGAACATTACTCCAGAAATTTGTAAGAAACTCGGATTGGCAGCAGATAC
CCGAGGGATTCTGGTAGTTGCTGTGGAGGCAGGCTCGCTGCAGCTTCTGCAGGCGTCTGCTCCTGGACAGCTTATCT
TAGCGGTGAATAGGCAGCGAGTCTGCTTCCGTTGAAAGATTAAATCAGGTTTTGAAAAACTCGAAAGGAGAGAAATGTT
CTCCTTATGGTTTCTCAAGGAGATGTGGTGGATTATCGTCTTGAATCAGACGAG

SEQ ID NO:142 - CT823 fragment protein sequence

SPMLGYSASKKDSKADICLAVSSGDQEVSDLLKEVSRGFSRVAAKATPGVVYIENFPKTGNQAIASPGNKRGFQEN
NPFDFYNDEFNRFFGLPSHREQRRPQRDAVRGTGFIVSEDDGYVVTNHHVVEDAGKIHVTLHDGQKYTAKIVGLDP
KTDLAVIKIQAEKLPFLTFGNSDQLQIGDWAIAIGNPFLQATVTVGVISAKGRNQLHIVDFEDFIQTDAAINPNS
GGPLLNINGQVIGVNTAIVSGSGGYIGIGFAIPSLMAKRVIDQLISDQVTRGFLGVTLQPIDSELATCYKLEKVVY
ALVTDVVKGSPAEEKAGLRQEDVIVAYNGKEVESLSALRNAISLMMPGTRVVLKIVREGKTIIEIPVTVTQIPTEDEVS
ALQKMGVVRVQNIPEICKLGLAADTRGILVVAVEAGSPAASAGVAPGQLILAVNRQVRASVEELNQLVKNKSGENV
LLMVSQGDVVRFIVLKSDE

SEQ ID NO:143 – TC0106 nucleotide sequence

ATGCTAACTAACTTTACCTTTTCGCAACTGTCTTTTGTGTTTTTCGTACACATTGTCCAGTGTCCCTGTTTTCTCGGCACC
CCAACTCGCGTAACGCTTCTAGTGGAGCCAAATAAAATCGGATCAGAAGCTTGGATAGAGCAAAAAGTCCGTCAAT
ATCCAGAACTTTTGTGGTTAGTTGAACCTTCTCCTGCAGGAACCTTCTTTAAACGCTCCTTCCGGGATGATCTTTTCT
CCCCTATTGTTCCAAAAGAAAGTCCCTGCTTTTGGATATCGCAGTACGCGAGTCTGATTCACTACACCTGCTTTTCCA
GGGCTCCCGCCAAGCTTATGCTCAGCTTGTCCAGCTGCAGGCTAATGAATCCCCATGACATTTAAACAGTTCCTTA
CCCTACATAAGCAGCTCTCCTTATTCTAAATTCCTAAAGAGTTTTATGATTCCGTCAAAAATTTAGAAAAGTCT
ATCATCTACGCCACTTAGGATGTTCAACAAAAGCTGTTGCCACATTTAAGCCTTATTTTTAGAAAAGCAGAAAAAGA
GGTCTTCTATACAAAAGCTTTGCTGTTCTGCATACTTTCCAGAATTGAGCCCTTCGTTTGTAGACTCTCTCCAG
AACAAAAAAGCTCTTCTCTCATTGAGAAAAGCTCGCTAATTATGATGAGTTACTTTCCCTGACAAAATGCCCTAGT
TTACAACACTACTATCTGCTGTACGCTCGCGACGCGCTTTTGGCTCTAGACTTGTATCTCTATGCTTTAGATTTTTG
TGGAGAACAGGGGATATCCTCTCAGTTTTCATATGGACTTTTTCTCCTTTACAGTCCATGTTGCAACAATATGCTACGG
TTGAAGAAGCCTTCTCCCGCTACTTTACTTACCAGGCTAATCGCCTAGGATTTGCGGGTCTTCTCGAACTGAAATG
GCCTTAGTTAGAAATAGTACTTTAATGAACCTATCCCTTCAGAAGCTGCTATTTTAAACAAGCTTTAAGTCTCT
TTCCTTGGAAAGTGTCTGAAAGCTTAGTGAATAGCTTTTATACAATAAGGGAGACTCTTTAGCTCTTTCTTTACGAG
GACTACCAACTCTTATATCTGAACTAACACGCGCTGCGCATGGAAATACGAATGCGGAAGCTCGAGCTCAGCAAAAT
TACGCCACAACGTTATCATTGGTAGCAAAAAGCTTGAAGCTCACAAAGAGATGCAAAAACAACAATTTCTCCCGA
AGAAGTCTTTTTAGATTTCTCTGAAACTGCTTCTCCTGTCAAGGATTGGACATCTTCTCTGAGAACGTTGCTGTT
AAATCCACTTGAATGGATCTGTACGATCCATCTATAA

SEQ ID NO:144 - TC0106 protein sequence

MLTNFTFRNCLLFFVTLSSVPVFSAPQPRVTLPSGANKIGSEAWIEQKVRQYPELLWLVEPSPAGTSLNAPSGMIFS
PLLFQKKVPAFDIAVRSLIHLHLILQGSRQAYAQLVQLQANESPMTFKQFLTLHKQLSLFLNSPKEFYDSVKILETA

IILRHLGCSTKAVATFKPYFSETQKEVFYTKALHVLHTFPELSPSFARLSPEQKTLFFSLRKLANYDELLSLTNAPS
LQLLSAVRSRRALLALDLYLYALDFCGEQGISSQFHMDFSPLQSMLOQYATVEEAFSRYFTYRANRLGFAGSSRTEM
ALVRIATLMNLSPEAAIILTTSFKLSLEDAESLVNSFYTNKGDSLALSLRGLPTLISELTRAHGNNTNAEARAQRI
YATTLISLVAKSLKAHKEMQNKQILPEEVVLDLDFSETASSCQGLDIFSENVAVQIHLNGSVSIHL

SEQ ID NO:145 - TC0106 fragment nucleotide sequence

TCAGAAGCTTGGATAGAGCAAAAAGTCCGTCAATATCCAGAACTTTTGTGGTTAGTTGAACCTTCTCCTGCAGGAAC
TTCTTTAAACGCTCCTTCGGGGATGATCTTTTCTCCCTATTGTTCCAAAAGAAAGTCCCTGCTTTTGATATCGCAG
TACGCAGTCTGATTACCTACACCTGCTTATCCAGGGCTCCCGCCAAGCTTATGCTCAGCTTGTCCAGCTGCAGGCT
AATGAATCCCCTATGACATTTAAACAGTTTCCCTACCTACATAAGCAGCTCTCCTTATTCCTAAATTCTCCTAAAGA
GTTTTATGATTCGTCAAAATTTAGAAAAGTGTATCTCCTACGCCACTTAGGATGTTCAACAAAAGCTGTTGCCA
CATTTAAGCCTTATTTTTCAGAAAACGCAAAAAGAGGCTTCTCTATACAAAAGCTTTGCATGTTCTGCATACTTTCCCA
GAATTGAGCCCTTCGTTTGTAGACTCTCTCCAGAACAAAAACGCTCTTCTTCTCATTGAGAAAAGCTCGCTAATTA
TGATGAGTTACTTTCCCTGACAAATGCCCTAGTTTACAACACTATCTGCTGTACGCTCGCGACGCGCGCTTTTGG
CTCTAGACTTGTATCTCTATGCTTTAGATTTTTGTGGAGAACAGGGGATATCCTCTCAGTTTCATATGGACTTTTCT
CCTTTACAGTCCATGTTGCAACAATATGCTACGGTTGAAGAAGCTTCTCCCGCTACTTTACTTACCGAGCTAATCG
CCTAGGATTTGCGGGTCTTCTCGAACTGAAATGGCCTTAGTTAGAATAGCTACTTTAATGAACCTATCCCTTCAG
AAGCTGCTATTTTAAACAACAAAGCTTTAAGTCTCTTCCCTTGGAAAGATGCTGAAAGCTTAGTGAATAGCTTTTATA
AATAAGGGGAGACTCTTTAGCTCTTTCTTTACGAGGACTACCAACTCTTATATCTGAACTAACACGCGCTGCGCATGG
AAATACGAATGCGGAAGCTCGAGCTCAGCAAAATTTACGCCACAACGTTATCATTGGTAGCAAAAAGCTTGAAGGCTC
ACAAAGAGATGCAAAAACAAATTTCTCCGAAGAAGTCTTTTTAGATTTCTCTGAAAAGCTTCTTCTGCTCAA
GGATTGGACATCTTCTCTGAGAACGTTGCTGTTCAATCCACTTGAATGGATCTGTCAGCATCCATCTA

SEQ ID NO:146 - TC0106 fragment protein sequence

SEAWIEQKVRQYPELLWLVEPSAGTSLNAPSGMIFSPLLFQKKVPAFDIAVRSLIHLHLLIQGSRQAYAQLVQLQA
NESPMTFKQFLTLHKQLSLFLNSPKFQYDSVKILETAIILRHLGCSTKAVATFKPYFSETQKEVFYTKALHVLHTFP
ELSPSFARLSPEQKTLFFSLRKLANYDELLSLTNAPSLQLLSAVRSRRALLALDLYLYALDFCGEQGISSQFHMDFS
PLQSMLOQYATVEEAFSRYFTYRANRLGFAGSSRTEMALVRIATLMNLSPEAAIILTTSFKLSLEDAESLVNSFYT
NKGDSLALSLRGLPTLISELTRAHGNNTNAEARAQRIYATTLISLVAKSLKAHKEMQNKQILPEEVVLDLDFSETASSCQ
GLDIFSENVAVQIHLNGSVSIHL

SEQ ID NO:147 - TC0431 nucleotide sequence

ATGCCCCACTCTCCTTTTTATATGTTGTTCAACCGCATTCTGTTTTTAATCCTAGATTGGGAGAGCGGCACCCTAT
TACTTTAGATTTCAAAAGAAAAGAATCGATTAGCTGATTTTATTGAAAACCTACCTTTAGAAAATTTTTGGAGCCC
CTTCTTTCTGGAAAATGCTTCTTTAGAAGCCTCTTATGTCTTGTCTAGGGAATCCACAAAAGATGGCACTCTTTTT
ACCGTTCTAGAACCCTAAACTATCTGCCTGCGTAGCTACTTGGCTTGTGGATTCTTATTCTATGGAGCCCGATAA
CGAGCTCTTAGAAGAAATTAACACACTTTGTTGAAAAGCTCTTGTGATGGCGTACAATATCGTGTAACCCGAGAGA
CTCTCCAAAACAAAGATGAAGCCCCAGAGTCTCTTTAGTTGCTGATGATATCGAACTTATCCGCAATGTAGATTTT
TTAGGACGTTCCGTTGATATTGTAATAATGGATCCCTTGAATATTCCTAATACCGTAAGCGAGGAGAATGCTCTCGA
TTACTCTTTACAAAGGGAAACCGCCAAACTTAGCCCTGACGGACGAGTTGGCATCCCTCAAGGGACAAAATTTTTGC
CAGCTCCCTCTCTTGAAGTTGAAATTAGCACTCTATTTTTGAGGAAACCTCTCTTTTGAACAAAACCTTTCTTCC
TCTATTACTTTTTGTGTAACACCTCTTACCTCTTTTTCTCCTTTGCAAGAACCCTCTCTAGTGGGAGCTGGACAGCA
GGAAATTTCTGTGACTAAAAGCACTTATCCCTAGCTATACCCCTAAACTTATTGATATTGTCAAACGACACAAAA
GAGACGCAAGATTCTAGTAAACAAGATCCAGTTGAGAAAATATGGAGAAGTCAATGCCAAAAGTCAAATCTTAAAA
GAAGGCTCTGTTGCTTGGATTTACAAGGATTTACAGGGGAGCTGTTTAACTACCAACTTCAAGTAGGATCTCATA
AATTCAGCCGTTGTTAATTTGATCCGGAATTTGCTAACGTCAAATCCCTCCCGAACAACCTTACGCTGTAAGAAAA
TTAAATCAGGGTCCAATGTAGTTTGGATGACCAACACATTTATCAAGTCGCGAGTAAAAAACATCTTTCTGTCT
TCACAACCTCCGAAGATATCTCCGTTATCTCAATCCGAAAGCTCCGATTTAAGTCTCTTTGAAGCAGCAGCGTTTT
AGCAAGCCTAATCTACGAGTTGTAAGAAAATACATATCATGCTAAGAATACTGTAACCTTGTCCACGGTATCGC
ACTCTCTGTATATTCTCAAAGAAGATGACGGGGCTAATGCTGCAGAAAACGCTTAGACAACAGTTTCCGAAACTGG
GTCGAAAATAAGTTGAACGCAAAATTTCCAGATTTCTGACTGCAATTTATTCAAAAATTCGGCACACATTACATCA
ATCGGCAACTTTTGGAGGATCTGGGTTCCAAGTTCTTAAATTTATCCTTTGAACAGGTAGAAGGCTCCGTAAGTA
AGATCTCCCTAGAAGCAGCAGCAGCAAAATCCTTATTAAAAAGCTCTGTGTCAAACAGCACGGAATCTGGCTACTCT
ACTTACGATTCCTTCTTCTCTCATAAGATTTCTAGGGGGCACTGTATTACCTCTGTTCATGATGGACAGTT
AGATTTTAAAGATTGGTCTGAAAGTGTCTGTTTGAACCTGTTCCATTACATTTCTTTACTCCCTTAAACAGACT
TGCTCACCCCTCTTATTTTCTGAAACGGATACAACCGAACTATCTAATAAACGTAATGCTCTCCAACAAGCGGTT
CGAGTTTACCTTAAAGACCATCGTTTCAGCTAAACAAGCGAAACGCTCCGTATTCAAGCGGGGATCAATAGTCTTC
TTCTGTTTACATTAAGAATCTGCTAATTCACCTCTTGTGTGAGTTCTCCTTACATGACGATTTGGTCTACTCTCC
CCTATCTCTTCCCAACATTAAGAGAGCGTTCTTACGAGCTCCCATCGTTTTTTATTTTTGTGTGGATAATAATGAA
CACGCCCTCCAAAATTTTAAACCAACATATTGCTTCATAGGTTCTTTACCTATTGACAAAAGATTTTTGGCAG
AGAAATTTGCTGAGAATCCTTATTTATCTTTCTATGGAAGGTTTGGAGAAGCTTATTTGATGGCGGTTATCCAGAAC
GTTGTGGATGGATTGTTGAAAAGTTAAATACTACTAAAGATCAAATTTCCGCGATGAGGATGAAGTGAACATAAG
CATGTTTATAGCGGAGAGTATCTGTCTACAATTCCTATTAAGGATTCCTTGCACACTCTCGCGTACATGCACCGA
ATCGAATGCTGTTTTTATTATCAAAAACCTTCGAGCTATTGA

SEQ ID NO:148 - TC0431 protein sequence

MPHSPFLYVYVQPHSVFNPRLGERHPITLDFI KEKNRLADFIENLPLEIFGAPSFLENASLEASYVLSRESTKDGTLF
TVLEPKLSACVATCLVDSSIPMEPDNELLEEIKHTLLKSSCDGVQYRVTRTELQNKDEAPRVSLVADDIELIRNVDF
LGRSVDIVKLDPLNIPNTVSEENALDYSFTRETAKLSPDGRVGIQGTIKLPAPSLEVEISTSI FEETSSFEQNFSS
SITFCVPLTSFSPLQEPPLVGAGQEQEILVTKKHLFPSYTPKLI DIVKRHKRDAKILVNKIQFEKLRSHAKSQILK
EGSVRLDLQGF TGELFNQQLQVGSHTIAAVLIDPEIANVKSLEPQTYAVRKIKSGFQCSLDDQHIYQVAVKKHLSLS
SQPPKISPLSQSESSDLSLFEAAAFSASLYEFVKKNTYHAKNTVTCSTVSHSLYILKEDDGANAAEKRLDNSFRNW
VENKLNANSPDSCFAIQKFGTHYITSATF GSGGFQVLKLSFEQVEGLRSKKISLEAAAAANLLKSSVSNSTESGYS
TYDSSSSSHTVFLGGTVLPSVHDGQLDFKDWSESVCLPVIHISLLPLTDLLTPLYFPE TDTTELSNKRNALQAVR
RVYVKDHRSAKQSERVFTAGINSPSSWFTLESANSPLVVSSPYMTYWSTL PYLFP TLKERSSAAPIVYFCVDNNE
HASQKILNQTYCFIGSLPIRQKIFGREFAENPYLSFYGRFGEAYFDGGYPERCGWIVEKLN TTKDQILRDEDEVQLK
HVYSGEYLS TIPIKDSHCTLSRTCTESNAVFIKKPSSY

SEQ ID NO:149 - TC0431 fragment nucleotide sequence

CCCCACTCTCCTTTTTATATGTTGTTCAACCGCATTCTGTTTTAATCCTAGATTGGGAGAGCGGCACCCATTAC
TTTAGATTTTCATCAAAGAAAAGAATCGATTAGCTGATTTTATTGAAAACCTACCTTTAGAAAATTTTTGGAGCCCTT
CTTTCTTGAAAAATGCTCTTTAGAAAGCCTCTTATGTCTGTCTAGGGAATCCACAAAAGATGGCATTCTTTTACC
GTTCTAGAACCCAAACTATCTGCCTGCGTAGCTACTTGCCTTGTGGATTCTTCTATTCCCTATGGAGCCGATAACGA
GCTCTTAGAAGAAATTAACACACTTTGTTGAAAAGCTCTTGTGATGGCGTACAATATCGTGTAACCCGAGAGACTC
TCCAAAACAAAGATGAAGCCCCAGAGTCTCTTTAGTTGCTGATGATATCGA ACTTATCCGCAATGTAGATTTTTTA
GGACGTTCCGTTGATATTGTA AAAATTGGATCCCTTGAATATTCCTAATACCGTAAGCGAGGAGAATGCTCTCGATTA
CTCTTTCAAGGGAAACCGCAAACCTTAGCCCTGACGGACGAGTTGGCATCCCTCAAGGGACAAAATTTTTGCCAG
CTCCCTCTCTTGAAGTTGAAATTAGCACCTCTATTTTTGAGGAAAACCTCTTCTTTTGAACAAAACCTTTTCTTCTCT
ATTACTTTTTGTGTACCACCTCTTACCTCTTTTTCTCCTTTGCAAGAACCTCTCTAGTGGGAGCTGGACAGCAGGA
AATTCTTGTGACTAAAAAGCACTTATTCCCTAGCTATACCCCTAAACTTATTGATATTGTCAAACGACACAAAAGAG
ACGCAAAGATTCTAGTAAACAAGATCCAGTTCGAGAAACTATGGAGAAGTCA TGCCAAAAGTCAAATCTTAAAAGAA
GGCTCTGTTGCTGGATTTACAAGGATTTACAGGGGAGCTGTTTAACTACCAACTCAAGTAGGATCTCATACAAT
TGCAGCCGTGTTAATTGATCCGGAATTGCTAACGTCAAAATCCCTCCCGAACAAAACCTTACGCTGTAAGAAAAATTA
AATCAGGGTTCCAATGTAGTTTGGATGACCAACACATTTATCAAGTCGCAGTAAAAAACATCTTTCTCTGTCTTCA
CAACCTCCGAAGATATCTCCGTTATCTCAATCCGAAAGCTCCGATTTAAGTCTCTTTGAAAGCAGCAGCGTTTTCAGC
AAGCCTAACTTACGAGTTCGTAAGAAAAATACATATCATGCTAAGAATACTGTA ACTTGCTCCACGGTATCGCACT
CTCTGTATATTTCAAGAAGATGACGGGGCTAATGCTGCAGAAAAACGCTTAGACAACAGTTTTCCGAAAACGGGTC
GAAAATAAGTTGAACGCAAATTTCTCCAGATTCTGTACTGCATTTATTCAAAAATTCGGCACACATTACATCACATC
GGCAACTTTTGGAGGATCTGGGTTCCAAGTTCTTAAATTATCCTTTGAAACAGGTAGAAGGCCCTCCGTAGTAAGAAGA
TCTCCCTAGAAGCAGCAGCAGCAAATTTCTTATTA AAAAGCTCTGTGTCAAACAGCACGGAATCTGGCTACTCTACT
TACGATTCCTCTTCTTCTCTCATAACAGTATTCCTAGGGGGCACTGTATTACCTCTGTTT CATGATGGACAGTTAGA
TTTTAAAGATTGGTCTGAAAAGTGTCTGTTTAGAACCTGTTCCCATTCACATTTCTTTACTCCCCTTAAACAGACTTGC
TCACCCCTCTTTATTTCTGAAACGGATACAACCGAATCTAATAAACGTAATGCTCTCCAACAAGCGGTTCCGA
GTTTACCTTAAAGACCATCGTTT CAGCTAAACAAGCGAACGCTCCGTAATTCACAGCGGGGATCAATAGTCTTCTTC
CTGGTTCACATTAGAACTGCTAATTCACCTCTTGTGTGAGTTCTCCTTACATGACGTATTGGTCTACTCTCCCT
ATCTCTTCCCACTTAAAAGAGCGTTCCTT CAGCAGCTCCCATCGTTTTTTATTTTTGTGTGGATAATAATGAACAC
GCCTCCCAAAAATTTTAAACCAAACATATTGCTT CATAGGTTCTTTACCTATTCGACAAAAGATTTTTGGCAGAGA
ATTTGCTGAGAATCCTTATTTATCTTTCTATGGAAGTTTGGAGAAGCTTATTTT GATGGCGTTATCCAGAACGTT
GTGGATGGATTGTTGAAAAGTTAAATACTACTAAAGATCAAATTTCTCGCGATGAGGATGAAGTGCAACTAAAGCAT
GTTTATAGCGGAGATATCTGTCTACAATTCCTATTAAGGATCCCATTCACACTCTCGCGTACATGCACCGAATC
GAATGCTGTTTTTATTATCAAAAACCTTCCGAGCTAT

SEQ ID NO:150 - TC0431 fragment protein sequence

PHSPFLYVYVQPHSVFNPRLGERHPITLDFI KEKNRLADFIENLPLEIFGAPSFLENASLEASYVLSRESTKDGTLF T
VLEPKLSACVATCLVDSSIPMEPDNELLEEIKHTLLKSSCDGVQYRVTRTELQNKDEAPRVSLVADDIELIRNVDFL
GRSVDIVKLDPLNIPNTVSEENALDYSFTRETAKLSPDGRVGIQGTIKLPAPSLEVEISTSI FEETSSFEQNFSSS
ITFCVPLTSFSPLQEPPLVGAGQEQEILVTKKHLFPSYTPKLI DIVKRHKRDAKILVNKIQFEKLRSHAKSQILKE
GSVRLDLQGF TGELFNQQLQVGSHTIAAVLIDPEIANVKSLEPQTYAVRKIKSGFQCSLDDQHIYQVAVKKHLSLS
QPPKISPLSQSESSDLSLFEAAAFSASLYEFVKKNTYHAKNTVTCSTVSHSLYILKEDDGANAAEKRLDNSFRNWV
ENKLNANSPDSCFAIQKFGTHYITSATF GSGGFQVLKLSFEQVEGLRSKKISLEAAAAANLLKSSVSNSTESGYST
YDSSSSSHTVFLGGTVLPSVHDGQLDFKDWSESVCLPVIHISLLPLTDLLTPLYFPE TDTTELSNKRNALQAVR
VYVKDHRSAKQSERVFTAGINSPSSWFTLESANSPLVVSSPYMTYWSTL PYLFP TLKERSSAAPIVYFCVDNNEH
ASQKILNQTYCFIGSLPIRQKIFGREFAENPYLSFYGRFGEAYFDGGYPERCGWIVEKLN TTKDQILRDEDEVQLKH
VYSGEYLS TIPIKDSHCTLSRTCTESNAVFIKKPSSY

SEQ ID NO:151 - TC0210 nucleotide sequence

ATGATGAAAAGATTATTATGTGTGTTGCTATCGACATCAGTTTTCTCTTCCGCCATGTTGGGCTATAGTGCGCCAAA
GAAAGATTCCAGTACTGGCATTGTCTTGCAGCATCTCAAAGTGATCGGGAACCTTCCCAAGAAGATTTGCTAAAAG
AAGTGTCTAGAGGATTTTCAAAGTCGCTGCTCAGGCAACTCCAGGAGTTGTGTATATAGAAAATTTTCTAAAAC
GGGAGTCAAGCTATTGCTTCTCCTGGGAATAAAAGGGGTTTTCAAGAGAATCCCTTTGATTATTTCAATGATGAGTT
TTTCAATCGATTTTTTGGTTTACCCTCGCATAGAGAGCAGCCTCGTCCCAACAGCGTGATGCTGTAAGAGGAACAG

GTTTTATTGTGTCAGAAGATGGGTACGTTGTGACCAACCATCACGTAGTGGAAGATGCGGGGAAAATTCATGTTACT
 TTACACGATGGACAAAAATACACCGCAAAAAATCATAGGATTAGATCCTAAAACGGATCTCGCTGTGATTAAGATCCA
 AGCAAAAAATCTCCCTTTTTAACTTTTTGGAACCTCTGATCAGCTTCAGATAGGGGATTGGTCAATAGCCATTGGAA
 ATCCTTTCCGATTACAAGCCACAGTAACCGTTGGCGTGATTAGTGCTAAGGGAAGAAACCAATTACATATTGTTGAT
 TTTGAAGATTTTATTTCAGACGGATGCAGCAATTAATCCCGGGAATTCAGGTGGTCCATTATTGAACATTGATGGACA
 GGTTATTGGAGTGAATACAGCAATCGTTAGCGGTAGCGGGGATACATTGGAATAGGATTTGCCATTCTAGCTTAA
 TGGCTAAACGAGTTATTGACCAACTCATTAGCGATGGACAGGTGACGAGAGGATTTTTAGGAGTAACCTTACAGCCT
 ATTGATTCCGGAGCTTGCCGCTTGTACAAATTAAGAAAAGGTGTACGGAGCCTTGATTACGGATGTTGTTAAGGGATC
 TCCTGCAGAAAAAGCAGGTTTGCGCCAGGAAGATGTCATTGTTGCTTACAATGGGAAAGAAGTGGAGTCTTTGAGTG
 CTTTACGTAATCGGATTTCTTTGATGATGCCAGGGACTCGTGTGCTTAAAAGTTGTGCGTGAAGGGAAATTCATT
 GAAATACCTGTCACTGTTACACAAATTCCTGCGGAGGATGGGGTATCTGCTCTTCAAAAAATGGGAGTTCGGGTACA
 GAATCTTACTCCAGAGATATGCAAGAAACTAGGATTAGCGTCTGATACTCGAGGGATTTTTGTAGTGTCCGTAGAAG
 CTGGTTCTCCTGCAGCTTCTGCAGGAGTGGTTCCAGGACAACCTTATTCTGGCTGTAAACAGACAGAGAGTTTCTTCT
 GTTGAAGAATTGAATCAGGTCTTGAAGAATGCAAAAGGAGAGAATGTTCTCCTTATGGTTTCTCAAGGAGAAGTCAT
 TCGATTCTGTTGTTTTAAAGTCTGATGAATAG

SEQ ID NO:152 - TC0210 protein sequence

MMKRLLCVLLSTSVMSSPMLGYSAPKKDSSTGICLAASQSDRELSQEDLLKEVSRGFSKVAAQATPGVVYIENFPKT
 GSQAIASPGNKRGFQENPFDFNDEFFNRFFGLPSHREQPRPQQRDAVRGTGFIVSEDEYVVVTHHVVVEDAGKIHVT
 LHDGQKYTAKIIGLDPKTDLAVIKIQAKNLPFLTFGNSDQLQIGDWSIAIGNPFLQATVTVGVISAKGRNQLHIVD
 FEDFIQTDAAINPNSGGPPLLNDGQVIGVNTAIVSGSGGYIGIGFAIPSLMAKRVIDQLISDGQVTRGFLGVTLQRP
 IDSELAACYKLEKVGALITDVVKGSPAEEKAGLRQEDVIVAYNGKEVESLSALRNAISLMMPGTRVVLKVVREGKFI
 EIPVTVTQIPAEDGVSALQKMGVRVQNLTPEICKKLGLASDTRGIFVVSVEAGSPAASAGVVPGLILAVNRQVRVSS
 VEELNQLVKNKAGENVLLMVSQGEVIRFVVLKSDE

SEQ ID NO:153 - TC0210 fragment nucleotide sequence

TCCCCATGTTGGGCTATAGTGCGCCAAAAGAAAGATTCCAGTACTGGCATTGTTGCTTGCAGCATCTCAAAGTGATCG
 GGAACCTTTCCCAAGAAGATTTGCTAAAAGAAGTGCTAGAGGATTTTCAAAGTCGCTGCTCAGGCAACTCCAGGAG
 TTGTGTATATAGAAAATTTTCTAAAACCTGGGAGTCAAGCTATTGCTTCTCCTGGGAATAAAAGGGGTTTTCAAGAG
 AATCCCTTTGATTATTTCAATGATGAGTTTTTCAATCGATTTTTTGGTTTACCCTCGCATAGAGAGCAGCCTCGTCC
 CCAACAGCGTGATGCTGTAAGAGGAACAGGTTTTATTGTGTCAGAAGATGGGTACGTTGTGACCAACCATCACGTAG
 TGGAAGATGCGGGGAAAAATTCATGTTACTTTACACGATGGACAAAAATACACCGCAAAAAATCATAGGATTAGATCCT
 AAAACGGATCTCGCTGTGATTAAGATCCAAGCAAAAAATCTCCCTTTTTAACTTTTGGAAACTCTGATCAGCTTCA
 GATAGGGGATTTGGTCAATAGCCATTGGAAATCCTTTCCGATTACAAGCCACAGTAACCGTTGGCGTGATTAGTGCTA
 AGGGAAGAAACCAATTACATATTGTTGATTTTTGAAGATTTTTATTAGACGGATGCAGCAATTAATCCCGGGAATTCA
 GGTGGTCCATTATTGAACATTGATGGACAGGTTATTGGAGTGAATACAGCAATCGTTAGCGGTAGCGGGGGATACAT
 TGGAATAGGATTTGCCATTCTAGCTTAATGGCTAAACGAGTATTGACCAACTCATTAGCGATGGACAGGTGACGA
 GAGGATTTTTAGGAGTAACTTACAGCCTATTGATTCGGAGCTTGGCGCTTGTACAAATAGAAAAGGTGTACGGA
 GCCTTGATTACGGATGTTGTTAAGGGATCTCCTGCAGAAAAAGCAGGTTTTGCGCCAGGAAGATGTCATTGTTGCTTA
 CAATGGGAAAGAAGTGGAGTCTTTGAGTGTCTTACGTAATGCGATTTCTTTGATGATGCCAGGGACTCGTGTGTCT
 TAAAAGTTGTGCGTGAAGGAAATTCATTGAAATACCTGTCACTGTTACACAAATTCCTGCGGAGGATGGGGTATCT
 GCTCTTCAAAAAATGGGAGTTCGGGTACAGAATCTTACTCCAGAGATATGCAAGAAACTAGGATTAGCGTCTGATAC
 TCGAGGGATTTTTGTAGTGTCCGTAGAAGCTGGTTCTCCTGCAGCTTCTGCAGGAGTGGTTCCAGGACAACCTTATTC
 TGGCTGTAACAGACAGAGAGTTTTCTTCTGTTGAAGAATTGAATCAGGTCTTGAAGAATGCAAAAGGAGAGAATGTT
 CTCCTTATGGTTTCTCAAGGAGAAGTCATTGATTCGTTGTTTTAAAGTCTGATGAA

SEQ ID NO:154 - TC0210 fragment protein sequence

SPMLGYSAPKKDSSTGICLAASQSDRELSQEDLLKEVSRGFSKVAAQATPGVVYIENFPKTGSQAIASPGNKRGFQ
 NPFDFYFNDEFFNRFFGLPSHREQPRPQQRDAVRGTGFIVSEDEYVVVTHHVVVEDAGKIHVTLHDGQKYTAKIIGLDP
 KTDLAVIKIQAKNLPFLTFGNSDQLQIGDWSIAIGNPFLQATVTVGVISAKGRNQLHIVDFEDFIQTDAAINPNS
 GGPLLNDGQVIGVNTAIVSGSGGYIGIGFAIPSLMAKRVIDQLISDGQVTRGFLGVTLQRPIDSELAACYKLEKVG
 ALITDVVKGSPAEEKAGLRQEDVIVAYNGKEVESLSALRNAISLMMPGTRVVLKVVREGKFI
 EIPVTVTQIPAEDGVSALQKMGVRVQNLTPEICKKLGLASDTRGIFVVSVEAGSPAASAGVVPGLILAVNRQVRVSS
 VEELNQLVKNKAGENVLLMVSQGEVIRFVVLKSDE

SEQ ID NO:155 – CT163 nucleotide sequence

ATGTTTGTGTCGTTTCGATAAATCCCGTTGCAGAGCGGATGTCCCCGATTTTTTTGAAAAGGACAGGAAACTTTCTTCT
 CCATTGTGTGGCAAGAGGGATCAATGTTTTATATCGTGTGAAACAAATCTCTAACTATCCTTCATGCTATTTCTCAC
 ATAAAGAGATTTCTGTGTTGTCGTCGTATTGCAACATTGTGATCTGTATTCTCACAGGGCCTCTGATGTTATTGGCC
 ACTGTGTTAGGATTTAGCGTATAGGTTTTCTTCTACTTACCAGACTTCTTTACAAGAACGCTTTCTGTTATAAATA
 TGAACAAAAGCAAGCTTTAGATGAATACCGTGATAGGGAAGAAAAAGTCATTACGCTTCAGAAAGTTTTGTAGAGGAT
 TTCTAGTTAGAAAATCAATTTGCTCAACCAAGAACTTTAAACAACGTGTAAGCAATGGGGGCAAAAACCTATTAGAAGGA
 GAAAAATTTCCAAAGGTTCCAGAGGACGGTCTCTGATATATATTTTCAAACAGTTTCTTTCTTTAGTAGCAAAAACA
 CAGTTGGGGCTCAAGATGCCAGGTCTCGTTGGCATCATATTTTTCTATGCGCAAAAGCGCTTGTCTTATTAGATATTA
 AGCGCATAACGAGCACCACGCGCTAGAGTTTTATCAAAACTTTATATTTGAAAGAAAACTTCTGTTTTACGAATTTCT
 GTAGATTCAATGTGTCCTATAAAAGAAAAATCCACAAGCTTTTCGATGAGGCGATCAAAGAACTCTTATTTCTATTTAA
 AGAAGTGCATTTCAAGGATTTTTGTTGTAGAAACAGAGTCTCCAACAGACGATTTCCCTTAGCCGTGAAAGTACACA

ACTATTGGGTATGCCACGATACGATAATTTACCTTTATTTATTCAAGAAGGAAAAGATGGCTCTCCAGAAGGGCGT
 ATAGGACTGGTTCGATCTAGAAACTTTTTCTTGGTCTCCACATCCATACCCCGTAGAAGAACTAGCTGTGATGTTTCC
 TATGCATAAAGAGCTTCTTATGACAGAGGGCGAAAAAACTACAAATCCCTTTCTCTACAAAAGGAGGTCGAGCGCTCTG
 TAGAGAAAAGGGCTTGGCTTTTTTGAACATATGCTAGGGCATCAAGATTTTTGTTCCCAAAAAAGCGTAACGCCATTG
 CGTAATTGTGCCCTTATATTCATCTAGAAGTATGGAGATTTCTCACTGAAAATTTTTGATTTTTTAAAAGCTGCTAT
 TCAACTAAATGGAGCACTCAATGTTCTGTTATCTCCAGATATTCGAGAGCGGTTGAGTGCTATTTCCGGATAAGCAAT
 GGTTGGCTATTAGCTCCCAGGTTACGTCATCGTTACTCGAGCAAGTTTCTACAAACATCTATCAGTCTCATACTGAA
 GAGGCTAAACGAGTAAATTTCTCAGGGACTTTTATCATGTGTCGATCTCCTATCTCCGGAAAAGCATCTTCATTA
 AAATCTCCCACAATTTCTAAACAAGAAATTCAGTTGCTTCCAGAGGAGAAAAGCAATCAGCGAGGCGCTTGTCTCTC
 TATGTTTACGTTGCAGTAATGGAAGAGCTAGTAGCAACAGGAAAATTTTATTCTTATGATTCTATGGATGATTTTTTT
 GAAGGGCAGTATTGTCGCATTCGTTATTAG

SEQ ID NO:156 – CT163 protein sequence

MFVVSFDKSRCRADVPDFFERTGNFLLHCVARGINVL YRVKQISNYPSCYF SHKEISCCRRIANIVICILTGPLMLLA
 TVLGLLAYRFSSTYQTSLQERFRYKYEQKQALDEYRDREKVVITLQKFCRGFLVRNHLLNQETLTTCKQWQKLLLEG
 EKFRVPEGRSLVYISKQFPVSLVAKHVGAQDARSRWHHIFSMRKALAYLDIKRIRAPRARVYQNFIFEEKLPVSRIS
 VDSMCLYKENPQAFDEAIKELLFLFKEVHFRDFVVE TESPTDDFPLAVKVHNYWVCPRYDNLPLFIQEGKDGSPGR
 IGLVDLETFWSPHYPVEELAVMFPMHKELLMTEAKKLQIPFSTKEVERSVKGLAFFEHMLGHQDFCSQKSVTPL
 RNCAPYIHLEVWRFSLKIFDILKAAIQNLGALNVLLSPDIRERLSAISDKQWLAISSQVTSLLLEQVSTNIYQSHTE
 EAKRVNSSGTFIMCRSPIFRKSIFIKNLPQFLNKKLQLLPEEKAISEALASLCLRAVMEELVATGNIYSYDSMDDFF
 EQYCRIRY

SEQ ID NO:157 – CT163 fragment nucleotide sequence

TTTGTGTCGTTGATAAAATCCCGTTGAGAGCGGATGTCCCGATTTTTTTTGAAGGACAGGAACTTTCTTCTCCA
 TTGTGTGGCAAGAGGGATCAATGTTTTATATCGTGTGAAACAAATCTCTAACTATCCTTCATGCTATTTCTCACATA
 AAGAGATTTTCGTGTTGTCGTCGATTGCAAACATTGTGATCTGTATTCTCACAGGGCCCTGATGTTATTGGCCACT
 GTGTTAGGATTATTAGCGTATAGGTTTTCTTCTACTTACCAGACTTCTTTACAAGAACGCTTTTCGTTATAAATATGA
 ACAAAAAGCAAGCTTTAGATGAATACCGTGATAGGGAAGAAAAGTCAATACGCTTCAGAAGTTTTGTAGAGGATTT
 TAGTTAGAAATCATTGCTCAACCAAGAAAACCTTAAACAACGTTAAGCAATGGGGGCAAAAACATTTAGAGGAGAA
 AAATTTCCAAAGGGTCCAGAAAGGACGGTCTCTTGTATATATTTTCAAAACAGTTTCTTTTAGTAGCAAAAACAGT
 TGGGGCTCAAGATGCCAGGTCTCGTTGGCATCATATTTTTTCTATGCGCAAAGCGCTTGCTTATTTAGATATTAAGC
 GCATACGAGCACCACGCGCTAGAGTTTATCAAACTTTATATTCGAAGAAAAACTTCTGTTTACGAATTTCTGTA
 GATTCAATGTGTCTCTATAAAGAAAATCCACAAGCTTTTCGATGAGGCGATCAAAGAACTCTTATTTCTATTTAAAGA
 AGTGCAATTTTCAGGGATTTTGTGTAGAAACAGAGTCTCCAACAGACGATTTCCCTTAGCCGTGAAAAGTACACAAC
 ATTGGGTATGCCACGATACGATAATTTACCTTTATTTTATTCAAGAAGGAAAAGATGGCTCTCCGAAAGGGCGTATA
 GGACTGGTTCGATCTAGAAAATTTTTCTTGGTCTCCACATCCATAACCCGTTAGAGAACTAGCTGTGATGTTTCTAT
 GCATAAAGAGCTTCTTATGACAGAGGGCGAAAAAACTACAAATCCCTTTCTCTACAAAAGGAGGTCGAGCGCTCTGTAG
 AGAAAGGGCTTGTCTTTTTTGAACATATGCTAGGGCATCAAGATTTTTGTTCCCAAAAAAGCGTAACGCCATTGCGT
 AATTGTGCCCCTTATATTCATCTAGAAGTATGGAGATTTCTCACTGAAAATTTTTGATATTTTTAAAAGCTGCTATTCA
 ACTAAATGGAGCACTCAATGTTCTGTTATCTCCAGATATTCGAGAGCGGTTGAGTGCTATTTCCGGATAAGCAATGGT
 TGGCTATTAGCTCCCAGGTTACGTCATCGTTACTCGAGCAAGTTTCTACAAACATCTATCAGTCTCATACTGAAGAG
 GCTAAACGAGTAAATTTCTCAGGGACTTTTATCATGTGTCGATCTCCTATCTTCCGGAAAAGCATCTTCATTA
 TCTCCACAATTTCTAAACAAGAAATTCAGTTGCTTCCAGAGGAGAAAAGCAATCAGCGAGGCGCTTGTCTCTAT
 GTTTACGTGCAGTAATGGAAGAGCTAGTAGCAACAGGAAAATTTTATTCTTATGATTCTATGGATGATTTTTTTGAA
 GGGCAGTATTGTCGCATTCGTTAT

SEQ ID NO:158 – CT163 fragment protein sequence

FVVSFDKSRCRADVPDFFERTGNFLLHCVARGINVL YRVKQISNYPSCYF SHKEISCCRRIANIVICILTGPLMLLAT
 VLGLLAYRFSSTYQTSLQERFRYKYEQKQALDEYRDREKVVITLQKFCRGFLVRNHLLNQETLTTCKQWQKLLLEGE
 KFPRVPEGRSLVYISKQFPVSLVAKHVGAQDARSRWHHIFSMRKALAYLDIKRIRAPRARVYQNFIFEEKLPVSRISV
 DSMCLYKENPQAFDEAIKELLFLFKEVHFRDFVVE TESPTDDFPLAVKVHNYWVCPRYDNLPLFIQEGKDGSPGRI
 GLVDLETFWSPHYPVEELAVMFPMHKELLMTEAKKLQIPFSTKEVERSVKGLAFFEHMLGHQDFCSQKSVTPLR
 NCAPYIHLEVWRFSLKIFDILKAAIQNLGALNVLLSPDIRERLSAISDKQWLAISSQVTSLLLEQVSTNIYQSHTEE
 AKRVNSSGTFIMCRSPIFRKSIFIKNLPQFLNKKLQLLPEEKAISEALASLCLRAVMEELVATGNIYSYDSMDDFFE
 EQYCRIRY

SEQ ID NO:159 - CT214 nucleotide sequence

ATGCGAACAGACTCTCTTTTCAATCCTCCCGACTCTACTAGAGGAGTTTTTTCAGTTTTTATAGAGACTCAGTGTGATCG
 AGCCGTGGCTCGGTCCAGACAAAGCCAATTTATAGGGTTAGTCTCTGCTGTAGCAGCTGCAGCATTATTATTGTTGC
 TTGTGGTTCGCTCTATCTGTTCCAGGATTTCCAGTTGCAGCTTCAATTTGTTGTAGGGGTTCTCTTTGCTTTATCGATC
 GTAGCATTAACAGCTTCGTTTTTGGTATATATAGCTAATGCTAAGCTTGTGCAATAAGAATTAATTTCTGAGTAG
 TGGTCTGCAAGATCACTTTTCCGAGTCTATTTTTAGGGACTCTCCGTAAGGACGTTGGTGCTAGTATTCGGCTTA
 TTTCCGGACAAGCAGATGATCCTCTCCCTAATCGGATTTGGGATCAAAAAAGCACTGAAAATGCGTGTCTTCAAAAA
 GGAATTTGGGACAGATTATAAAAAAATAAGCAGCATCTTGATAGAGTGAATAATGATTTCACTTTTGTCTGTGAGGG
 GATTAGCGCTTTAATTTCTACAGAAAAAGATGCTCCATTTCCCTATAGAACCCTTCTCAATTTAGCAGGTGTTTTTTAG
 TATCATTTTACCAGACAAGAATCCGATTCTAAAGATTACGCGTCATGCTGAGAAGATGTTACAGCCTCCTCAAGGC
 GGATTTCCCTAACGGGCTGGTTTTGGTTGTGTGGAGCTTTTCTGATCCTAAGAAATTTGCAGCTCCCTTTCTATCTTT

GATTGAGAAGACTCACCAAGGGATTTTGGTGAGTAAAGACTTGAAAGACAATAAGGAAAGAAAGCTAGCTTTAGAGG
 CTTCCCTTCTTTTCATTGAATATTTTCTTTCCGGTGGTGGTGGGGAATCCGGAGTACAATCAGTATATCACAAC
 GCTGTAGCTGAGAAATATAGGGATGTCTCTGTAAGAAATTTGATTTTATGATTTCTGGATACAGGGAATGTGATTC
 AGCTCTTGCTTTAGCAAGTAGTTATTCACAAGATCCGCTTGGGCTGCAGGGTTCAGAAAAGTTTACGTGAAGAAG
 AAAAAAGACTAAGAAAAAGTCACGTGAAGAAGTCTCTGTTTGTATCGTGATATAGATCCAGGCTGTTGTTAAGA
 GCCCTTCCCTAAGCGATTTGAATCCAAGTCTTCAGGTAGTCAAGGTAGTCTTAAAGAGCAGTTAAGCTCTTTGTTGAA
 AGCTTTAGACCAGAAAATTCCTTCAGGGATTTTAGGATTGATTGCAAAAGCTTCTTCTGCAGATCTCAAGGCTGATT
 TTGCAGGTATGCTTGAAGTTATTAAGCAATTACAAGCTTATTTCGATTCTTACCCACCTTTATGCGAAGACAATATT
 CTCTTGTGGTTAAGCGCTTCTTTAGAACAAAGTAGGCTTGCAAGAAATTTGAGAACCCTTTTACCTTCATCAGAAAA
 AAACTCTTAGAAAGAGTTCTCTCTACATTTTTATTAGGTTTGTATACTCGAGGAGCTTTTCTGTAGGGCAAGTGA
 ATCAGCTAGCTACTATTTGTAATACTCAGGACTCTACAGAATTCTGCCAGAGAGTAAGTGACCTTTTCGTTAATTA
 CGAGCTCTACCTGCATTATTTGGTTAA

SEQ ID NO:160 - CT214 protein sequence

MRTDSLFPDSTRGVFQFLETQCDRAVARSRQSQFIGLVSAVAAAAALLLLVVALSVPGFVPAASIVVGVLFALSI
 VALTASFLVYIANAKLVAIRIKFLSSGLQDHFSESSILGTLRKGRGASIP LISGQADDPLPNRIGIKKSTEMRVLQK
 GIGTDYKYYKQHLDRVNDFTFVCEGISALIPTEKDAFPPIEPSHLAGVFLVSPDKNPILKITRHAEKMLQPPQG
 GPNGLVWLCGALSDPKKFAAPFLSLIEKTHQGILVSKDLKDNKERKLALAEASLLSLNIFSSGWCLGNPEYNQYITTA
 AVAEKYRQDVSVRNCIYDFLDTGNVISALALASSYSQDSAWAAGLQKVLREEDKTKKKSREEVSVCLYRDIIDPGCCLR
 ALPKRFESKSSGSQSPKEQLSSLLKALDQKIPSGILGLIAKASSADLKADFAGMLEVIKQLQALFDSYPLCEDNII
 LLWLSASLEQVGLQKLRFLPSSEKLLERVLSTFLLGLYTRGVFSVGQVNLATICNTQDSTEFQVRVSDLSLIR
 RALPALFG

SEQ ID NO:161 - CT214 fragment nucleotide sequence

CGAACAGACTCTCTTTTCAATCCTCCGACTCTACTAGAGGAGTTTTTTCAGTTTTTTCAGACTCAGTGTGATCGAGC
 CGTGGCTCGGTCCAGACAAAAGCCAATTTATAGGGTAGTCTCTGCTGTAGCAGCTGCAGCATTATTATTGTTGCTTG
 TGGTCGCTCTATCTGTTCCAGGATTTCCAGTTGCAGCTTCAATTTGTTGTAGGGGTTCTCTTTGCTTTATCGATCGTA
 GCATTAACAGCTTCGTTTTTGGTATATATAGCTAATGCTAAGCTTGTGCAATAAGAATTAATTTCTTGAGTAGTGG
 TCTGCAAGATCACTTTTCCGAGTCATCTATTTTAGGACTCTCCGTAAGGACGTTGGTGTAGTATTCCGCTTATTT
 CCGGACAAGCAGATGATCTCTCCCTAATCGGATTGGGATCAAAAAAAGCACTGAAATGCGTGTCTTCAAAAAGGA
 ATTTGGGACAGATTATAAAAAATATAAGCAGCATCTTGATAGAGTGAATAATGATTTCACTTTTGTCTGTGAGGGGAT
 TAGCGCTTTAATTCCTACAGAAAAAGATGCTCCATTCCTATAGAACCTTCTCATTAGCAGGTGTTTTTTTAGTAT
 CATTTTACCCAGACAAGAATCCGATTCTAAAGATTACCGCTCATGCTGAGAAGATGTTACAGCCTCCTCAAGCGGA
 TTCCCTAACGGCTGGTTTTGTTGTGTGGAGCTCTTTCTGATCCTAAGAAATTTGACAGTCCCTTTCTATCTTTGAT
 TGAGAAGACTCACCAAGGGATTTTGGTGAGTAAAGACTTGAAAGACAATAAGGAAAAGAAAGCTAGCTTTAGAGGCTT
 CCTTTCTTTTCAATGAATATTTTCTTTTCCGGTTGGTGTTTGGGGAATCCGGAGTACAATCAGTATATCACAACCTGCT
 GTAGCTGAGAAATATAGGGATGTCTCTGTAAGAAATTTGATTTTATGATTTCTGGATACAGGGAAATGTGATTTACG
 TCTTGCTTTAGCAAGTAGTTATTCACAAGATTCGCTTGGGCTGCAGGGTTCAGAAAAGTTTTACGTGAAGAAGATA
 AAAAGACTAAGAAAAAGTCACGTGAAGAAGTCTCTGTTTGTATCGTGATATAGATCCAGGCTGTTGTTTAAAGGCC
 CTTCCCTAAGCGATTTGAATCCAAGTCTTCAGGTAGTCAAGGTAGTCTAAAGAGCAGTTAAGCTCTTTGTTGAAAGC
 TTTAGACCAGAAAATTCCTTCAGGGATTTTAGGATTGATTGCAAAAGCTTCTTCTGCAGATCTCAAGGCTGATTTTG
 CAGGTATGCTTGAAGTTATTAAGCAATTACAAGCTTTATTTCGATTCTTACCCACCTTTATGCGAAGACAATATTCTC
 TTGTGGTTAAGCGCTTCTTTAGAACAAAGTAGGCTTGCAAGAAATTTGAGAACCCTTTTACCTTCATCAGAAAAAAA
 ACTCTTAGAAAGAGTTCTCTCTACATTTTTATTAGGTTTGTATACTCGAGGAGTCTTTTCTGTAGGGCAAGTGAATC
 AGCTAGCTACTATTTGTAATACTCAGGACTCTACAGAATTCTGCCAGAGAGTAAGTGACCTTTTCGTTAATTAACGA
 GCTCTACCTGCATTATTTGGT

SEQ ID NO:162 - CT214 fragment protein sequence

RTDSLFPDSTRGVFQFLETQCDRAVARSRQSQFIGLVSAVAAAAALLLLVVALSVPGFVPAASIVVGVLFALSI
 VALTASFLVYIANAKLVAIRIKFLSSGLQDHFSESSILGTLRKGRGASIP LISGQADDPLPNRIGIKKSTEMRVLQK
 GIGTDYKYYKQHLDRVNDFTFVCEGISALIPTEKDAFPPIEPSHLAGVFLVSPDKNPILKITRHAEKMLQPPQG
 GPNGLVWLCGALSDPKKFAAPFLSLIEKTHQGILVSKDLKDNKERKLALAEASLLSLNIFSSGWCLGNPEYNQYITTA
 AVAEKYRQDVSVRNCIYDFLDTGNVISALALASSYSQDSAWAAGLQKVLREEDKTKKKSREEVSVCLYRDIIDPGCCLR
 ALPKRFESKSSGSQSPKEQLSSLLKALDQKIPSGILGLIAKASSADLKADFAGMLEVIKQLQALFDSYPLCEDNII
 LLWLSASLEQVGLQKLRFLPSSEKLLERVLSTFLLGLYTRGVFSVGQVNLATICNTQDSTEFQVRVSDLSLIR
 RALPALFG

SEQ ID NO:163 - CT721 nucleotide sequence

ATGGACGGGACAAAAATTCACGAAACACGCTCCTTCTCTTGGTTAAACAACCAACAAGCCATCCCTCCTTCCGAAAT
 GGTGAAGGAGGCTTTTCAACGTTACGCAGACGATTTTTCGTACAGCGCAAATACCTCCATTCTGACTTTACAAGCAG
 AAGCTGAAGCTTCTGCCCGCAAACCTCACAGGGTGTACAGGAGAAGGCTTTTACCTTTCATTTTATTCTTCATTACCCG
 AATGTACGGCCATTATCGTGGCCGCTCTTCTGAAAAACAAAATGCCTTCCAGGGGCGTAATCACCTTCTTGTTC
 TTTCTTGGCAGCAACAAATTTATCATTAATGCTCTCTGCGTGGCAAAACTTAGGGACAACCTATGATTGGGTAACCA
 GCAAAACGGCCGCTAAAAAGAAATCCGATCTAGCAGAAGCTCTTTCCCGGACCTTGCTGTTTTCCATATCTGCT
 GCGAATGGTATGACAGGATTTCTGGAAGCGATCCCTGAGCTTGTGCTGCTTATGTAAGAAGCGGGGTAATTTTCCA
 CATAGACCTGAGTGATATCTTAGGAAGATGCGCGTACCCGAGAAGCTCTATCAAGCAGATATCCTTACTTTTTCTT
 CACAGTCTCTTGGTGGGATTGGTCCCTCAGGAGCGATGTTTTATTTCTCCCGCTTAAACAAAATATTTTTCTTATGG

CTTCCTAGTAATCCACAAGTCCCTACCTGCCTGAGTTCTCTTGCAGCTTTTTCTCTTGCCTGTCAGGAACGTACAAC
CGCTTTCTCCTCTCTTGTGCTTTCTGCTATTTCTTCTCGAGCAGCTCTTAAACAGGCTCTTCCGCTATTCTCAAG
TCGAATTCCTTTTGGGAAGACAGTGGCCCTCGTCTCCCTAATGTCGCTGTCTTTGCTATTCTGGTATCCCTGCAGAG
TCCTTAGGATTTTTCTTTCCAGAAAAATTTTTGTAGGGTTAGGCTATGAACGCTTCCAGCCTCTATCGCAGAT
LPTNCAAAGTTCGGGCATCTCCCTTTCTTATGCCACAGCGCTTTACACGTATCTTTTACTGAACGTACTCCTACTA
CACACTTCTCTGCATTAGCAACCGCTTACAAGAAGGGATCTCTACCTACAACCACTGGTACTCAATCCTTATGA

SEQ ID NO:164 – CT721 protein sequence

MDGTKIHETRSFSLNNDQAIAPPSEMVKEAFQRYADVFSYSANTSILTLQAEAEASARKLTGCQEKAFTHFILHYP
NVTAIIVAALLENQNAFQGRNHLLVPSCEQRFIINALCRRQNLGTTYDQWVTSKNGRVKESDLAEALSPRTLLFSISA
ANGMTGFLEAIPELAALCKERGVIHFIDLSIDLGRCALPAELYQADILTFSSQSLGGIGPSGAMFISPALTKYFSLW
LPTNCAAAGTTCGGGCATCTCCCTTTCTTATGCCACAGCGCTTTACACGTATCTTTTACTGAACGTACTCCTACTA
CACACTTCTCTGCATTAGCAACCGCTTACAAGAAGGGATCTCTACCTACAACCACTGGTACTCAATCCTTATGA

SEQ ID NO:165 – CT721 fragment nucleotide sequence

GACGGGACAAAAATTCACGAAACACGCTCCTTCTCTTGGTTAAACAACCAACAAGCCATCCCTCCTTCCGAAATGGT
GAAGGAGGCTTTTCAACGTTACGCAGACGTATTTTCTGACAGCGCAAATACTCCATTCTGACTTTACAAGCAGAAG
CTGAAGCTTCTGCCCGCAAACCTCACAGGGTGTCCAGGAGAAGGCTTTTACCTTTCAATTTATTCTTATTACCCGAAT
GTCACGGCCATTATCGTGGCCGCTCTTCTGGAAAAACCAAATGCCTTCCAGGGGCGTAATCACCTTCTTGTTCCTTC
TTGCGAGCAACAATTTATCATTAAATGCTCTCTGCGTCCGCAAAACTTAGGGACAACCTATGATTGGGTAACCAGCA
AAAACGGCCGCGTAAAAGAAATCCGATCTAGCAGAAGCTTTTCCCGCGGACCTTGTGTTTTCCATATCTGCTGCG
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CCTAGTAATCCACAAGTCCCTACCTGCCTGAGTTCTCTTGCAGCTTTTTCTCTTGCCTGTGTCAGGAACGTACAACCGC
TTTCTCCTCTCTTGTGCTTTCTGCTATTTCTTCTCGAGCAGCTCTTAAACAGGCTCTTCCGCTATTCTCAAGTCCG
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ACTTCTCTGCATTAGCAACCGCTTACAAGAAGGGATCTCTACCTACAACCACTGGTACTCAATCCTTA

SEQ ID NO:166 – CT721 fragment protein sequence

DGTKIHETRSFSLNNDQAIAPPSEMVKEAFQRYADVFSYSANTSILTLQAEAEASARKLTGCQEKAFTHFILHYPN
VTAIIVAALLENQNAFQGRNHLLVPSCEQRFIINALCRRQNLGTTYDQWVTSKNGRVKESDLAEALSPRTLLFSISAA
ANGMTGFLEAIPELAALCKERGVIHFIDLSIDLGRCALPAELYQADILTFSSQSLGGIGPSGAMFISPALTKYFSLW
LPTNCAAAGTTCGGGCATCTCCCTTTCTTATGCCACAGCGCTTTACACGTATCTTTTACTGAACGTACTCCTACTACAC
ACTTCTCTGCATTAGCAACCGCTTACAAGAAGGGATCTCTACCTACAACCACTGGTACTCAATCCTTA

SEQ ID NO:167 – CT127 nucleotide sequence

ATGCCGCACCAAGTCTTATTGTCTCCTGTTTGGCATCTTTTATCGAATGCTGAAGGTATAGAGACGCAAGTACTGTT
TGGAGAAAGGATATGCAACCATAACCATCGACACTATGCCTATTCTCAACTAGTCTTTTCTTCTATATGGAAGCCAT
ACCTTGGCGACTCTCTACAGAATATTCCTCTATTCTCTTCCAACTGCAGCCTCCTAATGCTGTTGTCTGCTCTCAA
GAAGCTTTTTTAGATCTTGGCATATCCCTTACCTTTTGGCCGCTCCACATAGATAAACCAAAATCAAGTGTCT
CCTATCTCCTGCTAGCATAGCATTATTAATTTCCAATCCAGAAAGTAACTATGCAAAAAGCTTTCTGCTCTACCAAAG
AGATTCGTTTTTAAATTTCTTATTCTCTCCAAGAGATTAGTTTTCTTTCGAGAACAAATGATAGATACTCCGTAC
GTTTGGGGTGGCCGGTGCATTATAAACAGCTTCTCGTAATGGTGTAGATTGTTCCGGGTATATTCAACTACTTTA
CCAAGTACAGGAAGAAAATATCCCTCGCAATGCTAGAGATCAATACAGAGACTGTTCTCCAGTAAAAGATTTCTCGT
CTCTACTATAGGAGGACTTATCTTCTCAAGAAAGCAAGCAGCGGACAAAATCAACCATGATGATGAAAATCTCG
GAGCATGAATTCATTCTGCTGCGGAAAAAATAGGGAAAGTAGAAAAAGTAATCCTAGGAAAATAGGGCTTTCTTTAA
AGGGAATCTATTCTGCTCATTAGGTGAACCGCCTATAGAAGCTGTTTTTGGCGTTCCTAAAAATAGAAAAGCCTTCT
TTTGA

SEQ ID NO:168 – CT127 protein sequence

MPHQVLLSPVCDLLSNAEGIETQVLFGERICNHNHRHYAYSQLVFSSIWKPYPGDSLQNIPLFSSQLQPPNAVVCSD
EAFLDPWHIPLPFAAPLHIDNQNQVSLSPASIALLNNSRSNYAKAFKSTKEIRFLNSSFSRDLVSAEQLIDTPY
VWGGRCIHKQLPRNGVDCSGYIQLLYQVTGRNIPRNARDQYRDCSPVKDFSSLPIGGILFLKKASTGQINHVMMKIS
EHFIIHAAEKIGKVEKVLGNRAFFKGNLFCSLGEPPIEAVFGVPKNRKAFF

SEQ ID NO:169 – CT127 fragment nucleotide sequence

CCGCACCAAGTCTTATTGTCTCCTGTTTGGCATCTTTTATCGAATGCTGAAGGTATAGAGACGCAAGTACTGTTTGG
AGAAAGGATATGCAACCATAACCATCGACACTATGCCTATTCTCAACTAGTCTTTTCTTCTATATGGAAGCCATACC
CTGGCGACTCTCTACAGAATATTCCTCTATTCTCTTCCAACTGCAGCCTCCTAATGCTGTTGTCTGCTCTCAAGAA
GCTTTTTTATGATCCTTGGCATATCCCTTACCTTTTGGCCGCTCCGCTCCACATAGATAAACCAAAATCAAGTGTCCCT
ATCTCCTGCTAGCATAGCATTATTAATTTCCAATCCAGAAAGTAACTATGCAAAAAGCTTTCTGCTCTACCAAAGAGA
TTGTTTTTTTTAAATTTCTTATTCTCTCAAGAGATTTAGTTTTCTTTCGAGAACAAATGATAGATACTCCGTACGTT
TGGGGTGGCCGGTGCATTATAAACAGCTTCTCGTAATGGTGTAGATTGTTCCGGGTATATTCAACTACTTTACCA

AGTCACAGGAAGAAATATCCCTCGCAATGCTAGAGATCAATACAGAGACTGTTCTCCAGTAAAAGATTTCTCGTCTC
TACCTATAGGAGGACTTATCTTCTCAAGAAAGCAAGCACGGGACAAATCAACCATGTTATGATGAAAATCTCGGAG
CATGAATTCATTCATGCTGCGGAAAAAATAGGGAAAAGTAGAAAAAGTAATCCTAGGAAAATAGGGCTTTCTTTAAAGG
GAATCTATTCTGCTCATTAGGTGAACCGCCTATAGAAGCTGTTTTTGGCGTTCCTAAAAATAGAAAAGCCTTCTTT

SEQ ID NO:170 – CT127 fragment protein sequence

PHQVLLSPVCDLLSNAEGIETQVLFGERICNHNHRHYAYSQLVFSSIWKPYPGDSLQNIPLFSSQLQPPNAVVCSE
AFLDPWHIPLPFAAPLHIDNQNQVSLSPASIALLSNSRSNYAKAFCSTKEIRFLNSSFSRDLVSFAEQIIDTPYV
WGGRCIHKQLPRNGVDCSGYIQLLYQVTGRNIPRNARDQYRDCSPVKDFSSLPIGGLIFLKKASTGQINHVMMKISE
HEFIHAAEKIGKVEKVLGNRAFFKGNLFCSLGEPPIEAVFGVPKNRKAFF

CLAIMS

1. A protein comprising the amino acid sequence of any one of SEQ ID NO: 2, SEQ ID NO:4, SEQ ID NO:6, SEQ ID NO:8, SEQ ID NO:10, SEQ ID NO:12, SEQ ID NO:14, SEQ ID NO:16, SEQ ID NO: 18, SEQ ID NO:40, SEQ ID NO:42, SEQ ID NO:44, SEQ ID NO:46, SEQ ID NO: 48, SEQ ID NO:50, SEQ ID NO:52, SEQ ID NO:54, SEQ ID NO:136 or SEQ ID NO:140 for use in therapy or diagnosis.
2. A protein having 50% or greater sequence identity to a protein according to claim 1.
3. A protein comprising a fragment of the amino acid sequence of claim 1 or claim 2.
4. A protein according to claim 3, wherein the fragment comprises at least 8 consecutive amino acids of the amino acid sequence of claim 1 or claim 2.
5. An antibody which binds to a protein according to any one of claims 1 to 4 for use in therapy or diagnosis.
6. A nucleic acid molecule which encodes a protein or antibody according to any one of claims 1 to 5 for use in therapy or diagnosis.
7. A nucleic acid molecule according to claim 6, comprising the amino acid sequence of any one of SEQ ID NO: 1, SEQ ID NO:3, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:13, SEQ ID NO:15, SEQ ID NO:17, SEQ ID NO:39, SEQ ID NO:41, SEQ ID NO:43, SEQ ID NO:45, SEQ ID NO:47, SEQ ID NO:49, SEQ ID NO:51, SEQ ID NO:53, SEQ ID NO:135 or SEQ ID NO:139.
8. A nucleic acid molecule comprising of a fragment of the nucleotide sequence of any one of SEQ ID NO: 1, SEQ ID NO:3, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:13, SEQ ID NO:15, SEQ ID NO:17, SEQ ID NO:39, SEQ ID NO:41, SEQ ID NO:43 or SEQ ID NO:45, SEQ ID NO:47, SEQ ID NO:49, SEQ ID NO:51, SEQ ID NO:53, SEQ ID NO:135 or SEQ ID NO:139.
9. A nucleic acid molecule comprising a nucleotide sequence complementary to a nucleic acid molecule according to any one of claims 6 to 8.
10. A nucleic acid molecule comprising a nucleotide sequences having 50% or greater sequence identity to a nucleic acid molecule according to any one of claims 6 to 9.
11. A nucleic acid molecule which can hybridise to a nucleic acid molecule according to any one of claims 6 to 10 under high stringency conditions.
12. A vector comprising a nucleic acid according to any one of claims 6 to 11 for use in therapy or diagnosis.
13. A host cell comprising a nucleic acid or vector according to any one of claims 6 to 12 for use in therapy or diagnosis.

14. A composition comprising a protein, antibody, nucleic acid molecule, vector or host cell according to any preceding claim for use in therapy or diagnosis.
15. A protein, antibody, nucleic acid, vector, host cell or composition according to any preceding claim for use as a vaccine composition.
16. A protein, antibody, nucleic acid, vector, host cell or composition according to any preceding claim for use as a pharmaceutical.
17. A protein, antibody, nucleic acid, vector, host cell or composition according to any preceding claim, for use in the treatment, prevention or diagnosis of Chlamydia.
18. A protein, antibody, nucleic acid, vector, host cell or composition according claim 17, for use in the treatment, prevention or diagnosis of *Chlamydia trachomatis*.
19. A protein, antibody, nucleic acid, vector, host cell or composition according to any preceding claim, for raising an immune response in a mammal.
20. A protein, antibody, nucleic acid, vector, host cell or composition according to any preceding claim, for eliciting antibodies that are capable of neutralising Chlamydia infection.
21. A protein or nucleic acid according to any one of claims 1 to 19, wherein the immune response is a CD4+ Th1 cell-mediated response.
22. The use of a protein, antibody, nucleic acid, vector or host cell according to any one of claims 1 to 13 in the manufacture of a medicament for the treatment or prevention of infection due to *Chlamydia* bacteria, particularly *Chlamydia trachomatis*.
23. A method of treating, preventing or diagnosing Chlamydia in a patient, comprising administering a therapeutically effective amount of a protein, antibody, nucleic acid, vector, host cell or composition according to any preceding claim.
24. A protein, antibody, nucleic acid, vector, host cell or composition according to any preceding claim, for use as a medicament in combination with one or more additional Chlamydia antigens or their encoding nucleic acids.
25. A protein, antibody, nucleic acid, vector, host cell or composition according to claim 24, wherein the combination comprises CT279+CT601, CT372+CT443, CT733+CT153, CT456+CT381, CT279+CT153+CT733+CT601, CT279+CT601+CT372+CT443, CT823+CT733+CT043+CT456, CT387+CT812+CT869, CT387+CT812C+CT869 (or variants thereof).
26. A protein, antibody, nucleic acid, vector, host cell or composition according to claim 24 or claim 25, wherein the protein, antibody, nucleic acid, vector, host cell or composition and the one or more additional Chlamydia antigens or their encoding nucleic acids are a combined preparation for simultaneous, separate or sequential administration.
27. A method for diagnosing Chlamydia infection, comprising:

- (a) raising an antibody against a protein according to any one of claims 1 to 4;
- (b) contacting the antibody of step (a) with a biological sample suspected of being infected with Chlamydia under conditions suitable for the formation of antibody-antigen complexes; and
- (c) detecting said complexes, wherein detection of said complex is indicative of Chlamydia infection.

FIGURE 1

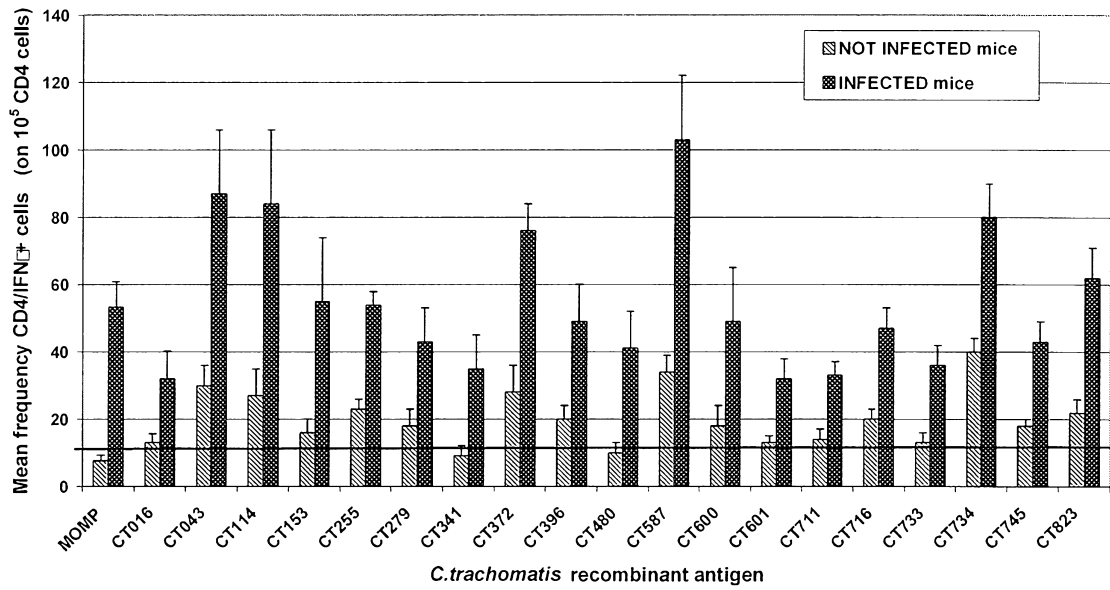


FIGURE 2

FIGURE 2A

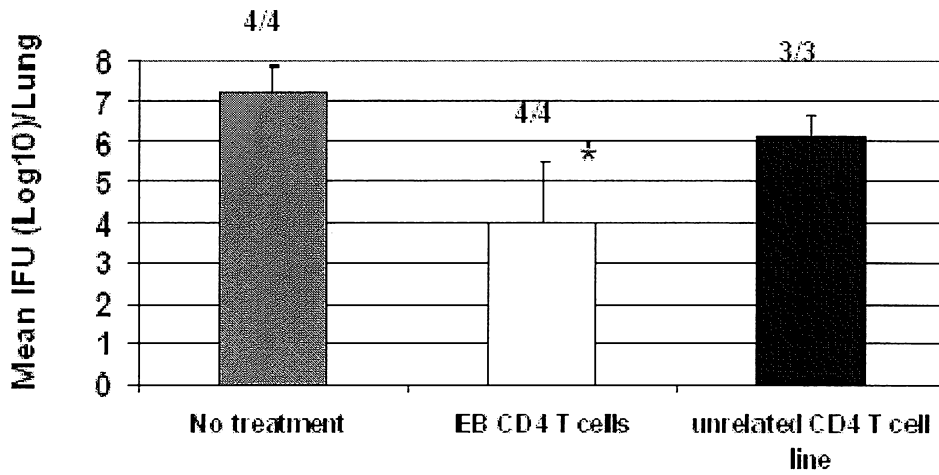


FIGURE 2B

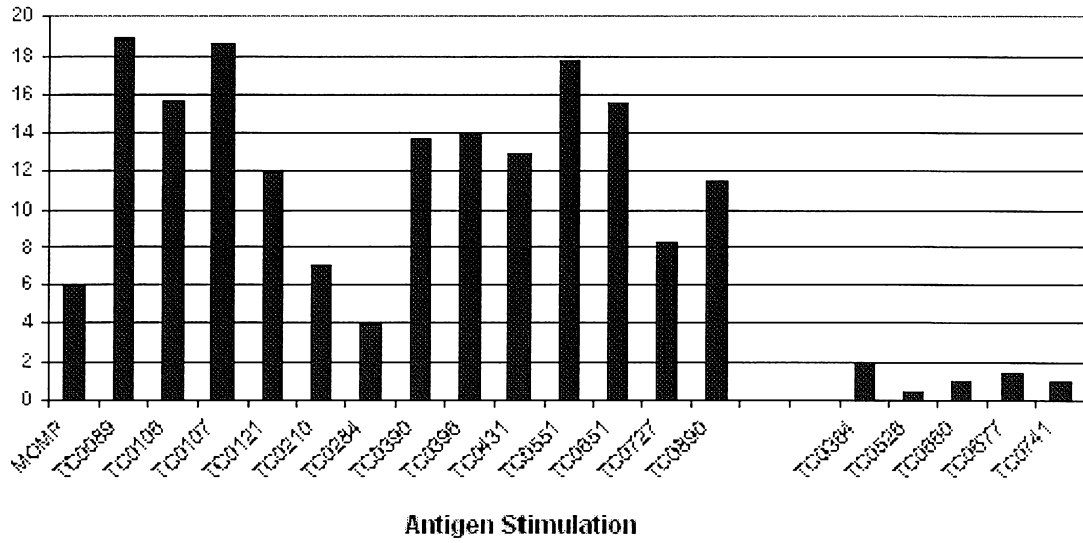
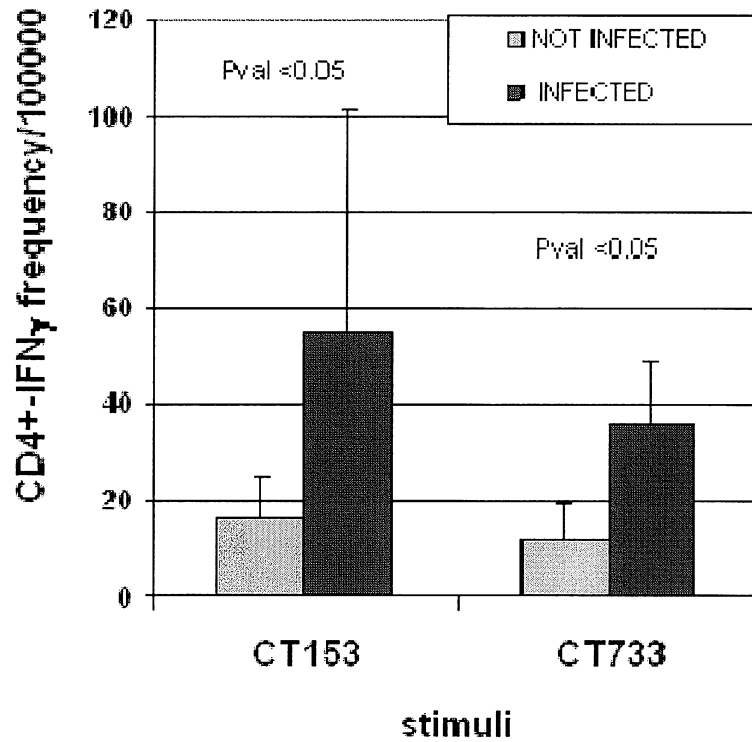


FIGURE 3



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FIGURE 4

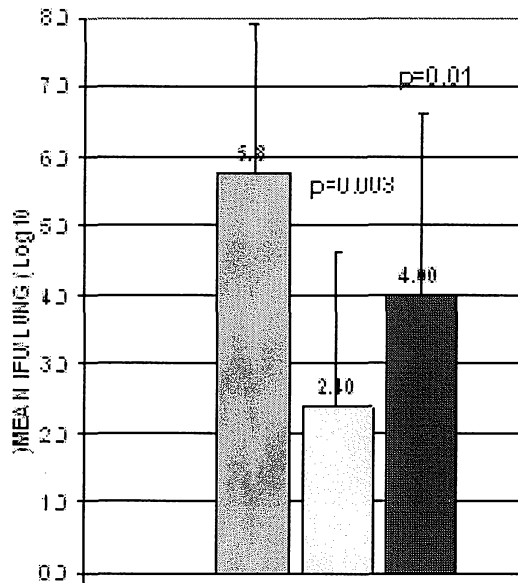


FIGURE 5

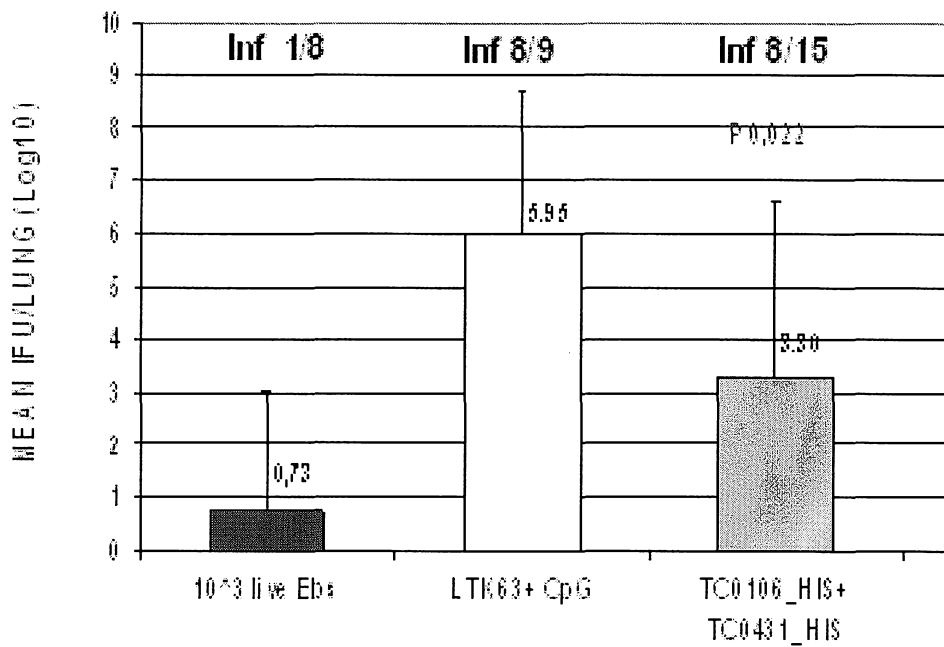


FIGURE 6

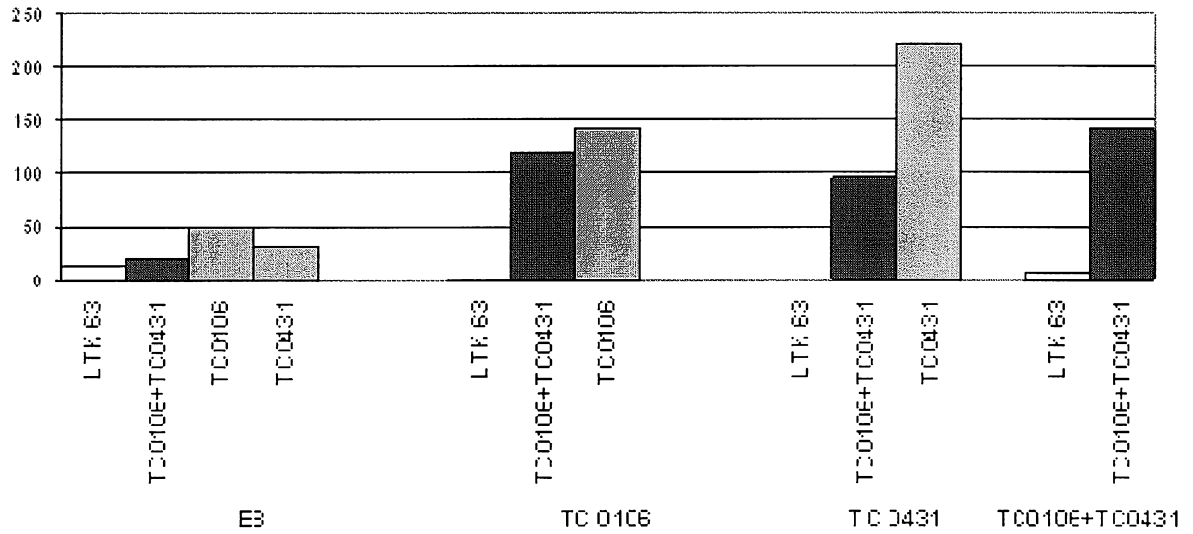
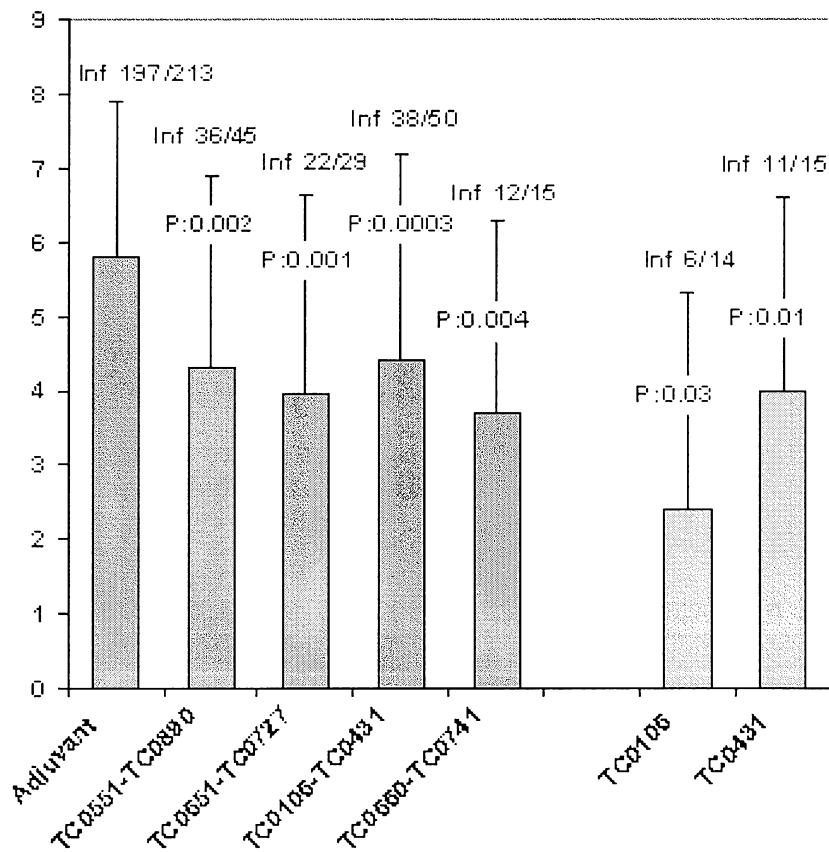


FIGURE 7



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FIGURE 8

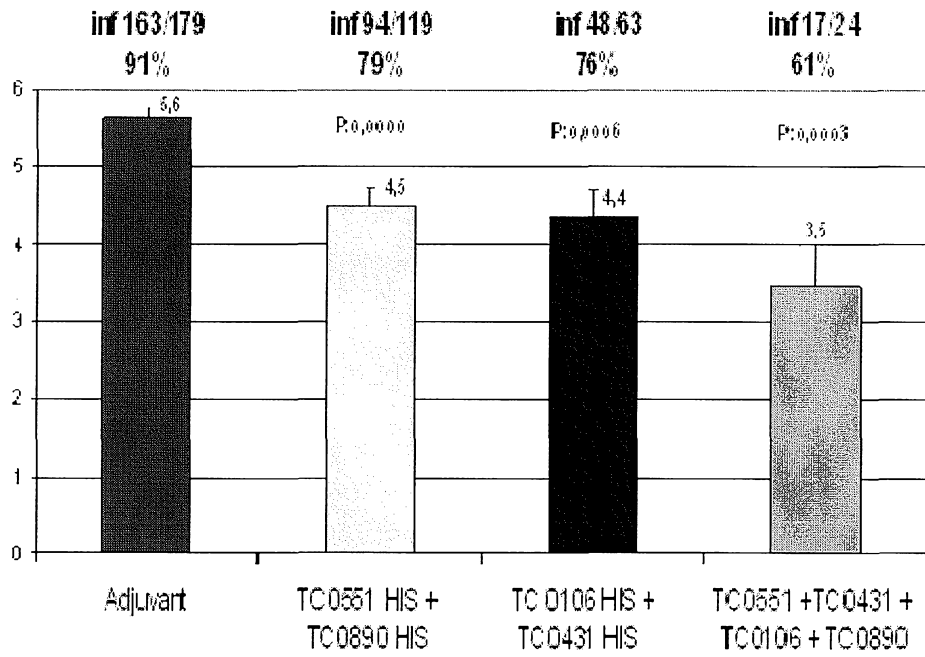


FIGURE 9

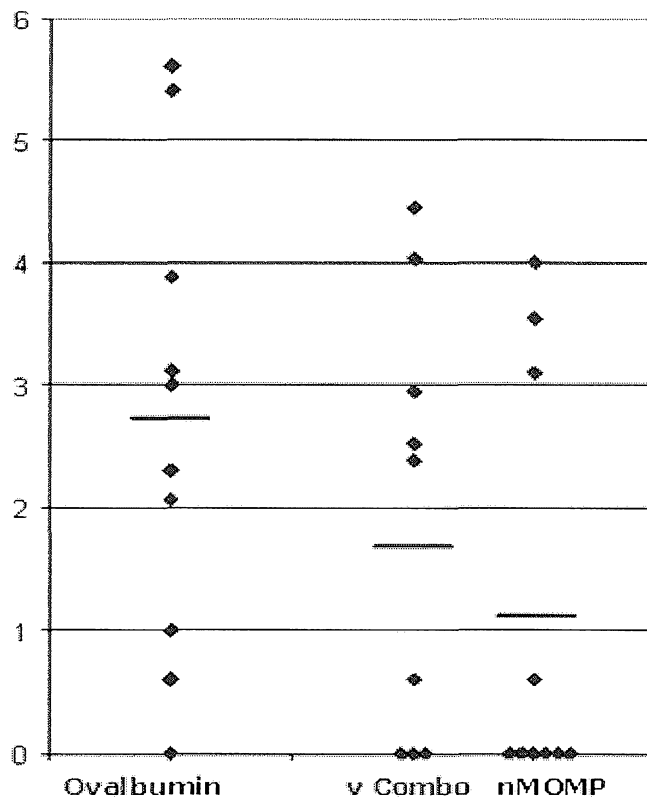


FIGURE 10

FIGURE 10A

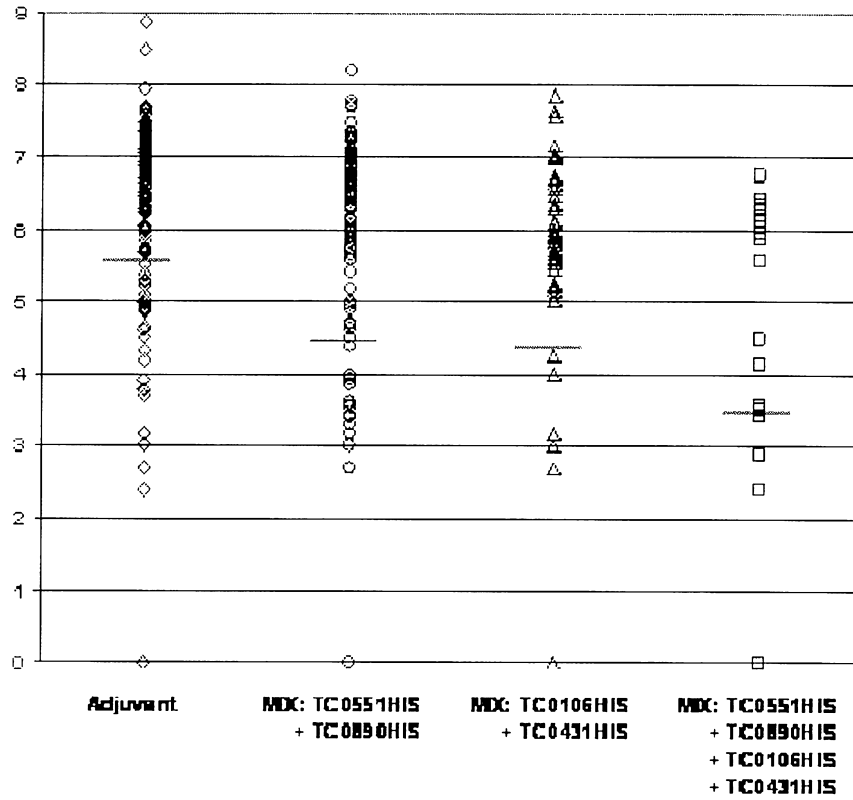


FIGURE 10B

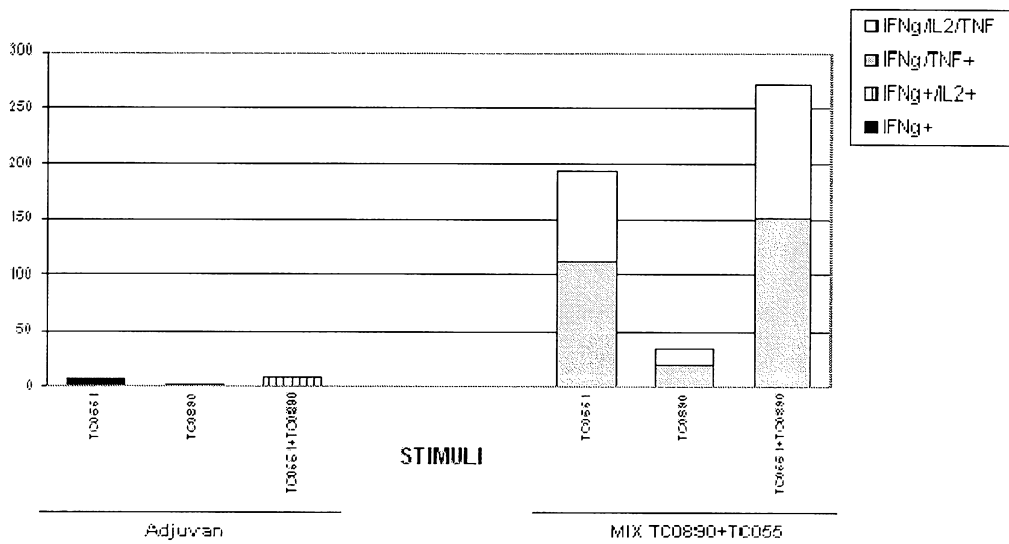
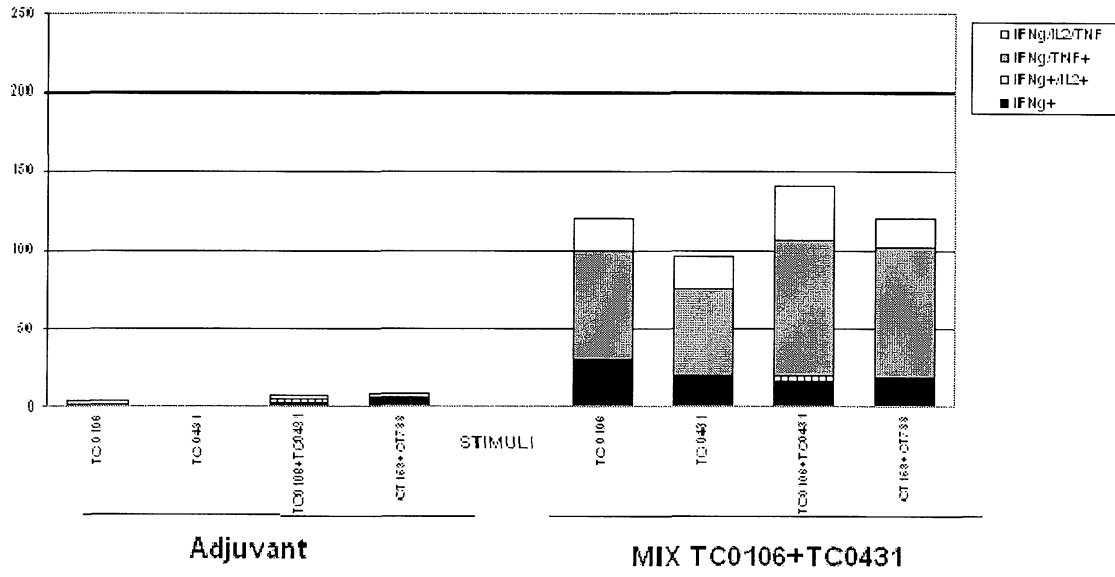


FIGURE 10C



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FIGURE 11

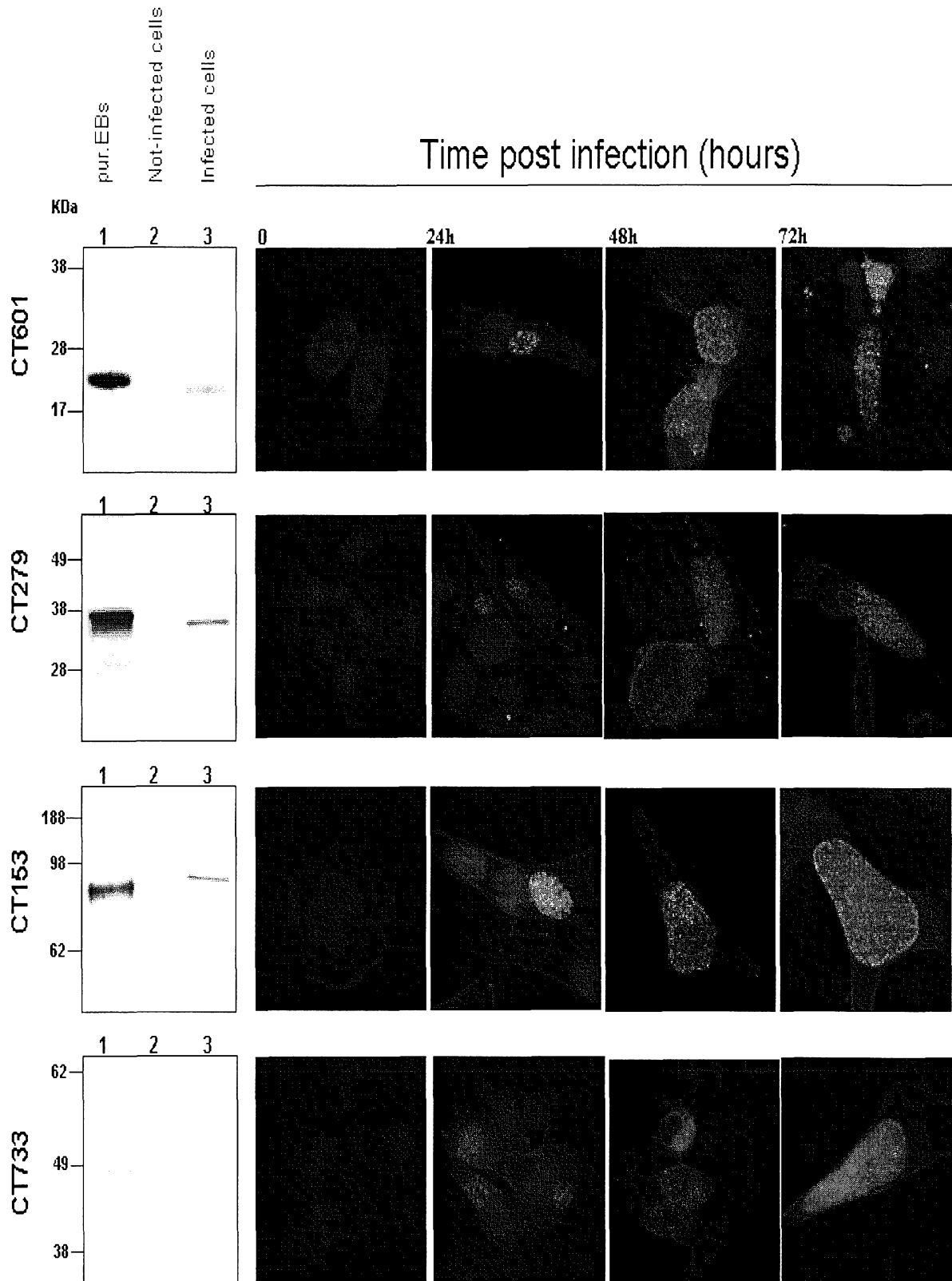


FIGURE 12

FIGURE 12A

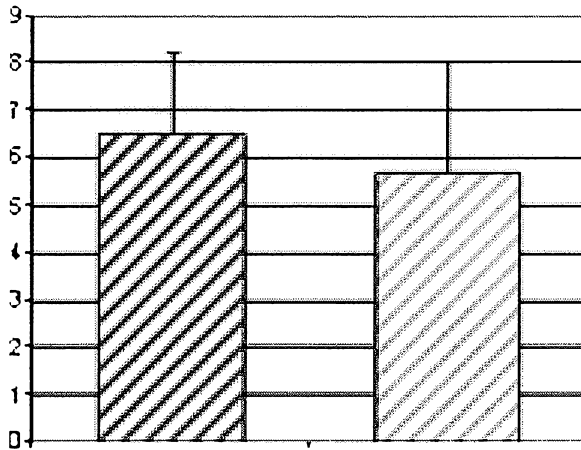


FIGURE 12B

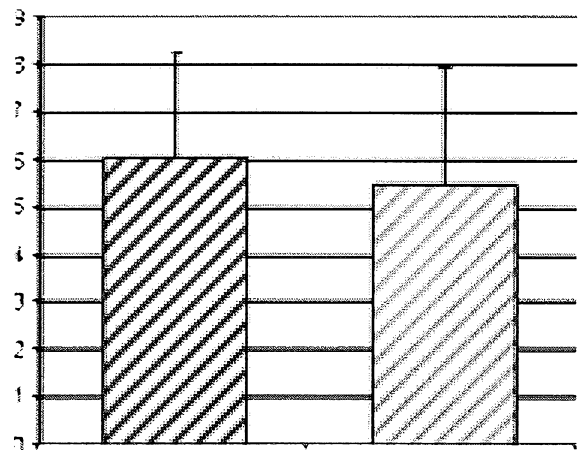


FIGURE 12C

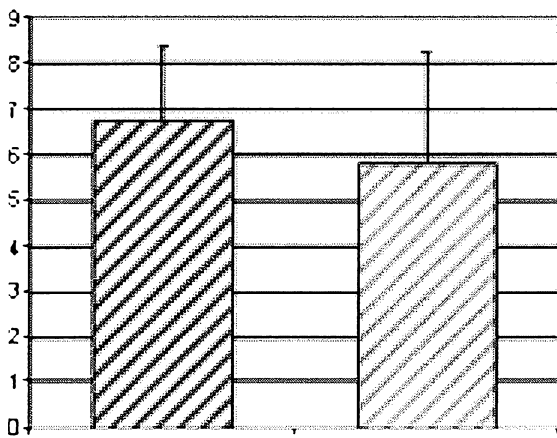


FIGURE 12D

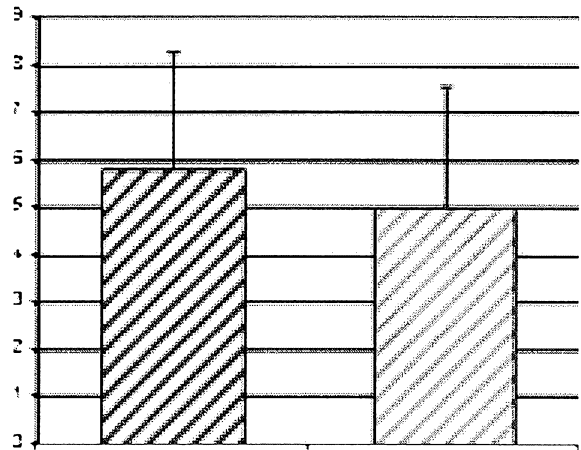
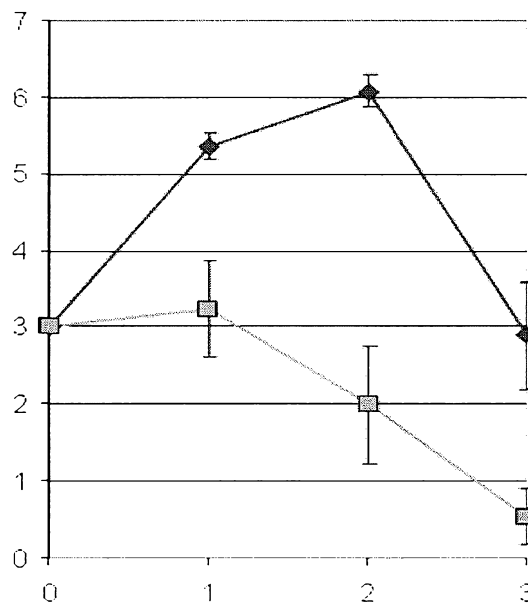


FIGURE 12E



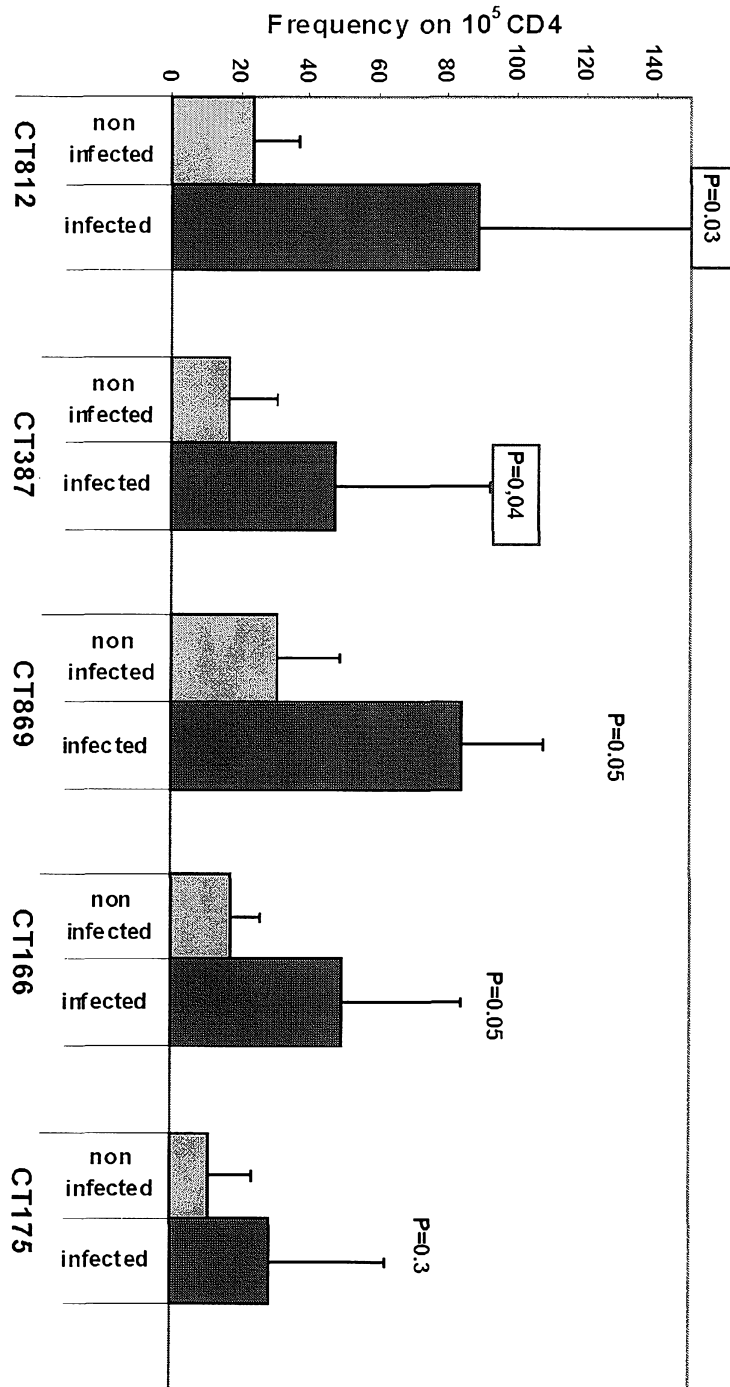
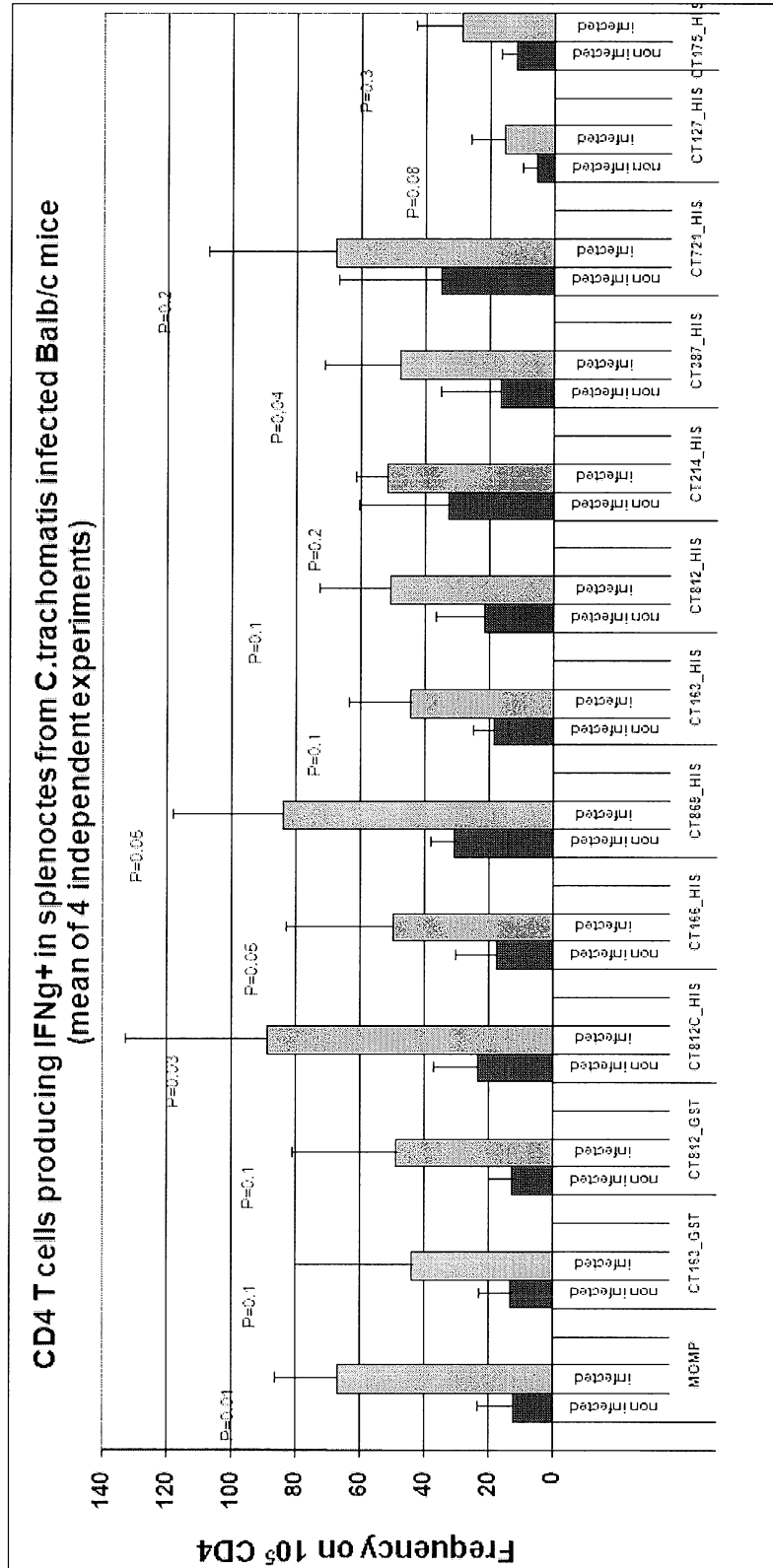


FIGURE 13A

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FIGURE 13B



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FIGURE 14

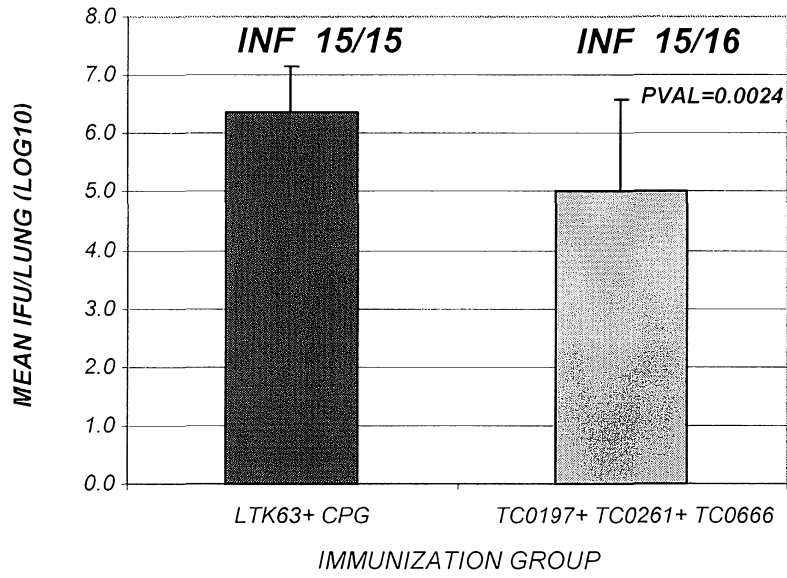


FIGURE 15

