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(54) **BIOINFORMATIC METHOD FOR IDENTIFYING SURFACE-ANCHORED PROTEINS FROM GRAM-POSITIVE BACTERIA AND PROTEINS OBTAINED THEREBY**

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(51) **Int. Cl.**
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(52) **U.S. Cl.** **424/190.1**; 424/185.1; 424/234.1
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See application file for complete search history.

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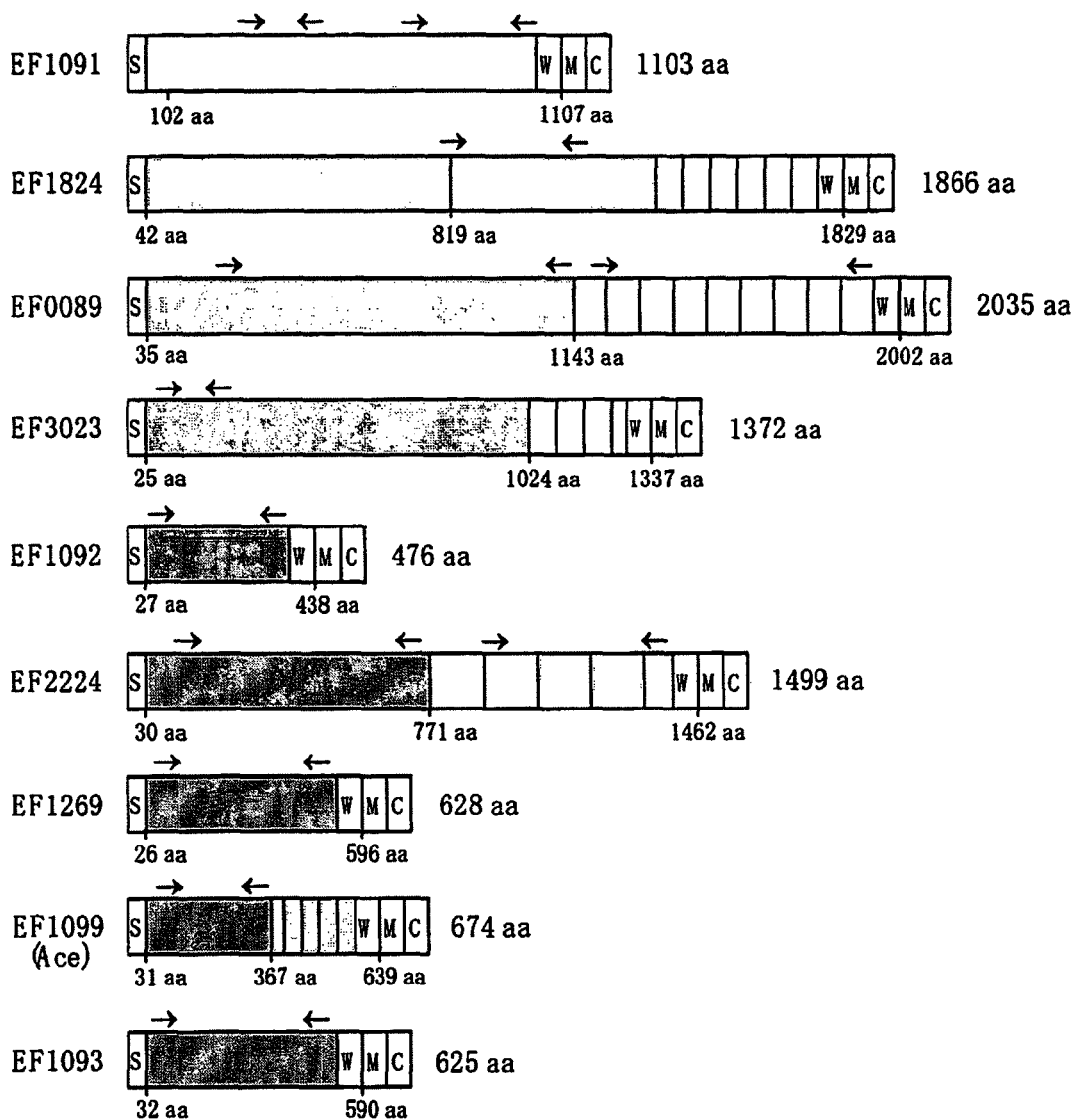
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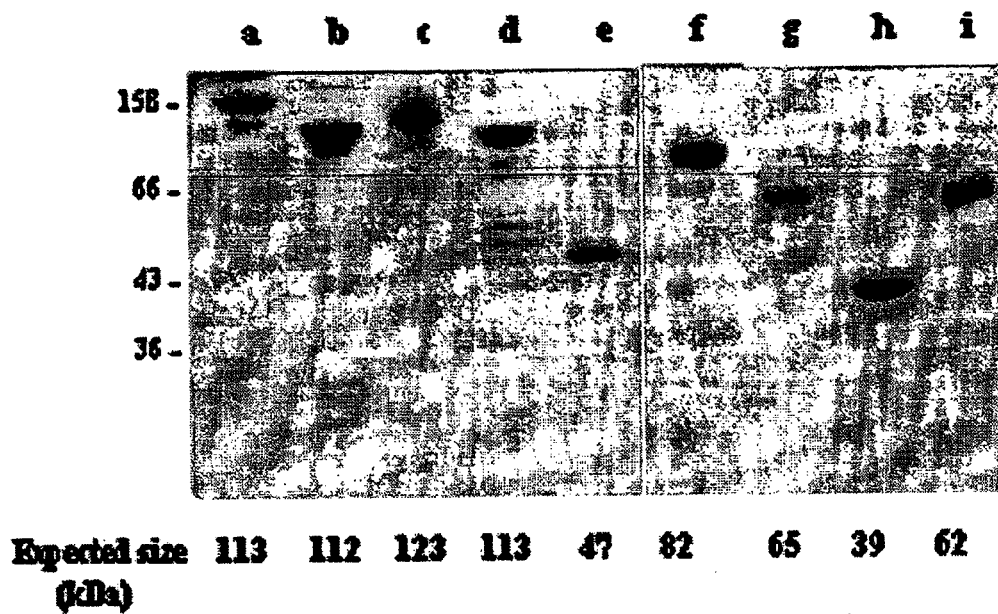
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(57) **ABSTRACT**
A bioinformatic method for identifying and isolating proteins and peptides with MSCRAMM®-like characteristics from Gram positive bacteria, such as *Enterococcus*, *Staphylococcus*, *Streptococcus* and *Bacillus* bacteria, and proteins and peptides obtained thereby are provided which can be utilized in methods to prevent and treat infections caused by Gram-positive bacteria. The method involves identifying from sequence information those proteins with a putative C-terminal LPXTG (SEQ ID NO:1) cell wall sorting signal and other structural similarities to MSCRAMM® proteins having the LPXTG-anchored cell wall proteins. The MSCRAMM® proteins and immunogenic regions therein that are identified and isolated using the present invention may be useful in the diagnosis, treatment or prevention of Gram positive bacterial infections.

4 Claims, 2 Drawing Sheets

Figure. 1





Coomassie-stained SDS-PAGE of the *E. coli*-expressed and purified A domains of *E. faecalis* LPxTG proteins. a, EF1091; b, EF1824; c, EF0089; d, EF3023; e, EF1092; f, EF2224; g, EF1269; h, Ace; i, EF1093.

FIG. 2

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**BIOINFORMATIC METHOD FOR
IDENTIFYING SURFACE-ANCHORED
PROTEINS FROM GRAM-POSITIVE
BACTERIA AND PROTEINS OBTAINED
THEREBY**

CROSS-REFERENCE TO RELATED
APPLICATIONS

The present application is a divisional application of U.S. patent application Ser. No. 10/661,809, filed Sep. 15, 2003 now U.S. Pat. No. 7,615,616, which claims the benefit of U.S. provisional application Ser. No. 60/410,303, filed Sep. 13, 2002.

STATEMENT REGARDING FEDERALLY
SPONSORED RESEARCH OR DEVELOPMENT

This Invention was made with Government support under Contracts 7R01-AR44415-04 and 2R01-AI20624-17 awarded by NIH. The government has certain rights in this invention.

FIELD OF THE INVENTION

The present invention relates to the fields of microbiology, molecular biology, and immunology and more particularly relates to surface-anchored proteins known as MSCRAMM®s, and to a bioinformatic method of identifying putative MSCRAMM® proteins, i.e., proteins that can bind to extracellular matrix molecules, from Gram positive bacteria having a recognizable cell wall sorting signal and the genes encoding those proteins through detecting structural features from potential proteins including immunoglobulin (Ig)-like fold regions. In addition, the invention relates to antibodies which recognize such proteins, including polyclonal and monoclonal antibodies as well as host cells transformed with nucleic acids encoding monoclonal antibodies, and the use of such antibodies in the diagnosis, treatment or prevention of Gram positive bacterial infections in humans and animals.

BACKGROUND OF THE INVENTION

There are numerous Gram positive bacteria which have been of interest in the fields of medicine and epidemiology because of their potential to cause a myriad of infectious diseases in humans and animals. One such Gram positive bacterium, *Enterococcus faecalis*, belongs to the commensal flora in mammalian intestines. It has also long been known as a major causative agent of bacterial endocarditis (Murray, 1990). During the last decades, *E. faecalis* has increasingly emerged as an opportunistic nosocomial pathogen, typically causing infections in hospitalized patients receiving antibiotic therapy. Clinical strains of this bacterium frequently harbor a multitude of acquired and intrinsically evolved resistance mechanisms toward the most commonly used antibiotics, which has complicated the treatment of enterococcal infections (Murray, 1990, 1999) (Tailor, 1993) (Huycke, 1998). Many of the antibiotic resistance genes are located in mobile genetic elements, e.g., small plasmids and transposons (Paulsen, 2003) This has raised fears for genetic transfer of resistance determinants from this organism to other bacterial species, e.g., the recently documented transfer of vancomycin resistance to *Staphylococcus aureus* (CDC, 2002). Still other Gram positive bacteria are known which commonly cause infections which are hard to control, includ-

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ing other bacteria from the *Enterococcus* genus, including *Enterococcus faecium*, as well as bacteria from species *Streptococcus*, such as *Streptococcus mutans* and *pneumoniae*, *Staphylococcus*, such as *Staphylococcus aureus* and *epidermidis*, and *Bacillus*, such as *Bacillus anthracis*.

The ability to adhere to mammalian tissue is a critical step in the colonization and onset of microbial infections. However, in light of the many unknown factors regarding microbial adherence, it remains a challenge to study and utilize information obtained regarding relatively little known adhesion mechanisms of Gram positive bacteria so as to provide a means for developing alternative antibacterial therapies. One such inroad into developing such therapies is the presence of the human extracellular matrix underneath epithelial and endothelial cells which is a complex, dynamic and multifunctional structure consisting mainly of collagens and other glycoproteins. As one of the outermost layers to external environment, it is a major adhesion target and entry point for pathogenic bacteria (Foster and Hook, 1998) (Westerlund and Korhonen, 1993). Numerous bacterial adhesins that specifically bind to ECM components have been characterized at the molecular level. A group of related cell surface proteins from Gram-positive bacteria, collectively designated MSCRAMM® proteins (microbial surface components recognizing adhesive matrix molecules) bind to major components of the ECM, such as collagens, fibronectin, laminin, fibrinogen, keratin, vitronectin and bone sialoprotein (Patti, 1994) (Foster and Hook, 1998) (Tung, 2000) (O'Brien, 2002). MSCRAMM® proteins are mosaic proteins that typically consist of an N-terminal signal sequence for Sec-dependent transport across the cytoplasmic membrane, followed by an N-terminal A domain which exhibits the binding activity in most cases and repetitive B domains that confer fibronectin binding in a group of fibronectin binding MSCRAMM® protein (Joh et al., 1994). Covalent attachment to the bacterial cell wall is mediated through a C-terminally located LPxTG motif preceded by a cell wall spanning domain and followed by a hydrophobic trans-membrane region and, finally, a cytosolic tail composed of a short sequence of positively charged amino acid residues (Schneewind et al., 1995) (Mazmanian et al., 2001).

In any event, it remains a distinct problem in the field of infectious diseases to develop new means of countering a wide range of bacterial infections in an efficient and effective manner without the potential of increasing the development of antibiotic-resistant bacterial strains. Moreover, in light of the potential problems caused by bacterial strains and antibiotic-resistant strains in general, particularly in hospitalized patients, it is increasingly important to develop methods to counteract such infections without utilizing antibiotics or increasing the likelihood that antibiotic-resistant strains will develop. It is thus highly desirable to develop new means for identifying, treating and preventing infectious diseases caused by Gram positive bacteria, and to develop means for identifying and isolating new MSCRAMM® proteins from such bacteria which will allow the generation of antibodies thereto which will lead to new methods for treating and preventing the spread of infections from Gram-positive bacteria.

SUMMARY OF THE INVENTION

Accordingly, it is an object of the present invention to provide a bioinformatic method of identifying and isolating MSCRAMM® proteins from Gram-positive bacteria which can be utilized in methods of treating or preventing infectious diseases arising from Gram-positive bacteria.

It is another object of the present invention to identify and isolate proteins obtained using the bioinformatic method of the present invention, and to identify therein effective antigenic domains such as the A domain, and to utilize these antigenic domains in methods of treating or preventing infectious diseases arising from Gram-positive bacteria.

It is further an object of the present invention to utilize the proteins and antigenic domains isolated and identified using the bioinformatic method of the present invention to generate antibodies which can recognize these proteins and antigenic regions which can thus be useful in diagnosing, treating or preventing diseases and infections caused by Gram positive bacteria

It is still further an object of the present invention to provide vaccines, kits and other therapeutic methods which utilize the proteins and antigenic domains identified and isolated using the bioinformatic method of the present invention which can be used as an alternative to conventional antibiotic therapy and can thus provide safe and effective modes of treating or preventing infections caused by Gram-positive bacteria.

These and other objects are provided by virtue of the present invention which utilizes a bioinformatic approach to identify proteins with MSCRAMM®-like characteristics among Gram positive bacteria, such as bacteria from *Enterococcus*, *Staphylococcus*, *Streptococcus* and *Bacillus*, among many others, the obtaining of said proteins and peptides therein, which can then be utilized in methods to prevent and treat infections caused by Gram-positive bacteria. In particular, the method involves looking for proteins with a putative C-terminal LPXTG (SEQ ID NO:1) cell wall sorting signal and structural similarities to MSCRAMM® proteins having the LPXTG-anchored cell wall proteins. In particular, the present invention provides a method for identifying and isolating MSCRAMM® proteins, i.e., proteins that can bind to extracellular matrix molecules, such as by locating regions that adopt an immunoglobulin-like fold, and includes the recombinant production of these proteins from nucleic acids identified in the present process which code for those proteins. These Ig fold-containing regions consist of several consecutive and overlapping matches to solved crystal structures (~150-500 aa) of the immunoglobulin superfamily (IgSF), which consist of one to four domains of equal size and Ig-type fold. The homologous Ig-fold regions are indicative of a "beads-in-a-string" arrangement of consecutive modules such like the ones found in fibronectin and other IgSF proteins (Leahy, 1996) (Sharma, 1999) (Hamburger, 1999) (Luo, 2000). For example, a tandem repeat of Ig folded subdomains (N2 and N3) is found in the crystal structure of the fibrinogen-binding domain of ClfA. The full-length A domains of ClfA and the similarly structured ClfB consist of an additional N-terminal subdomain, N1 (Deivanayagam, 2002) (Perkins, 2001). Based on sequence and secondary structure similarities, an analogous subdomain organization is also expected in other MSCRAMM® proteins including FnbpA, FnbpB, Ace and the Sdr proteins. The solved crystal structure of CNA minimum collagen-binding domain is made of a single Ig-type subdomain (N2) (Symersky, 1997) and the C-terminal repeat domains B1 and B2 each consist of a tandem repeat of Ig-folded subdomains (Deivanayagam, 2000). A similar modular structure is expected in the B3 and B4 repeats.

In accordance with the invention, novel MSCRAMM®-like protein surface-anchored proteins which can bind to major extracellular matrix proteins are obtained from Gram-positive bacteria such as those from the genera *Enterococcus*, *Streptococcus*, *Staphylococcus* and *Bacillus*, and such proteins are characterized in that they are (i) structurally homolo-

gous to the solved Ig-folded crystal structures of ClfA and CNA as well as to the predicted tertiary structures of other MSCRAMM® proteins, (ii) share a similar β -sheet rich secondary structure as is found in Ig-folded proteins and (iii) have a similar organization with a secretion signal, a non-repeated domain followed by repeats as well as a C-terminal cell wall anchor domain. Moreover, the binding of proteins identified by the present method has confirmed that they target and bind to various extracellular matrix (ECM) molecules including proteins and other components. For example, three of the isolated proteins bind to major ECM proteins; two to fibrinogen and at least one to collagen and laminin. The proteins of the present invention have also been shown to be present in most isolates and are expressed in vivo during infection.

Thus, in accordance with the present invention, a method is provided for identifying and isolating a module structure of multiple Ig-folded units which appears to be a general characteristic in the MSCRAMM® protein family. The length of the N-subdomains of MSCRAMM® proteins is typically ~150 aa, and the proteins identified by the present invention including those set forth below may accommodate more than three Ig-folded subdomains in their A domains.

These embodiments and other alternatives and modifications within the spirit and scope of the disclosed invention will become readily apparent to those skilled in the art from reading the present specification and/or the references cited herein, all of which are incorporated by reference.

BRIEF DESCRIPTION OF THE DRAWING FIGURES

FIG. 1 is a schematic representation of MSCRAMM® proteins identified in accordance with the present invention illustrating the different regions of the proteins and their immunoglobulin-like fold regions

FIG. 2 illustrates a Coomassie stained SDS-PAGE of the *E. coli*-expressed and purified A domains of the LPXTG-containing proteins of the present invention.

DETAILED DESCRIPTION OF THE PREFERRED EMBODIMENTS

In accordance with the present invention, there is provided a bioinformatic method for identifying and isolating proteins from Gram-positive bacteria, for example bacteria from genera such as *Enterococcus*, *Staphylococcus*, *Streptococcus* and *Bacillus*, in particular proteins which have MSCRAMM®-like characteristics, and utilizing the identified and isolated proteins to generate antibodies and diagnose, treat or prevent infections caused by Gram-positive bacteria. In general, the method involves looking for proteins with a putative C-terminal LPXTG (SEQ ID NO:1) cell wall sorting signal and/or other structural similarities to MSCRAMM® proteins (Microbial Surface Components Recognizing Adhesive Matrix Molecules) having LPXTG-containing cell wall-anchored proteins. In the preferred embodiment, the present invention provides a method for identifying and isolating MSCRAMM® proteins, i.e., surface proteins that bind to extracellular matrix molecules, such as proteins, carbohydrates and other components, of host cells, wherein those located proteins contain regions that adopt an immunoglobulin-like fold. These Ig fold-containing regions consist of several consecutive and overlapping matches to solved crystal structures (~150-500 aa) of the immunoglobulin superfamily (IgSF), which consist of one to four domains of equal size and Ig-type fold. The homologous Ig-fold regions are indicative

of a “beads-in-a-string” arrangement of consecutive modules such like the ones found in fibronectin and other IgSF proteins (Leahy, 1996) (Sharma, 1999) (Hamburger, 1999) (Luo, 2000). For example, a tandem repeat of Ig folded subdomains (N2 and N3) is found in the crystal structure of the fibrinogen-binding domain of ClfA. The full-length A domains of ClfA and the similarly structured ClfB consist of an additional N-terminal subdomain, N1 (Deivanayagam, 2002) (Perkins, 2001). Based on sequence and secondary structure similarities, an analogous subdomain organization is also expected in other MSCRAMM® proteins including FnbpA, FnbpB, Ace and the Sdr proteins. The solved crystal structure of CNA minimum collagen-binding domain is made of a single Ig-type subdomain (N2) (Symersky, 1997) and the C-terminal repeat domains B1 and B2 each consist of a tandem repeat of Ig-folded subdomains (Deivanayagam, 2000). A similar modular structure is expected in the B3 and B4 repeats.

In accordance with the invention novel MSCRAMM®-like protein surface-anchored proteins are obtained from Gram-positive bacteria such as those from the genera *Enterococcus*, *Streptococcus*, *Staphylococcus* and *Bacillus*, and such proteins are characterized in that they are (i) structurally homologous to the solved Ig-folded crystal structures of ClfA and CNA as well as to the predicted tertiary structures of other MSCRAMM® proteins, (ii) share a similar β -sheet rich secondary structure as is found in Ig-folded proteins and (iii) have a similar organization with a secretion signal, a non-repeated domain followed by repeats as well as a C-terminal cell wall anchor domain. Moreover, the binding of proteins identified by the present method has confirmed that they target and bind to various extracellular matrix molecules. For example, three of the isolated proteins bind to major ECM proteins; two to fibrinogen and at least one to collagen and laminin. The proteins of the present invention have also been shown to be present in most isolates and are expressed in vivo during infection.

In accordance with the present invention, a method is provided for identifying and isolating a module structure of multiple Ig-folded units which have the general characteristics of the MSCRAMM® protein family. The length of the N-subdomains of MSCRAMM® proteins is typically ~150 aa, and the proteins identified by the present invention including those set forth below may accommodate more than three Ig-folded subdomains in their A domains. The isolation and use of the MSCRAMM® proteins of the present invention or their A domains in the generation of antibodies that can bind thereto or in methods of diagnosing, treating or preventing disease will be similar to that as described with other MSCRAMM® proteins such as in U.S. Pat. Nos. 6,288,214; 6,177,084; 6,008,241; 6,086,895; 5,980,908; 5,866,541; 5,851,794; 5,840,846; 5,789,549; 5,770,702; 5,652,217; 5,648,240; 5,571,514; 5,440,014; 5,416,021 and 5,320,951; and WO 00/68242; all of said references incorporated herein by reference.

In accordance with the present invention, a series of steps is undertaken in order to identify and isolate the characteristic module structure of one or more surface-anchored MSCRAMM® protein family of Gram positive bacteria, including the step of locating immunoglobulin-like (or Ig-like) folds in the putative LPXTG-containing proteins. This method can be used with any presently known database containing sequence information from Gram positive bacterial species, e.g., amino acid and/or nucleic acid sequences, and involves the steps of locating proteins with the LPXTG (SEQ ID NO:1) motif, and then reviewing and analyzing the

sequence information so as to screen for proteins having particular structural similarities to MSCRAMM® as set forth below.

In the general process of the invention, the first part of the process is to search a database containing sequence information on one or more Gram positive bacteria so as to locate those proteins which contain the LPXTG (SEQ ID NO:1) motif contained in cell wall anchored proteins in annotated genomes of Gram-positive bacteria. This is done by initially obtaining the entire genome of amino acids sequences from one or more Gram positive bacteria of interest, such as from any of a number of web sites of sequencing centers, e.g., TIGR, NCBI, etc. In the preferred method, these sequences can be downloaded and stored in electronic memory before carrying out the identifying steps, such as in a local Silicon Graphics machine (SGI) or other suitable computer system. In the preferred method, this stored information is used to prepare a local searchable database, such as by using the program form “atdb” obtained from NCBI, and such a searchable database is installed locally on the SGI.

The LPXTG-motif is identified from the stored sequence information by any of a number of suitable programs. For example, these LPXTG-motif containing proteins can be identified using PHI-blast, which is obtained from NCBI and once again can be installed and stored locally on the SGI or other suitable computer system. The PHI-blast search uses a degenerate LPXTG pattern L-P—X-[TSA]-[GANS] (SEQ ID NO: 25), X being any amino acid. The exact templates for PHI-blast can vary depending on the particular organism, but in any case, the present system includes methods of identifying the LPXTG motif. For each organism, it is preferred to use at least two known cell wall anchored proteins of *S. aureus* with no sequence homology as well as known cell wall anchored proteins from the target organism if available.

Once LPXTG-containing proteins are identified obtained using a suitable system such as PHI-blast, these proteins are further analyzed so as to select for those that contain typical features of LPXTG-motif containing cell wall anchored proteins which have the properties of MSCRAMM®s. In the preferred process, these features will generally include a signal peptide at the N-terminus, the LPXTG-motif being close to the C-terminus, followed by a hydrophobic transmembrane segment, and several positively charged residues at the C-terminus. These are done as described below:

The signal peptides may be identified using any suitable identification method such as that method described in “Identification of prokaryotic and eukaryotic signal peptides and prediction of their cleavage sites”. Henrik Nielsen, Jacob Engelbrecht, Søren Brunak and Gunnar von Heijne, *Protein Engineering* 10, 1-6 (1997), incorporated herein by reference. In the present process, a preferred system is to use the SignalP prediction server, but other similar methods for identifying the signal peptide may also be used. Location of LPXTG-motif and the determination of positively charged amino acids residues at the C terminus are accomplished using visual examination of the sequence, although databases may also be used to determine the presence of these features.

In the preferred embodiment, the hydrophobic transmembrane segment after the LPXTG-motif may also be located using a conventional program which can predict the presence of such regions. An example of one such system is the TMHMM server available on the Internet which can be used for the prediction of transmembrane segments. However, a number of other suitable prediction servers are available either on the Internet or in stored computer programs, including the TMpred, the DAS system, and the HMMTOP.

By following the procedures set forth above, putative LPXTG-containing sequences that contain the above features can be selected as highly likely to be MSCRAMM® proteins, i.e., to have the ability to bind extracellular matrix components. Following these initial steps, it is contemplated that the LPXTG-containing proteins identified in this matter will turn out to be MSCRAMM® proteins at least about 90% of the time, as confirmed by expressing the putative protein or its A domain and determining if that protein or its A domain binds to extracellular matrix components. This can be done by simple binding assays which are routine in the art and which would be well within the abilities of one skilled in the art.

Additionally, the LPXTG-containing sequences as initially located, or as further selected using the signal peptide/C-terminal/transmembrane identifying characteristics as described above, can be further analyzed as indicated below to confirm the presence of immunoglobulin-like folds characteristic of MSCRAMM® proteins from Gram positive bacteria.

Similarly, in such a method, LPXTG-containing cell wall proteins may also be located using an annotated genomic nucleotide database such as the one located at the TIGR website (comprehensive microbial resource). With these databases, the term "LPXTG" or "cell wall" may be used to search for such proteins that are annotated as cell wall anchored proteins in the genome of interest.

Finally, LPXTG-motif containing cell wall anchored proteins may also be identified in un-annotated nucleotide genomes of Gram-positive bacteria. In this case, genome sequences are obtained from the web sites of sequencing centers, and the sequences may be stored as appropriate in computer memory such as a local Silicon Graphics machine (SGI). Gene prediction may be carried out using the program such as Glimmer 2.0 from TIGR, and this can be facilitated by UNIX C shell scripts which may be modified as desired to suit particular organisms or features. In the preferred process, the predicted genes are translated into amino acid sequences using a suitable translation program, preferably one that is capable of translating large batches of sequences. Finally, the translated amino acid sequences are formatted into a searchable database locally as described above, and subject to further analysis as described below.

In the preferred process of the present invention, steps are carried out by which the Immunoglobulin-like (Ig-like) fold in putative LPXTG-motif containing cell wall anchored proteins can be predicted and identified. In accordance with the invention, the amino acid sequences of putative LPXTG-motif containing cell wall anchored proteins are then analyzed to determine the presence of Ig-like folds which are characteristic of MSCRAMM® proteins. This can be done in a number of ways, such as by processing the putative MSCRAMM® using fold-recognition software, such as available using the web server 3D-PSSM. Additional methods of fold prediction are discussed in Kelley L A, MacCallum R M & Sternberg M J E. Enhanced Genome Annotation using Structural Profiles in the Program 3D-PSSM. J Mol. Biol. 2000 Jun. 2; 299(2):499-520, incorporated herein by reference. Using this method, the output of 3D-PSSM gives a probability E value indicating the likelihood of the submitted sequence adopting a similar 3D structure as the known and published MSCRAMM®s. In accordance with the invention, proteins that have an E value <0.25 to a published Ig-like fold structure, are considered to contain the predicted Ig-like folds, and such proteins are identified as useful MSCRAMM® proteins in accordance with the invention, i.e., proteins that recognize adhesin molecules on the extracellular matrix of host cells.

The present invention has thus been carried out so as to identify and produce proteins and A domains therefrom which have MSCRAMM®-like characteristics from such Gram positive bacteria, such as *Enterococcus*, *Streptococcus*, *Staphylococcus* and *Bacillus*. In the preferred process, proteins identified as set forth above or their antigenic A domains may be expressed, purified and characterized as set forth herein.

In accordance with the present invention, a bioinformatic approach was thus used to identify proteins with MSCRAMM®-like characteristics among Gram positive bacteria, and those predicted proteins have been shown to have MSCRAMM®-like characteristics. In one such case using *Enterococcus faecalis*, forty-two proteins with a putative C-terminal LPXTG cell wall sorting signal were identified in the *E. faecalis* genome. In accordance with the present method, these proteins were analyzed to determine the presence of Ig-like folds in the manner set forth above. Based on the present method, nine proteins were found to contain regions that adopt an immunoglobulin-like fold. The Ig fold-containing regions for these nine proteins are shown in FIG. 1 and consist of several consecutive and overlapping matches to solved crystal structures (~150-500 aa) of the immunoglobulin superfamily (IgSF), which consist of one to four domains of equal size and Ig-type fold. The homologous Ig-fold regions cover most of the enterococcal proteins and may indicate a similar "beads-in-a-string" arrangement of consecutive modules that are found in fibronectin and other IgSF proteins.

Further expression, purification and analysis of the A domains of these proteins was carried out. As shown in FIG. 2, the A regions of eight proteins expressed as N-terminal His6-tag fusion proteins migrated as expected in SDS-PAGE gels, while EF1091 showed a band approx. 160 kDa in size; a larger-size molecule than the expected 113 kDa. Some degradation was observed in proteins EF1091, EF1824, EF0089 and EF3023, possibly due to their relatively large sizes. They were nevertheless estimated to be >95% pure. The putative glucosyl hydrolase domain of EF1824 (amino acids 42-819), which was cloned and expressed separately from the rest of the protein, (FIG. 1) was found in the insoluble fraction of *E. coli* cytoplasm. Hence, purification by metal affinity chromatography under native, non-denaturing conditions employed for the other expressed proteins was not feasible. The purified proteins were further characterized with MALDI-TOF mass spectrometry. All nine proteins, including EF1091 with aberrant migration in SDS-PAGE, gave peaks that were in good agreement with the molecular weights calculated from amino acid sequences (Table 1), and thus indicated that full-size proteins had been produced with no post-translational processing.

TABLE 1

| Molecular size analysis | | |
|-------------------------|---------------------|-------------------|
| Protein | Molecular mass (Da) | |
| | Sequence prediction | Mass spectrometry |
| EF1091 | 113,021 | 113,025 |
| EF1824 | 111,893 | 111,901 |
| EF0089 | 122,853 | 122,857 |
| EF3023 | 113,338 | 113,323 |
| EF1092 | 47,291 | 47,295 |
| EF2224 | 82,194 | 82,199 |
| EF1269 | 64,776 | 64,776 |

TABLE 1-continued

| Molecular size analysis | | |
|-------------------------|---------------------|-------------------|
| Protein | Molecular mass (Da) | |
| | Sequence prediction | Mass spectrometry |
| EF1099 | 39,281 | 39,293 |
| EF1093 | 62,363 | 62,366 |

Secondary structure predictions and CD-measurements (Table 2) support finding of Ig-folded module-structures in the enterococcal proteins. Both methods show a similar high proportion of β -sheet (~50%) and coil and a minor quantity of α -helix, an identical situation as seen in MSCRAMM® proteins and in IgSF in general. The higher amount of α -helix in EF1824 and EF3023 probably reflects their relatively short predicted regions with Ig-folds and suggests the remainder of the proteins is structurally more distant to MSCRAMM® proteins.

TABLE 2

| Summary of secondary structure components | | | |
|-------------------------------------------|---------------------|-----------------|-----------------|
| Protein | α -Helix | β -Sheet | Other |
| | Sequence prediction | | |
| EF1091 | 0.10 \pm 0.05 | 0.33 \pm 0.08 | 0.53 \pm 0.06 |
| EF1824 | 0.45 \pm 0.04 | 0.16 \pm 0.04 | 0.39 \pm 0.08 |
| EF0089 | 0.07 \pm 0.07 | 0.44 \pm 0.14 | 0.49 \pm 0.08 |
| EF3023 | 0.24 \pm 0.09 | 0.29 \pm 0.10 | 0.47 \pm 0.12 |
| EF1092 | 0.15 \pm 0.05 | 0.36 \pm 0.06 | 0.49 \pm 0.10 |
| EF2224 | 0.15 \pm 0.10 | 0.32 \pm 0.05 | 0.54 \pm 0.10 |
| EF1269 | 0.09 \pm 0.10 | 0.42 \pm 0.12 | 0.49 \pm 0.10 |
| EF1099 | 0.04 \pm 0.07 | 0.47 \pm 0.07 | 0.49 \pm 0.07 |
| EF1093 | 0.09 \pm 0.06 | 0.41 \pm 0.11 | 0.51 \pm 0.11 |
| CD measurement | | | |
| EF1091 | 0.14 \pm 0.05 | 0.41 \pm 0.11 | 0.45 \pm 0.10 |
| EF1824 | 0.29 \pm 0.04 | 0.29 \pm 0.17 | 0.44 \pm 0.17 |
| EF0089 | 0.08 \pm 0.04 | 0.49 \pm 0.13 | 0.43 \pm 0.12 |
| EF3023 | 0.33 \pm 0.05 | 0.16 \pm 0.05 | 0.51 \pm 0.03 |
| EF1092 | 0.05 \pm 0.04 | 0.50 \pm 0.12 | 0.45 \pm 0.14 |
| EF2224 | 0.16 \pm 0.03 | 0.36 \pm 0.10 | 0.48 \pm 0.09 |
| EF1269 | 0.03 \pm 0.04 | 0.55 \pm 0.14 | 0.42 \pm 0.12 |
| EF1099 | 0.07 \pm 0.03 | 0.49 \pm 0.13 | 0.44 \pm 0.14 |
| EF1093 | 0.06 \pm 0.05 | 0.57 \pm 0.18 | 0.37 \pm 0.17 |

In addition to EF1099 (Ace), the primary sequence of EF1269 is clearly related to the MSCRAMM® protein family. Similarly to Ace, it has homologous N2 and N3 subdomains including the conserved TYTDYVD-motif and a connecting tyrosine residue between the two subdomains. The absence of N1 further resembles Ace. However, the rest of their sequences share little homology. Although the A domain of EF1269 is made of similar N2 and N3 subdomains as the fibrinogen-binding ClfA, ClfB, SdrG, and to a lesser extent, FnbpA and FnbpB, it failed to bind fibrinogen. In this respect, EF1269 resembles SdrD and SdrE, which contain N2 and N3 subdomains, but for which the ligand is yet to be found. This is strengthened by our finding that the highest similarity of the EF1269 N2 and N3 domains is to the corresponding region in SdrE (identity 26%). Further, two putative repeats (95 and 109 aa) with lower conservation (identity 20%), which make up the rest of the C-terminal EF1269 sequence, show relatedness to the B repeats of SdrE (25% identity over 375 to 531 aa of EF1269). Proteins EF1091, EF0089, EF1092, EF2224 and EF1093 are not simply orthologs of previously described

MSCRAMM® proteins, since they lack high sequence identity to streptococcal and staphylococcal adhesins. Yet, they share similar structural organization and an abundance of β -sheet rich secondary structures with similar predicted folding as MSCRAMM® proteins. The two remaining proteins, EF1824 and EF3023, have large regions related to known enzymes, glucosyl hydrolases and hyaluronan lyases, respectively, which sets these regions apart from MSCRAMM® proteins. Hyaluronidase activity could be significant for bacterial entry and spreading in hyaluronan-containing tissues during infection and/or potentially contribute to bacterial nutrition during commensal life in the human intestine. The large putative catalytic domains of EF1824 and EF3023 agree well with the above-discussed structural unrelatedness in these regions to MSCRAMM® proteins.

When screening binding to major ECM proteins, we found ligands for five of the MSCRAMM® proteins EF0089, EF1091, EF1092, EF1093, and EF2224. The presence of more than one fibrinogen-binding MSCRAMM® proteins in *E. faecalis* is consistent to findings in the related *S. aureus* in which four fibrinogen-binding MSCRAMM® proteins, ClfA, ClfB, FnbpA and FnbpB, have been described (McDevitt et al., 1994) (Ni Eidhin et al., 1998) (Wann et al., 2000) (Davis et al., 2001; Hartford et al., 2001). EF0089 and EF2224 have strong structural resemblance to MSCRAMM® proteins throughout their lengths: similar primary organization and homologous β -sheet rich secondary structure expected to form modular Ig-folded subdomains. Relatively low sequence identity to known fibrinogen binding adhesins may mean novel adaptations for ligand binding. Our initial results suggest EF2224 binds to the α - and β -chains of fibrinogen and thus resembles ClfB (Ni Eidhin et al., 1998). Mammalian tissue surfaces express a multitude of possible ligands for bacterial adherence. Here, we assessed binding to type I, III and IV collagens, laminin, fibronectin, fibrinogen and vitronectin.

In accordance with the invention, a PCR process may be used to amplify A domains from proteins identified and isolated using the present invention. Using PCR oligonucleotides such as those in Table 3, below, the A domains from EF0089, EF1091, EF1092, EF1093, EF1099, EF1269, EF1824, EF2224, and EF3023 were amplified from *E. faecalis* V583 or *E. faecalis* EF1 (EF1099) genomic DNA and subcloned into the *E. coli* expression vector pQE-30 (Qiagen). One liter culture of *E. coli* M15(pREP4) cultures harboring appropriate pQE-30 based constructs were grown to OD₆₀₀=0.6 with an initial 2% inoculation from overnight cultures. After 2-3 h induction with 0.4 mM isopropyl-beta-D-thiogalactoside (IPTG), cells were collected with centrifugation, resuspended in 10 mM Tris-Cl, 100 mM NaCl, pH 7.9 and stored at -80 C.

To lyse the cells and release the expressed protein, cells were passed twice through French Press with a gauge pressure setting at 1200 PSI to give an estimated internal cell pressure of 20,000 PSI. The lysate was centrifuged at RCF_{max} of 165,000 \times g and the supernatant was filtered through a 0.45 μ m filter. The volume was adjusted to 15 ml with 10 mM Tris-Cl, 100 mM NaCl, pH 7.9 and 0.2 M imidazole in the same buffer was added to increase the imidazole concentration to 6.5 mM in order to minimize non-specific binding. The sample was loaded to a nickel affinity chromatography column (HiTrap chelating, Pharmacia) connected to an FPLC system (Pharmacia) and previously equilibrated with 10 mM Tris-Cl, 100 mM NaCl, pH 7.9. Bound protein was eluted with a linear gradient of 0-100 mM imidazole in 10 mM Tris-Cl, 100 mM NaCl, pH 7.9 over 100-200 ml. Protein-containing fractions were analyzed in SDS-PAGE (FIG. 2)

and dialyzed against 25 mM Tris-Cl, 1 mM EDTA, pH 6.5-9 (depending on pI of protein purified) before applying the samples to an ion-exchange column (HiTrap Q, Pharmacia) for further purification. Bound protein was eluted with a linear gradient of 0-0.5 M NaCl in 25 mM Tris-Cl, 1 mM EDTA, pH 6.5-9 over 100 ml. Finally, protein samples were dialyzed extensively against PBS and stored at +4° C.

Alternatively EF1091, EF1092, and EF1093 were expressed in shake flasks or in bioreactors, the cells were harvested by centrifugation and the cell paste frozen at -80° C. Cells were lysed in 1xPBS (10 mL of buffer/1 g of cell paste) using 2 passes through a microfluidizer at 10,000 psi.

This may be due to different expression levels in physiological conditions or to highly immunogenic surface epitopes and, hence, a strong immune response. Interestingly, the three proteins (EF1091, EF1092 and EF1093) with the highest titers are organized as a putative operon in the *E. faecalis* genome. The operon is preceded by two promoter consensus regions and a ribosome binding site and thus, these proteins are likely co-transcribed. The next gene downstream, EF1094, codes for a putative LPxTG transpeptidase sortase and EF1099 (Ace) is closely linked. It remains to be seen what role this cluster of MSCRAMM®-like proteins and a putative sortase may have in the infection process.

TABLE 3

| Synthetic oligonucleotides used in this study (SEQ ID NOS: 26-43) | | | | | |
|----------------------------------------------------------------------|------------------|-----------------|-------|-------------------------------------------------------|--|
| Oligonucleotide | Location (aa) | Cloning site | | Oligonucleotide | |
| EF1091A | Fw | 102 | SphI | 5' - CCGCATGCCAAGAGCAAACAGCAAAGAAG - 3' | |
| | Rev | 1107 | SalI | 5' - CCGTCGACTTAAAGTACCAGAAGTGGTGGTTTC - 3' | |
| EF1824AI | Fw | 42 | SphI | 5' - CCGCATGCCAAGAGCAAACAGCAAAGAAG - 3' | |
| | Rev | 819 | SalI | 5' - GGGTCGACTTATTGTTCAAGGTTACTTCTGTC | |
| EF1824AII | Fw | 819 | BamHI | 5' - CCGGATCCGCAGCTAATAAAGAAGAATTTT TAG | |
| | Rev | 1829 | SalI | 5' - CCGTCGACTTAAAGTACCAGAAGTGGTGGTTTC - 3' | |
| EF0089A | Fw | 35 | SacI | 5' - CCGAGCTCGAAGAGGTTAACAGCGATGG - 3' | |
| | Rev | 1143 | PstI | 5' - CCCTGCGACTTACCCACCAATGTGATAACCC - 3' | |
| EF3023A | Fw | 25 | BamHI | 5' - CCGGATCCGAAGAAATAACTGATTATTTT TAC - 3' | |
| | Rev | 1024 | SacI | 5' - CCGAGCTCTTATTGTTCTGATTAATTTT TCTAAC - 3' | |
| EF1092A | Fw | 27 | SphI | 5' - CCGCATGCTCGCAAGCAAGCGTTCAAG - 3' | |
| | Rev | 438 | PstI | 5' - CCCTGCGACTTAGAAGCCTGACTCTTTT TACTTTT - 3' | |
| EF2224A | Fw | 30 | BamHI | 5' - CCGGATCCCAAGAAGTAACAAGTGATGCTG - 3' | |
| | Rev | 771 | SacI | 5' - CCGAGCTCTTAAAGTACTTGTTCGTC CCGCAAT - 3' | |
| EF1269A | Fw | 26 | BamHI | 5' - CCGGATCCGAAACAGGATATGCGCAAAC - 3' | |
| | Rev | 596 | SacI | 5' - CCGAGCTCTTATTCTTATTACGAATCGCCTG - 3' | |
| EF1093A | Fw | 32 | BamHI | 5' - GCGGGATCCGAAGAAAATGGGGAGAGCGC - 3' | |
| | Rev | 590 | SacI | 5' - GCGGAGCTCTTAGGTACCTTTGTGTTTGTGG - 3' | |

5' overhang cloning site in each oligonucleotide sequence is marked in bold, stop codon in italic Fw, oligonucleotide primer in forward direction; Rev, in reverse direction

Lysed cells were spun down at 17,000 rpm for 30 minutes to remove cell debris. Supernatant was passed over a 5-mL HiTrap Chelating (Pharmacia) column charged with 0.1M NiCl₂. After loading, the column was washed with 5 column volumes of 10 mM Tris, pH 8.0, 100 mM NaCl (Buffer A). Protein was eluted using a 0-100% gradient of 10 mM Tris, pH 8.0, 100 mM NaCl, 500 mM imidazole (Buffer B). Protein containing fractions were dialyzed in 1xPBS.

The nine enterococcal genes encoding the MSCRAMM® are ubiquitous among *E. faecalis* strains as summarized in Table 3. Seven of the nine genes were 100% preserved in all strains. The two genes, EF1824 and EF3023, with predicted encoded protein catalytic domains and relatively low proportion of MSCRAMM®-like protein characteristics, were present in 16/30 and 23/30 strains, respectively. Nine enterococcal proteins encoded by their respective gene showed elevated titers in infected individuals suggesting expression in vivo during an *E. faecalis* infection. Although these proteins have a high distribution in strains, there were clear differences in induced antigenic responses; proteins EF1091, EF1092, EF1093 and EF2224 exhibited the highest titers.

The presence of several MSCRAMM®-like proteins in *E. faecalis* including two that bind fibrinogen and the previously described collagen and laminin binding Ace, suggests that *E. faecalis* resembles *S. aureus* and other Gram-positive cocci by having an armory of ECM-binding adhesins. Since the introduction of antibiotic therapy, *E. faecalis* has shown an increasing tendency to emerge as an opportunistic pathogen capable of crossing the thin line from a harmless commensal to being able to invade host tissues and cause infections. A repertoire of adhesins may enhance its adaptability for colonizing and spreading in various human tissue types of susceptible human hosts.

Accordingly, the present invention allows for the identification and ultimate production of novel MSCRAMM®-like protein surface-anchored proteins from Gram positive bacteria which (i) are structurally homologous to the solved Ig-folded crystal structures of ClfA and CNA as well as to the predicted tertiary structures of other MSCRAMM® proteins, (ii) can share a similar β -sheet rich secondary structure as is found in Ig-folded proteins and (iii) have a similar organiza-

tion with a secretion signal, a non-repeated domain followed by repeats as well as a C-terminal cell wall anchor domain. Further, these proteins may bind to major ECM proteins such as fibrinogen, collagen and laminin, and due to the similarities in proteins from different Gram positive bacterial species, these proteins may provide antibodies which are cross-reactive and can bind to similar proteins found in different Gram positive bacterial species. Such antibodies, as described further below, may thus be useful in diagnosing or fighting a variety of different infections at the same time.

In addition to proteins identified and isolated using the present method, particular, the present invention contemplates the generation of antibodies from the MSCRAMM®-like proteins obtained using the present method, or from antigenic regions such as the A domains from these proteins. By "antibody" is meant any intact antibody molecule or fragments thereof that recognize antigen (e.g. Fab or F(ab')₂ fragments) and can be of polyclonal or monoclonal type, and the antibodies in accordance with the invention will be capable of recognizing the MSCRAMM® proteins of the invention and/or the specific antigenic epitopes from said proteins including their A domains. These antibodies will thus be effective in methods of diagnosing, monitoring, treating or preventing infection from Gram positive bacteria. By "epitope" is meant any antigenic determinant responsible for immunochemical binding with an antibody molecule. Epitopes usually reside within chemically active surface groupings of protein molecules (including amino acids and often also sugar side-chains) and have specific three-dimensional structural characteristics and specific charge characteristics. With reference to the proteins of the invention, or epitopes and peptides as described herein, it is understood that such terms also include those proteins and peptides which differ from a naturally occurring or recombinant protein by the substitution, deletion and/or addition of one or more amino acids but which retains the ability to be recognized by an antibody raised against the entire protein. An example is a carrier/antigen fusion polypeptide of the whole antigen or an immunoreactive fragment thereof, where the antigen or fragment can be embedded within the carrier polypeptide or linked to the carrier polypeptide at either end.

Accordingly, in accordance with the present invention, isolated and/or purified antibodies can be generated from the Gram-positive MSCRAMM® proteins of the present invention, or from particular epitopes such as those epitopic peptide sequences from the A domains from those proteins as described herein. These antibodies may be monoclonal or polyclonal and may be generated using any suitable method to raise such antibodies such as would be well known in this art. The antibodies in accordance with the invention will be particularly useful in inhibiting the binding of Gram positive bacteria to extracellular matrix components of the host cells and in diagnosing, treating or preventing infections of Gram positive bacteria.

For example, with regard to polyclonal antibodies, these may be generated using a number of suitable methods generally involving the injection of the isolated and/or purified or recombinantly produced proteins (or their immunogenic active peptides or epitopes) into a suitable host in order to generate the polyclonal antibodies which can then be recovered from the host. For example, in accordance with the invention, an isolated and purified MSCRAMM® protein or its A domain may be injected into rabbits in order to generate polyclonal antisera recognizing this protein.

In addition, monoclonal antibodies in accordance with the invention may be generated using a suitable hybridoma as would be readily understood by those of ordinary skill in the

art. In the preferred process, a protein in accordance with the invention is first identified and isolated using the bioinformatic method as described above. Next, the protein is isolated and/or purified in any of a number of suitable ways commonly known in the art, or after the protein is sequenced, the protein used in the monoclonal process may be produced by recombinant means as would be commonly used in the art and then purified for use. In one suitable purification process, the cell wall proteins of the invention are isolated and examined using polyacrylamide gel electrophoresis (PAGE) and Western-blot techniques, and other conventional techniques including those discussed herein. In one suitable process, monoclonal antibodies were generated from proteins isolated and purified as described above by mixing the protein with an adjuvant, and injecting the mixture into BALB/c mice.

Immunization protocols consisted of a first injection (using complete Freund's adjuvant), two subsequent booster injections (with incomplete Freund's adjuvant) at three-week intervals, and one final booster injection without adjuvant three days prior to fusion (all injections were subcutaneous). For hybridoma production, mice were sacrificed and their spleen removed aseptically. Antibody secreting cells isolated and mixed with myeloma cells (NS1) using drop-wise addition of polyethylene glycol. After the fusion, cells were diluted in selective medium (vitamin-supplemented DMEM/HAT) and plated at low densities in multiwell tissue culture dishes. Tissue supernatants from the resulting fusion were screened by both ELISA (using the total 2-ME extract to coat the wells of a microtiter plate) and immunoblot techniques. Cells from these positive wells were grown and single cell cloned by limiting dilution, and supernatants subjected to one more round of screening by both ELISA and immunoblot. Positive clones were identified, and monoclonal antibodies collected as hybridoma supernatants.

In accordance with the invention, antibodies are thus produced which are capable of recognizing and binding proteins obtained using the bioinformatic method of the present invention and/or its epitopes and active regions such as the A domain, and such antibodies can be utilized in many diagnostic and therapeutic applications such as the ones described in more detail below.

Vaccines, Humanized Antibodies and Adjuvants

The isolated antibodies of the present invention, or the isolated proteins or epitopes as described above, may also be utilized in the development of vaccines for active and passive immunization against bacterial infections, as described further below. In the case of active vaccines, said vaccines are prepared by providing an immunogenic amount of the proteins of the invention or their active regions or epitopes as set forth above, and the active vaccine in accordance with the invention will thus comprise an immunogenic amount of the protein or peptide and will be administered to a human or animal in need of such a vaccine. The vaccine may also comprise a suitable, pharmaceutically acceptable vehicle, excipient or carrier which will be those known and commonly used in the vaccine arts. As referred to above, an "immunogenic amount" of the antigen to be used in accordance with the invention is intended to mean a nontoxic but sufficient amount of the agent, such that an immunogenic response will be elicited in the host so that the desired prophylactic or therapeutic effect is produced. Accordingly, the exact amount of the antigen that is required will vary from subject to subject, depending on the species, age, and general condition of the subject, the severity of the condition being treated, the particular carrier or adjuvant being used and its mode of administration, and the like. Similarly, the "immunogenic

amount” of any such antigenic vaccine composition will vary based on the particular circumstances, and an appropriate immunogenic amount may be determined in each case of application by one of ordinary skill in the art using only routine experimentation. The dose should be adjusted to suit the individual to whom the composition is administered and will vary with age, weight and metabolism of the individual.

Further, when administered as pharmaceutical composition to a patient or used to coat medical devices or polymeric biomaterials *in vitro* and *in vivo*, the antibodies of the present invention may also be useful because these antibodies may be able to interfere with the ability of Gram positive bacteria to adhere to host cells and limit the extent and spread of the infection.

In addition, the antibody may be modified as necessary so that, in certain instances, it is less immunogenic in the patient to whom it is administered. For example, if the patient is a human, the antibody may be “humanized” by transplanting the complementarity determining regions of the hybridoma-derived antibody into a human monoclonal antibody as described, e.g., by Jones et al., *Nature* 321:522-525 (1986) or Tempest et al. *Biotechnology* 9:266-273 (1991) or “veneered” by changing the surface exposed murine framework residues in the immunoglobulin variable regions to mimic a homologous human framework counterpart as described, e.g., by Padlan, *Molecular* 1 mm. 28:489-498 (1991), these references incorporated herein by reference. Even further, under certain circumstances, it may be desirable to combine the monoclonal antibodies of the present invention with a suitable antibiotic when administered so as to further enhance the ability of the present compositions to fight or prevent infections.

In a preferred embodiment, the antibodies may also be used as a passive vaccine which will be useful in providing suitable antibodies to treat or prevent a Gram-positive bacterial infection. As would be recognized by one skilled in this art, a vaccine may be packaged for administration in a number of suitable ways, such as by parenteral (i.e., intramuscular, intradermal or subcutaneous) administration or nasopharyngeal (i.e., intranasal) administration. One such mode is where the vaccine is injected intramuscularly, e.g., into the deltoid muscle, however, the particular mode of administration will depend on the nature of the bacterial infection to be dealt with and the condition of the patient. The vaccine is preferably combined with a pharmaceutically acceptable vehicle, carrier or excipient to facilitate administration, and the carrier is usually water or a buffered saline, with or without a preservative. The vaccine may be lyophilized for resuspension at the time of administration or in solution.

The preferred dose for administration of an antibody composition in accordance with the present invention is that amount will be effective in preventing of treating a bacterial infection, and one would readily recognize that this amount will vary greatly depending on the nature of the infection and the condition of a patient. An “effective amount” of antibody or pharmaceutical agent to be used in accordance with the invention is intended to mean a nontoxic but sufficient amount of the agent, such that the desired prophylactic or therapeutic effect is produced. Accordingly, the exact amount of the antibody or a particular agent that is required will vary from subject to subject, depending on the species, age, and general condition of the subject, the severity of the condition being treated, the particular carrier or adjuvant being used and its mode of administration, and the like. Accordingly, the “effective amount” of any particular antibody composition will vary based on the particular circumstances, and an appropriate effective amount may be determined in each case of

application by one of ordinary skill in the art using only routine experimentation. The dose should be adjusted to suit the individual to whom the composition is administered and will vary with age, weight and metabolism of the individual. The compositions may additionally contain stabilizers or pharmaceutically acceptable preservatives, such as thimerosal (ethyl(2-mercaptobenzoate-S)mercury sodium salt) (Sigma Chemical Company, St. Louis, Mo.).

In addition, the antibody compositions of the present invention and the vaccines as described above may also be administered with a suitable adjuvant in an amount effective to enhance the immunogenic response against the conjugate. For example, suitable adjuvants may include alum (aluminum phosphate or aluminum hydroxide), which is used widely in humans, and other adjuvants such as saponin and its purified component Quil A, Freund’s complete adjuvant, and other adjuvants used in research and veterinary applications. Still other chemically defined preparations such as muramyl dipeptide, monophosphoryl lipid A, phospholipid conjugates such as those described by Goodman-Snitkoff et al. *J. Immunol.* 147:410-415 (1991) and incorporated by reference herein, encapsulation of the conjugate within a proteoliposome as described by Miller et al., *J. Exp. Med.* 176:1739-1744 (1992) and incorporated by reference herein, and encapsulation of the protein in lipid vesicles such as Novasome™ lipid vesicles (Micro Vesicular Systems, Inc., Nashua, N.H.) may also be useful.

Pharmaceutical Compositions

As would be recognized by one skilled in the art, the identified and isolated proteins or the invention, and the antibodies thereto capable of recognizing and binding to said proteins may also be formed into suitable pharmaceutical compositions for administration to a human or animal patient in order to treat or prevent a Gram-positive bacterial infection, such as those caused by *Enterococcus*, *Streptococcus*, *Staphylococcus*, etc. Pharmaceutical compositions containing the proteins or antibodies of the present invention as defined and described above may be formulated in combination with any suitable pharmaceutical vehicle, excipient or carrier that would commonly be used in this art, including such as saline, dextrose, water, glycerol, ethanol, other therapeutic compounds, and combinations thereof. As one skilled in this art would recognize, the particular vehicle, excipient or carrier used will vary depending on the patient and the patient’s condition, and a variety of modes of administration would be suitable for the compositions of the invention, as would be recognized by one of ordinary skill in this art. Suitable methods of administration of any pharmaceutical composition disclosed in this application include, but are not limited to, topical, oral, anal, vaginal, intravenous, intraperitoneal, intramuscular, subcutaneous, intranasal and intradermal administration.

For topical administration, the composition may be formulated in the form of an ointment, cream, gel, lotion, drops (such as eye drops and ear drops), or solution (such as mouth-wash). Wound or surgical dressings, sutures and aerosols may be impregnated with the composition. The composition may contain conventional additives, such as preservatives, solvents to promote penetration, and emollients. Topical formulations may also contain conventional carriers such as cream or ointment bases, ethanol, or oleyl alcohol.

Additional forms of compositions, and other information concerning compositions, methods and applications with regard to other microbial surface proteins and peptides of the present invention and antibodies thereto, will be found in other patent references relating to MSCRAMM®, includ-

ing, for example, in U.S. Pat. No. 6,288,214 (Hook et al.), incorporated herein by reference.

The compositions which are generated in accordance with the present invention may also be administered with a suitable adjuvant in an amount effective to enhance the immunogenic response in a patient. For example, suitable adjuvants may include alum (aluminum phosphate or aluminum hydroxide), which is used widely in humans, and other adjuvants such as saponin and its purified component Quil A, Freund's complete adjuvant, RIBI adjuvant, and other adjuvants used in research and veterinary applications. Still other chemically defined preparations such as muramyl dipeptide, monophosphoryl lipid A, phospholipid conjugates such as those described by Goodman-Snitkoff et al. *J. Immunol.* 147:410-415 (1991) and incorporated by reference herein, encapsulation of the conjugate within a proteoliposome as described by Miller et al., *J. Exp. Med.* 176:1739-1744 (1992) and incorporated by reference herein, and encapsulation of the protein in lipid vesicles such as Novasome™ lipid vesicles (Micro Vesicular Systems, Inc., Nashua, N.H.) may also be useful.

In any event, the compositions of the present invention will thus be useful for interfering with, modulating, or inhibiting binding interactions by Gram positive bacteria. Accordingly, the present invention will have particular applicability in developing compositions and methods of preventing or treating Gram positive bacterial infections, and in inhibiting binding and spreading of bacteria to host cells.

Methods:

Detecting and Diagnosing Infections

In accordance with the present invention, methods are provided for identifying and diagnosing infection from Gram positive bacteria through the use of the proteins, epitopes and peptides obtained by the bioinformatic method of the invention as described above and antibodies that recognize such proteins, epitopes and/or peptides. In accordance with the present invention, the antibodies of the invention as set forth above may be used in kits to diagnose such infections, and such kits may be of the type generally known in the art and commonly used to detect an antigen or microorganism of interest which will bind to the antibodies of the invention. These diagnostic kits will generally include the antibodies of the invention along with suitable means for detecting binding by that antibody such as would be readily understood by one skilled in this art. For example, the means for detecting binding of the antibody may comprise a detectable label that is linked to said antibody. These kits can then be used in diagnostic methods to detect the presence of a Gram positive bacterial infection wherein one obtains a sample suspected of being infected by one or more Gram positive bacteria, such as a sample taken from an individual, for example, from one's blood, saliva, urine, cerebrospinal fluid, genitourinary tract, tissues, bone, muscle, cartilage, or skin, and introduces to the sample one or more of the antibodies as set forth herein. After introduction of the antibodies, it is then determined through conventional means whether there has been binding by the antigens or microorganisms in the sample, such as through suitable labeling, or assays wherein the antibodies are bound to solid supports, and this binding is reflective of the presence of the target antigens or microorganisms in the sample.

Methods for Monitoring Levels of Antibodies or Antigens

In accordance with the present invention, it is also contemplated that another use of the invention may be in monitoring the level of Gram positive bacterial antigens, or antibodies recognizing said antigens in a human or animal patients suspected of containing said antigens or antibodies. In the pre-

ferred process, this may be carried out by first obtaining a biological sample from the human or animal patient, and this would include any suitable sample routinely monitored for infection, such as for example, from one's blood, serum, saliva, tissues, bone, muscle, cartilage, or skin. Next, one would introduce into the sample either (1) when monitoring levels of one's antibodies to Gram positive bacteria, a determinable level of a protein or its A domain to which such antibodies will bind; or (2), when monitoring levels of bacterial infestation is desired, introducing into said sample a measurable level of an antibody to a protein as set forth above. The next step in the process is, after allowing sufficient time and conditions so that the antigens and antibodies in the sample can achieve binding, then determining the level of antigen-antibody binding which will be reflective of the amount or level of the Gram positive bacteria, or antibodies thereto, which are located in the sample. In the desired process, levels may be monitored at regular time periods (e.g., hourly, daily, etc.) so as to track the progression/remission of a Gram positive bacterial infection such as during the period of hospitalization or treatment.

Assays for Detecting and Diagnosing Infections

In accordance with the present invention, the detection of Gram positive bacteria present in a biological fluid (e.g. blood, serum, plasma, saliva, urine, cerebrospinal fluid, genitourinary tract) or other biological material (e.g., tissues, bone, muscle, cartilage, or skin) can constitute a method for the diagnosis of acute or chronic infections caused by Gram positive bacteria. Because the antibodies as set forth above can recognize the epitopes found in several Gram positive bacteria, these antibodies can be used in assays to allow the diagnosis of a wide variety of Gram positive bacteria and disease conditions. Either monoclonal antibodies or polyclonal antibodies could be used in the assay, and in the case of the monoclonals such as those referred to above. The detected antigens identified by use of the present assays can be detected by a number of conventional means, including Western immunoblot and other similar tests.

With regard to the assays of the present invention, these assays may use the antibodies of the invention in labeled form, and all well-known methods of labeling antibodies are contemplated, including without limitation enzymatic conjugates, direct labeling with dye, radioisotopes, fluorescence, or particulate labels, such as liposome, latex, polystyrene, and colloid metals or nonmetals. Multiple antibody assay systems, such as antigen capture sandwich assays, are also within the scope of this invention. Further, competitive immunoassays involving labeled protein or assays using the labeled protein to detect serum antibodies are also contemplated forms of the diagnostic assays of the present invention. Beyond diagnostic assays which occur in solution, assays which involve immobilized antibody or protein are also considered within the scope of the invention. (See, for example, Miles et al., *Lancet* 2:492, 1968; Berry et al., *J. Virol. Met.* 34:91-100, 1991; Engvall et al., *G. Immunochemistry*, 8:871, 1971, Tom, *Liposomes and Immunology*, Elsevier/North Holland, New York, N.Y., 1980; Gribnau et al., *J. of Chromatogr.* 376:175-89, 1986 and all references cited therein). Examples of the types of labels which can be used in the present invention include, but are not limited to, enzymes, radioisotopes, fluorescent compounds, chemiluminescent compounds, bioluminescent compounds, particulates, and metal chelates. Those of ordinary skill in the art will know of other suitable labels for binding to the monoclonal or polyclonal antibody (or to an antigen) or will be able to ascertain the same by the use of routine experimentation. Furthermore,

the binding of these labels to the monoclonal or polyclonal antibody (or antigen) can be accomplished using standard techniques commonly known to those of ordinary skill in the art.

One of the ways in which an assay reagent (generally, a monoclonal antibody, polyclonal antibody or antigen) of the present invention can be detectably labeled is by linking the monoclonal antibody, polyclonal antibody, or antigen to an enzyme. This enzyme, in turn, when later exposed to its substrate, will react with the substrate in such a manner as to produce a chemical moiety which can be detected as, for example, by spectrophotometric or fluorometric means. Examples of enzymes which can be used to detectably label the reagents of the present invention include malate dehydrogenase, staphylococcal nuclease, delta-V-steroid isomerase, yeast alcohol dehydrogenase, alpha-glycerophosphate dehydrogenase, triose phosphate isomerase, horseradish peroxidase, alkaline phosphatase, asparaginase, glucose oxidase, beta-galactosidase, ribonuclease, urease, catalase, glucose-VI-phosphate dehydrogenase, glucoamylase and acetylcholine esterase.

The presence of the detectably labeled reagent of the present invention can also be detected by labeling the reagent with a radioactive isotope which can then be determined by such means as the use of a gamma counter or a scintillation counter. Isotopes which are particularly useful for the purpose of the present invention are .sup.3H, .sup.125 I, .sup.32 P, .sup.35 S, .sup.14 C, .sup.51 Cr, .sup.36 Cl, .sup.57 Co, .sup.58 Co, .sup.59 Fe and .sup.75 Se. It is also possible to detect the binding of the detectably labeled reagent of the present invention by labeling the monoclonal or polyclonal antibody with a fluorescent compound. When the fluorescently labeled reagent is exposed to light of the proper wave length, its presence can then be detected due to the fluorescence of the dye. Among the most commonly used fluorescent labeling compounds are fluorescein isothiocyanate, rhodamine, phycoerythrin, phycocyanin, allophycocyanin, o-phthaldehyde and fluorescamine. The reagents of the present invention also can be detectably labeled by coupling it to a chemiluminescent compound. The presence of the chemiluminescent-tagged reagent is then determined by detecting the presence of luminescence that arises during the course of the chemical reaction. Examples of particularly useful chemiluminescent labeling compounds are luminol, isoluminol, therromatic acridinium ester, imidazole, acridinium salt and oxalate ester. Likewise, a bioluminescent compound may be used to label the reagent of the present invention. Bioluminescence is a type of chemiluminescence found in biological systems in which a catalytic protein increases the efficiency of the chemiluminescent reaction. The presence of a bioluminescent reagent is determined by detecting the presence of luminescence. Important bioluminescent compounds for purposes of labeling are luciferin, luciferase and aequorin.

Another technique which may also result in greater sensitivity when used in conjunction with the present invention consists of coupling the monoclonal or polyclonal antibody of the present invention to low molecular weight haptens. The haptens can then be specifically detected by means of a second reaction. For example, it is common to use such haptens as biotin (reacting with avidin) or dinitrophenol, pyridoxal and fluorescamine (reacting with specific antihapten antibodies) in this manner. Any biological sample containing the detectable yet unknown amount of a Gram positive antigen can be used in the assay. Normally, the sample is preferably a liquid, such as, for example, urine, saliva, cerebrospinal fluid,

blood, serum and the like, or a solid or semi-solid, such as, for example, tissue, feces and the like.

The diagnostic assay of the present invention includes kit forms of such an assay. This kit would include antibodies as described above (raised against whole proteins or active immunoreactive fragments such as the A domain or immunogenic analogs thereof) which can be optionally immobilized, as well as any necessary reagents and equipment to prepare the biological sample for and to conduct analysis, e.g. preservatives, reaction media such as nontoxic buffers, microtiter plates, micropipettes, etc. The reagent (Abs and/or antigens) can be lyophilized or cryopreserved. As described above, depending on the assay format, the antibodies can be labeled, or the kit can further comprise labeled proteins, fragments or analogs thereof containing the relevant epitopes so as to enable the detection of antibodies to Gram positive bacteria in biological fluids and tissues. By analog is meant a protein or peptide which may differ from its naturally occurring or recombinant counterpart by the substitution, deletion and/or addition of one or more amino acids but which retains the ability to be recognized by an antibody raised against the entire protein. An example is a carrier/antigen fusion polypeptide of the whole antigen or an immunoreactive fragment thereof, where the antigen or fragment can be embedded within the carrier polypeptide or linked to the carrier polypeptide at either end. Accordingly, antibodies in accordance with the invention may also recognize such analogs. The types of immunoassays which can be incorporated in kit form are many. Typical examples of some of the immunoassays which can utilize the antibodies of the invention are radioimmunoassays (RIA) and immunometric, or sandwich, immunoassays.

By "immunometric assay" or "sandwich immunoassay", in meant to include simultaneous sandwich, forward sandwich and reverse sandwich immunoassays. These terms are well understood by those skilled in the art. Those of skill will also appreciate that the monoclonal antibodies, polyclonal antibodies and/or antigens of the present invention will be useful in other variations and forms of immunoassays which are presently known or which may be developed in the future. These are intended to be included within the scope of the present invention. In a forward sandwich immunoassay, a sample is first incubated with a solid phase immunoabsorbent containing monoclonal or polyclonal antibody(ies) against the antigen. Incubation is continued for a period of time sufficient to allow the antigen in the sample to bind to the immobilized antibody in the solid phase. After the first incubation, the solid phase immunoabsorbent is separated from the incubation mixture and washed to remove excess antigen and other interfering substances, such as non-specific binding proteins, which also may be present in the sample. Solid phase immunoabsorbent containing antigen bound to the immobilized antibody is subsequently incubated for a second time with soluble labeled antibody or antibodies. After the second incubation, another wash is performed to remove unbound labeled antibody(ies) from the solid phase immunoabsorbent and removing non-specifically bound labeled antibody(ies). Labeled antibody(ies) bound to the solid phase immunoabsorbent is then detected and the amount of labeled antibody detected serves as a direct measure of the amount of antigen present in the original sample.

Alternatively, labeled antibody which is not associated with the immunoabsorbent complex can also be detected, in which case the measure is in inverse proportion to the amount of antigen present in the sample. Forward sandwich assays are described, for example, in U.S. Pat. Nos. 3,867,517; 4,012,294 and 4,376,110, incorporated herein by reference.

In carrying out forward immunometric assays, the process may comprise, in more detail: (a) first forming a mixture of the sample with the solid phase bound antibody(ies) and incubating the mixture for a time and under conditions sufficient to allow antigen in the sample to bind to the solid phase bound antibody(ies), (b) adding to the mixture after said incubation of step (a) the detectably labeled antibody or antibodies and incubating the new resulting mixture for a time and under conditions sufficient to allow the labeled antibody to bind to the antigen-antibody complex on the solid phase immuno-adsorbent; (c) separating the solid phase immuno-adsorbent from the mixture after the incubation in step (b); and (d) detecting either the labeled antibody or antibodies bound to the antigen-antibody complex on the solid phase immuno-adsorbent or detecting the antibody not associated therewith.

In a reverse sandwich assay, the sample is initially incubated with labeled antibody(ies), after which the solid phase immuno-adsorbent containing multiple immobilized antibodies is added thereto, and a second incubation is carried out. The initial washing step of a forward sandwich assay is not required, although a wash is performed after the second incubation. Reverse sandwich assays have been described, for example, in U.S. Pat. Nos. 4,098,876 and 4,376,110. In carrying out reverse immunometric assays, the process may comprise, in more detail; (a) first forming a mixture of the sample with the soluble detectably labeled antibody for a time and under conditions sufficient to allow antigen in the sample to bind to the labeled antibody; (b) adding to the mixture after the incubation of step (a) the solid phase bound antibodies and incubating the new resulting mixture for a time and under conditions sufficient to allow antigen bound to the labeled antibody to bind to the solid phase antibodies; (c) separating the solid phase immuno-adsorbent from the incubating mixture after the incubation in step (b); and (d) detecting either the labeled antibody bound to the solid phase immuno-adsorbent or detecting the labeled antibody not associated therewith.

In a simultaneous sandwich assay, the sample, the immuno-adsorbent having multiple immobilized antibodies thereon and labeled soluble antibody or antibodies are incubated simultaneously in one incubation step. The simultaneous assay requires only a single incubation and does not include washing steps. The use of a simultaneous assay is by far the preferred one. This type of assay brings about ease of handling, homogeneity, reproducibility, and linearity of the assays and high precision. The sample containing antigen, solid phase immuno-adsorbent with immobilized antibodies and labeled soluble antibody or antibodies is incubated under conditions and for a period of time sufficient to allow antigen to bind to the immobilized antibodies and to the soluble antibody(ies). In general, it is desirable to provide incubation conditions sufficient to bind as much antigen as possible, since this maximizes the binding of labeled antibody to the solid phase, thereby increasing the signal. Typical conditions of time and temperature are two hours at 45 degrees C., or twelve hours at 37 degrees C. Antigen typically binds to labeled antibody more rapidly than to immobilized antibody, since the former is in solution whereas the latter is bound to the solid phase support. Because of this, labeled antibody may be employed in a lower concentration than immobilized antibody, and it is also preferable to employ a high specific activity for labeled antibody. For example, labeled antibody might be employed at a concentration of about 1-50 ng per assay, whereas immobilized antibody might have a concentration of 10-500 ng per assay per antibody. The labeled antibody might have a specific activity with, for instance, one

radioiodine per molecule, or as high as two or more radioiodines per molecule of antibody.

Of course, the specific concentrations of labeled and immobilized antibodies, the temperature and time of incubation as well as other assay conditions can be varied, depending on various factors including the concentration of antigen in the sample, the nature of the sample and the like. Those skilled in the art will be able to determine operative and optimal assay conditions for each determination by employing routine experimentation.

In carrying out the simultaneous immunometric assay on a sample containing a multivalent antigen, the process may comprise, in more detail: (a) simultaneously forming a mixture comprising the sample, together with the solid phase bound antibody and the soluble labeled antibody or antibodies; (b) incubating the mixture formed in step (a) for a time and under conditions sufficient to allow antigen in the sample to bind to both immobilized and labeled antibodies; (c) separating the solid phase immuno-adsorbent from the incubation mixture after the incubation; and (d) detecting either labeled antibody bound to the solid phase immuno-adsorbent or detecting labeled antibody not associated therewith. Other such steps as washing, stirring, shaking filtering and the like may of course be added to the assays, as is the custom or necessity for any particular situation.

There are many solid phase immuno-adsorbents which have been employed and which can be used in the present invention. Well-known immuno-adsorbents include nitrocellulose, glass, polystyrene, polypropylene, dextran, nylon and other materials; tubes, beads, and microtiter plates formed from or coated with such materials, and the like. The immobilized antibodies can be either covalently or physically bound to the solid phase immuno-adsorbent, by techniques such as covalent bonding via an amide or ester linkage, or by absorption. Those skilled in the art will know many other suitable solid phase immuno-adsorbents and methods for immobilizing antibodies thereon, or will be able to ascertain such, using no more than routine experimentation.

Kits

As indicated above, in accordance with the present invention, the antibodies of the invention as set forth above may be used in kits to diagnose a Gram positive infection. Such diagnostic kits are well known in the art and will generally be prepared so as to be suitable for determining the presence of epitopes or proteins that will bind to the antibodies of the invention. These diagnostic kits will generally include the antibodies of the invention along with suitable means for detecting binding by that antibody such as would be readily understood by one skilled in this art. For example, the means for detecting binding of the antibody may comprise a detectable label that is linked to said antibody. These kits can then be used in diagnostic methods to detect the presence of a bacterial infection wherein one obtains a biological sample suspected of having such an infection, such as a sample taken from an individual, for example, from one's blood, saliva, urine, cerebrospinal fluid, genitourinary tract, tissues, bone, muscle, cartilage, or skin, introduces to the sample one or more of the antibodies as set forth herein, and then determines if the antibodies bind to the sample which would indicate the presence of such microorganisms in the sample.

In addition, as set forth above, these kits can also be useful in methods of monitoring the level of antibodies or bacterial antigens in the serum of a human or animal patient. If monitoring the level of antigen is desired, the kit will include an antibody in accordance with the present invention as described above along with a means of determining the level

of binding to that antibody. When it is desired to measure the level of antibodies to Gram positive bacteria in a sample, the kit will preferably include an isolated protein or epitope (e.g., the A domain) such as described above, along with means for detecting binding of those antigens to antibodies present in the sample.

Treating or Protecting Against Infections

In accordance with the present invention, methods are provided for preventing or treating an infection caused by Gram positive bacteria which comprise administering an effective amount of the antibodies as described above to a human or animal patient in need of such treatment in amounts effective to treat or prevent the infection. Accordingly, in accordance with the invention, administration of an effective amount of the antibodies of the present invention in any of the conventional ways described above (e.g., topical, parenteral, intramuscular, etc.), and will thus provide an extremely useful method of treating or preventing bacterial infections in human or animal patients. As indicated above, by effective amount is meant that level of use, such as of an antibody titer, that will be sufficient to either prevent adherence of the bacteria, or to inhibit binding and colonization of such organisms to host cells and thus be useful in the treatment or prevention such infections. In addition, these antibodies also exhibit protective effects by a number of other mechanisms, including direct killing of the infectious microorganisms, increased opsonization, inhibition of morphological transition, etc., and thus an effective amount of antibodies will also include that amount by which any of the means to achieve a protective effect is obtained. As would be recognized by one of ordinary skill in this art, the level of antibody titer needed to be effective in treating or preventing infections will vary depending on the nature and condition of the patient, and/or the severity of the pre-existing infection.

Eliciting an Immune Response

In accordance with the present invention, a method is provided for eliciting an immunogenic reaction in a human or animal comprising administering to the human or animal an immunologically effective amount of a protein isolated using the bioinformatic method as described above, or a recombinantly produced version of such a protein, or an immunogenic fragment, region or epitope as described above so as to elicit an immunogenic response. As indicated above, an "immunogenic amount" of the antigen to be used in accordance with the invention to obtain an immunogenic reaction is intended to mean a nontoxic but sufficient amount of the agent, such that an immunogenic response will be elicited in the host so that the desired prophylactic or therapeutic effect is produced. Accordingly, the exact amount of the isolated protein that is required to elicit such a response will vary from subject to subject, depending on the species, age, and general condition of the subject, the severity of the condition being treated, the particular carrier or adjuvant being used and its mode of administration, and the like. The invention also contemplates methods of generating antibodies which recognize the proteins and epitopes as described above, and suitable methods of generating monoclonal and polyclonal antibodies are described in more detail above.

Coating Devices

In accordance with the invention, the antibodies and compositions as described above may also be utilized to treat or protect against outbreaks of bacterial infections on certain medical devices and other implanted materials such as prosthetic devices. Medical devices or polymeric biomaterials that may be advantageously coated with the antibodies and/or

compositions described herein include, but are not limited to, staples, sutures, replacement heart valves, cardiac assist devices, hard and soft contact lenses, intraocular lens implants (anterior chamber or posterior chamber), other implants such as corneal inlays, kerato-prostheses, vascular stents, epikeratophalia devices, glaucoma shunts, retinal staples, scleral buckles, dental prostheses, thyroplastic devices, laryngoplastic devices, vascular grafts, soft and hard tissue prostheses including, but not limited to, pumps, electrical devices including stimulators and recorders, auditory prostheses, pacemakers, artificial larynx, dental implants, mammary implants, penile implants, cranio/facial tendons, artificial joints, tendons, ligaments, menisci, and disks, artificial bones, artificial organs including artificial pancreas, artificial hearts, artificial limbs, and heart valves; stents, wires, guide wires, intravenous and central venous catheters, laser and balloon angioplasty devices, vascular and heart devices (tubes, catheters, balloons), ventricular assists, blood dialysis components, blood oxygenators, urethral/ureteral/urinary devices (Foley catheters, stents, tubes and balloons), airway catheters (endotracheal and tracheostomy tubes and cuffs), enteral feeding tubes (including nasogastric, intragastric and jejunal tubes), wound drainage tubes, tubes used to drain the body cavities such as the pleural, peritoneal, cranial, and pericardial cavities, blood bags, test tubes, blood collection tubes, vacutainers, syringes, needles, pipettes, pipette tips, and blood tubing.

It will be understood by those skilled in the art that the term "coated" or "coating", as used herein, means to apply the antibody or composition as defined above to a surface of the device, preferably an outer surface that would be exposed to an infection such as those caused by Gram positive bacteria. The surface of the device need not be entirely covered by the protein, antibody or active fragment.

As indicated above, the antibodies of the present invention, or active portions or fragments thereof, may also be useful for interfering with the physical interaction between bacteria responsible for infection and a mammalian host, and may also be useful in interfering with the ability of the bacteria to adhere to extracellular matrix proteins such as fibrinogen, collagen, laminin, etc. Accordingly, the antibodies of the invention may be useful both in treating patients and in preventing or reducing bacterial infections, or for reducing or eliminating infection and infestation of such organisms indwelling medical devices and prosthetics to make them safer for use.

In short, the antibodies of the present invention as described above can be extremely useful in detecting, treating or preventing infections by Gram positive bacteria in human and animal patients, or in preventing or reducing infection of medical devices and prosthesis that can be caused by such organisms. In particular, the present invention will be of importance in the treatment or prevention of such infections in highly susceptible groups such as premature newborns, AIDS and debilitated cancer patients, and are particularly frequent and severe after bone marrow transplantation.

EXAMPLES

The following examples are provided which exemplify aspects of the preferred embodiments of the present invention. It should be appreciated by those of skill in the art that the techniques disclosed in the examples which follow represent techniques discovered by the inventors to function well

in the practice of the invention, and thus can be considered to constitute preferred modes for its practice. However, those of skill in the art should, in light of the present disclosure, appreciate that many changes can be made in the specific embodiments which are disclosed and still obtain a like or similar result without departing from the spirit and scope of the invention.

Examples

Example 1

Method to Identify MSCRAMM® Proteins from Gram Positive Bacteria and Expression and Purification of their a Domains

A. Searching for LPXTG-Motif Containing Cell Wall Anchored Proteins in Annotate Genomes of Gram-Positive Bacteria.

1. Obtain the amino acid sequences of the entire genome of interest from web sites of sequencing centers. These sequences are stored in a local Silicon Graphics machine (SGI).

2. A local searchable database is established using the program format db obtained from NCBI and installed locally on the SGI.

3. LPXTG-motif containing proteins are identified using PHI-blast, which is obtained from NCBI and installed locally on the SGI. The PHI-blast search uses a degenerate LPXTG pattern L-P—X-[TSA]-[GANS], X being any amino acid. The templates for PHI-blast vary depend on the particular organism. For each organism, two known cell wall anchored proteins of *S. aureus* with no sequence homology were used as well as known cell wall anchored proteins from that particular organism if available.

4. The LPXTG-containing proteins obtained from PHI-blast were analyzed to select for those that contain typical features of LPXTG-motif containing cell wall anchored proteins: a signal peptide at the N-terminus, the LPXTG-motif being close to the C-terminus followed by a hydrophobic transmembrane segment, and several positively charged residues at the C-terminus. These are done as described below:

Signal peptide: we use the SignalP prediction server. The method has been described in "Identification of prokaryotic and eukaryotic signal peptides and prediction of their cleavage sites". Henrik Nielsen, Jacob Engelbrecht, Søren Brunak and Gunnar von Heijne, *Protein Engineering* 10, 1-6 (1997).

Location of LPXTG-motif: visual examination of the sequence.

A hydrophobic transmembrane segment after the LPXTG-motif: we use the TMHMM server for the prediction of transmembrane segments. Several other prediction web servers can also be used, among which are TMpred, DAS, and HMMTOP.

Positively charged residues at C-terminus: visual examination.

5. Sequences that contain the above features are putative LPXTG-motif containing cell wall anchored proteins.

6. The term "LPXTG" or "cell wall" are used to search for proteins that are annotated as cell wall anchored proteins in the genome of interest at TIGR website (comprehensive microbial resource, <http://www.tigr.org/tigr-scripts/CMR2/CMRHomePage.spl>).

B. Searching for LPXTG-Motif Containing Cell Wall Anchored Proteins in Un-Annotated Genomes of Gram-Positive Bacteria.

1. Obtain genome sequences from the web sites of sequencing centers. These sequences are stored in a local Silicon Graphics machine (SGI).

2. Gene prediction using the program Glimmer 2.0 from TIGR. This is facilitated by UNIX C shell scripts written in house.

3. The predicted genes are translated into amino acid sequences using a translation program written in house. This program is capable of translating large batch of sequences.

4. The translated amino acid sequences are formatted into a searchable database locally as in Section A.2. Subsequent analysis is as described in Section A.3-5.

C. Prediction of Immunoglobulin-Like (Ig-Like) Fold in Putative LPXTG-Motif Containing Cell Wall Anchored Proteins.

The amino acid sequences of putative LPXTG-motif containing cell wall anchored proteins are submitted to a Fold recognition web server 3D-PSSM. The method of prediction is described in Kelley L A, MacCallum R M & Sternberg M J E. Enhanced Genome Annotation using Structural Profiles in the Program 3D-PSSM. *J Mol Biol.* 2000 Jun. 2; 299(2):499-520

The output of 3D-PSSM gives a probability E value indicating the likelihood of the submitted sequence adopting a similar 3D structure as a published structure.

Proteins that have E value <0.25 to a published Ig-like fold structure, are considered containing predicted Ig-like folds. These should be considered MSCRAMM® proteins.

Accordingly, in accordance with the present invention, a bioinformatic approach was used to identify proteins with MSCRAMM®-like characteristics among Gram positive bacteria, particularly *Enterococcus faecalis*. Forty-two proteins with a putative C-terminal LPxTG cell wall sorting signal were identified in the *E. faecalis* genome. We then looked for structural similarities to MSCRAMM® proteins among LPxTG-anchored enterococcal proteins. Nine proteins were predicted to contain regions that adopt an immunoglobulin-like fold. The Ig fold-containing regions in FIG. 1 consist of several consecutive and overlapping matches to solved crystal structures (~150-500 aa) of the immunoglobulin superfamily (IgSF), which consist of one to four domains of equal size and Ig-type fold. The homologous Ig-fold regions cover most of the enterococcal proteins and may indicate a similar "beads-in-a-string" arrangement of consecutive modules that are found in fibronectin and other IgSF proteins (Leahy, 1996) (Sharma, 1999) (Hamburger, 1999) (Luo, 2000). A tandem repeat of Ig folded subdomains (N2 and N3) is found in the crystal structure of the fibrinogen-binding domain of ClfA. The full-length A domains of ClfA and the similarly structured ClfB consist of an additional N-terminal subdomain, N1 (Deivanayagam, 2002) (Perkins,-

2001). Based on sequence and secondary structure similarities, an analogous subdomain organization is also expected in other MSCRAMM® proteins including FnbpA, FnbpB, Ace and the Sdr proteins. The solved crystal structure of CNA minimum collagen-binding domain is made of a single Ig-type subdomain (N2) (Symersky, 1997) and the C-terminal repeat domains B1 and B2 each consist of a tandem repeat of Ig-folded subdomains (Deivanayagam, 2000). A similar modular structure is expected in the B3 and B4 repeats. Thus, a module structure of multiple Ig-folded units seems a general characteristic in the MSCRAMM® protein family. The length of the N-subdomains of MSCRAMM® proteins is typically ~150 aa suggesting that the large size of the A domains of EF1091 and EF0089 could accommodate more than three Ig-folded subdomains in their A domains.

Expression and Purification of Recombinant Enterococcal MSCRAMM® Protein Fragments

To further characterize the utility of this invention, the A-domains of EF1091, EF1092 and EF1093 proteins from *E. faecalis* as well as Efae 2926, Efae 2925 and Efae 2924 proteins from *E. faecium* were cloned, expressed and purified. In addition, EF1824 was cloned in two segments, EF1824AI (aa 43-819) and EF1824AII (aa 820-1829) because of the large size of the protein. EF1824AI was insoluble in *E. coli* cytoplasm and excluded from the assays. Bolded and underlined sequence represents the putative A-domains that were cloned.

EF1824AI: amino acid residues 43-819

(SEQ ID NO: 2)

QEQTAKEDVADSATSVGAIYSIEKAEKNFVITYASGKKAQISILNDHLFRYHLDP
TGKFEEYPTPNPKHVAKITAKTMADYGTQAFEQTNVTDGNGQFILENNGLKI
MFEKESALMKVLKDKKNQVILEETAPLSFKNDKATQTLKQSSQENYFGGGTQ
NGRFTHRGTAIQIVNTNWNVDGGVASPNPFYWSYTAGYGVVNTWPKGNFYD
GSHDPQKTTTTHEGTDFAFYFFNDSSAGILKDYELTGKPALMPEYGFYEAH
LNAYNRDYVVKVAEGTAGAVKFDGNFYKEYYQPGDLGNLNGTLESNGEKE
NYQFSARAVIDRYKKNMPLGWFLPNDGYGAGYQTDLSLDGVDVQNLKEFTFY
AQANGVEVGLWTSQNLHPADPKNPKKGERDIAKEVSVAGVKALKTDVAVWG
YGYSFGLNGVEDAANVFKETDGAVRPMIVSLDGWAGTQRHAGIWTGDQTC
GQWEYIRFHIPTYIGTSLSGQPNVGSMDMGIIFGGKNKEINIRDFQWKTFTPVQL
NMDGWSGNPKTPFAFDQEAIDLNRAYLKLKSMMPYNYSIKESVSDGLPMV
RAMALEFPNEGTAATKDSQYQYMWGNLLVAPIYNGNQDEAGNSIRDGIYLPD
EKQVWDLFTGEKYQGGVRLNGVKTPLWKVPVFKDGSIIIPMTNPNNPKKI
QRDQRSFLIYPNGTTSFNMYEDDGIISTSYEAGQSATTKINSQGPKNSEKGLDT
VTIEPTKGSYKDFVDERSTLDDLASEAPESVTAMVGGTEVTLKQ

EF1824AII: amino acid residues 820-1829

(SEQ ID NO: 3)

AANKEEFLAGTNLVYFDKFEQVNYQLSEASGEKLNQALSVKLAKQSVTAKDVQITVK
GFINKGTVDDGNTTVDDQLTIPANVAINEEKTTPSSLTLOWDQVTEATSVEVERDGTVF
GNIQTNTATFDGFSFLSEHTFRVRAVGKNGVSEWSEPIKGTQDDPYKETINQVKATS
NLPEQPGAELKLLTKDLSTGWHNTWSTGIANPSDGNFLSLKFDLGAEYQMDKLEVL
PRDNAGNGNLLQLYRTSKDGANWTEFSEPINWKQDALTKTIETKDQAYRFVEMKVL
KSVGNFGSGREMLFYKQPGTEGILHGDITNDGTIDENDAMSYRNYTGLESDVDFNGY
VEKGDNLKNGVVIDAYDISYVLRQLDGGIEIPDVEEIIAGGLSLAVVNEGKDTYLPDGLTT
FILKQDLKLNALSTKMSFDSSKFLVQGPATTNNTQOMENYSKYRKHNSDVENLYL
VLSNQGNKQLNGSMDLVTFKVKVKTETRVKRATTVQPLQFDMSQGLLVQGGFQQ
ATLSDFSVTVPTELVDKELLQALITLNQARVEKEYTPETWALFKPILDEAVAVLANEQA
TQTDVSAANLENLEKASQLEKMPDVANKADLEKAIQEGLAKKPSDGEFTEETKKVL
EESLAAQKVFQEKVTQEEIDQATKTLREAIAQLKEQPVAVDKETTLEQIAQARGRK
PEEGYQFTKETEKQLQEAIAQAAEIVAKETATKEEVSEALNALETAMAQLKEVPLVNK
DQLQEVVKRAQQVTPSEGHQFTASSLQELQKALLAAKNTLNKPAANQKMIDEAVAEI
TSAIDGLQEEVLVTDKKALEAMI AKAKAIKPSAGKEFTSESKARLTAIDQAEGLADKN
ARQEQIDIAEKNVKLTALDSLEQVLQTDKTKLKEKELLQKAEITLKPAGKQPTKASQEA
AEAIAKQAKALVEDPNATQEAVDKCLSLLSQAIEAMAEPISSNSTGNNGNHSTVSGTGG
VTSQKGTATGGTTTKTTTSGT

EF0089A: amino acid residues 36-1143

(SEQ ID NO: 4)

EEVNSDGQLTLGVEKQTSQEQMTLALQKKAQPVTOEYVHVHSANVSIKAAHWAAPN
NTRKIQVDDQKQIQIQLNQALADTLVLTNLNPTATEDVTFYSYQQQRALTLKGTGDTPT
ESTAITSPPAASANEGSTEEASTNSSVPRSEETVAATKAIKESKTTESITVPRVAGPT
DISDYFTGDETTIIDNFEDPIYLNPDGTPATPPYKEDVTHHWNFNWIPEDVREQMKAGD
YFEFQLPGNLKPNKPGSGDLVDAEGNVYGTYYTISEDGTVRFTFNERITSESDIHGDFSL
DTHLNDSDGRGPDGWDVIDIPTQEDLPPVVIPVDPTEQQIDKQGHFRTPNPSAITWTV
DINQAMKDQTNPTVTEWPTGNTFKSVKVELVMNLDGTLKEVGRELSPEDEYTVDKNG
NVTIKGDTNKAIRLEYQTTIDEAVIPDGGGDVFPKNHATLTSDDNPNGLDAEATVTATY
KMLDKRNIYDEANQEFTEINYNIGEQTIPKQAVITDMDGNLTFEIPDSLHLVSVT
FDDKGNVEVGAELVEGDKYKVVINGDGSFAIDFLHDVTVGAVKIDYKTKVPGVIEGVDVAV
NNRVVDVGTGQHSDDGTASQQNIKNTGAVDYQNSTIGWTLAVNQNNYLMENAVITDT
YEPVPGLTMPVNSLVKDTTGAQLTLGKDFMVEITRNADGETGFKVFSFIGAYAKTSD
AFHITYTTFDVTLELDANNPALDHYRNTAAIDWTDEAGNNHSEDSKPFKPLPAFDLNA
QKSGVYNAVTKETITWTIAVNLNNRLVDAFLTDFILTNQTYLAGSLKVIYEGNTKPDGVS
EKVKTPQLTYDIMEEPSEKNQNTWRVDFPNDSRTYVIEFKTSVDEKVIKESGSAYDNTA
SYTNQSSRDVTKVSIQHGGESVKKGGEYHKDDPDHVYWHVMINGAQSVLDDVVIT

-continued

DTSPSNQVLDPESLVIYGTNVTEGDTITPKDSVILEEGKDYTLEVTDDNETGQQKIVVKM
AHIEAPYYMEYRSLVTSAAAGSTDTVSNQVSI TGNSEVHVHDDNGDVVVDIDHSGGH
ATGTRGKIQLKKTAMDETTILAGAHFQIWDQAKTQVLRGTVDATGVITFGG

EF3023A: amino acid residues 26-1024

(SEQ ID NO: 5)

EEITDLFLQKEVTVYSGVEGGKIGENWKYPQFVGEKAVDGDDETRWSADKQDEQWLIY
DLGEVKNIGELVLQLHAESPVEILVSTDGESYQSI FKEENGKGGQPTKKYIDGNNVQA
RFVKYQQMKMWOHTNKQFYSSSIISFEAYEKRLPEAKLLTENLTISEKRKQQLAFEV
SPAGVDITEDQIEWSSSDPTIVTVDTQGNLTAVKSGEAKVTVKIKGTETISDIPVTVVAEN
KQYAEMLAKWKMLLGTQYDNDADVQYRAQIATESLALWQTLNQAADREYLWER
KPSDTVSAADYTTQFTNKKLALGYEPESELFEKPEVYDAIVKGFEMIDTKKYNQTYTT
GNWWDWQIGSAQPLTDTLLHDDLNTDAEKLKFTAPLMLYAKDPNIQWPIYRATG
ANLTDISITVLGTGLLLEDNORLVQVQEAVPSVLKSVSSGDGLYPDGLIOHGYPFYNG
SYGNELKGFGRIQITILQGSWEMNDPNISNLFNVVDKGYLQLMVNGKMPMSVSGRS
LSRAPETNPFTEFESGKETIANLTLIAKFAPENLRNDIYTSIQTWLQQSGSYHYFFKKP
RDFEALIDLKNNVNSASPAQATPMQSLNVYGSMDRVLQKNNEYAVGISMYSQRVGNY
EFNTENKKGWHTADGMLYLYNQDFAQFDEGYWATIDPYRLPGTTVDTRERLANGAYT
GKRSFQSWVSGSNNGQVASIGMFLDKSNEGMLVAKKSWFLDQGIINLGGGITGT
DASIETILDNRMIHQVEKLVNQGSKDKNSWISLSAANPLNNIGYVFPNSMNTLDVQIEE
RSGRYGDINEYFVNDKTYTNTFAKISKNYGKTVENGTYEYLVVQKTNEEIAALSKNKG
YTVLENTANLQAEAGNYVMNNTWNNQEIAGLYAYDPMVISEKIDNGVYRLTLANPL
QNNASVSI EFDKGI LEVVAADPEISVDQNIITLNSAGLNGSSRSIIVKTTPEVTKALEKLI
QEQ

EF2224A: amino acid residues 31-771

(SEQ ID NO: 6)

QEVTSDAEKTVEKDGKLVIGKIEDTSSQEDIKTVTYEVTNTRDVP IKDLILKQKNTNDSP I
KEVLDLTLSEERGPTSL EEQAKVETNEKDQTTDIKLLNLQPNSTRKITINGQITTKASNKL
LVSVLIEDNEKGTLVIDLPSKIDILADKESVSKKQETSETKVENQANETASSTNEMTATT
SNETKPEAGKAIESIQETALTQATESPEQPLKAQPTGFLVPPTPGRGFNTPIYQSVHK
GELFSTGNTNLKIANENTAAAQFLNTRGASSGYAINNFPLEFADVDNDPNTYNSRAY
IDLNGAKEIWAAGLFWASRYKGPAYGNTLSDDEEISAPVQPTTNGTVQVSPQRYHR
IDQDATNPGQRFQYNNNTGFSNYADVTSILQGDKSATGSYTLADIPMTSSLNGQYQYIN
FSGWSLFVVTKDQAKSKRAFSIYYGARGNAAGTNNFTMSNFLTAKQGNLDFIVTWF
VQGDYWTGDNAIKNSAGTWNISNTLNPVNNMNAATVTDNDEHMVDKYPGKFAF
DHPNFDLIDIDRMAIPEGVLNAGQNIINFRFTSSGDDYSTNAIGFAVNAETPEFEIKKEIV
EPKETYKVGETITRYVSLKNTKADSEAINSVSKDALDGRNLNYPGSLKIIISGPNSGEKTD
ASGDDQAEYDETNKQIIVRVNGATATQGGYKADTAETIYEFKARINERAKANELVFN
SATVEAVDILTSKAVNETSNIVEAKIADEQVT

EF1269A: amino acid residues 27-596

(SEQ ID NO: 7)

ETGYAQTEPTSTSETNQISATPNVVPRKQVGNIVTAIQITDKEGNPLGTINQYTDIYLRIE
FNLPDNTVNSGDTSYITLPEELRLEKNMTFNVDVDTGTVVAIAQTDVANKVTTLTYTQY
VENHANISGSLYFTSLIDFENVENESKIP IYVTVGEKIFAGDLDYQGEQDQVNEKFSKY
SWFIEDDPTEIYNVLRINPTGQTYTDLEVEDVLKTESLSYMKDTMKIERGQWTLDGNAI
WQFTPEEDITDQLAVQYGPDDRNFSVHFGNIGTNEYRITYKTKIDHLPKGETFTNYAK
LTENQTVVEEVEVSRVSTQGGGEANGEQYVVEIHKEDEAGQRLAGAEFELIRNSTNQ
VAKITTDQNGTAIVKGLKLDNYTLVETKAPTGYQLSQNKIPITPEDFGKNLVALKTVVNH
KISYQPVAAASFLAGVLLGKPLKDAEFQFELLDEKGTVLETVSNDTLGKIQFSPITFET
PGNYQYTTIREVNTQQTGVSYDTHNLQVQVTVVEALLGNLVATTQYDGGQVFTNHYTPE
KPIESTTPPTSGTTDTTNTSTTETTSITIEKQAIRNKE

EF1091: Nucleotide Sequence

(SEQ ID NO: 8)

0 ATGATAACAG ATGAGAATGA TAAAACGAAT ATTAATATCG AGTTAAATCT
50 TCTCAACCAA ACAGAGGAGC CATTACAACG AGAAATTC AA TTGAAAAATG
100 CACAGTTCAT GGATACTGCT GTAATTGAAA AAGACGGATA TTCTTACCAA
150 GTGACTAATG GTACGCTTTA TCTGACTTTG GACGCACAAG TAAAAAGCC
200 GGTACAGCTT TCGTTAGCTG TTGAGCAAAG TTCGCTTCAA ACAGCTCAGC
250 CACCTAAGTT ATTGTATGAA AACACGAAT ATGATGTTTC AGTTACTTCT
300 GAAAAATAA CAGTAGAGGA TTCTGCTAAA GAATCAACTG AACAGAAAA
350 AATAACTGTA CCAGAAAAA CGAAAAGAAC TAACAAAAAT GATTCGGCTC
400 CAGAAAAAC AGAACAGCCG ACCGCAACAG AAGAGGTAAC CAATCCATTT
450 CGAGAAGCAA GAATGGCGCC AGCTACTTTG AGAGCGAATC TGGCACTGCC
500 TTTAATTGCA CCACAATAA CGACGGATAA TTCTGGGACT TATCCGACAG
550 CTAATTGGCA GCCCAGAGGC AATCAAAATG TGTTAAACCA TCAAGGGAA
600 AAAGACGGTA GTGCACAATG GGACGGCAA ACGAGTTGGA ATGGGGACCC
650 TACTAATCGC ACAAAATCTT ATATTGAGTA TGGCGGTACA GGAGACCAAG
700 CCGATTATGC CATCCGAAA TATGCTAGAG AAACAACAAC ACCAGGGCTT
750 TTTGATGTAT ATCTTAATGT GCGTGGGAAT GTTCAGAAAG AAATCACGCC
800 ATTGGATTG GTCTTAGTGC TTGACTGGTC CGGTAGTATG AATGAAAAA
850 ATCGGATTGG TGAAGTCAA AAAGGAGTGA ACCGTTTGTG TGATACATTG
900 CGAGATAGCG GTATTACCAA TAACATCAAC ATGGGCTATG TTGGCTACTC
950 AAGTGACGGT TATAATAACA ACGCCATTCA AATGGGGCCG TTTGATACAG
1000 TCAAAAATCC AATTAATAA ATTACGCCAA GTAGCACTAG AGGAGGAACT
1050 TTCACTCAA AAGCATTAA AGATGCTGGT GATATGTTAG CAACGCCAAA
1100 TGGACATAAG AAAGTCATTG TACTTTTAA CGATGGCGTC CCAACCTTCT
1150 CTTATAAAGT GAGTCGAGTT CAAACAGAGG CGGATGGTCG CTTTTACGG

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1200 ACACAATTTA CGAATCGACA AGATCAAACA GGTAGCACTT CTTATATCTC
 1250 TGGTAGCTAT AATGCGCCAG ATCAAAACAA TATCAATAAA CGGATTAACA
 1300 GTACGTTTAT CGCCACGATA GGTGAGGCAA TGGTCTTAAA ACAACGTGGG
 1350 ATTGAAATAC ATGGATTGGG CATTCAAATG CAAAGCGATC CACGAGCTAA
 1400 TTTATCTAAA CAACAAGTTG AAGATAAAAT GCGTGAGATG GTGTCAGCCG
 1450 ATGAAAATGG AGACCTTTAT TATGAATCCG CGGATTATGC ACCAGACATF
 1500 TCTGATTATT TAGCGAAAAA AGCCGTTTCCG ATTTCCAGGAA CGGTTGTAAA
 1550 CGGAAAAGTA GTTGATCCAA TTGCTGAACC TTTTAAATAC GAGCCAAATA
 1600 CATTATCAAT GAAAAGTGTG GGTCTCTGTT AGGTTCAAAC ATTACCAGAA
 1650 GTGTCGCTAA CAGGCGCTAC AATTAATAGT AATGAGATTT ATTTGGGTAA
 1700 AGGCAAGAA ATTCAAATTC ATTATCAAGT ACGTATTCAA ACAGAGTCAG
 1750 AAAACTTCAA ACCTGATTTT TGGTATCAAA TGAATGGTCG GACAAACGTTT
 1800 CAGCCATTAG CCACGGCCCC TGAAAAAGTT GATTTTGGGG TTCTTCGGG
 1850 AAAAGCACCT GCGTGGAAGT TAAACGTGAA AAAAATCTGG GAAGAGTATG
 1900 ATCAAGACCC GACAAGTCGG CCAGATAATG TGATTTATGA AATTAGTAGA
 1950 AAGCAAGTAA CTGACACAGC CAACTGGCAA ACTGGGTATA TTAATTTATC
 2000 AAAACCAGAA AATGATACCA GCAATAGTTG GGAGCGCAA AATGTAACCC
 2050 AACTTTCCAA AACCGCGGAT GAAAGCTATC AAGAAGTCTT TGGGCTTCCC
 2100 CAATACAACA ATCAAGGACA AGCTTTCAT TATCAAACA CCGCTGAATT
 2150 AGCAGTTCCT GGTTCAGTC AAGAAAAAT CGACGATACT ACTTGAAAA
 2200 ACACGAAGCA GTTCAAGCCA TTAGATTTAA AAGTAATCAA AAATTCCTCC
 2250 TCAGGTGAGT AAAACTTAGT GGGAGCCGTC TTTGAATGA GTGGTAAAA
 2300 TGTTCAAACA ACATTAGTGG ACAATAAAGA TGGTAGCTAT TCCTTGCCAA
 2350 AAGATGTGCG CCTACAAAAA GGGGAACGCT ATACATTAAC TGAAGTAAAA
 2400 GCACCTGACG GACATGAGTT AGGCAAGAAA ACGACTTGGC AAATTGAGGT
 2450 GAGTGAGCAA GCAAAAGTAA GCATCGATGG ACAAGAAGTG ACCACCACAA
 2500 ATCAAGTTAT TCATTGGAA ATTGAAAATA AATTTTCTTC TTTGCCAATC
 2550 AGAATTAGAA AATACACCAT GCAAAATGGC AAACAAGTGA ACTTAGCAGA
 2600 GCGGACTTTT GCGTTGCAA GAAAAATGC TCAAGGAAGT TACCAAACCTG
 2650 TGGCAACTCA AAAAAAGAT ACTACAGGAT TGAGCTATTT TAAAAATAGT
 2700 GAACCTGGTG AGTATCGAAT GGTGGAACAA TCAGGACCAT TAGGCTACGA
 2750 CACTCTTGCT GGAAATTATG AATTTACTGT TGATAAATAT GGGAAATTC
 2800 ACTATGACAG CAAAAATAT GAAGAAAATG CGCCAGAATG GACACTGACA
 2850 CATCAAAATA ATTTGAAACC TTTTGACTTA ACAGTTAATA AAAAAAGCGA
 2900 TAATCAGACG CCACTTAAAG GAGCGAAATT CCGTTTAAACA GGACCAGATA
 2950 CGGATATTGA ATTACAAAAA GATGGCAAAG AACCGGATAC TTTTGTTTTT
 3000 GAAAACTTAA AACCGGGAA ATATGTTCTA ACAGAAACCT TTACGCCAGA
 3050 AGGATATCAG GGGTTAAAAG AACCAATCGA ATTAATAATT CGTGAAGATG
 3100 GTTCAGTCAC GATAGATGGG GAAAAAGTAG CAGATGTTTT AATTTCTGGA
 3150 GAGAAGAATA ATCAAATTAC TTTAGACGTT ACGAACCAAG CAAAGGTTCC
 3200 TTTACCTGAA ACTGGTGGCA TAGGACGCTT GTGGTTTTAC TTGATAGCGA
 3250 TTAGTACATT CGTGATAGCG GGTGTTTATC TCTTTATTAG ACGACCAGAA
 3300 GGGAGTGTG

EF1091 amino acid residues 63-1067

(SEQ ID NO: 9)

0 MITDENDKTN INIENLNLNQ TEQPLQREIQ LKNAOFMDTA VIEKDVYSYQ
 50 VTNGTLYLTL DAQVKKPVQL SLAVEQSSLQ TAQPPKLLYE NNEYDVSVT
 100 EKITVEDSAK ESTEPEKITV PENTKETNKN DSAPEKTEQP TATEEVTNPF
 150 AEARMAPATL RANLALPLIA PQYTTDNSGT YPTANWQPTG NONVNLHQGN
 200 KDGSAQNDGQ TSWNGDPTNR TNSYIEYGGT GDQADYAIRK YARETTTPGL
 250 FDVYLNVRGN VQKEITPLDL VLVVDWSGSM NENNRIGEVO KGVNRFVDTL
 300 ADSGITNNIN MGYVGYSSDG YNNNAIQMGF FDTVKNPIKN ITPSSTRGGT
 350 FTQKALRDAG DMLATPNGHK KVIIVLLTDGV PTFYSYKVSRY QTEADGRFYG
 400 TQFTNRQDQP GSTSYISGSY NAPDQNNINK RINSTFIATI GEAMVLKQRG
 450 IEIHGLGIQL QSDPRANLSK QQVEDKMRM VSADENGDLV YESADYAPDI
 500 SDYLAKRAVQ ISGTVVNGKV VDPIAEFPKY EPNTLSMKSV GPVQVQTLPE
 550 VSLTGATINS NEIYLGKQGE IQIHVQVRIQ TESENFKPDF WYQMNGRITF
 600 QPLATAPEKV DFGVPSGKAP GVKLVKKIW EYDQDPTSR PDNVIYEISR
 650 KQVTDATANWQ TGYIKLSKPE NDTSNWERK NVTQLSKTAD ESYQEVGLPL
 700 QYNNQQAQAFN YQTTRELAVP GYSQEKIDOT TWKNTKQFKP LDLKVIKNS
 750 SGEKNLVGAV FELSGKNVQT TLVDNKGDSY SLPKDVLQK GERYYLTFEVK
 800 APAGHELKPK TTWQIEVSEQ GKVSIDGQEV TTTNQVIPLE IENKFSLLPI
 850 RIRKYTMQNG KQVNLAEATF ALQRKNAQGS YQTVATQKTD TTGLSYFKIS
 900 EPGEYRMEVEQ SGPLGYDTLA GNYEFTVDKY GKIHYAGKNI EENAPEWTLT
 950 HQNLLKPFDL TVNKKADNQT PLKGAKFRLT GPDTDIELPK DGKETDTFVF
 1000 ENLKPGRYVL TETFTPEGYQ GLKEPIELII REDGSVTIDG EKVAVDLISG
 1050 EKNNOITLDV TNQAKVPLPE TGGIGRLWYF LIAISTFVIA GVYLFIRRPE
 1100 GSV

EF1092: Nucleotide Sequence

(SEQ ID NO: 10)

0 ATGAAAAACG CACGTTGGTT AAGTATTTGC GTCATGCTAC TCGCTCTTTT
 50 CGGGTTTTCA CAGCAAGCAT TAGCAGAGGC ATCGCAAGCA AGCGTTCAAG
 100 TTACGTTGCA CAAATTATTG TTCCCTGATG GTCAAATTACC AGAACAGCAG
 150 CAAAACACAG GGGGAAGAGG AACCGTCTTT CAAAATTATC GGGGCTTAAA
 200 TGACGCTACT TATCAAGTCT ATGATGTGAC GGATCCGTTT TATCAGCTTC
 250 GTTCTGAAGG AAAAACGGTC CAAGAGGCAC AGCGTCAATT AGCAGAAACC
 300 GGTGCAACAA ATAGAAAAAC GATCGCAGAA GATAAACAC AGACAATAAA
 350 TGGAGAAGAT GGAGTGGTT CTTTTTCATT AGCTAGCAAA GATTTCGACG

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400 AACGAGATAA AGCCTATTTA TTTGTTGAAG CGGAAGCACC AGAAGTGGTA
 450 AAGGAAAAAG CTAGCAACCT AGTAGTGATT TTGCCTGTTT AAGATCCACA
 500 AGGGCAATCG TTAACGCATA TTCATTTATA TCCAAAAAAT GAAGAAAAATG
 550 CCTATGACTT ACCACCACTT GAAAAACGG TACTCGATAA GCAACAAGGC
 600 TTTAATCAAG GAGAGCACAT TAACTATCAG TTAACGACTC AGATTCCAGC
 650 GAATATTTTA GGATATCAGG AATTCGGTTT GTCAGATAAG GCGGATACAA
 700 CGTTGACACT TTTACCAGAA TCAATTGAGG TAAAAGTGGC TGGAAAAACA
 750 GTTACTACAG GTTACACACT GACGACGCAA AAGCATGGAT TTACGCTTGA
 800 TTTTTCAATT AAAGACTTAC AAAACTTTGC AAATCAAACA ATGACTGTGT
 850 CGTATCAAAT GCGTTTAGAA AAGACCGCTG AACCTGCAC TGCAGTAAAC
 900 AACGAAGGAC AATTAGTCAC GGACAAACAT ACCTTGACTA AAAGAGCCAC
 950 AGTTCGTACA GCGGCAAGT CTTTTGTCAA AGTTGATAGT GAAAAATCGCA
 1000 AAATCACCTT GCCAGAGGCT GTTTTATCG TCAAAAAATCA AGCGGGGGAA
 1050 TACCTCAATG AAACAGCAA CGGGTATCGT TGGCAAAAAG AAAAAGCATT
 1100 AGCTAAAAAA TTCACGTCTA ATCAAGCCGG TGAATTTTCA GTTAAAGGCT
 1150 TAAAAGATGG CCAGTACTTC TTGGAAGAAA TCTCTGCACC AAAAGGTTAT
 1200 CTTCTGAATC AAACAGAAAT TCCTTTTACG GTGGGAAAAA ATCTTATGCT
 1250 AACGAACGGA CAACGAACAG CACCGTTACA TGTAATCAAT AAAAAAGTAA
 1300 AAGAGTCAGG CTTCTTACCA AAAACAAATG AAGAACGTTT TATTTGGTTG
 1350 ACGATTGCAG GCCTGCTAAT CATTGGGATG GTAGTCATTT GGCTATTTTA
 1400 TCAAAAACAA AAAAGAGGAG AGAGAAAA

EF1092 amino acid residues 28-438

(SEQ ID NO: 11)

0 MKNARWLSIC VMLLALFGFS QQALAFASQA SVQVTLHKLL FPDGQLPEQQ
 50 QNTGEEGTLN QNYRGLNDVT YQVYDVTDFF YQLRSEGKTV QEAQRQLAET
 100 GATNRKPIAE DKTQTINGED GVVVSLASK DSQQRDKAYL FVEAEAFEVY
 150 KEKASNLVVI LPVQDPQGS LTHIHLYPKN EENAYDLPL EKTVLDKQQ
 200 FNQGEHINYQ LTTQIPANIL GYQEFRLSDK ADTTLTLLPE SIEVKVAGKT
 250 VTTGYTLTTO KHGFTLDFSI KDLQNFANQT MIVSYQMRLE KTAEPDTAIN
 300 NEGQLVTDKH TLTKRATVRT GSKSFVKVDS ENAKITLPEA VFIKVNQAGE
 350 YLNETANGYR WQEKALAKK FTSNQAGEFS VKGLKDGQYF LEEISAPKGY
 400 LLNQTEIPFT VGKNSYATNG QRTAPLHVIN KVKESGFLP KTNERSIWL
 450 TIAGLLIIGM VVIWLFYOKO KRGERK

EF1093 (V583): Nucleotide Sequence

(SEQ ID NO: 12)

0 ATGAAGCAAT TAAAAAAGT TTGGTACACC GTTAGTACCT TGTACTAAT
 50 TTTGCCACTT TTCACAAGTG TATTAGGGAC AACCACTGCA TTTGCAGAAG
 100 AAAATGGGGA GAGCGCACAG CTCGTGATTC ACAAAGAA GAAATGACGGAT
 150 TTACCAGATC CGCTTATTCA AAATAGCGGG AAAGAAATGA GCGAGTTTGA
 200 TAAATATCAA GGAATGGCAG ATGTGACGTT TAGTATTAT AACGTGACGA
 250 ACGAATTTTA CGAGCAACGA GCGGCAGGCG CAAGCGTTGA TGCAGTAAA
 300 CAAGCTGTCC AAAGTTTAACT CCTGGGAAA CCTGTTGCTC AAGGAACCCAC
 350 CGATGCAAAAT GGGAAATGCA CTGTTCAAGT ACCTAAAAAA CAAAATGGTA
 400 AAGATGCAGT GTATACCATT AAAGAAGAAC CAAAAGAGGG TGTAGTTGCT
 450 GCTACGAATA TGGTGGTGGC GTTCCAGTT TACGAAATGA TCAAGCAAACT
 500 AGATGGTTCC TATAAATATG GAACAGAAGA ATTAGCGGTT GTTCATATTT
 550 ATCTAAAAA TGTGGTAGCC AATGATGGTA GTTACATGT GAAAAAAGTA
 600 GGAACTGCTG AAAATGAAAG ATTAATGGC GCAGAATTTG TTTATTTCTAA
 650 AAGCGAAGGC TCACCAGGCA CAGTAAAAA TATCCAAGGA GTCAAAGATG
 700 GATTATATAC ATGGACAACG GATAAAGAAC AAGCAAAACG CTTTATTACT
 750 GGGAAAAGTT ATGAAATTGG CGAAAATGAT TTCACAGAAG CAGAGAATGG
 800 AACGGGAGAA TTAACAGTTA AAAATCTTGA GGTGGTTCG TATATTTTAG
 850 AAGAAGTAAA AGCTCCAAAT AATGCAGAA TAATTGAAAA TCAACAAAA
 900 ACACCATTTA CRAATTGAAAC AAACAATCAA ACACCTGTTG AAAAAACAGT
 950 CAAAAATGAT ACCTCTAAAG TTGATAAAAC AACACCAAGC TTAGATGGTA
 1000 AAGATGTGGC AATTGGCGAA AAAATTTAAAT ATCAAATTTT TGTAAATAT
 1050 CCATTGGGGA TTGCAGACAA AGAAGGCGAC GCTAATAAAT ACGTCAAAT
 1100 CAATTTAGTT GATAAACATG ATGCAGCCTT AACTTTTATG AACGTGACTT
 1150 CTGGAGAGTA TGCTTATGCG TTATATGATG GGGATACAGT GATTGCTCCT
 1200 GAAAATTTATC AAGTGACTGA ACAAGCAAAT GGCTTCACTG TCGCCGTTAA
 1250 TCCAGCGTAT ATTCTACGC TAACACCAGG CGGCACACTA AAATTCGTTT
 1300 ACTTTATGCA TTTAAATGAA AAAGCAGATC CTACGAAAGG CTTTAAAAAT
 1350 GAGGCGAATG TTGATAACGG TCATACCGAC GACCAACAC CACCAACTGT
 1400 TGAAGTTGTG ACAGGTGGGA AACGTTTCAT TAAAGTCGAT GCGGATGTGA
 1450 CAGCGACACA AGCCTTGGCG GGAGCTTCCT TTGTCGTCCG TGATCAAAAC
 1500 AGCGACACAG CRAATTATTT GAAAATCGAT GAAACAAAG AAGCAGCAAC
 1550 TTGGGTGAAA ACAAAGCTG AAGCAACTAC TTTTACAACA ACGGCTGATG
 1600 GATTAGTTGA TATCAGAGG CTTAAATACG GTACCTATTA TTTAGAAGAA
 1650 ACTGTAGCTC CTGATGATTA TGTCTGTGTA ACAAATCGGA TTGAATTTGT
 1700 GGTCAATGCA CRAATCATATG GCACAACAGA AAACCTAGTT TCACCAGAAA
 1750 AAGTACCAAA CAAACACAAA GGTACCTTAC CTTCACAGG TGGCAAAGGA

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1800 ATCTACGTTT ACTTAGGAAG TGGCGCAGTC TTGCTACTTA TTGCAGGAGT
 1850 CTACTTTTGT AGACGTAGAA AAGAAAATGC T

EF1093 amino acid residues 33-592

(SEQ ID NO: 13)

0 MKQLKKVWYT VSTLLLILPL FTSVLGTTTA FAEENGESAQ LVIHKKKMTD
 50 LPDPLIQNSG KEMSEFDKYQ GLADVTFISIY NVTNEFYEQR AAGASVDAAK
 100 QAVQSLTPGK PVAQGTDDAN GNVTVQLPKK QNGKDAVYTI KEEPKEGVVA
 150 ATNMVVAFPV YEMIKQTDGS YKYGTEELAV VHIYPKNVVA NDGSLHVKKV
 200 GTAENEGLNG AEFVISKSEG SPGTVKYIQG VKDGLYTWT DKEQAKRFIT
 250 GKSYEIGEND FTEAENGTE LTVKNLEVGS YILEEVKAPN NAELIENQTK
 300 TPFTIEANNO TPVERTVKND TSKVDKTFPS LDGRDVAIGE KIRYQISVNI
 350 PLGIADKEGD ANKYVKFNLV DKHDAALTFD NVTSGEYAYA LYDGDTVIAP
 400 ENYQVTEQAN GFTVAVNPAY IPTLTPGGTL KFVYFMHLNE KADPTKGFKN
 450 EANVDNGHTD DQTPPTVEVV TGGKRFIKVD GDVTATQALA GASFVVRDQN
 500 SDTANYLKID ETTKAATWVK TKAEATFTT TADGLVDITG LKYGTYLLEE
 550 TVAPDDYVLL TNRIEFVVNE QSYGTTENLV SPEKVPNKHK GLPSTGGKG
 600 IYVVLGSGAV LLLIAGVYFA RRRKENA

Efae2926: Nucleotide Sequence

(SEQ ID NO: 14)

0 ATGACGACCA CAGGGAAGAA ACTGAAAGTT ATTTTCATGC TGATAATATT
 50 GAGTTTATCA AACTTTGTGC CATTATCTGC AATAGCAGAC ACTACAGATG
 100 ATCCAACAGT TTTAGAAACA ATTTTCAGCTG AAGTCATTTT GGATCAGTCT
 150 GGAATAAAGG CACCTGAACAT CAAGCTAAAT GCGAATAACA CCAGTGTGTA
 200 AAAGATAGAA AAAGAAATTG GTCTAGTCGA AAATTACTTA AGTGATGTGG
 250 AAAGAAAAGA AGGAGATGGC TATGCTTATC AGGTAAATAG CGGGAAAATF
 300 ACGTTGGAAA TCTCATCAA CACTAAACAA ACTATCGATC TGAGTTTTC
 350 AATCGATCCA GCACTTTACC ACAGCCAGGC AAACAAGCTG ATCGTCGATA
 400 ATAAAGAATA TGACATTATF GATGAGACAG AAAATAAGAA AGATACAGAT
 450 GTGTCAGTAC CAAAGCCAGA CGAAATAGAA GAAGAAATCAT CAAAAAGAAA
 500 CGAAAATCTC GTCAGCCCAT TTACATTGCC TACATTATCC TTGCCAGCTG
 550 TGAGTGTGCC ATCTAATCAA ACGATTCCTA CAGAATATAC AAACAGATGAT
 600 CAGGGCACTT ATCCTAAAGC CAGTTGGCAA CCTACAGGAA ATACAAATGT
 650 TCTTGATCAT CAAGGCAATA AAAACGGAAC AAATCAATGG GATGGTATAA
 700 ATPTCTGGAA TGGAGATCCT AATGATCGGA CCCATTCTGA TATCGAATAT
 750 GGAGGAACCG GTAATCAAGC AGACTATGCG ATACGAAAGT ATGCAAGGA
 800 AACAAGTACA CCCGGATTGT TTGATGTTTA TTTGAATGCT CGTGGAATG
 850 TACAAAAGA TATCACGCCT CTTGATCTCG TATTGGTTCG AGACTGGTCA
 900 GGAAATGTA ACGACAATA TCCGATCCGG GAAGTAAAGA TTGGTGTGCA
 950 TCGTTTTGTC GATACTTTAG CAGATAGCGG TATCACAGAC AAAATCAATA
 1000 TGGGATATGT CGGCTACTCA AGCGAAGGAT ATAGCTACAG TAACGGTGCA
 1050 GTACAGATGG GTTCAATTGA TTCAAGTAAA AATCAAGTAA AATCCATTAC
 1100 ACCTTCACGG ACAAATGGTG GTAATTTTAC ACAAAAGCA CTAAGAGATG
 1150 CAGGAAGCAT GCTATCCGTT CCAAATGGAC ATAAAAAGT GATCGTTTTG
 1200 CTGACGGATG GTGTACCAAC ATTTTCTTAT AAAGTACAGC GGGTACACGC
 1250 ACAATCAAGC AGCAATTATF ACGGAACTCA GTTTTCTAAT ACGCAAGATC
 1300 GGCCGGGAAA TACTTCTCTA ATCTCAAGAA TCTATGATGC ACCTGACCAA
 1350 AACAATCTAT CCAGAAGAA CGACAGTACG TTTATCGCAA CCATCGGAGA
 1400 AGCGATGGCA CTCAAAGAAC GAGGAATCGA AATACATGGT CTTGGCATCC
 1450 AACTTCAAGG CGATCCGGCA GCTGGTCTCT CAAAAAGCAGA AGTAGAGTCT
 1500 CGTATGCGAC AAATGGTTTC ATCAGATGAA AAAGCGGATC TTTACTATGA
 1550 ATCAGCTGAT CATGCAACAG ATATCTCTGA ATACCTAGCC AAAAAGCTG
 1600 TACAGATCTC AGCAACTGTA AGCAATGGAC AAATAAATGA TCCAATCGCA
 1650 GAACCATTCA TTTATCAGCC TGGTACACTT TCAGTCAAGA GTGTGGGGAC
 1700 AAGTCTTACA ACGGTCACTC CATCTATTTT CATAGAAAGA AATACCATCA
 1750 AGAGCAATCA GATCTATTTA GGAAAAGACC AAGAAATCCA AATCCATTAC
 1800 CAAGTGAGAA TCCAAAACAGA AAATGAGGAC TTCCATCCAA ATTTCTGGTA
 1850 TCAAATGAAC GGCAGGACAA CTTCCAGCC AAACATTGAT ACCAATGAAT
 1900 TAGCTGAATT CGGTATACCA TCTGCTAAAG CTCCCGGAGT CAGTCTTACC
 1950 ATCAAAAAGT TATGGGAAGA ATTTGACAAC AATCTAGCTG ATCGTCCAGA
 2000 TCAAGTFACT TTTGAGATTC AACGGGAACA TACGACAAAT GCTGCAGCTT
 2050 GGAAAACCGG ATATATTCGA ATCATTAAAC CAGCTAAAGA TACAACAAAT
 2100 ACGTGGGAAC GTGCAGACAT TGACAAAATA TCTGCAATA GCGGAGAAAG
 2150 TTATCAAGAG ATATTATCAC TACCTCAATA CAATAATCAA GGTCAAGCAT
 2200 TCAGTTACCA AACAATCAAA GAATTACCTG TACCAGGATA CGATTCTCAA
 2250 CAATAGATG CAATGACATG GAAAAATACT AAACAATTCA CACCGTTAAA
 2300 CTTGAAAATA ACGAAAATTT CCTCTACAGG TGAAAAGGAT CTTATTTGGC
 2350 CTGTTTTCAA ATTAACAGGA GATTCTATTG ATACTTTACT AACAGATCAT
 2400 GGCGACGGAA CCTATTCTCT TCCAGAAAAT GTCAAATGTC AAAAAGAAAT
 2450 GACCTATACG CTGACAGAAA CAAAAGCTCC AGAAGGGCAT GGATTAAGCA
 2500 AAAAGACTAC TTGGGAAATC AAGATCGCTT CTGATGGTAC GGTAACCATT
 2550 TGATGAAAAT CAAGTCACTAC TTCCGATGAT ACGATCCAGT TGACTATTGA
 2600 AAATCCTTTT GTTGAAGTTC CTGTAGCAGT ACGTAAGTAT GCGATGCAAG
 2650 GGACGGACAA AGAGATAAAT CTTAAAGGAG CAGCATTTTC CCTACAGAAA
 2700 AAAGAAGCAA ATGGTACTTA TCAGCCAATT GACAGCCAAA CAACGAATGA
 2750 AAAAGGTCTT GCCAGTTTTG ATTCCTCAC ACCTGGTAAA TATCGAGTCG
 2800 TTGAAACAGC TGGTCTCTCC GGATATGATA CTTCGCCGGG AAATATATGA
 2850 TTCCAAATCG ATAAATATGG AAAAATCATT TACACGGGAA AAAAATCCGA
 2900 GATGACAAAT AATGTATGGA CGCTCACTCA TCAAATCGA CTAAAAGCGT

-continued

2950 TTGATCTAAC GGTACACAAA AAAGAAGACA ACGGACAGAC ATTAAAAGGA
 3000 GCAAATTCAC GACTGCAGGG ACCAGAAATG GACTTAGAAT CGCCAAAAGA
 3050 TGGACAAGAA ACAGATACCT TTCTATTGCA AAATTTAAAA CCTGGAACTT
 3100 ATACGCTGAC CGAAACTTTT ACACCAGAAG GATACCAAGG TCTAAAAGAG
 3150 CCAGTTACTA TAGTTATACA CGAAGATGGG TCAATTCAGG TGGATGGACA
 3200 AGATCATGAA TCTGTCTGT CACCAGGAGC CAAAAACAAC CAGATTCTTT
 3250 TAGACATCAC GAATCAGGCA AAAGTACCAT TACCTGAAAC GGGAGGAATT
 3300 GGCCGTTTAC GAATCTATCT AGTAGGGATG ATTGGTTGTG CGTTTTCTAT
 3350 TTGGTATCTT TTTTGA AAAA AAGAAAGAGG GGGCAGC

Efae2926: amino acid residues 53-734

(SEQ ID NO: 15)

0 MTTTGKKLKV IFMLIILSLV NFVPLSAIAD TDDPTVLET ISAEVISDQS
 50 GKKALNIKLN ANNTSAEKIE KEIGLVENYL SDVERKEGDG YAYQVNSGKI
 100 LEISSNTKQ TIDLSPIDP ALYHSQANKL IVDNKEYDII DETENKDDTD
 150 VSVFKPDEIE EESKKNENS VSPFTLPTLS LPAVSVPSNQ TIPTEYTTDD
 200 QGTYPKASWO PTGNTNVLHD QGNKNGTNQW DGINSWNGDP NDRTHSYIEY
 250 GGTGNQADYA IRKYAKETST PGLFDVYLNA RGNVQKDITP LDLVLVVDWS
 300 GSNDNNRIG EVKIGVDRFV DFLADSGITD KINMGYVYGS SEGYSYNGA
 350 VOMGSFDSVK NOVKSITPSR TNGGTFTOKA LRDAGSMLSV PNGHKKVIVL
 400 LTGVPPTFSY KVQRVHAQSS SNYYGTQFSN TQDRPGNTSL ISRIYDAPDQ
 450 NLRSRIDST FIATIGEAMA LKERGIEIHG LGIQLQSDPA AGLSKAEVES
 500 RMQMVSSDE KGDLYESAD HATDISEYLA KKAQISATV SNGQINDPIA
 550 EPFIYQPTL SVKSVGTSPT TVTPSISIEG NTIKSNQIYL GKDQEIQIHY
 600 QVRIQTENED FHPNFYQMN GRITFQPNID TNELAEFGIP SAKAPGVSLH
 650 IKKLWEEFDN NLADRPDQVT FEIQREHTTN AAANKNGYIR IIPAKDNTN
 700 TWERADIDLK SANSGESYQE ILSLPQYNNQ GQAFSYOTIK ELPVPGYDSQ
 750 QIDAMTWKNT KQFTPLNLKI TKNSSTGEKD LIGAVFKLTG DSIDTLTLDH
 800 GDGTYSLPEN VKLQKEMTYT LETKAPEGH GLSKKTTWEI KIASDGTVTI
 850 DGKTVTTSDD TIQLTIENPF VEVPAVRKY AMQGTDEKIN LKGAAPSLQK
 900 KEANOTYQPI DSQTTNEKGL ASFDSLTPGK YRVVETAGPA GYDTPSPNYE
 950 FQIDKYGKII YTGKNTMTN NVWTLTHQNR LKAPDLTVHK KEDNGQTLKG
 1000 AKERLQGPDM DLESPKDGQE TDTFLFENLK PGTYTLTETF TPEGYQLKE
 1050 PVTIVIHEDG SIQVDGQDHE SVLSPGAKNN QISLDITNQA KVPLPETGGI
 1100 GRLGIYLVGM IGCAPFSIWYL FLKKERGGG

Efae2925: Nucleotide Sequence

(SEQ ID NO: 16)

0 ATGAAAAAAC TTGGTTGGCT TAGTATGTGT CTCTTCTTGT TACTATTTAA
 50 ACCAGCTTTT ACTCAGGTAG CAACAGAAAC AGAAACAGAA ATGGTTTACA
 100 TTACTTTTACA CAATTGCTT TCCCAAACG GCAACTGCC GAAAAATCAT
 150 CCAAATGACG GACCAAGAAA AGCTTTATTA CAAACGTATC GAGGATTAAA
 200 TGGTGTACACA TTCCAAGTTT ATGATGTGAC AGATTCTTTT TACCATCTAC
 250 GGGAAAAGGG CAAAACGGTA GAAGAAGCAC AAGCAGAGAT CGCAAAAAAC
 300 GGTGCCTCTT CCGGTATGTT TACCGCAGAA GCAACAAC TA CAACTCTTAA
 350 CAACGAAGAT GGTATCGCTT CTTTTCTCTT GGCCGCTAAA GATCAAGAAA
 400 AAAGAGATAA AGCGTATCTT TTCATTGAAT CCAAAGTACC AGAAGTCGTC
 450 AAAGAAAAGG CAGAGAATAT GGTAGTTGTT CTTCCTGTAC ATGGACAAAA
 500 CAATCAAAAA CTTTCAACTA TCCATTTGTA TCCTAAAAAT GAAGAAAACG
 550 ACTACCTCGA TCCACCTTTT GAGAAGGTAT TAGAAGAGCC TAGAAATGAT
 600 TTTACGATTG GTGAAAAAAT CACTTATTCC TTGCATACGA CAATTCCTGT
 650 AAATATCCTT GACTATCAAA AGTTCGAATT GTCAGATAGT GCGGATGAAG
 700 CATTAACGTT TTTACCTAAT AGTTTAAACGA TTTATCGAA TGGAGAAAAG
 750 CTGACAGAAG GCTTTGTGAT ACACAAGAAA CCTCACGGAT TTGATGTTTT
 800 ATTTTCGATC CCTTCGTGG AAAAATATGC TGGAAAAAAA CTGACCATTT
 850 CTTATCAGAT GCAGCTAAGC AGTACAGCAC AGGCGAACAA GGAATCAAC
 900 AACAACGGAA CACTGGATT TGGTTTTGGT GTCAGTACAA AGAAAAGTCTC
 950 TGTATATACA GGGAGTAAGC AATTTGTCAA AATCGAGACA AATAAACAG
 1000 ATAAACGATT AGCTGGCGCA GTATTCCTTA TTA AAAACA AGCAGGAAAT
 1050 TACCTCCAGC AAACAGCCAA CGGATACAAG TGGACAAAGA ACCAATCAGA
 1100 TGCGCTTCCG CTGATTTCCG ATAAAAATGG CGCTTTTCA ATTTCCGGGT
 1150 TGAAAACAGG AAGTTATCGA TAAAAAGAGA TCGAAGCACC TTCTGGTTAT
 1200 ATTTTAAAGT AAACAGAAAT TCCGTTTACC ATTTCAACTT TTCTTTCTGA
 1250 GGATAAAGG GCGGACAGTA TATTGAAAGT AGTCAATAAA AAAGAAAAATA
 1300 GCCGTCCATT TCTTCAAAA ACAACGAAA CGAAAAATAC ACTTTTAGGC
 1350 GTTGTTGGTA TGGTATTCGC AAGCTTTGCA ATCTGGTTGT TTATCAAAAA
 1400 AAGAACAGGA GTGAAAAAAT GA

Efae 2925: amino acid residues 30-429

(SEQ ID NO: 17)

0 MKKLGWLSMC LLLLLFKPAF TQVATETETE MVQITLHKLL FPNGQLPKNH
 50 PNDGQEKALL QYRGLNGVT FQVYDVTDSE YHLREKGTV EEAQAEIAKN
 100 GASSGMFTA EATTTLNNED GIASFSLAAK DQEKRDKAYL FIESKVFVV
 150 KEKAENMVVV LPVHGQNNQK LSTIHLYPKN EENDYPDPFF EKVLEEFPRND
 200 FTIGEKITYS LHTTIPVNIL DYQKFELSDS ADEALTFLPN SLTSSNGEK
 250 LTEGFVIHKK PHGFDVLSI PSLEKYAGKK LTISYQMLSL STAQANKEIN
 300 NGTLDGFGF VSTKRVSVYT GSKQFVKIET NKPDKRLAGA VFLIKNKAGN

-continued

350 YLOQTANGYK WTKNESDALH LISDKNGAFS ISGLKTGSYR LKEIEAPSGY
 400 ILSETEIPFT ISTFLESDKE ADSILKVVNKENSRRPLPK TNETKNTLLG
 450 VVGMVFASFA IWLFIKKRTG VKK

Efae 2924: Nucleotide sequence

(SEQ ID NO: 18)

0 ATGAAAAATC ATAAAAAAT AAACGTTATG TTAGGAGTCC TTTTCCTTAT
 50 TTTACCATTAC CTCACAAACA GCTTCGGCGC AAAAAAAGTG TTTGCAGAGG
 100 AGACAGCAGC TCAAGTCATC CTTTCATAAAA AGAAAAATGAC TGATTTACCC
 150 GATCCTTTAA TCCAAAACAG CGGGAAGAA ATGAGCGAAT TCGATCAATA
 200 CCAAGGATTA GCCGATATTT CATTTCAGT TTATAACGTC ACTCAAGAA
 250 TTTATGCGCA ACGAGATAAA GGAGCGTCCG TGGATGCAGC AAAACAAGCA
 300 GTCCAGTCTT TGACTCCTGG TACACCAGTT GCTTCAGGAA CGACAGATGC
 350 TGATGGAAAT GTCACCTTAT CTTTACCTAA AAAACAAAAT GGGAAAGATG
 400 CAGTCTACAC GATCAAGAA GAACCAAAAG ACGGAGTGTG AGCTGCCGCA
 450 AACATGGTTT TAGCTTTCCC TGTATATGAG ATGATCAAAC AAGCATATGG
 500 CTCTTATAAA TACGGGACAG AAGAACTAGA TACTATCCAT CTCTACCTA
 550 AAAATACAGT CGGTAATGAT GGAACGTGTA AAGTTACAAA AATCGGTACT
 600 GCCGAAAACG AAGCACTAAA TGGAGCAGAA TTTATTATTT CTAAGAAGA
 650 AGGAACACCA AGCGTCAAAA AATACATCCA AAGTGTCA CA GATGGATTGT
 700 AACTTGGAC AACTGATCAA ACCAAAGCCA AACATTTTCA TACTGGTCA
 750 TCTTATGACA TCGGCAACAA TGACTTTGCC GAGGCATCTA TTGAAAAGG
 800 CCAGTTGATC GTTAATCATT TAGAAGTTGG AAAATATAAT TTAGAAGAAG
 850 TAAAGTCCG TGATAATGCG GAAATGATG AAAAGCAAA AATCAGCCCT
 900 TTTGAGATCT TGGCAAATAG CCAAACACCA GTAGAAAAGA CCATCAAAA
 950 TGATACGTCT AAAGTTGATA AAACAACACC TCAATTGAAT GAAAAAGATG
 1000 TCGCAATCGG TGAATAAAT CAATATGAGA TTTCTGTCAA TATCCCATTA
 1050 GGTATCGCTG ATAAAGAAGG AACGCAAAAC AAGTACACAA CATTCAAAC
 1100 TATCGATACT CATGACGCTG CTTTAAACATT TGATAATGAT TCTTCAGGAA
 1150 CGTATGCTTA TGCTTATAT GATGGAAATA AAGAAATCGA CCCAGTAAAT
 1200 TATTTGTCA CTGAGCAAAAC AGACGGATTC ACGGTTTTCAG TTGATCCGAA
 1250 TTATATTCCT TCATTAACCTC CTGGCGGTAC ATTGAAATTC GTTACTATA
 1300 TGCATTTGAA CGAAAAGCA GATCCAAACA AAGGATTTTC TAACCAAGCA
 1350 AATGTCGATA ACGGGCATA CAAATGATCAA ACACCACCGT CAGTCGATG
 1400 CGTTACTGGG GGCAACGAT TTGTTAAAGT AGATGGTGC GTTACATCAG
 1450 ACCAAACACT TGCTGGAGCA GAATTCGTCG TTCGTGATCA AGATAGTGAC
 1500 ACAGCGAAAT ATTTATCGAT CGACCCATCC ACAAAGCCG TCAGCTGGGT
 1550 ATCGGCGAAA GAATCAGCAA CGGTTTTTAC AACCAAGT AACGGTTAA
 1600 TCGATGTGAC AGGTCTAAAA TATGGCAGT ACTATCTGGA AGAAAAGAAA
 1650 CGCCACAGAA AATATGTTCC ATTAACAAAC CGTGTAGCAT TTACTATCGA
 1700 TGAACAATCT TATGTAACAG CAGGACAGTT GATTCTCTCT GAAAAATAC
 1750 CAAATAAACA CAAAGGTACA CTTCTTCAA CAGGCGGTAA GGAATCTAT
 1800 GTGTATATCG GTGCAGGAGT AGTCCTTCTA CTGATTGCTG GACTGTACTT
 1850 TGCTAGACCG AAGCACGTC AGATTTAG

Efae 2924: amino acid residues 55-588

(SEQ ID NO: 19)

0 MKNHKINVM LGLVFLILPL LTNSFGAKKV FAEETAQVI LHKKMTDLP
 50 DPLIQNSGKE MSEFDQYQGL ADISFSVYV TQEFYQRDK GASVDAKQA
 100 VQSLTPGTFV ASGTTDADGN VTLSLPKKQN GKDAVYTIKE EPKDGVSAAA
 150 NMVLAFPVYE MIKQADGSYK YGTEELDTIH LYPKNTVND GTLKVKTIGT
 200 AENEALNGAE FIIKKEGTP SVKKYIQSVT DGLYTWTDQ TKAKHFTGH
 250 SYDIGNDFE EASIEKQLI VNHLEVGKYN LEEVKAPDNA EMIEKQITP
 300 FELLANSQTP VEKTIKNDTS KVDKTTPLN GKDVAIKEKI QYELSVNPL
 350 GIADKEGTQN KYTTFKLIDT HDAALTFDND SSGTYAYALY DGNKEIDPVN
 400 YSVTEQTDGF TVSDPNYIP SLTPGGTLKF VYMHLEKA DPTKGFNQ
 450 NVDNGHTNDQ TSPSDVVTG GKRFRKVDGD VTSQTLGAG EFVVRDQSD
 500 TAKYLSIDPS TRAVSWVSAK ESATVFTTS NGLIDVTGLK YGTYLEETK
 550 APEKYVPLTN RVAFTIDEQS YVTAGQLISP EKIPNKHKT LPSTGGKIY
 600 VYIGAGVLL LIAGLYFARR KHSQI

Protein Expression and Purification

Using PCR (the oligonucleotides used in the PCR reaction are shown in Table 3), the A domains from EF0089, EF1091, EF1092, EF1093, EF1099, EF1269, EF1824, EF2224, and EF3023 were amplified from *E. faecalis* V583 or *E. faecalis* EF1 (EF1099) genomic DNA and subcloned into the *E. coli* expression vector pQE-30 (Qiagen). One liter culture of *E. coli* M15(pREP4) cultures harboring appropriate pQE-30 based constructs were grown to OD₆₀₀=0.6 with an initial 2% inoculation from overnight cultures. After 2-3 h induction with 0.4 mM isopropyl-beta-d-thiogalactoside (IPTG), cells were collected with centrifugation, resuspended in 10 mM Tris-Cl, 100 mM NaCl, pH 7.9 and stored at -80 C.

To lyse the cells and release the expressed protein, cells were passed twice through French Press with a gauge pres-

sure setting at 1200 PSI to give an estimated internal cell pressure of 20,000 PSI. The lysate was centrifuged at RCF_{max} of 165,000xg and the supernatant was filtered through a 0.45 μm filter. The volume was adjusted to 15 ml with 10 mM Tris-Cl, 100 mM NaCl, pH 7.9 and 0.2 M imidazole in the same buffer was added to increase the imidazole concentration to 6.5 mM in order to minimize non-specific binding. The sample was loaded to a nickel affinity chromatography column (HiTrap chelating, Pharmacia) connected to an FPLC system (Pharmacia) and previously equilibrated with 10 mM Tris-Cl, 100 mM NaCl, pH 7.9. Bound protein was eluted with a linear gradient of 0-100 mM imidazole in 10 mM Tris-Cl, 100 mM NaCl, pH 7.9 over 100-200 ml. Protein-containing fractions were analyzed in SDS-PAGE (FIG. 2) and dialyzed against 25 mM Tris-Cl, 1 mM EDTA, pH 6.5-9

(depending on pI of protein purified) before applying the samples to an ion-exchange column (HiTrap Q, Pharmacia) for further purification. Bound protein was eluted with a linear gradient of 0-0.5 M NaCl in 25 mM Tris-Cl, 1 mM EDTA, pH 6.5-9 over 100 ml. Finally, protein samples were dialyzed extensively against PBS and stored at +4° C.

Alternatively EF1091, EF1092, and EF1093 were expressed in shake flasks or in bioreactors, the cells were harvested by centrifugation and the cell paste frozen at -80° C. Cells were lysed in 1xPBS (10 mL of buffer/1 g of cell paste) using 2 passes through a microfluidizer at 10,000 psi. Lysed cells were spun down at 17,000 rpm for 30 minutes to remove cell debris. Supernatant was passed over a 5-mL HiTrap Chelating (Pharmacia) column charged with 0.1M NiCl₂. After loading, the column was washed with 5 column volumes of 10 mM Tris, pH 8.0, 100 mM NaCl (Buffer A). Protein was eluted using a 0-100% gradient of 10 mM Tris, pH 8.0, 100 mM NaCl, 500 mM imidazole (Buffer B). Protein containing fractions were dialyzed in 1xPBS.

Example 3

MSCRAMM® Genes Common to *E. faecalis* and *E. faecium* PCR Analysis

Primers for flanking regions of sequences above were used to amplify 1 µg genomic DNA from each *E. faecalis* strain. PCR products from 5 *E. faecalis* strains in Table 1 were sequenced and compared to the TIGR database sequence. Primers used to amplify the enterococcal MSCRAMM® A-domain gene products are shown below.

| Protein | 5' Primer | 3' Primer |
|----------|------------------------------------------------------------|-------------------------------------------------------------|
| ACE40 | GAATTGAGCAAAAAGTTCAATC G (SEQ ID NO: 44) | GTCTGTCTTTTCACTTGTTTC TGTTG (SEQ ID NO: 51) |
| EF1091 | CAAGTAAAAAGCCGGTACAG C (SEQ ID NO: 45) | AAAGGAACCTTTGCTTGGTTC SEQ ID NO: 52) |
| EF1092 | TCGCAAGCAAGCGTTCAAG (SEQ ID NO: 46) | AAGCTGAGTCTTTACTTTT TTATTG SEQ ID NO: 53) |
| EF1093 | GAGAGCGCACAGCTCGTG (SEQ ID NO: 47) | GGTACCTTTGTGTTTGTTTGG TAC SEQ ID NO: 54) |
| Efae2924 | CGGGATCCAAAACAGCGGGA AAGAAATGAGCGA (SEQ ID NO: 48) | CCCAAGCTTTCATGTACCTTT GTGTTTATTGG (SEQ ID NO: 55) |
| Efae2925 | CGGGATCCGAAATGGTTCAGA TTACTTTACAC (SEQ ID NO: 49) | TCTGCAGTCAATTGACTACT TTCAATATACTGTC (SEQ ID NO: 56) |
| Efae2926 | CGGGATCCAAAGCACTGAACA TCAAGCTAAATGCG (SEQ ID NO: 50) | CCCAAGCTTTCAGAATGCTTG ACCTTGATTATTGTA (SEQ ID NO: 57) |

Homology Among Enterococcal MSCRAMM® Proteins

A blastp search was performed using the AA sequence listed above with the NCBI search engine. The accession number is given for each putative homologue found. Both percent identity and similarity refer to the percentage of AA that match the query sequence exactly while similarity includes conservative AA changes in the matching calculation.

TABLE 4

| Comparison of <i>E. faecium</i> homologues of <i>E. faecalis</i> MSCRAMM® protein | | | | |
|-----------------------------------------------------------------------------------|------------------------------------------|------------------|------------|--------------|
| <i>E. faecalis</i> Protein | <i>E. faecium</i> Protein Homologue Name | Accession Number | % Identity | % Similarity |
| EF1091 | Efae2926 | 00038011 | 60 | 75 |
| EF1092 | Efae2925 | 00038010 | 48 | 63 |
| EF1093 | Efae2924 | 00038009 | 74 | 83 |

The "A" domain amino acid sequence from each *E. faecalis* MSCRAMM® protein was used as a query in a blastp search. Results shown were scored by NCBI computers. Identity is calculated as exact matches between the subject and query sequences while similarity also includes conservative changes in sequence at the same position.

Example 4

Additional Gram Positive Amino Acid Sequences Predicted to Be MSCRAMM® Proteins

List of LPXTG-motif containing cell wall anchored proteins that contain predicted immunoglobulin-like fold. The sequencing center for each genome is indicated in the parenthesis. All the sequence except for those of CNA from *S. aureus* and *Staphylococcus epidermidis* can be obtained from TIGR website, comprehensive microbial resource section. The *S. epidermidis* RP64A genome is not annotated. However, the nucleotide coordinates of the genes encoding the listed *S. epidermidis* proteins can be obtained through TIGR website.

Streptococcus pneumoniae TIGR4 (TIGR)
SP0368
SP0462
SP0463
SP0464

Enterococcus faecalis V583 (TIGR)
EF2224
EF1099
EF1092
EF3023
EF1269
EF0089
EF1824
EF1091
EF1093
EF1075
EF1074
EF1651

Streptococcus mutans UA159 (University of Oklahoma)
SMU.610
SMU.987
SMU.63c

Staphylococcus aureus N315 (Juntendo University, Japan)
SA2447
SA2290
SA2291
SA2423
SA0742
SA0519
SA0520
SA0521

-continued

Bacillus anthracis Ames (TIGR)
BA0871
BA5258

Staphylococcus epidermidis strain RP62A (TIGR)
>SERP_GSE_14.6.AA 2402 residues

(SEQ ID NO: 20)

mknkqgflpnllynkygirklsagtaslligatlvfgingqvkaaednivi
sqngdntkndessskelvkseddktsststsdtnlesefdqnnmpssiee
stnrndedltnqrstetetekdthvksadtqtnnetnknkdnatntntes
isdestyqsddskttqhdnsntnqdtqstlnpntskessnkdeatsptpke
stsiektlnlsndanhqttdevnhsdsdntmstnpndtenedtqtlshd
espsqsdnftgftnlmatplnlrnrnprlnllaatedtkpktykknps
eysyllndlgydatvksensdlrhagisqsgdntgsviklnltkwlslqs
dfvnggkvnisafsqsfdytqi esitlndvkmtdttnngqnwsapingstvr
sgligsvtnhdivitlknstlsslgyennkpvyltwtvndgaiaees
iqvasitptltdskapntiqksdftagrmtnkikydsqnsiksivhtflpxn
enflqtdyravlyikeqvnkelipyidpnsvklyvsdpdgnpisdqryvn
gsidndglfdesskineisiknntsgqlsnartsldrnvfgtlgqrsy
tkaisnktqkvnleiepikieatdmsgavtnkvegplagmtfdeatnti
sgtpevsydyitvtvttdegnsetttftidvedttkptvesvadqteq
nteiepikieatdmsgavtnkvegplagmtfdeatntisgtpevsydy
itvtvttdegnsetiftidvedttkptvesiaggtqevnteiepikiea
kdnsqgtvtnkvegplagmtfdeatntisgtpevsydyvtvttdegn
setttftievkdtkptvesvadqteqnteiepikieardnsgqavtnk
vdglpdgvtfdeatntisgtpevsydyitvtvttdegnsetttftieve
dttkptvenvadqteqnteiepitie sednsgqvtvtnkvegplagmtf
ettntisgtpevsydyitvtvttdegnsetttftievdedttkptvenva
gdglpdgvtfdeatntisgtpevsydyitvtvttdegnsetttftieve
evgsyvtvtvttdegnsetttftidvedttkptvesvadqteqnteie
pitie sednsgqavtnkvegplagmtfdeatntisgtpevsydyvtvtt
desgnsetttftidvedttkptvesvadqteqnteiepitieatdmsg
qvtvtnkvegplagmtfdeatntisgtpevsydyitvtvttdegnset
ftinvedttkptvediadqteqnteiepikieatdmsgqavtnkvegpl
dgvtfdeatntisgtpevsydyitvtvttdegnsetttftidvedttk
tvesvvdqteqnteiepitieatdmsgqavtnkvegplagmtfdeatnt
isgtpevsydyitvtvttdegnsetiftidvedttkptvesiaggtqev
nteiepikieatdmsgqavtnkvegplagmtfdeatntisgtpevsydy
itvtvttdegnsetttftidvedttkptvesvadqteqnteiepitie
sednsgqavtnkvegplagmtfdeatntisgtpevsydyitvtvttdegn
setttftidvedttkptvesiaggtqevnteiepikieatdmsgqavtn
kvegplagmtfdeatntisgtpevsydyitvtvttdegnsetttftinv
edttkptvesvadqteqnteiepikieardnsgqavtnkvegplagmtf
deatntisgtpevsydyitvtvttdegnsetttftidvedttkptved
idqteintemtpikieatdmsgqavtnkvegplagmtfdeatntisgt
sevgkylititidkdgnatattltinvidttpeqptinkvtenstevn
grgepgtvtvvtfdpdknvegkvdsgnyhiqipsettlkqgqplqia
dkagnkseattntvndt tapeqptinkvtenstevsgrgepgtvtvvt
dgnkvegkvdsgnyhiqipsettlkqgqplqivkvvdeegnvspsitmv
qkednkselstvtgtvknkskshkaseqsgyhnksiekinkvnpkpt
ivekdmsydyrsykdnsknksatfeqqnvsdinnnqysrknvnpq
kksrkneinkdplqtgeenfknstlfgtlvaslgalllffkrkkkdende
ke

>SERP_GSE_2_50.AA 892 residues

(SEQ ID NO: 21)

lfglhneakaeeentvqdvksndmdelssndqssneekndvinnsqsi
ntdddngikkeetnsndaienskditqstnvdeneatflqktpdqntq
lkeevvkepssvessnsmtdaqppshsttineseasigtndneersvdf
anskieenteenkeentieqgnkvrreditsqpskyndekiesnqdel
lnlpineyenkrvrlstsaqpskrvtvnqlaaeggsnvnhlikvtdq
itegyddsdgiikahdaenliydvtfvddkvsqgdmtvndkntvpsd
ltdsfaipkikdngseiatatgydntnkqitytftdyvdkyenikahlk
tsyidkksvppnntkldveyktalssvntkityevqkpenrntanlqsmf
tnidkntvegtiyinlprysaketnvnisqngdegstiidstilkvy
kvgdnqnlpdsnriydyseyedvntndyaqlgnndvniinfngidspyi
kvisydpnkddyttiqgtvtmqttineytfgefrtasdyntiafstsgg
gggdllppektiykigydvvedvdkdqigntndnekplsnvltltpdgt
ksvrtdeegkyqfdglkngltykitfetpegypptlkhsgtntpaldegn
svvwtinqqdmtdisgyfypkyislynyvydtnkdgicqgdekigsv
kvtldkengniisttttdengkyqfdlnsgnyivhfdkpsgmtqtttds
gddeeqdadgeevhvtitdhdfsidngyaddsdssdsdsdsdsds
sdsdsdsdsdsdsdsdsdsdsdsdsdsdsdsdsdsdsdsdsdsdskt

-continued

dklpdgtanedhdskgtilgalgagllgkrrkrkrknkn

>SERP_GSE_9_28.AA 1973 residues

(SEQ ID NO: 22)

mkenkrknldkntrfssirkygygyatstvaigfiiiscfseakadsdk
heikshqqsmtnhlttlpsdnqentsnnefnrnhdishslslnksiqmde
lkklklykainlndkteesiklfdqsdvqaeslinnpsqsgqhdafykh
flnsagklrkketvsikheresesntyrlgdevrsqtfshirhkrnavsfr
nadqnlstqplkaneinpei qngnfsqvsggplptsskrlttvtnvndv
hsystdnppeypmfytttavnypnfmsngnpygvlgrttdgwnrvnid
skvagiyqdidvvpgselnvfnistpsvfdsgaagaklkisnveqnrvlf
dserlmgmpyptgklsamvnpndinrvrisflpvsstgrvsvqrssreh
gfgdnssyyhgsvsdvrinsgsvvkvvtqreyttrpnssndtfarati
nlsvenkghngskdtyyevilpqnsrlstrggsgnynnatnklsirdn
lnpgdrdisytdvfesspklinalnahlllyktnatfrngdqgrtgdniv
dlqsiiallmkdvlelelneidkfrldneadftidswsalqekmtegg
ilneqqnqvalenqasqetinnvtqslleilknnlkyktpsqpiiksnqi
pnitispadkacklittiyqntdnesasiqgnlnnqslvlnnpiqieidm
qtglvtidykavvpsvvgandktgnsdasaesritmpraekatpive
aneervnvviapngeatqiaikyrtpdgqeatlvaskngsswtlnkqidy
vniesngkvtigyqavqpeveiatetkgnsdasaesritmpraekatp
spiveaneehvntviapngeatqiaikyrtpdgqeatliaskngsswtln
kqidvnieensgkvtigyqavqleseviatetkgnsdasaesritmrlk
eatpshpiveaneehvntviapngeatqiaikyrtpdgqeatlvasknes
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>lcl|SEPN_5_124.AA 10203 residues

(SEQ ID NO: 23)

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GNNQNI PSEQLVSAQQLEKALELARTLPQ
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ASNRWRYNLRPRRNILRASDVQGNAYIT
KRLKDGQWIDLFNHNHKGHEMYMYWFLGSLDQPTPTGPVTFITINRDGSS
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LNLKLIKVEVNAPTGNRRVYRVSTYNTLND
EINKLIKQAFKAANSGLNLDNDITVSNFDFHRNVSSTVTRKGDLIKEF
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-continued

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 TKQALQQLIYAETS LNGFERLNHARPRALE
 YIKSLEKINNAQKSALEDKVTSQSHDLLELEHIVNEGTLNNDIMGELANA
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 IQMDDARALNGIERLKDAQTKAHNDIKD
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 RDAI NNNTDLTPSQKAHALADIDKTEKDALQHIENSNSIDDINNKEHAF
 NTLAHI I IWDTDQQLVFEPELSELQNALV
 TSEVVVHRDETTISLESII GAMTLTDELKVNIVSLPNTDKVADHLTAKVKV
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 IETVKRTDFEEDQFPDKRFTLNKAKKDI
 TDVNTQIQNGFKEIETIKGLT SNEKTQDPKQLTALQKEFLEKVEHAHNLV
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 DVHKALQGI EQILKVVNSIINQSFNDSLH
 NFNYLHSKFDARLREKDVANHVQETEFKEVLKGTGVEPGKINKETQQPK
 LHKNDNDSL FKHLDVDFGKTVGVI TLTGLL
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>Icl|SEPN_8_63_AA 1973 residues

(SEQ ID NO: 24)

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 KETVSIKHERSESNTYRLGDEVRSTQFSHIRHKRNAVSRNADQSNLST
 PLKANEINPEIQNGNFSQVSGGLPTSSKR
 LTVVTVNVDNWSYS TDPNPEYPMFYT TAVNYPNFMNSGNAPYGVILGRT
 TDGWRNRVIDSKVAGIYQDIDVVPGESELNV
 NFI STSPVFSGGAAGAKLKI SNVEQNRVLFDSRLNGMGPYPTGKLSAMVN
 IPNDINRVRISFLPVSSGRVSVQRSSREH
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 NLSVENKGNHNSKDTTYEVILPQNSRLIST
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 DCSGVKCNLKNQIKTQLPDTGYSDKASKSNILSVLLLGFGLSYSRKRK
 EKQ

Example 5

Immunization Strategies for Antibody Production Using Three Representative Enterococcal MSCRAMM® Proteins

Purified EF1091, EF1092, and EF1093 proteins were used to generate a panel of murine antibodies. Briefly, a group of Balb/C mice received a series of subcutaneous immunizations of 1-10 mg of protein in solution or mixed with adjuvant as described below in Table 5:

TABLE 5

| Immunization Scheme | | | | |
|---------------------|-----|-------------|-----------------|----------|
| Conventional | | | | |
| Injection | Day | Amount (µg) | Route | Adjuvant |
| Primary | 0 | 5 | Subcutaneous | FCA |
| Boost #1 | 14 | 1 | Intraperitoneal | RIBI |
| Boost #2 | 28 | 1 | Intraperitoneal | RIBI |
| Boost #3 | 42 | 1 | Intraperitoneal | RIBI |

At the time of sacrifice serum was collected and titered in ELISA assays against MSCRAMM® proteins ACE, EF1091, EF1092 and EF1093 (Table 6).

Serum ELISA

Immulon 2-HB high protein binding 96 well plates were coated with 100 ng/well of the purified A-domains of EF1091, EF1092 or EF1093 and incubated overnight at 2-8° C. Plates were washed four times (350 µl/well) with PBS/0.5% Tween 20 using the Skatron Skanwasher plate washer and then blocked with 1% bovine serum albumin (BSA) solution, 200 µl/well for 1-2 hour at room temperature. Following incubation, the plates were washed as before and 100 it of 1xPBS, 0.05% Tween 20, 0.1% BSA buffer was added to

each well of rows B-H of the 96-well plate. The negative control serum (preimmune Balb/C serum) and hyperimmune samples were then diluted 1:100 in 1×PBS, 0.05% Tween 20, 0.1% BSA buffer. 200 μ l of negative control serum was added in duplicate to wells A1 and A2 of the 96-well plate and 200 μ l of each diluted hyperimmune test serum were added in duplicate to wells A3 to A12. Two-fold serial dilutions were performed down the plate ending with Row H with the remaining 100 μ l being discarded. The plates were incubated for 1 hour at room temperature. The plates were again washed as before followed by the addition of 1:5000 dilution of a secondary antibody solution, Goat anti-mouse IgG (whole molecule)-AP conjugate (Sigma Cat. A-5153), to each well (100 μ l/well) and incubated for 1 hour at room temperature. Following incubation, the plates were washed 4 times (350 μ l/well) with PBS/0.5% Tween 20. The developing solution, 1 mg/ml 4-nitrophenyl phosphate (pNPP) in 1M Diethanolamine, pH9.8, 0.5 mM MgCl₂, was added to each well (100 μ l/well) and the plates incubated at 37° C. for 30 minutes. After incubation, the absorbance (A405_{nm}) of each well was measured using the Spectra MAX 190 plate reader (Molecular Devices Corp., Sunnyvale, Calif.). The data was analyzed using SOFTmax Pro v.3.1.2. software (Molecular Devices Corp.) The dilution of the hyperimmune sera where the absorbance was 2-fold above the negative control serum absorbance was used as the titre for that hyperimmune serum sample.

TABLE 6

| Antibody Titer at Sacrifice | |
|-----------------------------|---------------------------|
| Antigen | Polyclonal Antibody Titre |
| EF1091 | >12,800 |
| EF1092 | >12,800 |
| EF1093 | >12,800 |

Example 6

Antibody Reactivity Against *E. faecalis* MSCRAMM® Proteins

Antisera derived from Balb/c mice (as described in Example 3) was used to identify EF1091, EF1092 or EF1093 natively expressed on the surface of *E. faecalis* strains.

Flow Cytometry Analysis—Whole Cell Staining

Bacterial samples (Table 7) were collected, washed and incubated with polyclonal antisera or pre-immune sera (control) at a dilution of 1:2000 after blocking with rabbit IgG (50 mg/ml). Following incubation with sera, bacterial cells were incubated with Goat-F_{(ab')₂}-Anti-Mouse-F_{(ab')₂}-FITC which served as the detection antibody. After antibody labeling, bacterial cells were aspirated through the FACScaliber flow cytometer to analyze fluorescence emission (excitation: 488, emission: 570). For each bacterial strain, 10,000 events were collected and measured.

TABLE 7

| Whole Cell Staining of <i>E. faecalis</i> and <i>E. faecium</i> | | | |
|-----------------------------------------------------------------|--------|--------|---------------|
| | EF1091 | EF1092 | EF1093 |
| <i>E. faecalis</i> | | | |
| ATCC700802 | -- | -- | Not done (NA) |
| 687097 | -- | -- | ND |
| V583 | -- | -- | ND |
| CG110 | -- | -- | ND |
| OG1RF | + | + | + |
| TX2708 | -- | -- | ND |
| TX0020 | ND | ND | ND |
| TX0045 | -- | -- | ND |
| TX0002 | -- | -- | ND |
| TX0039 | -- | -- | ND |
| TX0052 | ND | ND | ND |
| TX0012 | -- | -- | ND |
| TX0017 | ND | ND | ND |
| TX0008 | ND | ND | ND |
| TX0024 | ND | ND | ND |
| <i>E. faecium</i> | | | |
| 935/01 | -- | -- | ND |
| TX0016 | ND | ND | ND |
| TX0054 | +/- | +/- | ND |
| TX0074 | + | + | ND |
| TX0078 | -- | -- | ND |
| TX0080 | +/- | +/- | ND |
| TX0081 | +/- | +/- | ND |
| TX2535 | ND | ND | ND |
| TX2555 | +/- | + | + |
| TX0110 | -- | -- | -- |
| TX0111 | ND | ND | ND |

Polyclonal antisera raised in mice against EF1091, EF1092 and EF1093 were shown to recognize the native protein expressed on the surface of *E. faecalis* strains as well as *E. faecium* strains in flow cytometry studies (Table 7).

Example 7

Immunization Strategies for Monoclonal Antibody Production

With the goal of generating and characterizing monoclonal antibodies (mAbs), strategies were formulated to generate mAbs against EF1091, EF 1092 and EF 1093 that were of high affinity, able to interrupt or restrict the binding of extracellular matrix proteins (ECM) and demonstrate therapeutic efficacy in vivo. *E. coli* expressed and purified EF1091, EF1092, and EF1093 proteins were used to generate a panel of murine monoclonal antibodies. Briefly, a group of Balb/C or SJL mice received a series of subcutaneous immunizations of 1-10 μ g of protein in solution or mixed with adjuvant as described below in Table 8:

TABLE 8

| Immunization Schemes | | | | |
|----------------------|-----|-------------------|--------------|----------|
| | Day | Amount (μ g) | Route | Adjuvant |
| RIMMS Injection | | | | |
| #1 | 0 | 5 | Subcutaneous | FCA/RIBI |
| #2 | 2 | 1 | Subcutaneous | FCA/RIBI |
| #3 | 4 | 1 | Subcutaneous | FCA/RIBI |
| #4 | 7 | 1 | Subcutaneous | FCA/RIBI |
| #5 | 9 | 1 | Subcutaneous | FCA/RIBI |

TABLE 8-continued

| Immunization Schemes | | | | |
|------------------------|-----|-------------|-----------------|----------|
| | Day | Amount (µg) | Route | Adjuvant |
| Conventional Injection | | | | |
| Primary | 0 | 5 | Subcutaneous | FCA |
| Boost #1 | 14 | 1 | Intraperitoneal | RIBI |
| Boost #2 | 28 | 1 | Intraperitoneal | RIBI |
| Boost #3 | 42 | 1 | Intraperitoneal | RIBI |

At the time of sacrifice (RIMMS) or seven days after a boost (conventional) serum was collected and titered in ELISA assays against immunizing MSCRAMM or on whole cells (*E. faecalis* and/or *E. faecium*). Three days after the final boost, the spleens or lymph nodes were removed, teased into a single cell suspension and the lymphocytes harvested. The lymphocytes were then fused to a P3X63Ag8.653 myeloma cell line (ATCC #CRL-1580). Cell fusion, subsequent plating and feeding were performed according to the Production of Monoclonal Antibodies protocol from *Current Protocols in Immunology* (Chapter 2, Unit 2.).

Example 8

Screening and Selection of Anti-EF1091 Monoclonal Antibodies

Any clones that were generated from the EF1091 fusion were then screened for specific anti-EF1091 antibody production using a standard ELISA assay. Positive clones were expanded and tested further for activity in a whole bacterial cell binding assay by flow cytometry and EF1091 binding by Biacore analysis (Table 9).

ELISA Analysis

Immulon 2-HB high-binding 96-well microtiter plates (Dynex) were coated with 1 µg/well of rEF1091 in 1×PBS, pH 7.4 and incubated for 2 hours at room temperature. All washing steps in ELISAs were performed three times with 1×PBS, 0.05% Tween-20 wash buffer. Plates were washed and blocked with a 1% BSA solution at room temperature for 1 hour before hybridoma supernatant samples were added to wells. Plates were incubated with samples and relevant controls such as media alone for one hour at room temperature, washed, and goat anti-mouse IgG-AP (Sigma) diluted 1:5000 in 1×PBS, 0.05% Tween-20, 0.1% BSA was used as a secondary reagent. Plates were developed by addition of 1 mg/ml solution of 4-nitrophenyl phosphate (pNPP) (Sigma), followed by incubation at 37° C. for 30 minutes. Absorbance was read at 405 nm using a SpectraMax 190 Plate Reader (Molecular Devices Corp.). Antibody supernatants that had an OD₄₀₅ ≥ 3 times above background (media alone, ~0.10 D) were considered positive.

Biacore Analysis

Throughout the analysis, the flow rate remained constant at 10 ml/min. Prior to the EF1091 injection, test antibody was adsorbed to the chip via RAM-Fc binding. At time 0, EF1091 at a concentration of 30 mg/ml was injected over the chip for 3 min followed by 2 minutes of dissociation. This phase of the analysis measured the relative association and disassociation kinetics of the mAb/EF1091 interaction.

Flow Cytometric Analysis

Bacterial samples were collected, washed and incubated with mAb or PBS alone (control) at a concentration of 2 mg/ml after blocking with rabbit IgG (50 mg/ml). Following incubation with antibody, bacterial cells were incubated with Goat-F_{(ab')₂}-Anti-Mouse-F_{(ab')₂}-FITC which served as the detection antibody. After antibody labeling, bacterial cells were aspirated through the FACScaliber flow cytometer to analyze fluorescence emission (excitation: 488, emission: 570). For each bacterial strain, 10,000 events were collected and measured.

TABLE 9

| Representative Examples of Hybridoma Supernatants | | | | |
|---------------------------------------------------|----------------------|---------------------|------------------|---------------------------------------------|
| Fusion-Clone | Immunization Antigen | ELISA Data (EF1091) | Biacore Analysis | Flow Cytometric <i>E. faecalis</i> Staining |
| 85-8 | EF 1091 | 0.70 | + | + |
| 85-25 | EF 1091 | 0.75 | + | + |
| 85-58 | EF 1091 | 0.76 | + | -- |
| 85-78 | EF 1091 | 0.83 | + | + |
| 85-81 | EF 1091 | 0.84 | + | + |
| 85-162 | EF 1091 | 0.78 | + | + |
| 85-310 | EF 1091 | 0.30 | -- | -- |
| 85-341 | EF 1091 | 0.31 | -- | -- |
| 85-359 | EF 1091 | 0.48 | -- | -- |
| 85-374 | EF 1091 | 0.39 | -- | -- |
| 85-380 | EF 1091 | 0.32 | -- | -- |
| 85-399 | EF 1091 | 0.98 | + | -- |
| 85-473 | EF 1091 | 0.55 | + | -- |
| 85-511 | EF 1091 | 0.85 | + | -- |
| 85-581 | EF 1091 | 0.88 | + | + |
| 85-586 | EF 1091 | 0.88 | + | + |
| 85-641 | EF 1091 | 0.45 | + | + |
| 85-661 | EF 1091 | 0.32 | -- | -- |
| 85-712 | EF 1091 | 0.30 | -- | -- |

Example 9

Binding of Enterococcal MSCRAMM® Proteins to Extracellular Matrix (ECM) Proteins

Understanding the potential extracellular matrix proteins that these MSCRAMMs expressed from *Enterococcus* bind to is of great biological importance with therapeutic implications.

ELISA based Extracellular Matrix Ligand Screening

To determine the binding activity of the recombinant proteins EF1091, EF1092 and EF1093 (Table 10) with extracellular matrix molecules, duplicate wells of a 96-well Costar micro-titer plate (Corning) were coated overnight at 4° C. with 2 µg of either human collagen type I, III, IV, V or VI (Rockland Immunochemicals), fibrinogen, fibronectin, plasminogen, vitronectin (Sigma) or elastin (CalBiochem) in 1004 of 1×PBS, pH 7.4 (Gibco). Wells were washed 4 times with 1×PBS, pH 7.4 containing 0.05% Tween 20 (1×PBST). Wells were then blocked with a 1% (w/v) solution of BSA in 1×PBS, pH 7.4 for 1 hour followed by 4 washes with 1×PBST. Next, 5 µg of recombinant protein in 1004 of 1×PBST containing 0.1% BSA (1×PBST-BSA) was added to each well. After incubation with the protein for 1 hour at room temperature, wells were washed 4 times with 1×PBS-T and 100 µl of mouse polyclonal antisera raised against the respective recombinant protein was added to each well at a dilution of 1:2000 in 1×PBST-BSA. Following the 1 hour incubation at room temperature with antisera, the wells were washed 4

times with 1xPBST. Finally, goat anti-mouse IgG-alkaline phosphatase conjugate (Sigma) was diluted 1:2000 with 1xPBST-BSA and 100 µl was added to each well. This incubation proceeded for 1 hour at room temperature and the wells were then washed 4 times with 1xPBST. The alkaline phosphatase was developed by adding 100 µl of a 1 mg/mL pNP solution (Sigma 104 tablets) to each well and incubating for 30 minutes at room temperature. Development was stopped by addition of 504 of 2M NaOH to each well. The absorbance at 405 nm (A_{405}) was measured using a SpectraMax 190 (Molecular Devices). Reactivity was noted as positive if the signal was 2.5x greater than background.

Alternatively, EF0089 and EF2224 binding to components of the ECM (Table 10) was tested by immobilizing 1 µg of each ECM protein (human laminin, fibronectin, fibrinogen, type I, III and IV collagens) in 100 µl PBS, or 3% acetic acid in the case of collagens, on microplate wells (96-well, 4HBX, Thermo Labsystems, Franklin, Ma) overnight at 4° C. Plates were washed once with PBS and blocked with 1% BSA in PBS for 1 h. Fifty µl of 5 and 10 µM concentrations of purified His-tag proteins in the blocking buffer were added and incubated at ambient temperature for 2 h. Plates were washed three times with 0.05% Tween20 in PBS and incubated 2 h with 1:3000 dilution of His6-tag monoclonal antibody (Amersham Biosciences Corp., Piscataway, N.J.) in blocking buffer. After three washes, 1:3000 dilution of alkaline phosphatase-conjugated anti-mouse antibody in blocking buffer was added to the wells and incubated 2 h. Finally, signal was detected with nitrobluetetrazolium (NBT) and 5-bromo-4-chloro-3-indolyl-phosphate (BCIP) in 0.1 M NaHCO₃, 1 mM MgCl₂, pH 9.8. Absorbance at 405 nm was measured with an ELISA reader

TABLE 10

| MSCRAMM® Protein Recognition of ECM Proteins | | | | | |
|----------------------------------------------|---------------------|--------|---------|---------|---------|
| ECM Proteins | EF0089 | EF2224 | EF 1091 | EF 1092 | EF 1093 |
| Fibrinogen | + | + | -- | -- | + |
| Fibronectin | -- | -- | -- | -- | -- |
| Collagen I | -- | -- | -- | -- | -- |
| Collagen III | -- | -- | -- | -- | -- |
| Collagen IV | -- | -- | -- | -- | -- |
| Collagen V | Not determined (ND) | ND | -- | -- | -- |

TABLE 10-continued

| MSCRAMM® Protein Recognition of ECM Proteins | | | | | |
|----------------------------------------------|--------|--------|---------|---------|---------|
| ECM Proteins | EF0089 | EF2224 | EF 1091 | EF 1092 | EF 1093 |
| Collagen VI | -- | -- | -- | + | -- |
| Vitronectin | -- | -- | -- | -- | -- |
| Elastin | ND | ND-- | -- | -- | -- |
| Plasminogen | ND | ND | + | + | + |

Example 10

Serum From Patients Infected With *E. faecalis* Contain Elevated Levels of Antibodies Against MSCRAMM® Proteins

The presence of antibodies against enterococcal proteins in human sera collected from hospitalized patients with and without a previous *E. faecalis* infection was tested by an ELISA assay described in (Arduino et al., 1994) (Nallapareddy et al., 2000b) with some modifications (Table 11). Briefly, 20 ng of each purified enterococcal protein in 100 µl PBS was coated on microplates (96 well, 4HBX, Thermo Labsystems, Franklin, Ma) overnight at 4 °C. The plates were blocked with 1% BSA, 0.01% Tween20 in PBS at ambient temperature for 1 h and 100 µl of the sera in blocking buffer were added. Each serum was tested in triplicate with serial dilutions from 1:100 to 1:6400. Plates were incubated for 2 h at ambient temperature and washed three times with 0.01% Tween20 in PBS. 100 µl of 1:3000 dilution of horse-radish peroxidase-conjugated anti human IgG was added and incubated 2 h. After three washes, signal was detected with 3,3',5,5'-tetramethylbenzidine (TMB) in the presence of H₂O₂ in 0.1 M citrate-acetate buffer, pH 6.0 at ambient temperature for 15 min. The reaction was stopped with 2 M H₂SO₄ and absorbance at 450 nm was recorded. Titers were determined after subtracting A_{450nm} values from appropriate controls. To determine a cut-off level for serum titers, four additional control sera from healthy individuals without a prior *E. faecalis* infection were assayed. The sum of average A_{450nm} values and two times the standard deviations for each dilution of the control sera were set as cut-off levels for positive titers.

TABLE 11

| | Infection | | | | | | No infection | | | |
|---------|--------------|---------|---------|---------|---------|---------|--------------|---------|---------|---------|
| ≥1:6400 | ••••• | • | •• | • | ••• | ••• | ••••••• | | | |
| 1:3200 | | • | | • | | | • | | | |
| 1:1600 | | | | | | | | | | |
| 1:800 | | | | | | | | | | |
| 1:400 | | | | | | | • | | | |
| 1:200 | | | •• | • | • | ••• | • | • | | |
| ≤1:100 | •• | ••••• | ••••• | ••••• | ••••• | •• | ••••••• | ••••••• | | |
| | EF1091 | EF1824 | EF0089 | EF3023 | EF1092 | EF2224 | EF1269 | EF1093 | EF1091 | EF1824 |
| | No infection | | | | | | | | | |
| ≥1:6400 | | | | | | | | | | • |
| 1:3200 | | | | | | | | | | |
| 1:1600 | | | | | | | | | | |
| 1:800 | | | | | | | | | | |
| 1:400 | | | | | | | | | | |
| 1:200 | | | | | • | • | • | • | | |
| ≤1:100 | ••••••• | ••••••• | ••••••• | ••••••• | ••••••• | ••••••• | ••••••• | ••••••• | ••••••• | ••••••• |
| | EF0089 | EF3023 | EF1092 | EF2224 | EF1269 | EF1093 | | | | |

The following references referred to in the above description are incorporated as is set forth in their entirety herein:

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 Asn Lys Asp Gln Leu Gln Glu Val Val Lys Arg Ala Gln Gln Val Thr
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 Pro Ser Glu Gly His Gln Phe Thr Ala Ser Ser Leu Gln Glu Leu Gln
 770 775 780
 Lys Ala Leu Leu Ala Ala Lys Asn Thr Leu Lys Asn Pro Ala Ala Asn
 785 790 795 800
 Gln Lys Met Ile Asp Glu Ala Val Ala Glu Leu Thr Ser Ala Ile Asp
 805 810 815
 Gly Leu Gln Glu Glu Val Leu Val Thr Asp Lys Lys Ala Leu Glu Ala
 820 825 830
 Met Ile Ala Lys Ala Lys Ala Ile Lys Pro Ser Ala Gly Lys Glu Phe
 835 840 845
 Thr Ser Glu Ser Lys Ala Arg Leu Thr Glu Ala Ile Asp Gln Ala Glu
 850 855 860
 Gly Ile Leu Ala Asp Lys Asn Ala Arg Gln Glu Gln Ile Asp Ile Ala
 865 870 875 880
 Glu Lys Asn Val Lys Thr Ala Leu Asp Ser Leu Glu Glu Gln Val Leu
 885 890 895
 Gln Thr Asp Lys Thr Lys Leu Lys Glu Leu Leu Gln Lys Ala Glu Thr
 900 905 910
 Leu Lys Pro Lys Ala Gly Lys Gln Phe Thr Lys Ala Ser Gln Glu Ala
 915 920 925
 Leu Ala Glu Ala Ile Lys Gln Ala Lys Ala Leu Val Glu Asp Pro Asn
 930 935 940
 Ala Thr Gln Glu Ala Val Asp Lys Cys Leu Ser Ile Leu Ser Gln Ala
 945 950 955 960
 Ile Glu Ala Met Ala Glu Glu Pro Ile Ser Ser Asn Ser Thr Gly Asn
 965 970 975
 Asn Gly Asn His Ser Thr Val Ser Gly Thr Gly Gly Val Thr Ser Gln
 980 985 990
 Gly Lys Gly Thr Ala Thr Gly Gly Thr Thr Thr Lys Thr Thr Thr Ser
 995 1000 1005

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Gly Thr
1010

<210> SEQ ID NO 4
<211> LENGTH: 1108
<212> TYPE: PRT
<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 4

Glu Glu Val Asn Ser Asp Gly Gln Leu Thr Leu Gly Glu Val Lys Gln
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Thr Ser Gln Gln Glu Met Thr Leu Ala Leu Gln Gly Lys Ala Gln Pro
20 25 30

Val Thr Gln Glu Val Val Val His Tyr Ser Ala Asn Val Ser Ile Lys
35 40 45

Ala Ala His Trp Ala Ala Pro Asn Asn Thr Arg Lys Ile Gln Val Asp
50 55 60

Asp Gln Lys Lys Gln Ile Gln Ile Glu Leu Asn Gln Gln Ala Leu Ala
65 70 75 80

Asp Thr Leu Val Leu Thr Leu Asn Pro Thr Ala Thr Glu Asp Val Thr
85 90 95

Phe Ser Tyr Gly Gln Gln Gln Arg Ala Leu Thr Leu Lys Thr Gly Thr
100 105 110

Asp Pro Thr Glu Ser Thr Ala Ile Thr Ser Ser Pro Ala Ala Ser Ala
115 120 125

Asn Glu Gly Ser Thr Glu Glu Ala Ser Thr Asn Ser Ser Val Pro Arg
130 135 140

Ser Ser Glu Glu Thr Val Ala Ser Thr Thr Lys Ala Ile Glu Ser Lys
145 150 155 160

Thr Thr Glu Ser Thr Thr Val Lys Pro Arg Val Ala Gly Pro Thr Asp
165 170 175

Ile Ser Asp Tyr Phe Thr Gly Asp Glu Thr Thr Ile Ile Asp Asn Phe
180 185 190

Glu Asp Pro Ile Tyr Leu Asn Pro Asp Gly Thr Pro Ala Thr Pro Pro
195 200 205

Tyr Lys Glu Asp Val Thr Ile His Trp Asn Phe Asn Trp Ser Ile Pro
210 215 220

Glu Asp Val Arg Glu Gln Met Lys Ala Gly Asp Tyr Phe Glu Phe Gln
225 230 235 240

Leu Pro Gly Asn Leu Lys Pro Asn Lys Pro Gly Ser Gly Asp Leu Val
245 250 255

Asp Ala Glu Gly Asn Val Tyr Gly Thr Tyr Thr Ile Ser Glu Asp Gly
260 265 270

Thr Val Arg Phe Thr Phe Asn Glu Arg Ile Thr Ser Glu Ser Asp Ile
275 280 285

His Gly Asp Phe Ser Leu Asp Thr His Leu Asn Asp Ser Asp Gly Arg
290 295 300

Gly Pro Gly Asp Trp Val Ile Asp Ile Pro Thr Gln Glu Asp Leu Pro
305 310 315 320

Pro Val Val Ile Pro Ile Val Pro Asp Thr Glu Gln Gln Ile Asp Lys
325 330 335

Gln Gly His Phe Asp Arg Thr Pro Asn Pro Ser Ala Ile Thr Trp Thr
340 345 350

Val Asp Ile Asn Gln Ala Met Lys Asp Gln Thr Asn Pro Thr Val Thr
355 360 365

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Glu Thr Trp Pro Thr Gly Asn Thr Phe Lys Ser Val Lys Val Tyr Glu
 370 375 380
 Leu Val Met Asn Leu Asp Gly Thr Ile Lys Glu Val Gly Arg Glu Leu
 385 390 395 400
 Ser Pro Asp Glu Tyr Thr Val Asp Lys Asn Gly Asn Val Thr Ile Lys
 405 410 415
 Gly Asp Thr Asn Lys Ala Tyr Arg Leu Glu Tyr Gln Thr Thr Ile Asp
 420 425 430
 Glu Ala Val Ile Pro Asp Gly Gly Gly Asp Val Pro Phe Lys Asn His
 435 440 445
 Ala Thr Leu Thr Ser Asp Asn Asn Pro Asn Gly Leu Asp Ala Glu Ala
 450 455 460
 Thr Val Thr Ala Thr Tyr Gly Lys Met Leu Asp Lys Arg Asn Ile Asp
 465 470 475 480
 Tyr Asp Glu Ala Asn Gln Glu Phe Thr Trp Glu Ile Asn Tyr Asn Tyr
 485 490 495
 Gly Glu Gln Thr Ile Pro Lys Asp Gln Ala Val Ile Thr Asp Thr Met
 500 505 510
 Gly Asp Asn Leu Thr Phe Glu Pro Asp Ser Leu His Leu Tyr Ser Val
 515 520 525
 Thr Phe Asp Asp Lys Gly Asn Glu Val Val Gly Ala Glu Leu Val Glu
 530 535 540
 Gly Lys Asp Tyr Lys Val Ile Asn Gly Asp Gly Ser Phe Ala Ile
 545 550 555 560
 Asp Phe Leu His Asp Val Thr Gly Ala Val Lys Ile Asp Tyr Lys Thr
 565 570 575
 Lys Val Asp Gly Ile Val Glu Gly Asp Val Ala Val Asn Asn Arg Val
 580 585 590
 Asp Val Gly Thr Gly Gln His Ser Glu Asp Asp Gly Thr Ala Ser Gln
 595 600 605
 Gln Asn Ile Ile Lys Asn Thr Gly Ala Val Asp Tyr Gln Asn Ser Thr
 610 615 620
 Ile Gly Trp Thr Leu Ala Val Asn Gln Asn Asn Tyr Leu Met Glu Asn
 625 630 635 640
 Ala Val Ile Thr Asp Thr Tyr Glu Pro Val Pro Gly Leu Thr Met Val
 645 650 655
 Pro Asn Ser Leu Val Val Lys Asp Thr Thr Thr Gly Ala Gln Leu Thr
 660 665 670
 Leu Gly Lys Asp Phe Met Val Glu Ile Thr Arg Asn Ala Asp Gly Glu
 675 680 685
 Thr Gly Phe Lys Val Ser Phe Ile Gly Ala Tyr Ala Lys Thr Ser Asp
 690 695 700
 Ala Phe His Ile Thr Tyr Thr Thr Phe Phe Asp Val Thr Glu Leu Asp
 705 710 715 720
 Ala Asn Asn Pro Ala Leu Asp His Tyr Arg Asn Thr Ala Ala Ile Asp
 725 730 735
 Trp Thr Asp Glu Ala Gly Asn Asn His His Ser Glu Asp Ser Lys Pro
 740 745 750
 Phe Lys Pro Leu Pro Ala Phe Asp Leu Asn Ala Gln Lys Ser Gly Val
 755 760 765
 Tyr Asn Ala Val Thr Lys Glu Ile Thr Trp Thr Ile Ala Val Asn Leu
 770 775 780

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Lys Gln Asp Glu Gln Trp Leu Ile Val Asp Leu Gly Glu Val Lys Asn
 50 55 60
 Ile Gly Glu Leu Val Leu Gln Leu His Ala Glu Ser Pro Val Tyr Glu
 65 70 75 80
 Ile Leu Val Ser Thr Asp Gly Glu Ser Tyr Gln Ser Ile Phe Lys Glu
 85 90 95
 Glu Asn Gly Lys Gly Gly Gln Pro Thr Lys Lys Tyr Ile Asp Gly Asn
 100 105 110
 Asn Val Gln Ala Arg Phe Val Lys Tyr Gln Gln Met Lys Met Trp Gln
 115 120 125
 His Thr Asn Lys Gln Phe Tyr Ser Ser Ser Ile Ile Ser Phe Glu Ala
 130 135 140
 Tyr Glu Lys Lys Arg Leu Pro Glu Ala Ile Lys Leu Leu Thr Glu Asn
 145 150 155 160
 Leu Thr Ile Ser Glu Lys Arg Lys Gln Gln Leu Ala Phe Glu Val Ser
 165 170 175
 Pro Ala Gly Val Asp Ile Thr Glu Asp Gln Ile Glu Trp Ser Ser Ser
 180 185 190
 Asp Pro Thr Ile Val Thr Val Asp Gln Thr Gly Asn Leu Thr Ala Val
 195 200 205
 Lys Ser Gly Glu Ala Lys Val Thr Val Lys Ile Lys Gly Thr Glu Ile
 210 215 220
 Ser Asp Thr Ile Pro Val Thr Val Val Ala Glu Asn Lys Gln Tyr Ala
 225 230 235 240
 Glu Met Arg Ala Lys Trp Lys Met Arg Leu Leu Gly Thr Thr Gln Tyr
 245 250 255
 Asp Asn Asp Ala Asp Val Gln Gln Tyr Arg Ala Gln Ile Ala Thr Glu
 260 265 270
 Ser Leu Ala Leu Trp Gln Thr Leu Asn Gln Ala Ala Asp Arg Glu Tyr
 275 280 285
 Leu Trp Glu Arg Lys Pro Ser Asp Thr Val Ser Ala Asp Tyr Thr Thr
 290 295 300
 Gln Phe Thr Asn Ile Lys Lys Leu Ala Leu Gly Tyr Tyr Glu Pro Ser
 305 310 315
 Ser Glu Leu Phe Glu Lys Pro Glu Val Tyr Asp Ala Ile Val Lys Gly
 325 330 335
 Ile Glu Phe Met Ile Asp Thr Lys Lys Tyr Asn Gly Thr Tyr Tyr Thr
 340 345 350
 Gly Asn Trp Trp Asp Trp Gln Ile Gly Ser Ala Gln Pro Leu Thr Asp
 355 360 365
 Thr Leu Ile Leu Leu His Asp Asp Leu Leu Asn Thr Asp Ala Glu Lys
 370 375 380
 Leu Asn Lys Phe Thr Ala Pro Leu Met Leu Tyr Ala Lys Asp Pro Asn
 385 390 395 400
 Ile Gln Trp Pro Ile Tyr Arg Ala Thr Gly Ala Asn Leu Thr Asp Ile
 405 410 415
 Ser Ile Thr Val Leu Gly Thr Gly Leu Leu Leu Glu Asp Asn Gln Arg
 420 425 430
 Leu Val Gln Val Gln Glu Ala Val Pro Ser Val Leu Lys Ser Val Ser
 435 440 445
 Ser Gly Asp Gly Leu Tyr Pro Asp Gly Ser Leu Ile Gln His Gly Tyr
 450 455 460

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| | | | | | | | | | | | | | | | |
|------------|------------|------------|-----|------------|------------|------------|------------|-----|------------|------------|------------|-----|-----|------------|------------|
| Phe 465 | Pro | Tyr | Asn | Gly | Ser 470 | Tyr | Gly | Asn | Glu | Leu 475 | Leu | Lys | Gly | Phe | Gly 480 |
| Arg | Ile | Gln | Thr | Ile 485 | Leu | Gln | Gly | Ser | Asp 490 | Trp | Glu | Met | Asn | Asp 495 | Pro |
| Asn | Ile | Ser | Asn | Leu 500 | Phe | Asn | Val | Val | Asp 505 | Lys | Gly | Tyr | Leu | Gln | Leu |
| Met | Val | Asn 515 | Gly | Lys | Met | Pro | Ser | Met | Val 520 | Ser | Gly | Arg | Ser | Ile | Ser |
| Arg | Ala 530 | Pro | Glu | Thr | Asn | Pro | Phe | Thr | Thr | Glu | Phe 540 | Glu | Ser | Gly | Lys |
| Glu 545 | Thr | Ile | Ala | Asn | Leu 550 | Thr | Leu | Ile | Ala | Lys | Phe 555 | Ala | Pro | Glu | Asn 560 |
| Leu | Arg | Asn | Asp | Ile 565 | Tyr | Thr | Ser | Ile | Gln 570 | Thr | Trp | Leu | Gln | Gln | Ser 575 |
| Gly | Ser | Tyr | Tyr | His 580 | Phe | Phe | Lys | Lys | Pro | Arg | Asp | Phe | Glu | Ala | Leu 590 |
| Ile | Asp | Leu 595 | Lys | Asn | Val | Val | Asn | Ser | Ala | Ser | Pro | Ala | Gln | Ala | Thr |
| Pro | Met | Gln | Ser | Leu | Asn 615 | Val | Tyr | Gly | Ser | Met | Asp 620 | Arg | Val | Leu | Gln |
| Lys 625 | Asn | Asn | Glu | Tyr | Ala 630 | Val | Gly | Ile | Ser | Met | Tyr 635 | Ser | Gln | Arg | Val 640 |
| Gly | Asn | Tyr | Glu | Phe 645 | Gly | Asn | Thr | Glu | Asn | Lys | Lys | Gly | Trp | His | Thr 655 |
| Ala | Asp | Gly | Met | Leu 660 | Tyr | Leu | Tyr | Asn | Gln | Asp | Phe | Ala | Gln | Phe | Asp 670 |
| Glu | Gly | Tyr | Trp | Ala | Thr | Ile | Asp 680 | Pro | Tyr | Arg | Leu | Pro | Gly | Thr | Thr 685 |
| Val | Asp | Thr | Arg | Glu | Leu 695 | Ala | Asn | Gly | Ala | Tyr | Thr 700 | Gly | Lys | Arg | Ser |
| Pro 705 | Gln | Ser | Trp | Val | Gly 710 | Gly | Ser | Asn | Asn | Gly | Gln 715 | Val | Ala | Ser | Ile 720 |
| Gly | Met | Phe | Leu | Asp 725 | Lys | Ser | Asn | Glu | Gly | Met | Asn 730 | Leu | Val | Ala | Lys 735 |
| Lys | Ser | Trp | Phe | Leu 740 | Leu | Asp | Gly | Gln | Ile | Ile | Asn | Leu | Gly | Ser | Gly 750 |
| Ile | Thr | Gly | Thr | Thr | Asp | Ala | Ser | Ile | Glu | Thr | Ile 765 | Leu | Asp | Asn | Arg 765 |
| Met 770 | Ile | His | Pro | Gln | Glu 775 | Val | Lys | Leu | Asn | Gln | Gly 780 | Ser | Asp | Lys | Asp |
| Asn 785 | Ser | Trp | Ile | Ser | Leu 790 | Ser | Ala | Ala | Asn | Pro | Leu 795 | Asn | Asn | Ile | Gly 800 |
| Tyr | Val | Phe | Pro | Asn 805 | Ser | Met | Asn | Thr | Leu 810 | Asp | Val | Gln | Ile | Glu | Glu 815 |
| Arg | Ser | Gly | Arg | Tyr | Gly | Asp | Ile | Asn | Glu | Tyr | Phe | Val | Asn | Asp | Lys 830 |
| Thr | Tyr | Thr | Asn | Thr | Phe | Ala | Lys 840 | Ile | Ser | Lys | Asn | Tyr | Gly | Lys | Thr 845 |
| Val | Glu | Asn | Gly | Thr | Tyr | Glu 855 | Tyr | Leu | Thr | Val | Val 860 | Gly | Lys | Thr | Asn |
| Glu 865 | Glu | Ile | Ala | Ala | Leu 870 | Ser | Lys | Asn | Lys | Gly | Tyr 875 | Thr | Val | Leu | Glu 880 |
| Asn | Thr | Ala | Asn | Leu | Gln | Ala | Ile | Glu | Ala | Gly | Asn | Tyr | Val | Met | Met |

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| 885 | | | | | 890 | | | | | 895 | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Asn | Thr | Trp | Asn | Asn | Asp | Gln | Glu | Ile | Ala | Gly | Leu | Tyr | Ala | Tyr | Asp |
| | | | 900 | | | | | 905 | | | | | 910 | | |
| Pro | Met | Ser | Val | Ile | Ser | Glu | Lys | Ile | Asp | Asn | Gly | Val | Tyr | Arg | Leu |
| | | 915 | | | | | 920 | | | | | 925 | | | |
| Thr | Leu | Ala | Asn | Pro | Leu | Gln | Asn | Asn | Ala | Ser | Val | Ser | Ile | Glu | Phe |
| | 930 | | | | | 935 | | | | | 940 | | | | |
| Asp | Lys | Gly | Ile | Leu | Glu | Val | Val | Ala | Ala | Asp | Pro | Glu | Ile | Ser | Val |
| 945 | | | | | 950 | | | | | 955 | | | | | 960 |
| Asp | Gln | Asn | Ile | Ile | Thr | Leu | Asn | Ser | Ala | Gly | Leu | Asn | Gly | Ser | Ser |
| | | | 965 | | | | | | 970 | | | | | 975 | |
| Arg | Ser | Ile | Ile | Val | Lys | Thr | Thr | Pro | Glu | Val | Thr | Lys | Glu | Ala | Leu |
| | | 980 | | | | | | 985 | | | | | 990 | | |
| Glu | Lys | Leu | Ile | Gln | Glu | Gln | | | | | | | | | |
| | | 995 | | | | | | | | | | | | | |

<210> SEQ ID NO 6

<211> LENGTH: 741

<212> TYPE: PRT

<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 6

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

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| | | | | | | | | | | | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Asn | Thr | Ala | Ala | Ala | Gln | Thr | Phe | Leu | Asn | Thr | Arg | Gly | Ala | Ser | Ser |
| | | 260 | | | | | | 265 | | | | | 270 | | |
| Gly | Tyr | Ala | Ile | Asn | Asn | Phe | Pro | Leu | Glu | Phe | Ala | Asp | Val | Asp | Asn |
| | | 275 | | | | 280 | | | | | | 285 | | | |
| Asp | Pro | Asn | Thr | Tyr | Asn | Ser | Ser | Arg | Ala | Tyr | Ile | Asp | Leu | Asn | Gly |
| | 290 | | | | | 295 | | | | | 300 | | | | |
| Ala | Lys | Glu | Ile | Ala | Trp | Ala | Gly | Leu | Phe | Trp | Ser | Ala | Ser | Arg | Tyr |
| 305 | | | | | 310 | | | | | 315 | | | | | 320 |
| Lys | Gly | Pro | Ala | Tyr | Gly | Thr | Asn | Leu | Ser | Asp | Glu | Glu | Ile | Ser | Ala |
| | | | | 325 | | | | | 330 | | | | | 335 | |
| Pro | Val | Gln | Phe | Thr | Thr | Pro | Asn | Gly | Thr | Val | Gln | Arg | Val | Ser | Pro |
| | | | 340 | | | | | 345 | | | | | 350 | | |
| Gln | Arg | Tyr | His | Arg | Ile | Asp | Gln | Asp | Ala | Thr | Asn | Pro | Gly | Gln | Arg |
| | | 355 | | | | 360 | | | | | | 365 | | | |
| Phe | Gly | Tyr | Asn | Asn | Thr | Gly | Phe | Ser | Asn | Tyr | Ala | Asp | Val | Thr | Ser |
| | 370 | | | | | 375 | | | | | 380 | | | | |
| Ile | Leu | Gln | Gly | Asp | Lys | Ser | Ala | Thr | Gly | Ser | Tyr | Thr | Leu | Ala | Asp |
| 385 | | | | | 390 | | | | | 395 | | | | | 400 |
| Ile | Pro | Met | Thr | Ser | Ser | Leu | Asn | Gly | Gln | Tyr | Gln | Tyr | Tyr | Asn | Phe |
| | | | 405 | | | | | | 410 | | | | | 415 | |
| Ser | Gly | Trp | Ser | Leu | Phe | Val | Val | Thr | Lys | Asp | Gln | Ala | Ser | Lys | Ser |
| | | 420 | | | | | | 425 | | | | | 430 | | |
| Arg | Ala | Phe | Ser | Ile | Tyr | Tyr | Gly | Ala | Arg | Gly | Asn | Ala | Ala | Gly | Thr |
| | | 435 | | | | | 440 | | | | | 445 | | | |
| Asn | Asn | Glu | Phe | Thr | Met | Ser | Asn | Phe | Leu | Thr | Ala | Lys | Gln | Gly | Asn |
| | 450 | | | | | 455 | | | | | 460 | | | | |
| Leu | Asp | Pro | Ile | Val | Thr | Trp | Phe | Thr | Val | Gln | Gly | Asp | Lys | Tyr | Trp |
| 465 | | | | 470 | | | | | | 475 | | | | | 480 |
| Thr | Gly | Asp | Asn | Ala | Gln | Ile | Lys | Asn | Ser | Ala | Gly | Thr | Trp | Val | Asn |
| | | | 485 | | | | | | 490 | | | | | 495 | |
| Ile | Ser | Asn | Thr | Leu | Asn | Pro | Val | Asn | Asn | Ala | Met | Asn | Ala | Thr | Val |
| | | | 500 | | | | | 505 | | | | | 510 | | |
| Thr | Asp | Asn | Asp | Glu | His | Met | Val | Asp | Lys | Tyr | Pro | Gly | Lys | Phe | Ala |
| | | 515 | | | | 520 | | | | | | 525 | | | |
| Pro | Asp | His | Pro | Asn | Phe | Leu | Asp | Ile | Asp | Ile | Asp | Arg | Met | Ala | Ile |
| | 530 | | | | | 535 | | | | | 540 | | | | |
| Pro | Glu | Gly | Val | Leu | Asn | Ala | Gly | Gln | Asn | Gln | Ile | Asn | Phe | Arg | Thr |
| 545 | | | | | 550 | | | | | 555 | | | | | 560 |
| Thr | Ser | Ser | Gly | Asp | Asp | Tyr | Ser | Thr | Asn | Ala | Ile | Gly | Phe | Ala | Val |
| | | | 565 | | | | | | 570 | | | | | 575 | |
| Asn | Ala | Glu | Thr | Pro | Glu | Phe | Glu | Ile | Lys | Lys | Glu | Ile | Val | Glu | Pro |
| | | | 580 | | | | | 585 | | | | | 590 | | |
| Lys | Glu | Thr | Tyr | Lys | Val | Gly | Glu | Thr | Ile | Thr | Tyr | Arg | Val | Ser | Leu |
| | 595 | | | | | | 600 | | | | | 605 | | | |
| Lys | Asn | Thr | Lys | Ala | Asp | Ser | Glu | Ala | Ile | Asn | Ser | Val | Ser | Lys | Asp |
| | 610 | | | | | 615 | | | | | 620 | | | | |
| Ala | Leu | Asp | Gly | Arg | Leu | Asn | Tyr | Leu | Pro | Gly | Ser | Leu | Lys | Ile | Ile |
| 625 | | | | | 630 | | | | | 635 | | | | | 640 |
| Ser | Gly | Pro | Asn | Ser | Gly | Glu | Lys | Thr | Asp | Ala | Ser | Gly | Asp | Asp | Gln |
| | | | 645 | | | | | | 650 | | | | | 655 | |
| Ala | Glu | Tyr | Asp | Glu | Thr | Asn | Lys | Gln | Ile | Ile | Val | Arg | Val | Gly | Asn |
| | | | 660 | | | | | 665 | | | | | 670 | | |
| Gly | Ala | Thr | Ala | Thr | Gln | Gly | Gly | Ser | Tyr | Lys | Ala | Asp | Thr | Ala | Glu |

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      675              680              685
Thr  Ile  Tyr  Glu  Phe  Lys  Ala  Arg  Ile  Asn  Glu  Arg  Ala  Lys  Ala  Asn
   690              695              700

Glu  Leu  Val  Pro  Asn  Ser  Ala  Thr  Val  Glu  Ala  Val  Asp  Ile  Leu  Thr
   705              710              715              720

Ser  Ala  Lys  Val  Asn  Glu  Thr  Ser  Asn  Ile  Val  Glu  Ala  Lys  Ile  Ala
              725              730              735

Asp  Glu  Gln  Val  Thr
              740

<210> SEQ ID NO 7
<211> LENGTH: 570
<212> TYPE: PRT
<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 7

Glu  Thr  Gly  Tyr  Ala  Gln  Thr  Glu  Pro  Thr  Ser  Thr  Ser  Glu  Thr  Asn
 1              5              10              15

Gln  Ile  Ser  Ala  Thr  Pro  Asn  Val  Val  Pro  Arg  Lys  Gln  Val  Gly  Asn
 20              25              30

Ile  Val  Thr  Ala  Ile  Gln  Leu  Thr  Asp  Lys  Glu  Gly  Asn  Pro  Leu  Gly
 35              40              45

Thr  Ile  Asn  Gln  Tyr  Thr  Asp  Ile  Tyr  Leu  Arg  Ile  Glu  Phe  Asn  Leu
 50              55              60

Pro  Asp  Asn  Thr  Val  Asn  Ser  Gly  Asp  Thr  Ser  Val  Ile  Thr  Leu  Pro
 65              70              75              80

Glu  Glu  Leu  Arg  Leu  Glu  Lys  Asn  Met  Thr  Phe  Asn  Val  Val  Asp  Asp
 85              90              95

Thr  Gly  Thr  Val  Val  Ala  Ile  Ala  Gln  Thr  Asp  Val  Ala  Asn  Lys  Thr
 100             105             110

Val  Thr  Leu  Thr  Tyr  Thr  Asp  Tyr  Val  Glu  Asn  His  Ala  Asn  Ile  Ser
 115             120             125

Gly  Ser  Leu  Tyr  Phe  Thr  Ser  Leu  Ile  Asp  Phe  Glu  Asn  Val  Glu  Asn
 130             135             140

Glu  Ser  Lys  Ile  Pro  Ile  Tyr  Val  Thr  Val  Glu  Gly  Glu  Lys  Ile  Phe
 145             150             155             160

Ala  Gly  Asp  Leu  Asp  Tyr  Gln  Gly  Glu  Gly  Asp  Asp  Val  Asn  Glu  Lys
 165             170             175

Phe  Ser  Lys  Tyr  Ser  Trp  Phe  Ile  Glu  Asp  Asp  Pro  Thr  Glu  Ile  Tyr
 180             185             190

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

Val  Glu  Asp  Val  Leu  Lys  Thr  Glu  Ser  Leu  Ser  Tyr  Met  Lys  Asp  Thr
 210             215             220

Met  Lys  Ile  Glu  Arg  Gly  Gln  Trp  Thr  Leu  Asp  Gly  Asn  Ala  Ile  Trp
 225             230             235             240

Gln  Phe  Thr  Pro  Glu  Glu  Asp  Ile  Thr  Asp  Gln  Leu  Ala  Val  Gln  Tyr
 245             250             255

Gly  Pro  Asp  Asp  Arg  Asn  Phe  Ser  Val  His  Phe  Gly  Asn  Ile  Gly  Thr
 260             265             270

Asn  Glu  Tyr  Arg  Ile  Thr  Tyr  Lys  Thr  Lys  Ile  Asp  His  Leu  Pro  Glu
 275             280             285

Lys  Gly  Glu  Thr  Phe  Thr  Asn  Tyr  Ala  Lys  Leu  Thr  Glu  Asn  Gln  Thr
 290             295             300

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| | | | | | | | | | | | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Val | Val | Glu | Glu | Val | Glu | Val | Ser | Arg | Val | Ser | Gln | Thr | Gly | Gly | Gly |
| 305 | | | | | 310 | | | | | 315 | | | | | 320 |
| Glu | Ala | Asn | Gly | Glu | Gln | Tyr | Val | Val | Glu | Ile | His | Lys | Glu | Asp | Glu |
| | | | 325 | | | | | | 330 | | | | | 335 | |
| Ala | Gly | Gln | Arg | Leu | Ala | Gly | Ala | Glu | Phe | Glu | Leu | Ile | Arg | Asn | Ser |
| | | | 340 | | | | | 345 | | | | | 350 | | |
| Thr | Asn | Gln | Thr | Val | Ala | Lys | Ile | Thr | Thr | Asp | Gln | Asn | Gly | Thr | Ala |
| | 355 | | | | | 360 | | | | | 365 | | | | |
| Ile | Val | Lys | Gly | Leu | Leu | Lys | Asp | Asn | Tyr | Thr | Leu | Val | Glu | Thr | Lys |
| | 370 | | | | | 375 | | | | | 380 | | | | |
| Ala | Pro | Thr | Gly | Tyr | Gln | Leu | Ser | Gln | Asn | Lys | Ile | Pro | Ile | Thr | Pro |
| 385 | | | | | 390 | | | | | 395 | | | | | 400 |
| Glu | Asp | Phe | Gly | Lys | Asn | Leu | Val | Ala | Leu | Lys | Thr | Val | Val | Asn | His |
| | | | 405 | | | | | | 410 | | | | | 415 | |
| Lys | Ile | Ser | Tyr | Gln | Pro | Val | Ala | Ala | Ser | Phe | Leu | Ala | Gly | Lys | Val |
| | | | 420 | | | | | 425 | | | | | 430 | | |
| Leu | Leu | Gly | Lys | Pro | Leu | Lys | Asp | Ala | Glu | Phe | Gln | Phe | Glu | Leu | Leu |
| | | 435 | | | | | 440 | | | | | 445 | | | |
| Asp | Glu | Lys | Gly | Thr | Val | Leu | Glu | Thr | Val | Ser | Asn | Asp | Thr | Leu | Gly |
| | 450 | | | | | 455 | | | | | 460 | | | | |
| Lys | Ile | Gln | Phe | Ser | Pro | Leu | Thr | Phe | Glu | Thr | Pro | Gly | Asn | Tyr | Gln |
| 465 | | | | | 470 | | | | | 475 | | | | | 480 |
| Tyr | Thr | Ile | Arg | Glu | Val | Asn | Thr | Gln | Gln | Thr | Gly | Val | Ser | Tyr | Asp |
| | | | 485 | | | | | 490 | | | | | | 495 | |
| Thr | His | Asn | Leu | Gln | Val | Gln | Val | Thr | Val | Glu | Ala | Leu | Leu | Gly | Asn |
| | | 500 | | | | | | 505 | | | | | | 510 | |
| Leu | Val | Ala | Thr | Thr | Gln | Tyr | Asp | Gly | Gly | Gln | Val | Phe | Thr | Asn | His |
| | | 515 | | | | | 520 | | | | | 525 | | | |
| Tyr | Thr | Pro | Glu | Lys | Pro | Ile | Glu | Ser | Thr | Thr | Pro | Pro | Thr | Ser | Gly |
| | 530 | | | | | 535 | | | | | 540 | | | | |
| Thr | Thr | Asp | Thr | Thr | Thr | Asn | Ser | Thr | Thr | Glu | Thr | Thr | Ser | Ile | Thr |
| 545 | | | | | 550 | | | | | 555 | | | | | 560 |
| Ile | Glu | Lys | Gln | Ala | Ile | Arg | Asn | Lys | Glu | | | | | | |
| | | | 565 | | | | | 570 | | | | | | | |

<210> SEQ ID NO 8

<211> LENGTH: 3309

<212> TYPE: DNA

<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 8

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| acagagcagc cattacaacg agaaattcaa ttgaaaaatg cacagttcat ggatactgct | 120 |
| gtaattgaaa aagacggata ttcttacc aa gtgactaatg gtacgcttta tctgactttg | 180 |
| gacgcacaag taaaaaagcc ggtacagctt tcggttagctg ttgagcaaag ttcgcttcaa | 240 |
| acagctcagc cacctaagtt attgatatgaa aacaacgaat atgatgttcc agttacttct | 300 |
| gaaaaaataa cagtagagga ttctgctaaa gaatcaactg aaccagaaaa aataactgta | 360 |
| ccagaaaata cgaagaagaac taacaaaaat gattcggctc cagaaaaaac agaacagccg | 420 |
| accgcaacag aagaggtaac caatccattt gcagaagcaa gaatggcgcc agctactttg | 480 |
| agagcgaatc tggcactgcc ttttaattgca ccacaataca cgacggataa ttctgggact | 540 |
| tatccgacag ctaattggca gccacaggc aatcaaaatg tgttaaacca tcaagggat | 600 |

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| | | | | | | |
|-------------|------------|-------------|------------|------------|------------|------|
| aaagacggta | gtgcacaatg | ggacggccaa | acgagttgga | atggggaccc | tactaatcgc | 660 |
| acaaattctt | atattgagta | tggcggtaga | ggagaccaag | ccgattatgc | catccgaaaa | 720 |
| tatgctagag | aaacaacaac | accagggctt | tttgatgtat | atcttaatgt | gcgtgggaat | 780 |
| gttcagaaa | aatcacgcc | attggatttg | gtcttagtcg | ttgactggtc | cggtagtatg | 840 |
| aatgaaaaca | atcggattgg | tgaagttaa | aaaggagtga | accgttttgt | tgatacattg | 900 |
| gcagatagcg | gtattacca | taacatcaac | atgggctatg | ttggctactc | aagtgacggt | 960 |
| tataataaca | acgccattca | aatggggccg | tttgatacag | tcaaaaatcc | aattaaaaat | 1020 |
| attacgcaa | gtagcactag | aggaggaact | ttcactcaaa | aagcattaag | agatgctggt | 1080 |
| gatatgtag | caacgcaaaa | tggacataag | aaagtcattg | tacttttaac | ggatggcgtc | 1140 |
| ccaaccttct | cttataaagt | gagtcgagtt | caaacagagg | cggatggtcg | cttttacggg | 1200 |
| acacaattta | cgaatcgaca | agatcaacca | ggtagcactt | cttatatctc | tggtagctat | 1260 |
| aatgcgcag | atcaaaaaca | tatcaataaa | cggattaaca | gtacgtttat | cgccacgata | 1320 |
| ggtagaggca | tggtcttaaa | acaacgtggg | attgaaatac | atggattggg | cattcaattg | 1380 |
| caaagcgatc | cacgagctaa | tttatctaaa | caacaagttg | aagataaaat | gcgtgagatg | 1440 |
| gtgtcagccg | atgaaaatgg | agacctttat | tatgaatccg | cggattatgc | accagacatt | 1500 |
| tctgattatt | tagcgaaaaa | agccgttcag | atctcaggaa | cggttgtaaa | cggaaaagta | 1560 |
| gttgatccaa | ttgctgaacc | ttttaatac | gagccaaata | cattatcaat | gaaaagtgtg | 1620 |
| ggtcctgttc | aggttcaaac | attaccagaa | gtgtcgctaa | caggcgctac | aattaatagt | 1680 |
| aatgagattt | atctgggtaa | agggaagaa | attcaaattc | attatcaagt | acgtattcaa | 1740 |
| acagagtcag | aaaacttcaa | acctgatttt | tggtatcaaa | tgaatggtcg | gacaacgttt | 1800 |
| cagccattag | ccacggcccc | tgaaaaagtt | gattttgggg | ttccttcggg | aaaagcacct | 1860 |
| ggcgtgaagt | taaactgtaa | aaaaatctgg | gaagagtatg | atcaagaccc | gacaagtccg | 1920 |
| ccagataatg | tgatttatga | aattagtaga | aagcaagtaa | ctgacacagc | caactggcaa | 1980 |
| actgggtata | ttaaatatc | aaaaccagaa | aatgatacca | gcaatagttg | ggagcgcaaa | 2040 |
| aatgtaacc | aactttccaa | aaccgcgat | gaaagctatc | aagaagtctc | tgggcttccc | 2100 |
| caatacaaca | atcaaggaca | agctttcaat | tatcaaaaca | cccgtgaatt | agcagttcct | 2160 |
| ggttacagtc | aagaaaaaat | cgacgatact | acttggaaaa | acacgaagca | gttcaagcca | 2220 |
| ttagatttaa | aagtaatcaa | aaattcttcc | tcaggtgaga | aaaacttagt | gggagccgtc | 2280 |
| tttgaattga | gtggtaaaaa | tgttcaaaaca | acattagtg | acaataaaga | tggtagctat | 2340 |
| tccttgccaa | aagatgtgcg | cctacaaaaa | ggggaacgct | atacattaac | tgaagtaaaa | 2400 |
| gcacctgcag | gacatgagtt | aggcaagaaa | acgacttggc | aaattgaggt | gagttagcaa | 2460 |
| ggcaaaagta | gcatcgatgg | acaagaagtg | accaccacaa | atcaagttat | tccattggaa | 2520 |
| attgaaaata | aattttcttc | tttgccaatc | agaattagaa | aatacaccat | gcaaaatggc | 2580 |
| aaacaagtga | acttagcaga | ggcgactttt | gcgttgcaaa | gaaaaaatgc | tcaaggaagt | 2640 |
| tacaaaactg | tggcaactca | aaaacagat | actacaggat | tgagctat | ttaaattag | 2700 |
| gaacctgggt | agtatcgaat | ggtggaacaa | tcaggacat | taggctacga | cactcttgc | 2760 |
| ggaaattatg | aatttactgt | tgataaatat | gggaaaatc | actatgcagg | caaaaatatt | 2820 |
| gaagaaaatg | cgccagaatg | gacactgaca | catcaaaata | atgtgaaacc | ttttgactta | 2880 |
| acagtttaata | aaaagccga | taatcagacg | ccacttaag | gagcgaat | ccgtttaaca | 2940 |
| ggaccagata | cggatattga | attacaaaaa | gatggcaaa | aaacggatc | ttttgtttt | 3000 |

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gaaaacttaa aaccaggga atagtctta acagaaacct ttacgccaga aggatatcag 3060
gggttaaaag aaccaatcga attaataatt cgtgaagatg gttcagtcac gatagatggg 3120
gaaaaagtag cagatgtttt aatttctgga gagaagaata atcaaattac tttagacgtt 3180
acgaaccaag caaagggtcc tttacctgaa actggtggca taggacgctt gtgggtttac 3240
ttgatagcga ttagtacatt cgtgatagcg ggtgtttatc tctttattag acgaccagaa 3300
gggagtgtg 3309

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<210> SEQ ID NO 9

<211> LENGTH: 1103

<212> TYPE: PRT

<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 9

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

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| | | | | | | | | | | | | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| 305 | | | | 310 | | | | | | 315 | | | | | | 320 |
| Tyr | Asn | Asn | Asn | Ala | Ile | Gln | Met | Gly | Pro | Phe | Asp | Thr | Val | Lys | Asn | |
| | | | | 325 | | | | | | 330 | | | | | 335 | |
| Pro | Ile | Lys | Asn | Ile | Thr | Pro | Ser | Ser | Thr | Arg | Gly | Gly | Thr | Phe | Thr | |
| | | | 340 | | | | | 345 | | | | | 350 | | | |
| Gln | Lys | Ala | Leu | Arg | Asp | Ala | Gly | Asp | Met | Leu | Ala | Thr | Pro | Asn | Gly | |
| | | 355 | | | | | 360 | | | | | 365 | | | | |
| His | Lys | Lys | Val | Ile | Val | Leu | Leu | Thr | Asp | Gly | Val | Pro | Thr | Phe | Ser | |
| | 370 | | | | | 375 | | | | | 380 | | | | | |
| Tyr | Lys | Val | Ser | Arg | Val | Gln | Thr | Glu | Ala | Asp | Gly | Arg | Phe | Tyr | Gly | |
| | 385 | | | | 390 | | | | | 395 | | | | | 400 | |
| Thr | Gln | Phe | Thr | Asn | Arg | Gln | Asp | Gln | Pro | Gly | Ser | Thr | Ser | Tyr | Ile | |
| | | | | 405 | | | | | 410 | | | | | | 415 | |
| Ser | Gly | Ser | Tyr | Asn | Ala | Pro | Asp | Gln | Asn | Asn | Ile | Asn | Lys | Arg | Ile | |
| | | | 420 | | | | | 425 | | | | | 430 | | | |
| Asn | Ser | Thr | Phe | Ile | Ala | Thr | Ile | Gly | Glu | Ala | Met | Val | Leu | Lys | Gln | |
| | | | 435 | | | | 440 | | | | | 445 | | | | |
| Arg | Gly | Ile | Glu | Ile | His | Gly | Leu | Gly | Ile | Gln | Leu | Gln | Ser | Asp | Pro | |
| | 450 | | | | | 455 | | | | | 460 | | | | | |
| Arg | Ala | Asn | Leu | Ser | Lys | Gln | Gln | Val | Glu | Asp | Lys | Met | Arg | Glu | Met | |
| | 465 | | | | 470 | | | | | 475 | | | | | 480 | |
| Val | Ser | Ala | Asp | Glu | Asn | Gly | Asp | Leu | Tyr | Tyr | Glu | Ser | Ala | Asp | Tyr | |
| | | | | 485 | | | | | 490 | | | | | 495 | | |
| Ala | Pro | Asp | Ile | Ser | Asp | Tyr | Leu | Ala | Lys | Lys | Ala | Val | Gln | Ile | Ser | |
| | | | 500 | | | | | 505 | | | | | 510 | | | |
| Gly | Thr | Val | Val | Asn | Gly | Lys | Val | Val | Asp | Pro | Ile | Ala | Glu | Pro | Phe | |
| | | 515 | | | | | 520 | | | | | 525 | | | | |
| Lys | Tyr | Glu | Pro | Asn | Thr | Leu | Ser | Met | Lys | Ser | Val | Gly | Pro | Val | Gln | |
| | 530 | | | | | 535 | | | | | 540 | | | | | |
| Val | Gln | Thr | Leu | Pro | Glu | Val | Ser | Leu | Thr | Gly | Ala | Thr | Ile | Asn | Ser | |
| | 545 | | | | 550 | | | | | 555 | | | | | 560 | |
| Asn | Glu | Ile | Tyr | Leu | Gly | Lys | Gly | Gln | Glu | Ile | Gln | Ile | His | Tyr | Gln | |
| | | | | 565 | | | | | 570 | | | | | 575 | | |
| Val | Arg | Ile | Gln | Thr | Glu | Ser | Glu | Asn | Phe | Lys | Pro | Asp | Phe | Trp | Tyr | |
| | | | 580 | | | | | 585 | | | | | 590 | | | |
| Gln | Met | Asn | Gly | Arg | Thr | Thr | Phe | Gln | Pro | Leu | Ala | Thr | Ala | Pro | Glu | |
| | | 595 | | | | | 600 | | | | | 605 | | | | |
| Lys | Val | Asp | Phe | Gly | Val | Pro | Ser | Gly | Lys | Ala | Pro | Gly | Val | Lys | Leu | |
| | 610 | | | | | 615 | | | | | | 620 | | | | |
| Asn | Val | Lys | Lys | Ile | Trp | Glu | Glu | Tyr | Asp | Gln | Asp | Pro | Thr | Ser | Arg | |
| | 625 | | | | 630 | | | | | 635 | | | | | 640 | |
| Pro | Asp | Asn | Val | Ile | Tyr | Glu | Ile | Ser | Arg | Lys | Gln | Val | Thr | Asp | Thr | |
| | | | | 645 | | | | | 650 | | | | | 655 | | |
| Ala | Asn | Trp | Gln | Thr | Gly | Tyr | Ile | Lys | Leu | Ser | Lys | Pro | Glu | Asn | Asp | |
| | | | 660 | | | | | 665 | | | | | 670 | | | |
| Thr | Ser | Asn | Ser | Trp | Glu | Arg | Lys | Asn | Val | Thr | Gln | Leu | Ser | Lys | Thr | |
| | | 675 | | | | | 680 | | | | | 685 | | | | |
| Ala | Asp | Glu | Ser | Tyr | Gln | Glu | Val | Leu | Gly | Leu | Pro | Gln | Tyr | Asn | Asn | |
| | 690 | | | | | 695 | | | | | 700 | | | | | |
| Gln | Gly | Gln | Ala | Phe | Asn | Tyr | Gln | Thr | Thr | Arg | Glu | Leu | Ala | Val | Pro | |
| | 705 | | | | 710 | | | | | 715 | | | | | 720 | |
| Gly | Tyr | Ser | Gln | Glu | Lys | Ile | Asp | Asp | Thr | Thr | Trp | Lys | Asn | Thr | Lys | |
| | | | | 725 | | | | | 730 | | | | | 735 | | |

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Gln Phe Lys Pro Leu Asp Leu Lys Val Ile Lys Asn Ser Ser Ser Gly
 740 745 750
 Glu Lys Asn Leu Val Gly Ala Val Phe Glu Leu Ser Gly Lys Asn Val
 755 760 765
 Gln Thr Thr Leu Val Asp Asn Lys Asp Gly Ser Tyr Ser Leu Pro Lys
 770 775 780
 Asp Val Arg Leu Gln Lys Gly Glu Arg Tyr Thr Leu Thr Glu Val Lys
 785 790 795 800
 Ala Pro Ala Gly His Glu Leu Gly Lys Lys Thr Thr Trp Gln Ile Glu
 805 810 815
 Val Ser Glu Gln Gly Lys Val Ser Ile Asp Gly Gln Glu Val Thr Thr
 820 825 830
 Thr Asn Gln Val Ile Pro Leu Glu Ile Glu Asn Lys Phe Ser Ser Leu
 835 840 845
 Pro Ile Arg Ile Arg Lys Tyr Thr Met Gln Asn Gly Lys Gln Val Asn
 850 855 860
 Leu Ala Glu Ala Thr Phe Ala Leu Gln Arg Lys Asn Ala Gln Gly Ser
 865 870 875 880
 Tyr Gln Thr Val Ala Thr Gln Lys Thr Asp Thr Thr Gly Leu Ser Tyr
 885 890 895
 Phe Lys Ile Ser Glu Pro Gly Glu Tyr Arg Met Val Glu Gln Ser Gly
 900 905 910
 Pro Leu Gly Tyr Asp Thr Leu Ala Gly Asn Tyr Glu Phe Thr Val Asp
 915 920 925
 Lys Tyr Gly Lys Ile His Tyr Ala Gly Lys Asn Ile Glu Glu Asn Ala
 930 935 940
 Pro Glu Trp Thr Leu Thr His Gln Asn Asn Leu Lys Pro Phe Asp Leu
 945 950 955 960
 Thr Val Asn Lys Lys Ala Asp Asn Gln Thr Pro Leu Lys Gly Ala Lys
 965 970 975
 Phe Arg Leu Thr Gly Pro Asp Thr Asp Ile Glu Leu Pro Lys Asp Gly
 980 985 990
 Lys Glu Thr Asp Thr Phe Val Phe Glu Asn Leu Lys Pro Gly Lys Tyr
 995 1000 1005
 Val Leu Thr Glu Thr Phe Thr Pro Glu Gly Tyr Gln Gly Leu Lys
 1010 1015 1020
 Glu Pro Ile Glu Leu Ile Ile Arg Glu Asp Gly Ser Val Thr Ile
 1025 1030 1035
 Asp Gly Glu Lys Val Ala Asp Val Leu Ile Ser Gly Glu Lys Asn
 1040 1045 1050
 Asn Gln Ile Thr Leu Asp Val Thr Asn Gln Ala Lys Val Pro Leu
 1055 1060 1065
 Pro Glu Thr Gly Gly Ile Gly Arg Leu Trp Phe Tyr Leu Ile Ala
 1070 1075 1080
 Ile Ser Thr Phe Val Ile Ala Gly Val Tyr Leu Phe Ile Arg Arg
 1085 1090 1095
 Pro Glu Gly Ser Val
 1100

<210> SEQ ID NO 10

<211> LENGTH: 1428

<212> TYPE: DNA

<213> ORGANISM: Staphylococcus epidermidis

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<400> SEQUENCE: 10

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ttccctgatg gtcaattacc agaacagcag caaaacacag gggaagaggg aacgctgctt    180
caaaattatc ggggcttaaa tgacgtcact tatcaagtct atgatgtgac ggatccgttt    240
tatcagcttc gttctgaagg aaaaacggtc caagaggcac agcgtcaatt agcagaaacc    300
ggtgcaacaa atagaaaacc gatcgagaa gataaaacac agacaataaa tggagaagat    360
ggagtgggtt ctttttcatt agctagcaaa gattcgcagc aacgagataa agcctattta    420
tttgttgaag cggagacc agaagtggta aaggaaaaag ctacgaacct agtagtgatt    480
ttgcctgttc aagatccaca agggcaatcg ttaacgcata ttcatttata tccaaaaaat    540
gaagaaaatg cctatgactt accaccactt gaaaaaacgg tactcgataa gcaacaaggc    600
ttaaatacaag gagagccat taactatcag ttaacgactc agattccagc gaatatttta    660
ggatatcagg aattccgttt gtcagataag gcggatacaa cgttgacact ttaccagaa    720
tcaattgagg taaaagtggc tggaaaaaca gttactacag gttacacact gacgacgcaa    780
aagcatggat ttacgcttga tttttcaatt aaagacttac aaaactttgc aaatcaaaaca    840
atgactgtgt cgtatcaaat gcgtttagaa aagaccgctg aacctgacac tgcgattaac    900
aacgaaggac aattagtccac ggacaacat accttgacta aaagagccac agttcgtaca    960
ggcggcaagt cttttgtcaa agttgatagt gaaaatgcga aaatcacctt gccagaggct   1020
gtttttatcg tcaaaaatca agcgggggaa tacctcaatg aaacagcaaa cgggtatcgt   1080
tggcaaaaag aaaaagcatt agctaaaaaa ttcacgtcta atcaagccgg tgaattttca   1140
gttaaaggct taaaagatgg ccagtacttc ttggaagaaa tctctgcacc aaaaggttat   1200
cttctgaatc aaacagaaat tccttttacg gtgggaaaaa attcttatgc aacgaacgga   1260
caacgaacag caccgttaca tgtaatcaat aaaaaagtaa aagagtcagg cttcttacca   1320
aaaacaaatg aagaacgttc tatttggttg acgattgcag gcctgctaatt cattgggatg   1380
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<210> SEQ ID NO 11

<211> LENGTH: 476

<212> TYPE: PRT

<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 11

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Phe Gly Phe Ser Gln Gln Ala Leu Ala Glu Ala Ser Gln Ala Ser Val
 20          25          30
Gln Val Thr Leu His Lys Leu Leu Phe Pro Asp Gly Gln Leu Pro Glu
 35          40          45
Gln Gln Gln Asn Thr Gly Glu Glu Gly Thr Leu Leu Gln Asn Tyr Arg
 50          55          60
Gly Leu Asn Asp Val Thr Tyr Gln Val Tyr Asp Val Thr Asp Pro Phe
 65          70          75          80
Tyr Gln Leu Arg Ser Glu Gly Lys Thr Val Gln Glu Ala Gln Arg Gln
 85          90          95
Leu Ala Glu Thr Gly Ala Thr Asn Arg Lys Pro Ile Ala Glu Asp Lys
100         105         110

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| | | | | | | | | | | | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Thr | Gln | Thr | Ile | Asn | Gly | Glu | Asp | Gly | Val | Val | Ser | Phe | Ser | Leu | Ala |
| | 115 | | | | | | 120 | | | | | 125 | | | |
| Ser | Lys | Asp | Ser | Gln | Gln | Arg | Asp | Lys | Ala | Tyr | Leu | Phe | Val | Glu | Ala |
| | 130 | | | | | 135 | | | | | 140 | | | | |
| Glu | Ala | Pro | Glu | Val | Val | Lys | Glu | Lys | Ala | Ser | Asn | Leu | Val | Val | Ile |
| 145 | | | | | 150 | | | | | 155 | | | | | 160 |
| Leu | Pro | Val | Gln | Asp | Pro | Gln | Gly | Gln | Ser | Leu | Thr | His | Ile | His | Leu |
| | | | | 165 | | | | | 170 | | | | | 175 | |
| Tyr | Pro | Lys | Asn | Glu | Glu | Asn | Ala | Tyr | Asp | Leu | Pro | Pro | Leu | Glu | Lys |
| | | | 180 | | | | | 185 | | | | | 190 | | |
| Thr | Val | Leu | Asp | Lys | Gln | Gln | Gly | Phe | Asn | Gln | Gly | Glu | His | Ile | Asn |
| | | 195 | | | | | 200 | | | | | 205 | | | |
| Tyr | Gln | Leu | Thr | Thr | Gln | Ile | Pro | Ala | Asn | Ile | Leu | Gly | Tyr | Gln | Glu |
| | 210 | | | | | 215 | | | | | 220 | | | | |
| Phe | Arg | Leu | Ser | Asp | Lys | Ala | Asp | Thr | Thr | Leu | Thr | Leu | Leu | Pro | Glu |
| 225 | | | | | 230 | | | | | 235 | | | | | 240 |
| Ser | Ile | Glu | Val | Lys | Val | Ala | Gly | Lys | Thr | Val | Thr | Thr | Gly | Tyr | Thr |
| | | | | 245 | | | | | 250 | | | | | 255 | |
| Leu | Thr | Thr | Gln | Lys | His | Gly | Phe | Thr | Leu | Asp | Phe | Ser | Ile | Lys | Asp |
| | | | 260 | | | | | 265 | | | | | 270 | | |
| Leu | Gln | Asn | Phe | Ala | Asn | Gln | Thr | Met | Thr | Val | Ser | Tyr | Gln | Met | Arg |
| | | 275 | | | | | 280 | | | | | 285 | | | |
| Leu | Glu | Lys | Thr | Ala | Glu | Pro | Asp | Thr | Ala | Ile | Asn | Asn | Glu | Gly | Gln |
| | 290 | | | | | 295 | | | | | 300 | | | | |
| Leu | Val | Thr | Asp | Lys | His | Thr | Leu | Thr | Lys | Arg | Ala | Thr | Val | Arg | Thr |
| 305 | | | | | 310 | | | | | 315 | | | | | 320 |
| Gly | Gly | Lys | Ser | Phe | Val | Lys | Val | Asp | Ser | Glu | Asn | Ala | Lys | Ile | Thr |
| | | | | 325 | | | | | 330 | | | | | 335 | |
| Leu | Pro | Glu | Ala | Val | Phe | Ile | Val | Lys | Asn | Gln | Ala | Gly | Glu | Tyr | Leu |
| | | | 340 | | | | | 345 | | | | | 350 | | |
| Asn | Glu | Thr | Ala | Asn | Gly | Tyr | Arg | Trp | Gln | Lys | Glu | Lys | Ala | Leu | Ala |
| | | 355 | | | | | 360 | | | | | 365 | | | |
| Lys | Lys | Phe | Thr | Ser | Asn | Gln | Ala | Gly | Glu | Phe | Ser | Val | Lys | Gly | Leu |
| | 370 | | | | 375 | | | | | | 380 | | | | |
| Lys | Asp | Gly | Gln | Tyr | Phe | Leu | Glu | Glu | Ile | Ser | Ala | Pro | Lys | Gly | Tyr |
| 385 | | | | | 390 | | | | | 395 | | | | | 400 |
| Leu | Leu | Asn | Gln | Thr | Glu | Ile | Pro | Phe | Thr | Val | Gly | Lys | Asn | Ser | Tyr |
| | | | 405 | | | | | | 410 | | | | | 415 | |
| Ala | Thr | Asn | Gly | Gln | Arg | Thr | Ala | Pro | Leu | His | Val | Ile | Asn | Lys | Lys |
| | | | 420 | | | | | 425 | | | | | 430 | | |
| Val | Lys | Glu | Ser | Gly | Phe | Leu | Pro | Lys | Thr | Asn | Glu | Glu | Arg | Ser | Ile |
| | | 435 | | | | | 440 | | | | | 445 | | | |
| Trp | Leu | Thr | Ile | Ala | Gly | Leu | Leu | Ile | Ile | Gly | Met | Val | Val | Ile | Trp |
| | 450 | | | | | 455 | | | | | 460 | | | | |
| Leu | Phe | Tyr | Gln | Lys | Gln | Lys | Arg | Gly | Glu | Arg | Lys | | | | |
| 465 | | | | | 470 | | | | | 475 | | | | | |

<210> SEQ ID NO 12

<211> LENGTH: 1881

<212> TYPE: DNA

<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 12

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60

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ctcgtgattc acaaaaagaa aatgacggat ttaccagatc cgcttattca aaatagcggg 180
aaagaaatga gcgagtttga taaatatcaa ggactggcag atgtgacggt tagtatttat 240
aacgtgacga acgaatttta cgagcaacga ggggcaggcg caagcgttga tgcagctaaa 300
caagctgtcc aaagtttaac tcctgggaaa cctgttgctc aaggaaccac cgatgcaaat 360
gggaatgca ctgttcagtt acctaaaaaa caaaatggta aagatgcagt gtataccatt 420
aaagaagaac caaaagaggg tgtagttgct gctacgaata tgggtggggc gttcccagtt 480
tacgaaatga tcaagcaaac agatggttcc tataaatatg gaacagaaga attagcgggt 540
gttcataatt atcctaaaaa tgtggttagc aatgatggta gtttacatgt gaaaaagta 600
ggaactgctg aaaatgaagg attaaatggc gcagaatttg ttatttctaa aagcgaaggc 660
tcaccaggca cagtaaaata tatccaagga gtcaaatgat gattatatac atggacaacg 720
gataaagaac aagcaaacg ctttattact gggaaaagtt atgaaattgg cgaatgat 780
ttcacagaag cagagaatgg aacgggagaa ttaacagtta aaaatcttga ggttggttcg 840
tatattttag aagaagtaaa agctccaaat aatgcagaat taattgaaaa tcaacaaaa 900
acaccattta caattgaagc aaacaatcaa acacctgttg aaaaaacagt caaaaatgat 960
acctctaagc ttgataaaac aacaccaagc ttagatggta aagatgtggc aattggcgaa 1020
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gctaataaat acgtcaaat caatttagtt gataaacatg atgcagcctt aacttttgat 1140
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gacaaacac caccaactgt tgaagttgtg acaggtggga aacgtttcat taaagtcgat 1440
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<210> SEQ ID NO 13

<211> LENGTH: 627

<212> TYPE: PRT

<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 13

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Met Lys Gln Leu Lys Lys Val Trp Tyr Thr Val Ser Thr Leu Leu Leu
1           5           10           15

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Ile Leu Pro Leu Phe Thr Ser Val Leu Gly Thr Thr Thr Ala Phe Ala
           20           25           30

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Glu Glu Asn Gly Glu Ser Ala Gln Leu Val Ile His Lys Lys Lys Met
           35           40           45

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| | | | | | | | | | | | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Thr | Asp | Leu | Pro | Asp | Pro | Leu | Ile | Gln | Asn | Ser | Gly | Lys | Glu | Met | Ser |
| 50 | | | | | | 55 | | | | | 60 | | | | |
| Glu | Phe | Asp | Lys | Tyr | Gln | Gly | Leu | Ala | Asp | Val | Thr | Phe | Ser | Ile | Tyr |
| 65 | | | | | 70 | | | | | 75 | | | | | 80 |
| Asn | Val | Thr | Asn | Glu | Phe | Tyr | Glu | Gln | Arg | Ala | Ala | Gly | Ala | Ser | Val |
| | | | | 85 | | | | | | 90 | | | | | 95 |
| Asp | Ala | Ala | Lys | Gln | Ala | Val | Gln | Ser | Leu | Thr | Pro | Gly | Lys | Pro | Val |
| | | | 100 | | | | | | 105 | | | | | 110 | |
| Ala | Gln | Gly | Thr | Thr | Asp | Ala | Asn | Gly | Asn | Val | Thr | Val | Gln | Leu | Pro |
| | | | 115 | | | | | 120 | | | | | 125 | | |
| Lys | Lys | Gln | Asn | Gly | Lys | Asp | Ala | Val | Tyr | Thr | Ile | Lys | Glu | Glu | Pro |
| | | | | | | 135 | | | | | | | 140 | | |
| Lys | Glu | Gly | Val | Val | Ala | Ala | Thr | Asn | Met | Val | Val | Ala | Phe | Pro | Val |
| | | | | | 150 | | | | | 155 | | | | | 160 |
| Tyr | Glu | Met | Ile | Lys | Gln | Thr | Asp | Gly | Ser | Tyr | Lys | Tyr | Gly | Thr | Glu |
| | | | | 165 | | | | | | 170 | | | | | 175 |
| Glu | Leu | Ala | Val | Val | His | Ile | Tyr | Pro | Lys | Asn | Val | Val | Ala | Asn | Asp |
| | | | 180 | | | | | | 185 | | | | | | 190 |
| Gly | Ser | Leu | His | Val | Lys | Lys | Val | Gly | Thr | Ala | Glu | Asn | Glu | Gly | Leu |
| | | | 195 | | | | | | 200 | | | | 205 | | |
| Asn | Gly | Ala | Glu | Phe | Val | Ile | Ser | Lys | Ser | Glu | Gly | Ser | Pro | Gly | Thr |
| | | | | | | 215 | | | | | | | 220 | | |
| Val | Lys | Tyr | Ile | Gln | Gly | Val | Lys | Asp | Gly | Leu | Tyr | Thr | Trp | Thr | Thr |
| | | | | | 230 | | | | | | 235 | | | | 240 |
| Asp | Lys | Glu | Gln | Ala | Lys | Arg | Phe | Ile | Thr | Gly | Lys | Ser | Tyr | Glu | Ile |
| | | | | 245 | | | | | | 250 | | | | | 255 |
| Gly | Glu | Asn | Asp | Phe | Thr | Glu | Ala | Glu | Asn | Gly | Thr | Gly | Glu | Leu | Thr |
| | | | 260 | | | | | | 265 | | | | | 270 | |
| Val | Lys | Asn | Leu | Glu | Val | Gly | Ser | Tyr | Ile | Leu | Glu | Glu | Val | Lys | Ala |
| | | | 275 | | | | | | 280 | | | | | 285 | |
| Pro | Asn | Asn | Ala | Glu | Leu | Ile | Glu | Asn | Gln | Thr | Lys | Thr | Pro | Phe | Thr |
| | | | | | | 295 | | | | | | | 300 | | |
| Ile | Glu | Ala | Asn | Asn | Gln | Thr | Pro | Val | Glu | Lys | Thr | Val | Lys | Asn | Asp |
| | | | | | 310 | | | | | | 315 | | | | 320 |
| Thr | Ser | Lys | Val | Asp | Lys | Thr | Thr | Pro | Ser | Leu | Asp | Gly | Lys | Asp | Val |
| | | | | 325 | | | | | | 330 | | | | | 335 |
| Ala | Ile | Gly | Glu | Lys | Ile | Lys | Tyr | Gln | Ile | Ser | Val | Asn | Ile | Pro | Leu |
| | | | | 340 | | | | | 345 | | | | | 350 | |
| Gly | Ile | Ala | Asp | Lys | Glu | Gly | Asp | Ala | Asn | Lys | Tyr | Val | Lys | Phe | Asn |
| | | | | 355 | | | | | 360 | | | | | 365 | |
| Leu | Val | Asp | Lys | His | Asp | Ala | Ala | Leu | Thr | Phe | Asp | Asn | Val | Thr | Ser |
| | | | | | | 375 | | | | | | | 380 | | |
| Gly | Glu | Tyr | Ala | Tyr | Ala | Leu | Tyr | Asp | Gly | Asp | Thr | Val | Ile | Ala | Pro |
| | | | | | 390 | | | | | | 395 | | | | 400 |
| Glu | Asn | Tyr | Gln | Val | Thr | Glu | Gln | Ala | Asn | Gly | Phe | Thr | Val | Ala | Val |
| | | | | 405 | | | | | | 410 | | | | | 415 |
| Asn | Pro | Ala | Tyr | Ile | Pro | Thr | Leu | Thr | Pro | Gly | Gly | Thr | Leu | Lys | Phe |
| | | | | 420 | | | | | 425 | | | | | 430 | |
| Val | Tyr | Phe | Met | His | Leu | Asn | Glu | Lys | Ala | Asp | Pro | Thr | Lys | Gly | Phe |
| | | | | 435 | | | | | 440 | | | | | 445 | |
| Lys | Asn | Glu | Ala | Asn | Val | Asp | Asn | Gly | His | Thr | Asp | Asp | Gln | Thr | Pro |
| | | | | | | 455 | | | | | | 460 | | | |
| Pro | Thr | Val | Glu | Val | Val | Thr | Gly | Gly | Lys | Arg | Phe | Ile | Lys | Val | Asp |

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| | | | | | | | |
|-----------------------------------------------------------------|-------------------------------------------------|-----|--|-----|--|-----|--|
| 465 | | 470 | | 475 | | 480 | |
| Gly Asp Val Thr | Ala Thr Gln Ala Leu Ala Gly Ala Ser Phe Val Val | 485 | | 490 | | 495 | |
| Arg Asp Gln Asn Ser Asp Thr Ala Asn Tyr Leu Lys Ile Asp Glu Thr | | 500 | | 505 | | 510 | |
| Thr Lys Ala Ala Thr Trp Val Lys Thr Lys Ala Glu Ala Thr Thr Phe | | 515 | | 520 | | 525 | |
| Thr Thr Thr Ala Asp Gly Leu Val Asp Ile Thr Gly Leu Lys Tyr Gly | | 530 | | 535 | | 540 | |
| Thr Tyr Tyr Leu Glu Glu Thr Val Ala Pro Asp Asp Tyr Val Leu Leu | | 545 | | 550 | | 555 | |
| Thr Asn Arg Ile Glu Phe Val Val Asn Glu Gln Ser Tyr Gly Thr Thr | | 565 | | 570 | | 575 | |
| Glu Asn Leu Val Ser Pro Glu Lys Val Pro Asn Lys His Lys Gly Thr | | 580 | | 585 | | 590 | |
| Leu Pro Ser Thr Gly Gly Lys Gly Ile Tyr Val Tyr Leu Gly Ser Gly | | 595 | | 600 | | 605 | |
| Ala Val Leu Leu Leu Ile Ala Gly Val Tyr Phe Ala Arg Arg Arg Lys | | 610 | | 615 | | 620 | |
| Glu Asn Ala | | | | | | | |
| 625 | | | | | | | |

<210> SEQ ID NO 14
 <211> LENGTH: 3387
 <212> TYPE: DNA
 <213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 14

| | |
|--------------------------------------------------------------------|------|
| atgacgacca caggaagaa actgaaagt attttcatgc tgataatatt gagtttatca | 60 |
| aactttgtgc cattatctgc aatagcagac actacagatg atccaacagt tttagaaca | 120 |
| atttcagctg aagtcatttc ggatcagtct ggaaaaaaaa cactgaacat caagctaaat | 180 |
| gcgaataaca ccagtgtgta aaagatagaa aaagaaattg gtctagtoga aaattactta | 240 |
| agtgatgtgg aaagaaaaga aggagatggc tatgcttacc aggtaaatag cgggaaaatt | 300 |
| acgttggaaa tctcatcaaa cactaaacaa actatcgatc tgagttttcc aatcgatcca | 360 |
| gcactttacc acagccaggc aaacaagtct atcgtcgata ataaagaata tgacattatt | 420 |
| gatgagacag aaaataagaa agatacagat gtgtcagtac caaagccaga cgaatagaa | 480 |
| gaagaatcat caaaagaaaa cgaaaattct gtcagcccat ttacattgcc tacattatcc | 540 |
| ttgccagctg tgagtgtgcc atctaataca acgattccta cagaatatac aacagatgat | 600 |
| cagggcactt atcctaaagc cagttggcaa cctacaggaa atacaaatgt tcttgatcat | 660 |
| caaggcaata aaaacggaac aaatcaatgg gatggtataa attcttggaa tggagatcct | 720 |
| aatgatcgga ccattctgta tatcgaatat ggaggaacog gtaatcaagc agactatgcg | 780 |
| atacgaaagt atgcaagga aacaagtaca cccggattgt ttgatgttta tttgaatgct | 840 |
| cgtggaaatg tacaaaaaga tatcacgcct cttgatctcg tattggctgt agactggtca | 900 |
| ggaagtatga acgacaataa tcggatcggg gaagtaaaaga ttgggttoga tcgttttgtc | 960 |
| gatactttag cagatagogg tatcacagac aaaatcaata tgggatattg cggetactca | 1020 |
| agcgaaggat atagctacag taacgggtgca gtacagatgg gttcatttga ttcagtgaaa | 1080 |
| aatcaagtaa aatccattac accttcacgg acaaatggtg gtacttttac acaaaaagca | 1140 |
| ctaagagatg caggaagcat gctatccggt ccaaatggac ataaaaaagt gatcgttttg | 1200 |

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ctgacggatg gtgtaccaac attttcctat aaagtacagc gggtagacgc acaatcaagc 1260
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atctcaagaa tctatgatgc acctgaccaa aacaatctat ccagaagaat cgacagtacg 1380
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cgtatgagac aaatggtttc atcagatgaa aaaggcagtc tttactatga atcagctgat 1560
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caagtggaaa tccaacagaa aaatgaggac ttccatccaa atttctggta tcaaatgaac 1860
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aatctagctg atcgtccaga tcaagttact tttgagattc aacgggaaca tacgacaaat 2040
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acgtgggaac gtgcagacat tgacaaatta tctgcaaata gccgagaaaag ttatcaagag 2160
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gaattacctg taccaggata cgattctcaa caaatagatg caatgacatg gaaaaatact 2280
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ggcgacggaa cctattctct tccagaaaat gtcaaatgc aaaaagaaat gacctatagc 2460
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cagatttctt tagacatcac gaatcaggca aaagtaccat tacctgaaac gggaggaatt 3300
ggccgtttag gaatctatct agtagggatg attggttgtg cgttttctat ttggtatctt 3360
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<210> SEQ ID NO 15

<211> LENGTH: 1129

<212> TYPE: PRT

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<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 15

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Leu Ser Leu Ser Asn Phe Val Pro Leu Ser Ala Ile Ala Asp Thr Thr
20          25          30

Asp Asp Pro Thr Val Leu Glu Thr Ile Ser Ala Glu Val Ile Ser Asp
35          40          45

Gln Ser Gly Lys Lys Ala Leu Asn Ile Lys Leu Asn Ala Asn Asn Thr
50          55          60

Ser Ala Glu Lys Ile Glu Lys Glu Ile Gly Leu Val Glu Asn Tyr Leu
65          70          75          80

Ser Asp Val Glu Arg Lys Glu Gly Asp Gly Tyr Ala Tyr Gln Val Asn
85          90          95

Ser Gly Lys Ile Thr Leu Glu Ile Ser Ser Asn Thr Lys Gln Thr Ile
100         105         110

Asp Leu Ser Phe Pro Ile Asp Pro Ala Leu Tyr His Ser Gln Ala Asn
115        120        125

Lys Leu Ile Val Asp Asn Lys Glu Tyr Asp Ile Ile Asp Glu Thr Glu
130        135        140

Asn Lys Lys Asp Thr Asp Val Ser Val Pro Lys Pro Asp Glu Ile Glu
145        150        155        160

Glu Glu Ser Ser Lys Glu Asn Glu Asn Ser Val Ser Pro Phe Thr Leu
165        170        175

Pro Thr Leu Ser Leu Pro Ala Val Ser Val Pro Ser Asn Gln Thr Ile
180        185        190

Pro Thr Glu Tyr Thr Thr Asp Asp Gln Gly Thr Tyr Pro Lys Ala Ser
195        200        205

Trp Gln Pro Thr Gly Asn Thr Asn Val Leu Asp His Gln Gly Asn Lys
210        215        220

Asn Gly Thr Asn Gln Trp Asp Gly Ile Asn Ser Trp Asn Gly Asp Pro
225        230        235        240

Asn Asp Arg Thr His Ser Tyr Ile Glu Tyr Gly Gly Thr Gly Asn Gln
245        250        255

Ala Asp Tyr Ala Ile Arg Lys Tyr Ala Lys Glu Thr Ser Thr Pro Gly
260        265        270

Leu Phe Asp Val Tyr Leu Asn Ala Arg Gly Asn Val Gln Lys Asp Ile
275        280        285

Thr Pro Leu Asp Leu Val Leu Val Val Asp Trp Ser Gly Ser Met Asn
290        295        300

Asp Asn Asn Arg Ile Gly Glu Val Lys Ile Gly Val Asp Arg Phe Val
305        310        315        320

Asp Thr Leu Ala Asp Ser Gly Ile Thr Asp Lys Ile Asn Met Gly Tyr
325        330        335

Val Gly Tyr Ser Ser Glu Gly Tyr Ser Tyr Ser Asn Gly Ala Val Gln
340        345        350

Met Gly Ser Phe Asp Ser Val Lys Asn Gln Val Lys Ser Ile Thr Pro
355        360        365

Ser Arg Thr Asn Gly Gly Thr Phe Thr Gln Lys Ala Leu Arg Asp Ala
370        375        380

Gly Ser Met Leu Ser Val Pro Asn Gly His Lys Lys Val Ile Val Leu
385        390        395        400

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|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Leu | Thr | Asp | Gly | Val | Pro | Thr | Phe | Ser | Tyr | Lys | Val | Gln | Arg | Val | His |
| | | | | 405 | | | | | 410 | | | | | 415 | |
| Ala | Gln | Ser | Ser | Ser | Asn | Tyr | Tyr | Gly | Thr | Gln | Phe | Ser | Asn | Thr | Gln |
| | | | 420 | | | | | 425 | | | | | 430 | | |
| Asp | Arg | Pro | Gly | Asn | Thr | Ser | Leu | Ile | Ser | Arg | Ile | Tyr | Asp | Ala | Pro |
| | | 435 | | | | | 440 | | | | | 445 | | | |
| Asp | Gln | Asn | Asn | Leu | Ser | Arg | Arg | Ile | Asp | Ser | Thr | Phe | Ile | Ala | Thr |
| | 450 | | | | | 455 | | | | | 460 | | | | |
| Ile | Gly | Glu | Ala | Met | Ala | Leu | Lys | Glu | Arg | Gly | Ile | Glu | Ile | His | Gly |
| 465 | | | | | 470 | | | | | 475 | | | | | 480 |
| Leu | Gly | Ile | Gln | Leu | Gln | Ser | Asp | Pro | Ala | Ala | Gly | Leu | Ser | Lys | Ala |
| | | | | 485 | | | | 490 | | | | | | 495 | |
| Glu | Val | Glu | Ser | Arg | Met | Arg | Gln | Met | Val | Ser | Ser | Asp | Glu | Lys | Gly |
| | | | 500 | | | | | 505 | | | | | 510 | | |
| Asp | Leu | Tyr | Tyr | Glu | Ser | Ala | Asp | His | Ala | Thr | Asp | Ile | Ser | Glu | Tyr |
| | | 515 | | | | | 520 | | | | | 525 | | | |
| Leu | Ala | Lys | Lys | Ala | Val | Gln | Ile | Ser | Ala | Thr | Val | Ser | Asn | Gly | Gln |
| | 530 | | | | | 535 | | | | | 540 | | | | |
| Ile | Asn | Asp | Pro | Ile | Ala | Glu | Pro | Phe | Ile | Tyr | Gln | Pro | Gly | Thr | Leu |
| 545 | | | | | 550 | | | | | 555 | | | | | 560 |
| Ser | Val | Lys | Ser | Val | Gly | Thr | Ser | Pro | Thr | Thr | Val | Thr | Pro | Ser | Ile |
| | | | 565 | | | | | 570 | | | | | | 575 | |
| Ser | Ile | Glu | Gly | Asn | Thr | Ile | Lys | Ser | Asn | Gln | Ile | Tyr | Leu | Gly | Lys |
| | | 580 | | | | | 585 | | | | | | 590 | | |
| Asp | Gln | Glu | Ile | Gln | Ile | His | Tyr | Gln | Val | Arg | Ile | Gln | Thr | Glu | Asn |
| | | 595 | | | | | 600 | | | | | 605 | | | |
| Glu | Asp | Phe | His | Pro | Asn | Phe | Trp | Tyr | Gln | Met | Asn | Gly | Arg | Thr | Thr |
| | 610 | | | | | 615 | | | | | 620 | | | | |
| Phe | Gln | Pro | Asn | Ile | Asp | Thr | Asn | Glu | Leu | Ala | Glu | Phe | Gly | Ile | Pro |
| 625 | | | | | 630 | | | | | 635 | | | | | 640 |
| Ser | Ala | Lys | Ala | Pro | Gly | Val | Ser | Leu | His | Ile | Lys | Lys | Leu | Trp | Glu |
| | | | 645 | | | | | 650 | | | | | | 655 | |
| Glu | Phe | Asp | Asn | Asn | Leu | Ala | Asp | Arg | Pro | Asp | Gln | Val | Thr | Phe | Glu |
| | | 660 | | | | | 665 | | | | | | 670 | | |
| Ile | Gln | Arg | Glu | His | Thr | Thr | Asn | Ala | Ala | Ala | Trp | Lys | Asn | Gly | Tyr |
| | | 675 | | | | | 680 | | | | | 685 | | | |
| Ile | Arg | Ile | Ile | Lys | Pro | Ala | Lys | Asp | Thr | Thr | Asn | Thr | Trp | Glu | Arg |
| | 690 | | | | | 695 | | | | | 700 | | | | |
| Ala | Asp | Ile | Asp | Lys | Leu | Ser | Ala | Asn | Ser | Gly | Glu | Ser | Tyr | Gln | Glu |
| 705 | | | | | 710 | | | | | 715 | | | | | 720 |
| Ile | Leu | Ser | Leu | Pro | Gln | Tyr | Asn | Asn | Gln | Gly | Gln | Ala | Phe | Ser | Tyr |
| | | | 725 | | | | | 730 | | | | | | 735 | |
| Gln | Thr | Ile | Lys | Glu | Leu | Pro | Val | Pro | Gly | Tyr | Asp | Ser | Gln | Gln | Ile |
| | | | 740 | | | | | 745 | | | | | 750 | | |
| Asp | Ala | Met | Thr | Trp | Lys | Asn | Thr | Lys | Gln | Phe | Thr | Pro | Leu | Asn | Leu |
| | | 755 | | | | | 760 | | | | | 765 | | | |
| Lys | Ile | Thr | Lys | Asn | Ser | Ser | Thr | Gly | Glu | Lys | Asp | Leu | Ile | Gly | Ala |
| | 770 | | | | | 775 | | | | | 780 | | | | |
| Val | Phe | Lys | Leu | Thr | Gly | Asp | Ser | Ile | Asp | Thr | Leu | Leu | Thr | Asp | His |
| 785 | | | | | 790 | | | | | 795 | | | | | 800 |
| Gly | Asp | Gly | Thr | Tyr | Ser | Leu | Pro | Glu | Asn | Val | Lys | Leu | Gln | Lys | Glu |
| | | | 805 | | | | | | 810 | | | | | 815 | |
| Met | Thr | Tyr | Thr | Leu | Thr | Glu | Thr | Lys | Ala | Pro | Glu | Gly | His | Gly | Leu |

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| 820 | | | 825 | | | 830 | | | | | | | | | |
|-----|------|-----|-----|-----|-----|------|------|-----|-----|-----|------|-----|------|-----|-----|
| Ser | Lys | Lys | Thr | Thr | Trp | Glu | Ile | Lys | Ile | Ala | Ser | Asp | Gly | Thr | Val |
| | | 835 | | | | | 840 | | | | | | 845 | | |
| Thr | Ile | Asp | Gly | Lys | Thr | Val | Thr | Thr | Ser | Asp | Asp | Thr | Ile | Gln | Leu |
| | 850 | | | | | 855 | | | | | 860 | | | | |
| Thr | Ile | Glu | Asn | Pro | Phe | Val | Glu | Val | Pro | Val | Ala | Val | Arg | Lys | Tyr |
| | 865 | | | | 870 | | | | | 875 | | | | | 880 |
| Ala | Met | Gln | Gly | Thr | Asp | Lys | Glu | Ile | Asn | Leu | Lys | Gly | Ala | Ala | Phe |
| | | | | 885 | | | | | 890 | | | | | | 895 |
| Ser | Leu | Gln | Lys | Lys | Glu | Ala | Asn | Gly | Thr | Tyr | Gln | Pro | Ile | Asp | Ser |
| | | | 900 | | | | | 905 | | | | | | 910 | |
| Gln | Thr | Thr | Asn | Glu | Lys | Gly | Leu | Ala | Ser | Phe | Asp | Ser | Leu | Thr | Pro |
| | | | 915 | | | | 920 | | | | | 925 | | | |
| Gly | Lys | Tyr | Arg | Val | Val | Glu | Thr | Ala | Gly | Pro | Ala | Gly | Tyr | Asp | Thr |
| | 930 | | | | | 935 | | | | | 940 | | | | |
| Ser | Pro | Gly | Asn | Tyr | Glu | Phe | Gln | Ile | Asp | Lys | Tyr | Gly | Lys | Ile | Ile |
| | 945 | | | | 950 | | | | | 955 | | | | | 960 |
| Tyr | Thr | Gly | Lys | Asn | Thr | Glu | Met | Thr | Asn | Asn | Val | Trp | Thr | Leu | Thr |
| | | | | 965 | | | | | 970 | | | | | | 975 |
| His | Gln | Asn | Arg | Leu | Lys | Ala | Phe | Asp | Leu | Thr | Val | His | Lys | Lys | Glu |
| | | | 980 | | | | | 985 | | | | | | 990 | |
| Asp | Asn | Gly | Gln | Thr | Leu | Lys | Gly | Ala | Lys | Phe | Arg | Leu | Gln | Gly | Pro |
| | | | 995 | | | | 1000 | | | | | | 1005 | | |
| Glu | Met | Asp | Leu | Glu | Ser | Pro | Lys | Asp | Gly | Gln | Glu | Thr | Asp | Thr | |
| | 1010 | | | | | 1015 | | | | | 1020 | | | | |
| Phe | Leu | Phe | Glu | Asn | Leu | Lys | Pro | Gly | Thr | Tyr | Thr | Leu | Thr | Glu | |
| | 1025 | | | | | 1030 | | | | | 1035 | | | | |
| Thr | Phe | Thr | Pro | Glu | Gly | Tyr | Gln | Gly | Leu | Lys | Glu | Pro | Val | Thr | |
| | 1040 | | | | | 1045 | | | | | 1050 | | | | |
| Ile | Val | Ile | His | Glu | Asp | Gly | Ser | Ile | Gln | Val | Asp | Gly | Gln | Asp | |
| | 1055 | | | | | 1060 | | | | | 1065 | | | | |
| His | Glu | Ser | Val | Leu | Ser | Pro | Gly | Ala | Lys | Asn | Asn | Gln | Ile | Ser | |
| | 1070 | | | | | 1075 | | | | | 1080 | | | | |
| Leu | Asp | Ile | Thr | Asn | Gln | Ala | Lys | Val | Pro | Leu | Pro | Glu | Thr | Gly | |
| | 1085 | | | | | 1090 | | | | | 1095 | | | | |
| Gly | Ile | Gly | Arg | Leu | Gly | Ile | Tyr | Leu | Val | Gly | Met | Ile | Gly | Cys | |
| | 1100 | | | | | 1105 | | | | | 1110 | | | | |
| Ala | Phe | Ser | Ile | Trp | Tyr | Leu | Phe | Leu | Lys | Lys | Glu | Arg | Gly | Gly | |
| | 1115 | | | | | 1120 | | | | | 1125 | | | | |

Ser

<210> SEQ ID NO 16

<211> LENGTH: 1422

<212> TYPE: DNA

<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 16

```

atgaaaaaac ttggttggt tagtatgtgt ctcttctgt tactatntaa accagctttt    60
actcaggtag caacagaaac agaaacagaa atggttcaga ttactttaca caaattgctt    120
ttccccaaac ggcaactgcc gaaaaatcat ccaaatgacg gacaagaaaa agctttatta    180
caaacgtatc gaggattaaa tgggtgcaca ttccaagttt atgatgtcac agattctttt    240
taccatctac gggaaaaggg caaacggta gaagaagcac aagcagagat cgcaaaaaac    300

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ggtgcgcttt ccggtatggt taccgcagaa gcaacaacta caactcttaa caacgaagat 360
ggatcgcgtt ctttttctct ggccgctaaa gatcaagaaa aaagagataa agcgtatcct 420
ttcattgaat ccaaagtacc agaagtcgtc aaagaaaagg cagagaatat ggtagttggt 480
cttctgttac atggacaaaa caatcaaaaa ctttcaacta tccatttgta tcttaaaaa 540
gaagaaaacg actaccctga tccacctttt gagaaggat tagaagagcc tagaaatgat 600
tttacgattg gtgaaaaaat cacttattcc ttgcatacga caattcctgt aaatatacct 660
gactatcaaa agttcgaatt gtcagatagt gcggatgaag cattaacggt tttaccta 720
agttaacga tttcatcgaa tggagaaaag ctgacagaag gctttgtcat acacaagaaa 780
cctcacggat ttgatgtttt attttcgatc ccttcggttg aaaaatatgc tggaaaaaaa 840
ctgaccattt cttatcagat gcagctaagc agtacagcac aggcgaacaa ggaatcaac 900
aacaacggaa cactggattt tggttttggt gtcagtacaa agaaagtctc tgtatataca 960
gggagtaagc aatttgctaa aatcgagaca aataaaccag ataaacgatt agctggcgca 1020
gtattcctta ttaaaaaaaa agcaggaaat tacctccagc aaacagccaa cggatacaag 1080
tggacaaaaga acgaatcaga tgcgcttcac ctgatttccg ataaaaatgg cgctttttca 1140
atctccgggt tgaaaacagg aagttatcga ttaaaagaga tcgaagcacc ttctggttat 1200
attttaagtg aaacagaaat tccggttacc atttcaactt ttctttctga ggataaagag 1260
gcggacagta tattgaaagt agtcaataaa aaagaaaata gccgtccatt tcttcaaaa 1320
acaaacgaaa cgaaaaatac acttttaggc gttgttggtg tggtattcgc aagctttgca 1380
atctggttgt ttatcaaaaa aagaacagga gtgaaaaaat ga 1422

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<210> SEQ ID NO 17

<211> LENGTH: 473

<212> TYPE: PRT

<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 17

```

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

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Tyr Pro Lys Asn Glu Glu Asn Asp Tyr Pro Asp Pro Pro Phe Glu Lys
 180 185 190
 Val Leu Glu Glu Pro Arg Asn Asp Phe Thr Ile Gly Glu Lys Ile Thr
 195 200 205
 Tyr Ser Leu His Thr Thr Ile Pro Val Asn Ile Leu Asp Tyr Gln Lys
 210 215 220
 Phe Glu Leu Ser Asp Ser Ala Asp Glu Ala Leu Thr Phe Leu Pro Asn
 225 230 235 240
 Ser Leu Thr Ile Ser Ser Asn Gly Glu Lys Leu Thr Glu Gly Phe Val
 245 250 255
 Ile His Lys Lys Pro His Gly Phe Asp Val Leu Phe Ser Ile Pro Ser
 260 265 270
 Leu Glu Lys Tyr Ala Gly Lys Lys Leu Thr Ile Ser Tyr Gln Met Gln
 275 280 285
 Leu Ser Ser Thr Ala Gln Ala Asn Lys Glu Ile Asn Asn Asn Gly Thr
 290 295 300
 Leu Asp Phe Gly Phe Gly Val Ser Thr Lys Lys Val Ser Val Tyr Thr
 305 310 315 320
 Gly Ser Lys Gln Phe Val Lys Ile Glu Thr Asn Lys Pro Asp Lys Arg
 325 330 335
 Leu Ala Gly Ala Val Phe Leu Ile Lys Asn Lys Ala Gly Asn Tyr Leu
 340 345 350
 Gln Gln Thr Ala Asn Gly Tyr Lys Trp Thr Lys Asn Glu Ser Asp Ala
 355 360 365
 Leu His Leu Ile Ser Asp Lys Asn Gly Ala Phe Ser Ile Ser Gly Leu
 370 375 380
 Lys Thr Gly Ser Tyr Arg Leu Lys Glu Ile Glu Ala Pro Ser Gly Tyr
 385 390 395 400
 Ile Leu Ser Glu Thr Glu Ile Pro Phe Thr Ile Ser Thr Phe Leu Ser
 405 410 415
 Glu Asp Lys Glu Ala Asp Ser Ile Leu Lys Val Val Asn Lys Lys Glu
 420 425 430
 Asn Ser Arg Pro Phe Leu Pro Lys Thr Asn Glu Thr Lys Asn Thr Leu
 435 440 445
 Leu Gly Val Val Gly Met Val Phe Ala Ser Phe Ala Ile Trp Leu Phe
 450 455 460
 Ile Lys Lys Arg Thr Gly Val Lys Lys
 465 470

<210> SEQ ID NO 18

<211> LENGTH: 1878

<212> TYPE: DNA

<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 18

```

atgaaaaatc ataaaaaat aaacgttatg ttaggagtcc ttttccttat tttaccatta    60
ctcaciaaaca gcttcggcgc aaaaaaagtg ttgagcaggg agacagcagc tcaagtcate    120
cttcataaaa agaaaatgac tgatttacc gatcctttaa tccaaaacag cgggaaagaa    180
atgagcgaat tcgatcaata ccaaggatta gccgatattt cattttcagt ttataacgtc    240
actcaagaat tttatgcgca acgagataaa ggagcgtcgg tggatgcagc aaaacaagca    300
gtccagtcct tgactcctgg tacaccagtt gcttcaggaa cgacagatgc tgatggaat    360
gtcactttat ctttacctaa aaaacaaaat gggaaagatg cagtctacac gatcaaagaa    420

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gaacccaaaag acggagtgtc agctgccgca aacatggttt tagctttccc tgtatatgag 480
atgatcaaac aagcagatgg ctcttataaa tacgggacag aagaactaga tactatccat 540
ctctacccta aaaatacagt cggtaatgat ggaacgttga aagttacaaa aatcgggtact 600
gccgaaaaacg aagcactaaa tggagcagaa tttattattt ctaaagaaga aggaacacca 660
agcgtcaaaa aatacatcca aagtgtcaca gatggattgt acacttggac aactgatcaa 720
accaaagcca aacatttcat tactggtcac tcttatgaca tcggcaacaa tgactttgcc 780
gaggcatcta ttgaaaaagg ccagttgatc gttaatcatt tagaagttgg aaaatataat 840
ttagaagaag taaaagctcc tgataatgag gaaatgattg aaaagcaaac aatcacgcct 900
tttgagatcc tggcaaatag ccaaacacca gtagaaaaga ccatcaaaaa tgatacgtct 960
aaagttgata aaacaacacc tcaattgaat ggaaaagatg tcgcaatcgg tgaaaaaatt 1020
caatatgaga tttctgtcaa tatcccatta ggtatcgtg ataagaagg aacgcaaac 1080
aagtacacaa cattcaaact tatcgatact catgacgctg ctttaacatt tgataatgat 1140
tcttcaggaa cgtatgctta tgccttatat gatggaaata aagaatcga ccagtaaat 1200
tattctgtca ctgagcaaac agacggattc acggtttcag ttgatccgaa ttatattcct 1260
tcattaactc ctggcgttac attgaaatc gtttactata tgcattttaa cgaaaaagca 1320
gatccaacca aaggattttc taaccaagca aatgtcgata acgggcatac aatgatcaa 1380
acaccaccgt cagtcgatgt cgttactggg ggcaaacgat ttgttaaagt agatggtgac 1440
gttacatcag accaaacact tgcctggagca gaattcgtcg ttcgtgatca agatagtgac 1500
acagcgaat atttatcgat cgaccatcc acaaaagccg tcagctgggt atcggcgaaa 1560
gaatcagcaa cggtttttac aaccacaagt aacggtttaa tcgatgtgac aggtctaaaa 1620
tatggcagct actatctgga agaaacgaaa gcgccagaaa aatatgttcc attaacaaac 1680
cgtgtagcat ttactatcga tgaacaatct tatgtaacag caggacagtt gatttctcct 1740
gaaaaaatac caaataaaca caaaggtaca cttccttcaa caggcggtaa gggaatctat 1800
gtgtatatcg gtgcaggagt agtccttcta ctgattgctg gactgtactt tgctagacgc 1860
aagcacagtc agatttag 1878

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<210> SEQ ID NO 19

<211> LENGTH: 625

<212> TYPE: PRT

<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 19

```

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

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| | | | | | | | | | | | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Gly | Thr | Thr | Asp | Ala | Asp | Gly | Asn | Val | Thr | Leu | Ser | Leu | Pro | Lys | Lys |
| | 115 | | | | | | 120 | | | | | 125 | | | |
| Gln | Asn | Gly | Lys | Asp | Ala | Val | Tyr | Thr | Ile | Lys | Glu | Glu | Pro | Lys | Asp |
| | 130 | | | | | 135 | | | | | 140 | | | | |
| Gly | Val | Ser | Ala | Ala | Ala | Asn | Met | Val | Leu | Ala | Phe | Pro | Val | Tyr | Glu |
| 145 | | | | | 150 | | | | | 155 | | | | | 160 |
| Met | Ile | Lys | Gln | Ala | Asp | Gly | Ser | Tyr | Lys | Tyr | Gly | Thr | Glu | Glu | Leu |
| | | | | 165 | | | | | 170 | | | | | | 175 |
| Asp | Thr | Ile | His | Leu | Tyr | Pro | Lys | Asn | Thr | Val | Gly | Asn | Asp | Gly | Thr |
| | | | 180 | | | | | 185 | | | | | | 190 | |
| Leu | Lys | Val | Thr | Lys | Ile | Gly | Thr | Ala | Glu | Asn | Glu | Ala | Leu | Asn | Gly |
| | | 195 | | | | | 200 | | | | | 205 | | | |
| Ala | Glu | Phe | Ile | Ile | Ser | Lys | Glu | Glu | Gly | Thr | Pro | Ser | Val | Lys | Lys |
| | 210 | | | | | 215 | | | | | 220 | | | | |
| Tyr | Ile | Gln | Ser | Val | Thr | Asp | Gly | Leu | Tyr | Thr | Trp | Thr | Thr | Asp | Gln |
| 225 | | | | | 230 | | | | | | 235 | | | | 240 |
| Thr | Lys | Ala | Lys | His | Phe | Ile | Thr | Gly | His | Ser | Tyr | Asp | Ile | Gly | Asn |
| | | | | 245 | | | | | 250 | | | | | 255 | |
| Asn | Asp | Phe | Ala | Glu | Ala | Ser | Ile | Glu | Lys | Gly | Gln | Leu | Ile | Val | Asn |
| | | 260 | | | | | | 265 | | | | | | 270 | |
| His | Leu | Glu | Val | Gly | Lys | Tyr | Asn | Leu | Glu | Glu | Val | Lys | Ala | Pro | Asp |
| | 275 | | | | | | 280 | | | | | 285 | | | |
| Asn | Ala | Glu | Met | Ile | Glu | Lys | Gln | Thr | Ile | Thr | Pro | Phe | Glu | Ile | Leu |
| | 290 | | | | 295 | | | | | | 300 | | | | |
| Ala | Asn | Ser | Gln | Thr | Pro | Val | Glu | Lys | Thr | Ile | Lys | Asn | Asp | Thr | Ser |
| 305 | | | | | 310 | | | | | | 315 | | | | 320 |
| Lys | Val | Asp | Lys | Thr | Thr | Pro | Gln | Leu | Asn | Gly | Lys | Asp | Val | Ala | Ile |
| | | | | 325 | | | | | 330 | | | | | 335 | |
| Gly | Glu | Lys | Ile | Gln | Tyr | Glu | Ile | Ser | Val | Asn | Ile | Pro | Leu | Gly | Ile |
| | | | 340 | | | | | 345 | | | | | | 350 | |
| Ala | Asp | Lys | Glu | Gly | Thr | Gln | Asn | Lys | Tyr | Thr | Thr | Phe | Lys | Leu | Ile |
| | | 355 | | | | | 360 | | | | | 365 | | | |
| Asp | Thr | His | Asp | Ala | Ala | Leu | Thr | Phe | Asp | Asn | Asp | Ser | Ser | Gly | Thr |
| | 370 | | | | | 375 | | | | | 380 | | | | |
| Tyr | Ala | Tyr | Ala | Leu | Tyr | Asp | Gly | Asn | Lys | Glu | Ile | Asp | Pro | Val | Asn |
| 385 | | | | | 390 | | | | | 395 | | | | | 400 |
| Tyr | Ser | Val | Thr | Glu | Gln | Thr | Asp | Gly | Phe | Thr | Val | Ser | Val | Asp | Pro |
| | | | | 405 | | | | | 410 | | | | | 415 | |
| Asn | Tyr | Ile | Pro | Ser | Leu | Thr | Pro | Gly | Gly | Thr | Leu | Lys | Phe | Val | Tyr |
| | | 420 | | | | | | 425 | | | | | 430 | | |
| Tyr | Met | His | Leu | Asn | Glu | Lys | Ala | Asp | Pro | Thr | Lys | Gly | Phe | Ser | Asn |
| | 435 | | | | | | 440 | | | | | 445 | | | |
| Gln | Ala | Asn | Val | Asp | Asn | Gly | His | Thr | Asn | Asp | Gln | Thr | Pro | Pro | Ser |
| | 450 | | | | | 455 | | | | | 460 | | | | |
| Val | Asp | Val | Val | Thr | Gly | Gly | Lys | Arg | Phe | Val | Lys | Val | Asp | Gly | Asp |
| 465 | | | | | 470 | | | | | 475 | | | | | 480 |
| Val | Thr | Ser | Asp | Gln | Thr | Leu | Ala | Gly | Ala | Glu | Phe | Val | Val | Arg | Asp |
| | | | | 485 | | | | | 490 | | | | | 495 | |
| Gln | Asp | Ser | Asp | Thr | Ala | Lys | Tyr | Leu | Ser | Ile | Asp | Pro | Ser | Thr | Lys |
| | | | 500 | | | | | 505 | | | | | 510 | | |
| Ala | Val | Ser | Trp | Val | Ser | Ala | Lys | Glu | Ser | Ala | Thr | Val | Phe | Thr | Thr |
| | 515 | | | | | | 520 | | | | | 525 | | | |
| Thr | Ser | Asn | Gly | Leu | Ile | Asp | Val | Thr | Gly | Leu | Lys | Tyr | Gly | Thr | Tyr |

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530          535          540
Tyr Leu Glu Glu Thr Lys Ala Pro Glu Lys Tyr Val Pro Leu Thr Asn
545          550          555          560
Arg Val Ala Phe Thr Ile Asp Glu Gln Ser Tyr Val Thr Ala Gly Gln
565          570          575
Leu Ile Ser Pro Glu Lys Ile Pro Asn Lys His Lys Gly Thr Leu Pro
580          585          590
Ser Thr Gly Gly Lys Gly Ile Tyr Val Tyr Ile Gly Ala Gly Val Val
595          600          605
Leu Leu Leu Ile Ala Gly Leu Tyr Phe Ala Arg Arg Lys His Ser Gln
610          615          620

Ile
625

<210> SEQ ID NO 20
<211> LENGTH: 2402
<212> TYPE: PRT
<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 20
Met Lys Asn Lys Gln Gly Phe Leu Pro Asn Leu Leu Asn Lys Tyr Gly
1          5          10          15
Ile Arg Lys Leu Ser Ala Gly Thr Ala Ser Leu Leu Ile Gly Ala Thr
20          25          30
Leu Val Phe Gly Ile Asn Gly Gln Val Lys Ala Ala Glu Thr Asp Asn
35          40          45
Ile Val Ser Gln Asn Gly Asp Asn Lys Thr Asn Asp Ser Glu Ser Ser
50          55          60
Asp Lys Glu Leu Val Lys Ser Glu Asp Asp Lys Thr Ser Ser Thr Ser
65          70          75          80
Thr Asp Thr Asn Leu Glu Ser Glu Phe Asp Gln Asn Asn Asn Pro Ser
85          90          95
Ser Ile Glu Glu Ser Thr Asn Arg Asn Asp Glu Asp Thr Leu Asn Gln
100         105         110
Arg Thr Ser Thr Glu Thr Glu Lys Asp Thr His Val Lys Ser Ala Asp
115         120         125
Thr Gln Thr Thr Asn Glu Thr Thr Asn Lys Asn Asp Asp Asn Ala Thr
130         135         140
Thr Asn His Thr Glu Ser Ile Ser Asp Glu Ser Thr Tyr Gln Ser Asp
145         150         155         160
Asp Ser Lys Thr Thr Gln His Asp Asn Ser Asn Thr Asn Gln Asp Thr
165         170         175
Gln Ser Thr Leu Asn Pro Thr Ser Lys Glu Ser Ser Asn Lys Asp Glu
180         185         190
Ala Thr Ser Pro Thr Pro Lys Glu Ser Thr Ser Ile Glu Lys Thr Asn
195         200         205
Leu Ser Asn Asp Ala Asn His Gln Thr Thr Asp Glu Val Asn His Ser
210         215         220
Asp Ser Asp Asn Met Thr Asn Ser Thr Pro Asn Asp Thr Glu Asn Glu
225         230         235         240
Leu Asp Thr Thr Gln Leu Thr Ser His Asp Glu Ser Pro Ser Pro Gln
245         250         255
Ser Asp Asn Phe Thr Gly Phe Thr Asn Leu Met Ala Thr Pro Leu Asn
260         265         270

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|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Leu | Arg | Asn | Asp | Asn | Pro | Arg | Ile | Asn | Leu | Leu | Ala | Ala | Thr | Glu | Asp |
| | | 275 | | | | | 280 | | | | | 285 | | | |
| Thr | Lys | Pro | Lys | Thr | Tyr | Lys | Lys | Pro | Asn | Asn | Ser | Glu | Tyr | Ser | Tyr |
| | 290 | | | | | 295 | | | | | 300 | | | | |
| Leu | Leu | Asn | Asp | Leu | Gly | Tyr | Asp | Ala | Thr | Thr | Val | Lys | Glu | Asn | Ser |
| 305 | | | | | 310 | | | | | 315 | | | | | 320 |
| Asp | Leu | Arg | His | Ala | Gly | Ile | Ser | Gln | Ser | Gln | Asp | Asn | Thr | Gly | Ser |
| | | | 325 | | | | | 330 | | | | | | 335 | |
| Val | Ile | Lys | Leu | Asn | Leu | Thr | Lys | Trp | Leu | Ser | Leu | Gln | Ser | Asp | Phe |
| | | | 340 | | | | | 345 | | | | | | 350 | |
| Val | Asn | Gly | Gly | Lys | Val | Asn | Leu | Ser | Phe | Ala | Gln | Ser | Asp | Phe | Tyr |
| | | 355 | | | | 360 | | | | | | 365 | | | |
| Thr | Gln | Ile | Glu | Ser | Ile | Thr | Leu | Asn | Asp | Val | Lys | Met | Asp | Thr | Thr |
| | 370 | | | | | 375 | | | | | 380 | | | | |
| Asn | Asn | Gly | Gln | Asn | Trp | Ser | Ala | Pro | Ile | Asn | Gly | Ser | Thr | Val | Arg |
| 385 | | | | | 390 | | | | | 395 | | | | | 400 |
| Ser | Gly | Leu | Ile | Gly | Ser | Val | Thr | Asn | His | Asp | Ile | Val | Ile | Thr | Leu |
| | | | 405 | | | | | 410 | | | | | | 415 | |
| Lys | Asn | Ser | Gln | Thr | Leu | Ser | Ser | Leu | Gly | Tyr | Ser | Asn | Asn | Lys | Pro |
| | | | 420 | | | | | 425 | | | | | | 430 | |
| Val | Tyr | Leu | Thr | His | Thr | Trp | Thr | Thr | Asn | Asp | Gly | Ala | Ile | Ala | Glu |
| | | 435 | | | | | 440 | | | | | 445 | | | |
| Glu | Ser | Ile | Gln | Val | Ala | Ser | Ile | Thr | Pro | Thr | Leu | Asp | Ser | Lys | Ala |
| | 450 | | | | | 455 | | | | | 460 | | | | |
| Pro | Asn | Thr | Ile | Gln | Lys | Ser | Asp | Phe | Thr | Ala | Gly | Arg | Met | Thr | Asn |
| 465 | | | | | 470 | | | | | 475 | | | | | 480 |
| Lys | Ile | Lys | Tyr | Asp | Ser | Ser | Gln | Asn | Ser | Ile | Lys | Ser | Val | His | Thr |
| | | | 485 | | | | | 490 | | | | | | 495 | |
| Phe | Lys | Pro | Asn | Glu | Asn | Phe | Leu | Gln | Thr | Asp | Tyr | Arg | Ala | Val | Leu |
| | | | 500 | | | | | 505 | | | | | 510 | | |
| Tyr | Ile | Lys | Glu | Gln | Val | Asn | Lys | Glu | Leu | Ile | Pro | Tyr | Ile | Asp | Pro |
| | | 515 | | | | | 520 | | | | | 525 | | | |
| Asn | Ser | Val | Lys | Leu | Tyr | Val | Ser | Asp | Pro | Asp | Gly | Asn | Pro | Ile | Ser |
| | 530 | | | | | 535 | | | | | 540 | | | | |
| Gln | Asp | Arg | Tyr | Val | Asn | Gly | Ser | Ile | Asp | Asn | Asp | Gly | Leu | Phe | Asp |
| 545 | | | | | 550 | | | | | 555 | | | | | 560 |
| Ser | Ser | Lys | Ile | Asn | Glu | Ile | Ser | Ile | Lys | Asn | Asn | Asn | Thr | Ser | Gly |
| | | | 565 | | | | | 570 | | | | | | 575 | |
| Gln | Leu | Ser | Asn | Ala | Arg | Thr | Ser | Leu | Asp | Arg | Asn | Val | Phe | Phe | Gly |
| | | | 580 | | | | | 585 | | | | | 590 | | |
| Thr | Leu | Gly | Gln | Ser | Arg | Ser | Tyr | Thr | Ile | Ser | Tyr | Lys | Leu | Lys | Asp |
| | 595 | | | | | | 600 | | | | | 605 | | | |
| Gly | Tyr | Thr | Leu | Glu | Ser | Val | Ala | Ser | Lys | Val | Ser | Ala | Arg | Glu | Thr |
| | 610 | | | | | 615 | | | | | 620 | | | | |
| Phe | Asp | Ser | Trp | Met | Glu | Val | Asp | Tyr | Leu | Asp | Ser | Tyr | Asp | Ser | Gly |
| 625 | | | | | 630 | | | | | 635 | | | | | 640 |
| Ala | Pro | Asn | Lys | Arg | Leu | Leu | Gly | Ser | Tyr | Ala | Ser | Ser | Tyr | Ile | Asp |
| | | | 645 | | | | | 650 | | | | | | 655 | |
| Met | Ile | Asp | Arg | Ile | Pro | Pro | Val | Ala | Pro | Lys | Ala | Asn | Ser | Ile | Thr |
| | | | 660 | | | | | 665 | | | | | 670 | | |
| Thr | Glu | Asp | Thr | Ser | Ile | Lys | Gly | Thr | Ala | Glu | Val | Asp | Thr | Asn | Ile |
| | 675 | | | | | | 680 | | | | | 685 | | | |
| Asn | Leu | Thr | Phe | Asn | Asp | Gly | Arg | Thr | Leu | Asn | Gly | Lys | Val | Asp | Ser |

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| 690 | | 695 | | 700 | |
|---------|-------------|---------|-------------|---------|---------------------|
| Asn Gly | Asn Phe Ser | Ile Ala | Ile Pro Ser | Tyr Tyr | Val Leu Thr Gly |
| 705 | | 710 | | 715 | 720 |
| Lys Glu | Thr Ile Lys | Ile Thr | Ser Ile Asp | Lys Gly | Asp Asn Val Ser |
| | 725 | | 730 | | 735 |
| Pro Ala | Ile Thr | Ile Ser | Val Ile Asp | Lys Thr | Pro Pro Ala Val Lys |
| | 740 | | 745 | | 750 |
| Ala Ile | Ser Asn Lys | Thr Gln | Lys Val Asn | Thr Glu | Ile Glu Pro Ile |
| | 755 | | 760 | | 765 |
| Lys Ile | Glu Ala Thr | Asp Asn | Ser Gly Gln | Ala Val | Thr Asn Lys Val |
| | 770 | | 775 | | 780 |
| Glu Gly | Leu Pro Ala | Gly Met | Thr Phe Asp | Glu Ala | Thr Asn Thr Ile |
| | | 790 | | 795 | 800 |
| Ser Gly | Thr Pro Ser | Glu Val | Gly Ser Tyr | Asp Ile | Thr Val Thr Thr |
| | 805 | | 810 | | 815 |
| Thr Asp | Glu Asn Gly | Asn Ser | Glu Thr Thr | Thr Phe | Thr Ile Asp Val |
| | 820 | | 825 | | 830 |
| Glu Asp | Thr Thr Lys | Pro Thr | Val Glu Ser | Val Ala | Asp Gln Thr Gln |
| | 835 | | 840 | | 845 |
| Glu Val | Asn Thr Glu | Ile Glu | Pro Ile Lys | Ile Glu | Ala Thr Asp Asn |
| | 850 | | 855 | | 860 |
| Ser Gly | Arg Ala Val | Thr Asn | Lys Val Asp | Gly Leu | Pro Asp Gly Val |
| | | 870 | | 875 | 880 |
| Thr Phe | Asp Glu Ala | Thr Asn | Thr Ile Ser | Gly Thr | Pro Ser Glu Val |
| | 885 | | 890 | | 895 |
| Gly Ser | Tyr Asp | Ile Thr | Val Thr Thr | Thr Asp | Glu Ser Gly Asn Val |
| | 900 | | 905 | | 910 |
| Thr Glu | Thr Ile Phe | Thr Ile | Asp Val Glu | Asp Thr | Thr Lys Pro Thr |
| | 915 | | 920 | | 925 |
| Val Glu | Ser Ile Ala | Gly Gln | Thr Gln Glu | Val Asn | Thr Glu Ile Glu |
| | 930 | | 935 | | 940 |
| Pro Ile | Lys Ile Glu | Ala Lys | Asp Asn Ser | Gly Gln | Thr Val Thr Asn |
| | | 950 | | 955 | 960 |
| Lys Val | Asp Gly Leu | Pro Asp | Gly Val Thr | Phe Asp | Glu Ala Thr Asn |
| | 965 | | 970 | | 975 |
| Thr Ile | Ser Gly Thr | Pro Ser | Glu Val Gly | Ser Tyr | Asp Val Thr Val |
| | 980 | | 985 | | 990 |
| Thr Thr | Thr Asp Glu | Ser Gly | Asn Ser Glu | Thr Thr | Thr Phe Thr Ile |
| | 995 | | 1000 | | 1005 |
| Glu Val | Lys Asp Thr | Thr Lys | Pro Thr Val | Glu Ser | Val Ala Asp |
| | 1010 | | 1015 | | 1020 |
| Gln Thr | Gln Glu Val | Asn Thr | Glu Ile Glu | Pro Ile | Lys Ile Glu |
| | 1025 | | 1030 | | 1035 |
| Ala Arg | Asp Asn Ser | Gly Gln | Ala Val Thr | Asn Lys | Val Asp Gly |
| | 1040 | | 1045 | | 1050 |
| Leu Pro | Asp Gly Val | Thr Phe | Asp Glu Ala | Thr Asn | Thr Ile Ser |
| | 1055 | | 1060 | | 1065 |
| Gly Thr | Pro Ser Glu | Val Gly | Ser Tyr Asp | Ile Thr | Val Thr Thr |
| | 1070 | | 1075 | | 1080 |
| Thr Asp | Glu Ser Gly | Asn Val | Thr Glu Thr | Thr Phe | Thr Ile Glu |
| | 1085 | | 1090 | | 1095 |
| Val Glu | Asp Thr Thr | Lys Pro | Thr Val Glu | Asn Val | Ala Asp Gln |
| | 1100 | | 1105 | | 1110 |

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|------|-----|-----|-----|-----|-----|------|-----|-----|-----|-----|-----|------|-----|-----|
| Thr | Gln | Glu | Val | Asn | Thr | Glu | Ile | Thr | Pro | Ile | Thr | Ile | Glu | Ser |
| 1115 | | | | | | 1120 | | | | | | 1125 | | |
| Glu | Asp | Asn | Ser | Gly | Gln | Thr | Val | Thr | Asn | Lys | Val | Asp | Gly | Leu |
| 1130 | | | | | | 1135 | | | | | | 1140 | | |
| Pro | Asp | Gly | Val | Thr | Phe | Asp | Glu | Thr | Thr | Asn | Thr | Ile | Ser | Gly |
| 1145 | | | | | | 1150 | | | | | | 1155 | | |
| Thr | Pro | Ser | Lys | Val | Gly | Ser | Tyr | Asp | Ile | Thr | Val | Thr | Thr | Thr |
| 1160 | | | | | | 1165 | | | | | | 1170 | | |
| Asp | Glu | Ser | Gly | Asn | Ala | Thr | Glu | Thr | Thr | Phe | Thr | Ile | Glu | Val |
| 1175 | | | | | | 1180 | | | | | | 1185 | | |
| Glu | Asp | Thr | Thr | Lys | Pro | Thr | Val | Glu | Asn | Val | Ala | Gly | Gln | Thr |
| 1190 | | | | | | 1195 | | | | | | 1200 | | |
| Gln | Glu | Ile | Asn | Thr | Glu | Ile | Glu | Pro | Ile | Lys | Ile | Glu | Ala | Thr |
| 1205 | | | | | | 1210 | | | | | | 1215 | | |
| Asp | Asn | Ser | Gly | Gln | Ala | Val | Thr | Asn | Lys | Val | Glu | Gly | Leu | Pro |
| 1220 | | | | | | 1225 | | | | | | 1230 | | |
| Ala | Gly | Val | Thr | Phe | Asp | Glu | Ala | Thr | Asn | Thr | Ile | Ser | Gly | Thr |
| 1235 | | | | | | 1240 | | | | | | 1245 | | |
| Pro | Ser | Glu | Val | Gly | Ser | Tyr | Thr | Val | Thr | Val | Thr | Thr | Met | Asp |
| 1250 | | | | | | 1255 | | | | | | 1260 | | |
| Glu | Ser | Gly | Asn | Ala | Thr | Glu | Thr | Thr | Phe | Thr | Ile | Asp | Val | Glu |
| 1265 | | | | | | 1270 | | | | | | 1275 | | |
| Asp | Thr | Thr | Lys | Pro | Thr | Val | Glu | Ser | Val | Ala | Asp | Gln | Thr | Gln |
| 1280 | | | | | | 1285 | | | | | | 1290 | | |
| Glu | Val | Asn | Thr | Glu | Ile | Thr | Pro | Ile | Thr | Ile | Glu | Ser | Glu | Asp |
| 1295 | | | | | | 1300 | | | | | | 1305 | | |
| Asn | Ser | Asp | Gln | Ala | Val | Thr | Asn | Lys | Val | Asp | Gly | Leu | Pro | Asp |
| 1310 | | | | | | 1315 | | | | | | 1320 | | |
| Gly | Val | Thr | Phe | Asp | Glu | Ala | Thr | Asn | Thr | Ile | Ser | Gly | Thr | Pro |
| 1325 | | | | | | 1330 | | | | | | 1335 | | |
| Ser | Glu | Val | Gly | Ser | Tyr | Thr | Val | Thr | Val | Thr | Thr | Thr | Asp | Glu |
| 1340 | | | | | | 1345 | | | | | | 1350 | | |
| Ser | Gly | Asn | Ala | Thr | Glu | Thr | Thr | Phe | Thr | Ile | Asp | Val | Glu | Asp |
| 1355 | | | | | | 1360 | | | | | | 1365 | | |
| Thr | Thr | Lys | Pro | Thr | Val | Lys | Ser | Val | Ser | Asp | Gln | Thr | Gln | Glu |
| 1370 | | | | | | 1375 | | | | | | 1380 | | |
| Val | Asn | Thr | Glu | Ile | Thr | Pro | Ile | Lys | Ile | Glu | Ala | Thr | Asp | Asn |
| 1385 | | | | | | 1390 | | | | | | 1395 | | |
| Ser | Gly | Gln | Thr | Val | Thr | Asn | Lys | Val | Asp | Gly | Leu | Pro | Asp | Gly |
| 1400 | | | | | | 1405 | | | | | | 1410 | | |
| Ile | Thr | Phe | Asp | Glu | Ala | Thr | Asn | Thr | Ile | Ser | Gly | Thr | Pro | Ser |
| 1415 | | | | | | 1420 | | | | | | 1425 | | |
| Glu | Val | Gly | Ser | Tyr | Asp | Ile | Thr | Val | Thr | Thr | Thr | Asp | Glu | Ser |
| 1430 | | | | | | 1435 | | | | | | 1440 | | |
| Gly | Asn | Ala | Thr | Glu | Thr | Thr | Phe | Thr | Ile | Asn | Val | Glu | Asp | Thr |
| 1445 | | | | | | 1450 | | | | | | 1455 | | |
| Thr | Lys | Pro | Thr | Val | Glu | Asp | Ile | Ala | Asp | Gln | Thr | Gln | Glu | Val |
| 1460 | | | | | | 1465 | | | | | | 1470 | | |
| Asn | Thr | Glu | Ile | Glu | Pro | Ile | Lys | Ile | Glu | Ala | Thr | Asp | Asn | Gly |
| 1475 | | | | | | 1480 | | | | | | 1485 | | |
| Gly | Gln | Ala | Val | Thr | Asn | Lys | Val | Asp | Gly | Leu | Pro | Asp | Gly | Val |
| 1490 | | | | | | 1495 | | | | | | 1500 | | |

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| | | | | | | | | | | | | | | |
|------|-----|-----|-----|-----|-----|------|-----|-----|-----|-----|------|-----|-----|-----|
| Thr | Phe | Asp | Glu | Ala | Thr | Asn | Thr | Ile | Ser | Gly | Thr | Pro | Ser | Glu |
| 1505 | | | | | | 1510 | | | | | 1515 | | | |
| Val | Gly | Ser | Tyr | Asp | Ile | Ile | Val | Thr | Thr | Thr | Asp | Glu | Asn | Gly |
| 1520 | | | | | | 1525 | | | | | 1530 | | | |
| Asn | Ser | Glu | Thr | Thr | Thr | Phe | Thr | Ile | Asp | Val | Glu | Asp | Thr | Thr |
| 1535 | | | | | | 1540 | | | | | 1545 | | | |
| Lys | Pro | Thr | Val | Glu | Ser | Val | Val | Asp | Gln | Thr | Gln | Glu | Val | Asn |
| 1550 | | | | | | 1555 | | | | | 1560 | | | |
| Thr | Glu | Ile | Thr | Pro | Ile | Lys | Ile | Glu | Ala | Thr | Asp | Asn | Ser | Gly |
| 1565 | | | | | | 1570 | | | | | 1575 | | | |
| Gln | Ala | Val | Ala | Asn | Lys | Val | Asp | Gly | Leu | Pro | Asn | Gly | Val | Thr |
| 1580 | | | | | | 1585 | | | | | 1590 | | | |
| Phe | Asp | Glu | Thr | Thr | Asn | Thr | Ile | Ser | Gly | Thr | Pro | Ser | Glu | Val |
| 1595 | | | | | | 1600 | | | | | 1605 | | | |
| Gly | Ser | Tyr | Asp | Ile | Ile | Val | Thr | Thr | Thr | Asp | Glu | Ser | Gly | Asn |
| 1610 | | | | | | 1615 | | | | | 1620 | | | |
| Val | Thr | Glu | Thr | Ile | Phe | Thr | Ile | Asp | Val | Glu | Asp | Thr | Thr | Lys |
| 1625 | | | | | | 1630 | | | | | 1635 | | | |
| Pro | Thr | Val | Glu | Ser | Ile | Ala | Gly | Gln | Thr | Gln | Glu | Val | Asn | Thr |
| 1640 | | | | | | 1645 | | | | | 1650 | | | |
| Glu | Ile | Glu | Pro | Ile | Lys | Ile | Glu | Ala | Thr | Asp | Asn | Ser | Gly | Gln |
| 1655 | | | | | | 1660 | | | | | 1665 | | | |
| Ala | Val | Thr | Asn | Lys | Val | Asp | Gly | Leu | Pro | Asn | Gly | Val | Thr | Phe |
| 1670 | | | | | | 1675 | | | | | 1680 | | | |
| Asp | Glu | Ala | Thr | Asn | Thr | Ile | Ser | Gly | Thr | Pro | Ser | Glu | Val | Gly |
| 1685 | | | | | | 1690 | | | | | 1695 | | | |
| Ile | Tyr | Thr | Val | Thr | Val | Thr | Thr | Thr | Asp | Glu | Ser | Gly | Asn | Ala |
| 1700 | | | | | | 1705 | | | | | 1710 | | | |
| Thr | Glu | Thr | Thr | Phe | Thr | Ile | Asp | Val | Glu | Asp | Thr | Thr | Lys | Pro |
| 1715 | | | | | | 1720 | | | | | 1725 | | | |
| Thr | Val | Glu | Ser | Val | Ala | Asp | Gln | Thr | Gln | Glu | Val | Asn | Thr | Glu |
| 1730 | | | | | | 1735 | | | | | 1740 | | | |
| Ile | Thr | Pro | Ile | Thr | Ile | Glu | Ser | Glu | Asp | Asn | Ser | Gly | Gln | Ala |
| 1745 | | | | | | 1750 | | | | | 1755 | | | |
| Val | Thr | Asn | Lys | Val | Glu | Gly | Leu | Pro | Ala | Gly | Met | Thr | Phe | Asp |
| 1760 | | | | | | 1765 | | | | | 1770 | | | |
| Glu | Thr | Thr | Asn | Thr | Ile | Ser | Gly | Thr | Pro | Ser | Glu | Val | Gly | Ser |
| 1775 | | | | | | 1780 | | | | | 1785 | | | |
| Tyr | Thr | Val | Thr | Val | Thr | Thr | Thr | Asp | Glu | Ser | Gly | Asn | Glu | Thr |
| 1790 | | | | | | 1795 | | | | | 1800 | | | |
| Glu | Thr | Thr | Phe | Thr | Ile | Asp | Val | Glu | Asp | Thr | Thr | Lys | Pro | Thr |
| 1805 | | | | | | 1810 | | | | | 1815 | | | |
| Val | Glu | Ser | Ile | Ala | Asn | Gln | Thr | Gln | Glu | Val | Asn | Thr | Glu | Ile |
| 1820 | | | | | | 1825 | | | | | 1830 | | | |
| Thr | Pro | Ile | Lys | Ile | Glu | Ala | Thr | Asp | Asn | Ser | Gly | Gln | Ala | Val |
| 1835 | | | | | | 1840 | | | | | 1845 | | | |
| Thr | Asn | Lys | Val | Asp | Gly | Leu | Pro | Asn | Gly | Val | Thr | Phe | Asp | Glu |
| 1850 | | | | | | 1855 | | | | | 1860 | | | |
| Thr | Thr | Asn | Thr | Ile | Ser | Gly | Thr | Pro | Ser | Glu | Val | Gly | Ser | Tyr |
| 1865 | | | | | | 1870 | | | | | 1875 | | | |
| Asp | Ile | Lys | Val | Thr | Thr | Thr | Asp | Glu | Ser | Gly | Asn | Ala | Thr | Glu |
| 1880 | | | | | | 1885 | | | | | 1890 | | | |
| Thr | Thr | Phe | Thr | Ile | Asn | Val | Glu | Asp | Thr | Thr | Lys | Pro | Thr | Val |

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| | | | | | | | | | | | | | | | | |
|------|-----|-----|-----|-----|-----|------|-----|-----|-----|-----|------|-----|-----|-----|--|------|
| 1895 | | | | | | 1900 | | | | | | | | | | 1905 |
| Glu | Ser | Val | Ala | Asp | Gln | Thr | Gln | Glu | Ile | Asn | Thr | Glu | Ile | Glu | | |
| 1910 | | | | | | 1915 | | | | | 1920 | | | | | |
| Pro | Ile | Lys | Ile | Glu | Ala | Arg | Asp | Asn | Ser | Gly | Gln | Ala | Val | Thr | | |
| 1925 | | | | | | 1930 | | | | | 1935 | | | | | |
| Asn | Lys | Val | Asp | Gly | Leu | Pro | Asp | Gly | Val | Thr | Phe | Asp | Glu | Ala | | |
| 1940 | | | | | | 1945 | | | | | 1950 | | | | | |
| Thr | Asn | Thr | Ile | Ser | Gly | Thr | Pro | Ser | Glu | Val | Gly | Ser | Tyr | Asp | | |
| 1955 | | | | | | 1960 | | | | | 1965 | | | | | |
| Ile | Thr | Val | Thr | Thr | Thr | Asp | Glu | Ser | Gly | Asn | Ala | Thr | Glu | Thr | | |
| 1970 | | | | | | 1975 | | | | | 1980 | | | | | |
| Thr | Phe | Thr | Ile | Asp | Val | Glu | Asp | Thr | Thr | Lys | Pro | Thr | Val | Glu | | |
| 1985 | | | | | | 1990 | | | | | 1995 | | | | | |
| Asp | Ile | Thr | Asp | Gln | Thr | Gln | Glu | Ile | Asn | Thr | Glu | Met | Thr | Pro | | |
| 2000 | | | | | | 2005 | | | | | 2010 | | | | | |
| Ile | Lys | Ile | Glu | Ala | Thr | Asp | Asn | Ser | Gly | Gln | Ala | Val | Thr | Asn | | |
| 2015 | | | | | | 2020 | | | | | 2025 | | | | | |
| Lys | Val | Glu | Gly | Leu | Pro | Asp | Gly | Val | Thr | Phe | Asp | Glu | Ala | Thr | | |
| 2030 | | | | | | 2035 | | | | | 2040 | | | | | |
| Asn | Thr | Ile | Ser | Gly | Thr | Pro | Ser | Glu | Val | Gly | Lys | Tyr | Leu | Ile | | |
| 2045 | | | | | | 2050 | | | | | 2055 | | | | | |
| Thr | Ile | Thr | Thr | Ile | Asp | Lys | Asp | Gly | Asn | Thr | Ala | Thr | Thr | Thr | | |
| 2060 | | | | | | 2065 | | | | | 2070 | | | | | |
| Leu | Thr | Ile | Asn | Val | Ile | Asp | Thr | Thr | Thr | Pro | Glu | Gln | Pro | Thr | | |
| 2075 | | | | | | 2080 | | | | | 2085 | | | | | |
| Ile | Asn | Lys | Val | Thr | Glu | Asn | Ser | Thr | Glu | Val | Asn | Gly | Arg | Gly | | |
| 2090 | | | | | | 2095 | | | | | 2100 | | | | | |
| Glu | Pro | Gly | Thr | Val | Val | Glu | Val | Thr | Phe | Pro | Asp | Gly | Asn | Lys | | |
| 2105 | | | | | | 2110 | | | | | 2115 | | | | | |
| Val | Glu | Gly | Lys | Val | Asp | Ser | Asp | Gly | Asn | Tyr | His | Ile | Gln | Ile | | |
| 2120 | | | | | | 2125 | | | | | 2130 | | | | | |
| Pro | Ser | Glu | Thr | Thr | Leu | Lys | Gly | Gly | Gln | Pro | Leu | Gln | Val | Ile | | |
| 2135 | | | | | | 2140 | | | | | 2145 | | | | | |
| Ala | Ile | Asp | Lys | Ala | Gly | Asn | Lys | Ser | Glu | Ala | Thr | Thr | Thr | Asn | | |
| 2150 | | | | | | 2155 | | | | | 2160 | | | | | |
| Val | Ile | Asp | Thr | Thr | Ala | Pro | Glu | Gln | Pro | Thr | Ile | Asn | Lys | Val | | |
| 2165 | | | | | | 2170 | | | | | 2175 | | | | | |
| Thr | Glu | Asn | Ser | Thr | Glu | Val | Ser | Gly | Arg | Gly | Glu | Pro | Gly | Thr | | |
| 2180 | | | | | | 2185 | | | | | 2190 | | | | | |
| Val | Val | Glu | Val | Thr | Phe | Pro | Asp | Gly | Asn | Lys | Val | Glu | Gly | Lys | | |
| 2195 | | | | | | 2200 | | | | | 2205 | | | | | |
| Val | Asp | Ser | Asp | Gly | Asn | Tyr | His | Ile | Gln | Ile | Pro | Ser | Asp | Glu | | |
| 2210 | | | | | | 2215 | | | | | 2220 | | | | | |
| Arg | Phe | Lys | Val | Gly | Gln | Gln | Leu | Ile | Val | Lys | Val | Val | Asp | Glu | | |
| 2225 | | | | | | 2230 | | | | | 2235 | | | | | |
| Glu | Gly | Asn | Val | Ser | Glu | Pro | Ser | Ile | Thr | Met | Val | Gln | Lys | Glu | | |
| 2240 | | | | | | 2245 | | | | | 2250 | | | | | |
| Asp | Lys | Asn | Ser | Glu | Lys | Leu | Ser | Thr | Val | Thr | Gly | Thr | Val | Thr | | |
| 2255 | | | | | | 2260 | | | | | 2265 | | | | | |
| Lys | Asn | Asn | Ser | Lys | Ser | Leu | Lys | His | Lys | Ala | Ser | Glu | Gln | Gln | | |
| 2270 | | | | | | 2275 | | | | | 2280 | | | | | |
| Ser | Tyr | His | Asn | Lys | Ser | Glu | Lys | Ile | Lys | Asn | Val | Asn | Lys | Pro | | |
| 2285 | | | | | | 2290 | | | | | 2295 | | | | | |

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Thr Lys Ile Val Glu Lys Asp Met Ser Thr Tyr Asp Tyr Ser Arg
 2300 2305 2310
 Tyr Ser Lys Asp Ile Ser Asn Lys Asn Asn Lys Ser Ala Thr Phe
 2315 2320 2325
 Glu Gln Gln Asn Val Ser Asp Ile Asn Asn Asn Gln Tyr Ser Arg
 2330 2335 2340
 Asn Lys Val Asn Gln Pro Val Lys Lys Ser Arg Lys Asn Glu Ile
 2345 2350 2355
 Asn Lys Asp Leu Pro Gln Thr Gly Glu Glu Asn Phe Asn Lys Ser
 2360 2365 2370
 Thr Leu Phe Gly Thr Leu Val Ala Ser Leu Gly Ala Leu Leu Leu
 2375 2380 2385
 Phe Phe Lys Arg Arg Lys Lys Asp Glu Asn Asp Glu Lys Glu
 2390 2395 2400

<210> SEQ ID NO 21

<211> LENGTH: 892

<212> TYPE: PRT

<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 21

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

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| 260 | | | | | 265 | | | | | 270 | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Asp | Val | Thr | Phe | Glu | Val | Asp | Asp | Lys | Val | Lys | Ser | Gly | Asp | Thr | Met |
| | 275 | | | | | | 280 | | | | | 285 | | | |
| Thr | Val | Asn | Ile | Asp | Lys | Asn | Thr | Val | Pro | Ser | Asp | Leu | Thr | Asp | Ser |
| | 290 | | | | | 295 | | | | | 300 | | | | |
| Phe | Ala | Ile | Pro | Lys | Ile | Lys | Asp | Asn | Ser | Gly | Glu | Ile | Ile | Ala | Thr |
| 305 | | | | | 310 | | | | | 315 | | | | | 320 |
| Gly | Thr | Tyr | Asp | Asn | Thr | Asn | Lys | Gln | Ile | Thr | Tyr | Thr | Phe | Thr | Asp |
| | | | | 325 | | | | | 330 | | | | | 335 | |
| Tyr | Val | Asp | Lys | Tyr | Glu | Asn | Ile | Lys | Ala | His | Leu | Lys | Leu | Thr | Ser |
| | | | 340 | | | | | 345 | | | | | 350 | | |
| Tyr | Ile | Asp | Lys | Ser | Lys | Val | Pro | Asn | Asn | Asn | Thr | Lys | Leu | Asp | Val |
| | | 355 | | | | | 360 | | | | | 365 | | | |
| Glu | Tyr | Lys | Thr | Ala | Leu | Ser | Val | Asn | Lys | Thr | Ile | Thr | Val | Glu | |
| | 370 | | | | | 375 | | | | | 380 | | | | |
| Tyr | Gln | Lys | Pro | Asn | Glu | Asn | Arg | Thr | Ala | Asn | Leu | Gln | Ser | Met | Phe |
| 385 | | | | | 390 | | | | | 395 | | | | | 400 |
| Thr | Asn | Ile | Asp | Thr | Lys | Asn | His | Thr | Val | Glu | Gln | Thr | Ile | Tyr | Ile |
| | | | | 405 | | | | | 410 | | | | | 415 | |
| Asn | Pro | Leu | Arg | Tyr | Ser | Ala | Lys | Glu | Thr | Asn | Val | Asn | Ile | Ser | Gly |
| | | | 420 | | | | | 425 | | | | | 430 | | |
| Asn | Gly | Asp | Glu | Gly | Ser | Thr | Ile | Ile | Asp | Asp | Ser | Thr | Ile | Ile | Lys |
| | | 435 | | | | | 440 | | | | | 445 | | | |
| Val | Tyr | Lys | Val | Gly | Asp | Asn | Gln | Asn | Leu | Pro | Asp | Ser | Asn | Arg | Ile |
| | 450 | | | | | 455 | | | | | 460 | | | | |
| Tyr | Asp | Tyr | Ser | Glu | Tyr | Glu | Asp | Val | Thr | Asn | Asp | Asp | Tyr | Ala | Gln |
| 465 | | | | | 470 | | | | | 475 | | | | | 480 |
| Leu | Gly | Asn | Asn | Asn | Asp | Val | Asn | Ile | Asn | Phe | Gly | Asn | Ile | Asp | Ser |
| | | | | 485 | | | | | 490 | | | | | 495 | |
| Pro | Tyr | Ile | Ile | Lys | Val | Ile | Ser | Lys | Tyr | Asp | Pro | Asn | Lys | Asp | Asp |
| | | | 500 | | | | | 505 | | | | | 510 | | |
| Tyr | Thr | Thr | Ile | Gln | Gln | Thr | Val | Thr | Met | Gln | Thr | Thr | Ile | Asn | Glu |
| | | 515 | | | | | 520 | | | | | 525 | | | |
| Tyr | Thr | Gly | Glu | Phe | Arg | Thr | Ala | Ser | Tyr | Asp | Asn | Thr | Ile | Ala | Phe |
| | 530 | | | | | 535 | | | | | 540 | | | | |
| Ser | Thr | Ser | Ser | Gly | Gln | Gly | Gln | Gly | Asp | Leu | Pro | Pro | Glu | Lys | Thr |
| 545 | | | | | 550 | | | | | 555 | | | | | 560 |
| Tyr | Lys | Ile | Gly | Asp | Tyr | Val | Trp | Glu | Asp | Val | Asp | Lys | Asp | Gly | Ile |
| | | | | 565 | | | | | 570 | | | | | 575 | |
| Gln | Asn | Thr | Asn | Asp | Asn | Glu | Lys | Pro | Leu | Ser | Asn | Val | Leu | Val | Thr |
| | | | 580 | | | | | 585 | | | | | 590 | | |
| Leu | Thr | Tyr | Pro | Asp | Gly | Thr | Ser | Lys | Ser | Val | Arg | Thr | Asp | Glu | Glu |
| | | 595 | | | | | 600 | | | | | 605 | | | |
| Gly | Lys | Tyr | Gln | Phe | Asp | Gly | Leu | Lys | Asn | Gly | Leu | Thr | Tyr | Lys | Ile |
| | 610 | | | | | 615 | | | | | 620 | | | | |
| Thr | Phe | Glu | Thr | Pro | Glu | Gly | Tyr | Thr | Pro | Thr | Leu | Lys | His | Ser | Gly |
| 625 | | | | | 630 | | | | | 635 | | | | | 640 |
| Thr | Asn | Pro | Ala | Leu | Asp | Ser | Glu | Gly | Asn | Ser | Val | Trp | Val | Thr | Ile |
| | | | | 645 | | | | | 650 | | | | | 655 | |
| Asn | Gly | Gln | Asp | Asp | Met | Thr | Ile | Asp | Ser | Gly | Phe | Tyr | Gln | Thr | Pro |
| | | | 660 | | | | | 665 | | | | | 670 | | |
| Lys | Tyr | Ser | Leu | Gly | Asn | Tyr | Val | Trp | Tyr | Asp | Thr | Asn | Lys | Asp | Gly |
| | | 675 | | | | | 680 | | | | | 685 | | | |

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Ile Gln Gly Asp Asp Glu Lys Gly Ile Ser Gly Val Lys Val Thr Leu
690          695          700

Lys Asp Glu Asn Gly Asn Ile Ile Ser Thr Thr Thr Thr Asp Glu Asn
705          710          715

Gly Lys Tyr Gln Phe Asp Asn Leu Asn Ser Gly Asn Tyr Ile Val His
725          730          735

Phe Asp Lys Pro Ser Gly Met Thr Gln Thr Thr Thr Asp Ser Gly Asp
740          745          750

Asp Asp Glu Gln Asp Ala Asp Gly Glu Glu Val His Val Thr Ile Thr
755          760          765

Asp His Asp Asp Phe Ser Ile Asp Asn Gly Tyr Tyr Asp Asp Asp Ser
770          775          780

Asp Ser Asp Ser Asp Ser Asp Ser Asp Ser Asp Ser Asp Ser Asp
785          790          795          800

Ser Asp Ser Asp Ser Asp Ser Asp Ser Asp Ser Asp Ser Asp Ser Asp
805          810          815

Ser Asp Ser Asp Ser Asp Ser Asp Ser Asp Ser Asp Ser Asp Ser Asp
820          825          830

Ser Asp Ser Asp Ser Asp Ser Gly Leu Asp Asn Ser Ser Asp Lys Asn
835          840          845

Thr Lys Asp Lys Leu Pro Asp Thr Gly Ala Asn Glu Asp His Asp Ser
850          855          860

Lys Gly Thr Leu Leu Gly Ala Leu Phe Ala Gly Leu Gly Ala Leu Leu
865          870          875          880

Leu Gly Lys Arg Arg Lys Asn Arg Lys Asn Lys Asn
885          890

<210> SEQ ID NO 22
<211> LENGTH: 1973
<212> TYPE: PRT
<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 22

Met Lys Glu Asn Lys Arg Lys Asn Asn Leu Asp Lys Asn Asn Thr Arg
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Phe Ser Ile Arg Lys Tyr Gln Gly Tyr Gly Ala Thr Ser Val Ala Ile
20          25          30

Ile Gly Phe Ile Ile Ile Ser Cys Phe Ser Glu Ala Lys Ala Asp Ser
35          40          45

Asp Lys His Glu Ile Lys Ser His Gln Gln Ser Met Thr Asn His Leu
50          55          60

Thr Thr Leu Pro Ser Asp Asn Gln Glu Asn Thr Ser Asn Asn Glu Phe
65          70          75          80

Asn Asn Arg Asn His Asp Ile Ser His Leu Ser Leu Asn Lys Ser Ile
85          90          95

Gln Met Asp Glu Leu Lys Lys Leu Ile Lys Gln Tyr Lys Ala Ile Asn
100         105         110

Leu Asn Asp Lys Thr Glu Glu Ser Ile Lys Leu Phe Gln Ser Asp Leu
115         120         125

Val Gln Ala Glu Ser Leu Ile Asn Asn Pro Gln Ser Gln Gln His Val
130         135         140

Asp Ala Phe Tyr His Lys Phe Leu Asn Ser Ala Gly Lys Leu Arg Lys
145         150         155         160

Lys Glu Thr Val Ser Ile Lys His Glu Arg Ser Glu Ser Asn Thr Tyr

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| 165 | | | | | 170 | | | | | 175 | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Arg | Leu | Gly | Asp | Glu | Val | Arg | Ser | Gln | Thr | Phe | Ser | His | Ile | Arg | His |
| | | | 180 | | | | | 185 | | | | | 190 | | |
| Lys | Arg | Asn | Ala | Val | Ser | Phe | Arg | Asn | Ala | Asp | Gln | Ser | Asn | Leu | Ser |
| | | 195 | | | | | 200 | | | | | 205 | | | |
| Thr | Asp | Pro | Leu | Lys | Ala | Asn | Glu | Ile | Asn | Pro | Glu | Ile | Gln | Asn | Gly |
| | 210 | | | | | 215 | | | | | 220 | | | | |
| Asn | Phe | Ser | Gln | Val | Ser | Gly | Gly | Pro | Leu | Pro | Thr | Ser | Ser | Lys | Arg |
| 225 | | | | | 230 | | | | | 235 | | | | | 240 |
| Leu | Thr | Val | Val | Thr | Asn | Val | Asp | Asn | Trp | His | Ser | Tyr | Ser | Thr | Asp |
| | | | | 245 | | | | | 250 | | | | | 255 | |
| Pro | Asn | Pro | Glu | Tyr | Pro | Met | Phe | Tyr | Thr | Thr | Thr | Ala | Val | Asn | Tyr |
| | | | 260 | | | | | 265 | | | | | | 270 | |
| Pro | Asn | Phe | Met | Ser | Asn | Gly | Asn | Ala | Pro | Tyr | Gly | Val | Ile | Leu | Gly |
| | | 275 | | | | | 280 | | | | | 285 | | | |
| Arg | Thr | Thr | Asp | Gly | Trp | Asn | Arg | Asn | Val | Ile | Asp | Ser | Lys | Val | Ala |
| | 290 | | | | | 295 | | | | | 300 | | | | |
| Gly | Ile | Tyr | Gln | Asp | Ile | Asp | Val | Val | Pro | Gly | Ser | Glu | Leu | Asn | Val |
| 305 | | | | | 310 | | | | | 315 | | | | | 320 |
| Asn | Phe | Ile | Ser | Thr | Ser | Pro | Val | Phe | Ser | Asp | Gly | Ala | Ala | Gly | Ala |
| | | | | 325 | | | | | 330 | | | | | 335 | |
| Lys | Leu | Lys | Ile | Ser | Asn | Val | Glu | Gln | Asn | Arg | Val | Leu | Phe | Asp | Ser |
| | | | 340 | | | | | 345 | | | | | 350 | | |
| Arg | Leu | Asn | Gly | Met | Gly | Pro | Tyr | Pro | Thr | Gly | Lys | Leu | Ser | Ala | Met |
| | | 355 | | | | | 360 | | | | | 365 | | | |
| Val | Asn | Ile | Pro | Asn | Asp | Ile | Asn | Arg | Val | Arg | Ile | Ser | Phe | Leu | Pro |
| | 370 | | | | | 375 | | | | | 380 | | | | |
| Val | Ser | Ser | Thr | Gly | Arg | Val | Ser | Val | Gln | Arg | Ser | Ser | Arg | Glu | His |
| 385 | | | | | 390 | | | | | 395 | | | | | 400 |
| Gly | Phe | Gly | Asp | Asn | Ser | Ser | Tyr | Tyr | His | Gly | Gly | Ser | Val | Ser | Asp |
| | | | | 405 | | | | | 410 | | | | | 415 | |
| Val | Arg | Ile | Asn | Ser | Gly | Ser | Tyr | Val | Val | Ser | Lys | Val | Thr | Gln | Arg |
| | | | 420 | | | | | 425 | | | | | 430 | | |
| Glu | Tyr | Thr | Thr | Arg | Pro | Asn | Ser | Ser | Asn | Asp | Thr | Phe | Ala | Arg | Ala |
| | | 435 | | | | | 440 | | | | | 445 | | | |
| Thr | Ile | Asn | Leu | Ser | Val | Glu | Asn | Lys | Gly | His | Asn | Gln | Ser | Lys | Asp |
| | 450 | | | | | 455 | | | | | 460 | | | | |
| Thr | Tyr | Tyr | Glu | Val | Ile | Leu | Pro | Gln | Asn | Ser | Arg | Leu | Ile | Ser | Thr |
| 465 | | | | | 470 | | | | | 475 | | | | | 480 |
| Arg | Gly | Gly | Ser | Gly | Asn | Tyr | Asn | Asn | Ala | Thr | Asn | Lys | Leu | Ser | Ile |
| | | | | 485 | | | | | 490 | | | | | 495 | |
| Arg | Leu | Asp | Asn | Leu | Asn | Pro | Gly | Asp | Arg | Arg | Asp | Ile | Ser | Tyr | Thr |
| | | | 500 | | | | | 505 | | | | | 510 | | |
| Val | Asp | Phe | Glu | Ser | Ser | Ser | Pro | Lys | Leu | Ile | Asn | Leu | Asn | Ala | His |
| | | 515 | | | | | | 520 | | | | 525 | | | |
| Leu | Leu | Tyr | Lys | Thr | Asn | Ala | Thr | Phe | Arg | Gly | Asn | Asp | Gly | Gln | Arg |
| | 530 | | | | | 535 | | | | | 540 | | | | |
| Thr | Gly | Asp | Asn | Ile | Val | Asp | Leu | Gln | Ser | Ile | Ala | Leu | Leu | Met | Asn |
| 545 | | | | | 550 | | | | | 555 | | | | | 560 |
| Lys | Asp | Val | Leu | Glu | Thr | Glu | Leu | Asn | Glu | Ile | Asp | Lys | Phe | Ile | Arg |
| | | | | 565 | | | | | 570 | | | | | 575 | |
| Asp | Leu | Asn | Glu | Ala | Asp | Phe | Thr | Ile | Asp | Ser | Trp | Ser | Ala | Leu | Gln |
| | | | 580 | | | | | 585 | | | | | 590 | | |

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Glu Lys Met Thr Glu Gly Gly Asn Ile Leu Asn Glu Gln Gln Asn Gln
 595 600 605
 Val Ala Leu Glu Asn Gln Ala Ser Gln Glu Thr Ile Asn Asn Val Thr
 610 615 620
 Gln Ser Leu Glu Ile Leu Lys Asn Asn Leu Lys Tyr Lys Thr Pro Ser
 625 630 635 640
 Gln Pro Ile Ile Lys Ser Asn Asn Gln Ile Pro Asn Ile Thr Ile Ser
 645 650 655
 Pro Ala Asp Lys Ala Asp Lys Leu Thr Ile Thr Tyr Gln Asn Thr Asp
 660 665 670
 Asn Glu Ser Ala Ser Ile Ile Gly Asn Lys Leu Asn Asn Gln Trp Ser
 675 680 685
 Leu Asn Asn Asn Ile Pro Gly Ile Glu Ile Asp Met Gln Thr Gly Leu
 690 695 700
 Val Thr Ile Asp Tyr Lys Ala Val Tyr Pro Glu Ser Val Val Gly Ala
 705 710 715 720
 Asn Asp Lys Thr Gly Asn Ser Asp Ala Ser Ala Glu Ser Arg Ile Thr
 725 730 735
 Met Pro Arg Lys Glu Ala Thr Pro Leu Ser Pro Ile Val Glu Ala Asn
 740 745 750
 Glu Glu Arg Val Asn Val Val Ile Ala Pro Asn Gly Glu Ala Thr Gln
 755 760 765
 Ile Ala Ile Lys Tyr Arg Thr Pro Asp Gly Gln Glu Ala Thr Leu Val
 770 775 780
 Ala Ser Lys Asn Gly Ser Ser Trp Thr Leu Asn Lys Gln Ile Asp Tyr
 785 790 795 800
 Val Asn Ile Glu Glu Asn Ser Gly Lys Val Thr Ile Gly Tyr Gln Ala
 805 810 815
 Val Gln Pro Glu Ser Glu Val Ile Ala Thr Glu Thr Lys Gly Asn Ser
 820 825 830
 Asp Glu Ser Ala Glu Ser Arg Val Thr Met Pro Arg Lys Glu Ala Thr
 835 840 845
 Pro His Ser Pro Ile Val Glu Ala Asn Glu Glu His Val Asn Val Thr
 850 855 860
 Ile Ala Pro Asn Gly Glu Ala Thr Gln Ile Ala Ile Lys Tyr Arg Thr
 865 870 875 880
 Pro Asp Gly Gln Glu Thr Thr Leu Ile Ala Ser Lys Asn Gly Ser Ser
 885 890 895
 Trp Thr Leu Asn Lys Gln Ile Asp Tyr Val Asn Ile Glu Glu Asn Ser
 900 905 910
 Gly Lys Val Thr Ile Gly Tyr Gln Ala Val Gln Leu Glu Ser Glu Val
 915 920 925
 Ile Ala Thr Glu Thr Lys Gly Asn Ser Asp Ala Ser Ala Glu Ser Arg
 930 935 940
 Ile Thr Met Leu Arg Lys Glu Ala Thr Pro His Ser Pro Ile Val Glu
 945 950 955 960
 Ala Asn Glu Glu His Val Asn Val Thr Ile Ala Pro Asn Gly Glu Ala
 965 970 975
 Thr Gln Ile Ala Ile Lys Tyr Arg Thr Pro Asp Gly Gln Glu Ala Thr
 980 985 990
 Leu Val Ala Ser Lys Asn Glu Ser Ser Trp Thr Leu Asn Lys Gln Ile
 995 1000 1005

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| | | | | | | | | | | | | | | |
|------|-----|-----|-----|-----|-----|------|-----|-----|-----|-----|------|-----|-----|-----|
| Asp | His | Val | Asn | Ile | Asp | Glu | Asn | Ser | Gly | Lys | Val | Thr | Ile | Gly |
| 1010 | | | | | | 1015 | | | | | 1020 | | | |
| Tyr | Gln | Ala | Val | Gln | Pro | Glu | Ser | Glu | Ile | Ile | Ala | Thr | Glu | Thr |
| 1025 | | | | | | 1030 | | | | | 1035 | | | |
| Lys | Gly | Asn | Ser | Asp | Ala | Ser | Ala | Glu | Ser | Arg | Ile | Thr | Met | Pro |
| 1040 | | | | | | 1045 | | | | | 1050 | | | |
| Arg | Lys | Glu | Ala | Thr | Pro | Ile | Pro | Pro | Thr | Leu | Glu | Ala | Ser | Val |
| 1055 | | | | | | 1060 | | | | | 1065 | | | |
| Gln | Glu | Ala | Ser | Val | Thr | Val | Thr | Pro | Asn | Glu | Asn | Ala | Thr | Lys |
| 1070 | | | | | | 1075 | | | | | 1080 | | | |
| Val | Phe | Ile | Lys | Tyr | Leu | Asp | Ile | Asn | Asp | Glu | Ile | Ser | Thr | Ile |
| 1085 | | | | | | 1090 | | | | | 1095 | | | |
| Ile | Ala | Ser | Lys | Ile | Asn | Gln | Gln | Trp | Thr | Leu | Asn | Lys | Asp | Asn |
| 1100 | | | | | | 1105 | | | | | 1110 | | | |
| Phe | Gly | Ile | Lys | Ile | Asn | Pro | Leu | Thr | Gly | Lys | Val | Ile | Ile | Ser |
| 1115 | | | | | | 1120 | | | | | 1125 | | | |
| Tyr | Val | Ala | Val | Gln | Pro | Glu | Ser | Asp | Val | Ile | Ala | Ile | Glu | Ser |
| 1130 | | | | | | 1135 | | | | | 1140 | | | |
| Gln | Gly | Asn | Ser | Asp | Leu | Ser | Glu | Glu | Ser | Arg | Ile | Ile | Met | Pro |
| 1145 | | | | | | 1150 | | | | | 1155 | | | |
| Thr | Lys | Glu | Glu | Pro | Pro | Glu | Pro | Pro | Ile | Leu | Glu | Ser | Asp | Ser |
| 1160 | | | | | | 1165 | | | | | 1170 | | | |
| Ile | Glu | Ala | Lys | Val | Asn | Ile | Phe | Pro | Asn | Asp | Glu | Ala | Thr | Arg |
| 1175 | | | | | | 1180 | | | | | 1185 | | | |
| Ile | Val | Ile | Met | Tyr | Thr | Ser | Leu | Glu | Gly | Gln | Glu | Ala | Thr | Leu |
| 1190 | | | | | | 1195 | | | | | 1200 | | | |
| Val | Ala | Ser | Lys | Asn | Glu | Ser | Ser | Trp | Thr | Leu | Asn | Lys | Gln | Ile |
| 1205 | | | | | | 1210 | | | | | 1215 | | | |
| Asp | His | Val | Asn | Ile | Asp | Glu | Asn | Ser | Gly | Lys | Val | Thr | Ile | Gly |
| 1220 | | | | | | 1225 | | | | | 1230 | | | |
| Tyr | Gln | Ala | Val | Gln | Pro | Glu | Ser | Glu | Val | Ile | Ala | Thr | Glu | Thr |
| 1235 | | | | | | 1240 | | | | | 1245 | | | |
| Lys | Gly | Asn | Ser | Asp | Ala | Ser | Ala | Glu | Ser | Arg | Val | Thr | Met | Pro |
| 1250 | | | | | | 1255 | | | | | 1260 | | | |
| Arg | Lys | Glu | Ala | Thr | Pro | His | Ser | Pro | Ile | Val | Glu | Thr | Asn | Glu |
| 1265 | | | | | | 1270 | | | | | 1275 | | | |
| Glu | Arg | Val | Asn | Val | Val | Ile | Ala | Pro | Asn | Gly | Glu | Ala | Thr | Gln |
| 1280 | | | | | | 1285 | | | | | 1290 | | | |
| Ile | Ala | Ile | Lys | Tyr | Arg | Thr | Pro | Asp | Gly | Gln | Glu | Thr | Thr | Leu |
| 1295 | | | | | | 1300 | | | | | 1305 | | | |
| Ile | Ala | Ser | Lys | Asn | Gly | Ser | Ser | Trp | Thr | Leu | Asn | Lys | Gln | Ile |
| 1310 | | | | | | 1315 | | | | | 1320 | | | |
| Asp | His | Val | Asn | Ile | Asp | Glu | Asn | Ser | Gly | Lys | Val | Thr | Ile | Gly |
| 1325 | | | | | | 1330 | | | | | 1335 | | | |
| Tyr | Gln | Ala | Val | Gln | Pro | Glu | Ser | Glu | Ile | Ile | Ala | Thr | Glu | Thr |
| 1340 | | | | | | 1345 | | | | | 1350 | | | |
| Lys | Gly | Asn | Ser | Asp | Ala | Ser | Ala | Glu | Ser | Arg | Ile | Thr | Met | Pro |
| 1355 | | | | | | 1360 | | | | | 1365 | | | |
| Arg | Lys | Glu | Ala | Ile | Pro | His | Ser | Pro | Ile | Val | Glu | Ala | Asn | Glu |
| 1370 | | | | | | 1375 | | | | | 1380 | | | |
| Glu | His | Val | Asn | Val | Thr | Ile | Ala | Pro | Asn | Gly | Glu | Thr | Thr | Gln |
| 1385 | | | | | | 1390 | | | | | 1395 | | | |
| Ile | Ala | Val | Lys | Tyr | Arg | Thr | Pro | Asp | Gly | Gln | Glu | Ala | Thr | Leu |

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| | | |
|-----------------------------|---------------------------------|------|
| 1400 | 1405 | 1410 |
| Ile Ala Ser Lys Asn Glu Ser | Ser Trp Thr Leu Asn Lys Gln Ile | |
| 1415 | 1420 | 1425 |
| Asp His Val Asn Ile Asp Glu | Asn Ser Gly Lys Val Thr Ile Gly | |
| 1430 | 1435 | 1440 |
| Tyr Gln Ala Val Gln Pro Glu | Ser Glu Val Ile Ala Thr Glu Thr | |
| 1445 | 1450 | 1455 |
| Lys Gly Asn Ser Asp Ala Ser | Ala Glu Ser Arg Ile Thr Met Pro | |
| 1460 | 1465 | 1470 |
| Val Lys Glu Lys Thr Pro Ala | Pro Pro Ile Ser Ile Ile Asn Glu | |
| 1475 | 1480 | 1485 |
| Ser Asn Ala Ser Val Glu Ile | Ile Pro Gln Val Asn Val Thr Gln | |
| 1490 | 1495 | 1500 |
| Leu Ser Leu Gln Tyr Ile Asp | Ala Lys Gly Gln Gln Gln Asn Leu | |
| 1505 | 1510 | 1515 |
| Ile Ala Thr Leu Asn Gln Asn | Gln Trp Thr Leu Asn Lys Asn Val | |
| 1520 | 1525 | 1530 |
| Ser His Ile Thr Val Asp Lys | Asn Thr Gly Lys Val Leu Ile Asn | |
| 1535 | 1540 | 1545 |
| Tyr Gln Ala Val Tyr Pro Glu | Ser Glu Val Ile Ala Arg Glu Ser | |
| 1550 | 1555 | 1560 |
| Lys Gly Asn Ser Asp Ser Ser | Asn Val Ser Met Val Ile Met Pro | |
| 1565 | 1570 | 1575 |
| Arg Lys Thr Ala Thr Pro Lys | Pro Pro Ile Ile Lys Val Asp Glu | |
| 1580 | 1585 | 1590 |
| Met Asn Ala Ser Leu Ala Ile | Ile Pro Tyr Lys Asn Asn Thr Ala | |
| 1595 | 1600 | 1605 |
| Ile Asn Ile His Tyr Ile Asp | Lys Lys Gly Ile Lys Ser Met Val | |
| 1610 | 1615 | 1620 |
| Thr Ala Ile Lys Asn Asn Asp | Gln Trp Gln Leu Asp Glu Lys Ile | |
| 1625 | 1630 | 1635 |
| Lys Tyr Val Lys Ile Asp Ala | Lys Thr Gly Thr Val Ile Ile Asn | |
| 1640 | 1645 | 1650 |
| Tyr Gln Ile Val Gln Glu Asn | Ser Glu Ile Ile Ala Thr Ala Ile | |
| 1655 | 1660 | 1665 |
| Asn Gly Asn Ser Asp Lys Ser | Glu Glu Val Lys Val Leu Met Pro | |
| 1670 | 1675 | 1680 |
| Ile Lys Glu Phe Thr Pro Leu | Ala Pro Leu Leu Glu Thr Asn Tyr | |
| 1685 | 1690 | 1695 |
| Lys Lys Ala Thr Val Ser Ile | Leu Pro Gln Ser Asn Ala Thr Lys | |
| 1700 | 1705 | 1710 |
| Leu Asp Phe Lys Tyr Arg Asp | Lys Lys Gly Asp Ser Lys Ile Ile | |
| 1715 | 1720 | 1725 |
| Ile Val Lys Arg Phe Lys Asn | Ile Trp Lys Ala Asn Glu Gln Ile | |
| 1730 | 1735 | 1740 |
| Ser Gly Val Thr Ile Asn Pro | Glu Phe Gly Gln Val Val Ile Asn | |
| 1745 | 1750 | 1755 |
| Tyr Gln Ala Val Tyr Pro Glu | Ser Asp Ile Leu Ala Ala Gln Tyr | |
| 1760 | 1765 | 1770 |
| Val Gly Asn Ser Asp Ala Ser | Glu Trp Ala Lys Val Lys Met Pro | |
| 1775 | 1780 | 1785 |
| Lys Lys Glu Leu Ala Pro His | Ser Pro Ser Leu Ile Tyr Asp Asn | |
| 1790 | 1795 | 1800 |

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Arg Asn Asn Lys Ile Leu Ile Ala Pro Asn Ser Asn Ala Thr Glu
 1805 1810 1815
 Met Glu Leu Ser Tyr Val Asp Lys Asn Asn Gln Ser Leu Lys Val
 1820 1825 1830
 Lys Ala Leu Lys Ile Asn Asn Arg Trp Lys Phe Asp Ser Ser Val
 1835 1840 1845
 Ser Asn Ile Ser Ile Asn Pro Asn Thr Gly Lys Ile Val Leu Gln
 1850 1855 1860
 Pro Gln Phe Leu Leu Thr Asn Ser Lys Ile Ile Val Phe Ala Lys
 1865 1870 1875
 Lys Gly Asn Ser Asp Ala Ser Ile Ser Val Ser Leu Arg Val Pro
 1880 1885 1890
 Ala Val Lys Lys Ile Glu Leu Glu Pro Met Phe Asn Val Pro Val
 1895 1900 1905
 Leu Val Ser Leu Asn Lys Lys Arg Ile Gln Phe Asp Asp Cys Ser
 1910 1915 1920
 Gly Val Lys Asn Cys Leu Asn Lys Gln Ile Ser Lys Thr Gln Leu
 1925 1930 1935
 Pro Asp Thr Gly Tyr Ser Asp Lys Ala Ser Lys Ser Asn Ile Leu
 1940 1945 1950
 Ser Val Leu Leu Leu Gly Phe Gly Phe Leu Ser Tyr Ser Arg Lys
 1955 1960 1965
 Arg Lys Glu Lys Gln
 1970

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<212> TYPE: PRT

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<400> SEQUENCE: 23

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 Thr Tyr Asn Phe Gln Ile Thr Ser Asn Asn Lys Glu Lys Thr Ser Arg
 35 40 45
 Ile Gly Val Ala Ile Ala Leu Asn Asn Arg Asp Lys Leu Gln Lys Phe
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 Ser Ile Arg Lys Tyr Ala Ile Gly Thr Phe Ser Thr Val Ile Ala Thr
 65 70 75 80
 Leu Val Phe Met Gly Ile Asn Thr Asn His Ala Ser Ala Asp Glu Leu
 85 90 95
 Asn Gln Asn Gln Lys Leu Ile Lys Gln Leu Asn Gln Thr Asp Asp Asp
 100 105 110
 Asp Ser Asn Thr His Ser Gln Glu Ile Glu Asn Asn Lys Gln Asn Ser
 115 120 125
 Ser Gly Lys Thr Glu Ser Leu Arg Ser Ser Thr Ser Gln Asn Gln Ala
 130 135 140
 Asn Ala Arg Leu Ser Asp Gln Phe Lys Asp Thr Asn Glu Thr Ser Gln
 145 150 155 160
 Gln Leu Pro Thr Asn Val Ser Asp Asp Ser Ile Asn Gln Ser His Ser
 165 170 175
 Glu Ala Asn Met Asn Asn Glu Pro Leu Lys Val Asp Asn Ser Thr Met

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| 180 | | | | | 185 | | | | | 190 | | | | | |
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| Gln | Ala | His | Ser | Lys | Ile | Val | Ser | Asp | Ser | Asp | Gly | Asn | Ala | Ser | Glu |
| | | 195 | | | | | | 200 | | | | 205 | | | |
| Asn | Lys | His | His | Lys | Leu | Thr | Glu | Asn | Val | Leu | Ala | Glu | Ser | Arg | Ala |
| | 210 | | | | | 215 | | | | | 220 | | | | |
| Ser | Lys | Asn | Asp | Lys | Glu | Lys | Glu | Asn | Leu | Gln | Glu | Lys | Asp | Lys | Ser |
| | 225 | | | | | 230 | | | | | 235 | | | | 240 |
| Gln | Gln | Val | His | Pro | Pro | Leu | Asp | Lys | Asn | Ala | Leu | Gln | Ala | Phe | Phe |
| | | | | 245 | | | | | 250 | | | | | 255 | |
| Asp | Ala | Ser | Tyr | His | Asn | Tyr | Arg | Met | Ile | Asp | Arg | Asp | Arg | Ala | Asp |
| | | | 260 | | | | | 265 | | | | | | 270 | |
| Ala | Thr | Glu | Tyr | Gln | Lys | Val | Lys | Ser | Thr | Phe | Asp | Tyr | Val | Asn | Asp |
| | | 275 | | | | | | 280 | | | | | | 285 | |
| Leu | Leu | Gly | Asn | Asn | Gln | Asn | Ile | Pro | Ser | Glu | Gln | Leu | Val | Ser | Ala |
| | 290 | | | | | 295 | | | | | 300 | | | | |
| Tyr | Gln | Gln | Leu | Glu | Lys | Ala | Leu | Glu | Leu | Ala | Arg | Thr | Leu | Pro | Gln |
| | 305 | | | | | 310 | | | | | 315 | | | | 320 |
| Gln | Ser | Thr | Thr | Glu | Lys | Arg | Gly | Arg | Arg | Ser | Thr | Arg | Ser | Val | Val |
| | | | | 325 | | | | | 330 | | | | | 335 | |
| Glu | Asn | Arg | Ser | Ser | Arg | Ser | Asp | Tyr | Leu | Asp | Ala | Arg | Thr | Glu | Tyr |
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| Tyr | Val | Ser | Lys | Asp | Asp | Asp | Asp | Ser | Gly | Phe | Pro | Pro | Gly | Thr | Phe |
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| Phe | His | Ala | Ser | Asn | Arg | Arg | Trp | Pro | Tyr | Asn | Leu | Pro | Arg | Ser | Arg |
| | 370 | | | | | 375 | | | | | 380 | | | | |
| Asn | Ile | Leu | Arg | Ala | Ser | Asp | Val | Gln | Gly | Asn | Ala | Tyr | Ile | Thr | Thr |
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| Lys | Arg | Leu | Lys | Asp | Gly | Tyr | Gln | Trp | Asp | Ile | Leu | Phe | Asn | Ser | Asn |
| | | | | 405 | | | | | 410 | | | | | 415 | |
| His | Lys | Gly | His | Glu | Tyr | Met | Tyr | Tyr | Trp | Phe | Gly | Leu | Pro | Ser | Asp |
| | | | 420 | | | | | 425 | | | | | | 430 | |
| Gln | Thr | Pro | Thr | Gly | Pro | Val | Thr | Phe | Thr | Ile | Ile | Asn | Arg | Asp | Gly |
| | | 435 | | | | | | 440 | | | | | | 445 | |
| Ser | Ser | Thr | Ser | Thr | Gly | Gly | Val | Gly | Phe | Gly | Ser | Gly | Ala | Pro | Leu |
| | | 450 | | | | 455 | | | | | 460 | | | | |
| Pro | Gln | Phe | Trp | Arg | Ser | Ala | Gly | Ala | Ile | Asn | Ser | Ser | Val | Ala | Asn |
| | 465 | | | | | 470 | | | | | 475 | | | | 480 |
| Asp | Phe | Lys | His | Gly | Ser | Ala | Thr | Asn | Tyr | Ala | Phe | Tyr | Asp | Gly | Val |
| | | | | 485 | | | | | 490 | | | | | 495 | |
| Asn | Asn | Phe | Ser | Asp | Phe | Ala | Arg | Gly | Gly | Glu | Leu | Tyr | Phe | Asp | Arg |
| | | | 500 | | | | | | 505 | | | | | 510 | |
| Glu | Gly | Ala | Thr | Gln | Thr | Asn | Lys | Tyr | Tyr | Gly | Asp | Glu | Asn | Phe | Ala |
| | | 515 | | | | | | 520 | | | | | 525 | | |
| Leu | Leu | Asn | Ser | Glu | Lys | Pro | Asp | Gln | Ile | Arg | Gly | Leu | Asp | Thr | Ile |
| | 530 | | | | | 535 | | | | | 540 | | | | |
| Tyr | Ser | Phe | Lys | Gly | Ser | Gly | Asp | Val | Ser | Tyr | Arg | Ile | Ser | Phe | Lys |
| | 545 | | | | | 550 | | | | | 555 | | | | 560 |
| Thr | Gln | Gly | Ala | Pro | Thr | Ala | Arg | Leu | Tyr | Tyr | Ala | Ala | Gly | Ala | Arg |
| | | | | 565 | | | | | 570 | | | | | 575 | |
| Ser | Gly | Glu | Tyr | Lys | Gln | Ala | Thr | Asn | Tyr | Asn | Gln | Leu | Tyr | Val | Glu |
| | | | 580 | | | | | | 585 | | | | | 590 | |
| Pro | Tyr | Lys | Asn | Tyr | Arg | Asn | Arg | Val | Gln | Ser | Asn | Val | Gln | Val | Lys |
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 Leu Gln Arg Thr Thr Asp Val Pro Ile Leu Asp Ser Asp Gly Ser Gly
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 645 650 655
 Gly Thr Val Leu Gly Ile Tyr Pro Ser Tyr Leu Pro Tyr Asn Gln Glu
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 Arg Trp Gln Gly Ala Asn Ala Met Asn Ala Tyr Gln Ile Glu Glu Leu
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 Phe Ser Gln Glu Asn Leu Gln Asn Ala Ala Arg Ser Gly Arg Pro Ile
 690 695 700
 Gln Phe Leu Val Gly Phe Asp Val Glu Asp Ser His His Asn Pro Glu
 705 710 715 720
 Thr Leu Leu Pro Val Asn Leu Tyr Val Lys Pro Glu Leu Lys His Thr
 725 730 735
 Ile Glu Leu Tyr His Asp Asn Glu Lys Gln Asn Arg Lys Glu Phe Ser
 740 745 750
 Val Ser Lys Arg Ala Gly His Gly Val Phe Gln Ile Met Ser Gly Thr
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 Leu His Asn Thr Val Gly Ser Gly Ile Leu Pro Tyr Gln Gln Glu Ile
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 Thr Asn Asn Ala Thr Glu Lys Asn Leu Ala Leu Val Gly His Ile Asp
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 Pro Gly Asn Tyr Phe Ile Thr Val Lys Phe Gly Asp Lys Val Glu Gln
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 Phe Glu Ile Arg Ser Lys Pro Thr Pro Pro Arg Ile Ile Thr Thr Ala
 850 855 860
 Asn Glu Leu Arg Gly Asn Ser Asn His Lys Pro Glu Ile Arg Val Thr
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 Asp Ile Pro Asn Asp Thr Thr Ala Lys Ile Lys Leu Val Met Gly Gly
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 Thr Asp Gly Asp His Asp Pro Glu Ile Asn Pro Tyr Thr Val Pro Glu
 900 905 910
 Asn Tyr Thr Val Val Ala Glu Ala Tyr His Asp Asn Asp Pro Ser Lys
 915 920 925
 Asn Gly Val Leu Thr Phe Arg Ser Ser Asp Tyr Leu Lys Asp Leu Pro
 930 935 940
 Leu Ser Gly Glu Leu Lys Ala Ile Val Tyr Tyr Asn Gln Tyr Val Gln
 945 950 955 960
 Ser Asn Phe Ser Asn Ser Val Pro Phe Ser Ser Asp Thr Thr Pro Pro
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 Thr Ile Asn Glu Pro Ala Gly Leu Val His Lys Tyr Tyr Arg Gly Asp
 980 985 990
 His Val Glu Ile Thr Leu Pro Val Thr Asp Asn Thr Gly Gly Ser Gly
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 Leu Arg Asp Val Asn Val Asn Leu Pro Gln Gly Trp Thr Lys Thr
 1010 1015 1020

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| | | | | | | | | | |
|---------|---------|---------|---------|---------|---------|---------|---------|--|--|
| Phe Thr | Ile Asn | Pro Asn | Asn Asn | Asn Thr | Glu Gly | Thr Leu | Lys Leu | | |
| 1025 | | | 1030 | | | 1035 | | | |
| Ile Gly | Asn Ile | Pro Ser | Asn Asn | Glu Ala | Tyr Asn | Thr Thr | Tyr His | | |
| 1040 | | | 1045 | | | 1050 | | | |
| Phe Asn | Ile Thr | Ala Thr | Asp Asn | Asn Ser | Gly Asn | Thr Thr | Asn Pro | | |
| 1055 | | | 1060 | | | 1065 | | | |
| Ala Lys | Thr Phe | Ile Leu | Asn Asn | Val Gly | Lys Leu | Ala Asp | Asp Leu | | |
| 1070 | | | 1075 | | | 1080 | | | |
| Asn Pro | Val Gly | Leu Ser | Arg Asn | Asp Gln | Leu Gln | Leu Val | Thr Asp | | |
| 1085 | | | 1090 | | | 1095 | | | |
| Pro Ser | Ser Leu | Ser Asn | Ser Asn | Glu Arg | Glu Glu | Val Lys | Arg Lys | | |
| 1100 | | | 1105 | | | 1110 | | | |
| Ile Ser | Glu Ala | Asn Ala | Asn Asn | Ile Arg | Ser Tyr | Leu Leu | Gln Asn | | |
| 1115 | | | 1120 | | | 1125 | | | |
| Asn Pro | Ile Leu | Ala Gly | Val Asn | Asn Gly | Asp Val | Thr Phe | Tyr Tyr | | |
| 1130 | | | 1135 | | | 1140 | | | |
| Arg Asp | Gly Ser | Val Asp | Val Asn | Ile Asp | Ala Glu | Asn Val | Ile Thr | | |
| 1145 | | | 1150 | | | 1155 | | | |
| Tyr Glu | Pro Glu | Arg Lys | Ser Asn | Ile Phe | Ser Glu | Asn Gly | Asn Thr | | |
| 1160 | | | 1165 | | | 1170 | | | |
| Asn Lys | Lys Glu | Ala Val | Ile Asn | Thr Ile | Ala Arg | Gly Gln | Asn Tyr | | |
| 1175 | | | 1180 | | | 1185 | | | |
| Thr Ile | Gly Pro | Asn Leu | Arg Asn | Lys Tyr | Phe Ser | Leu Ser | Asn Gly | | |
| 1190 | | | 1195 | | | 1200 | | | |
| Ser Asp | Leu Pro | Asn Arg | Asp Asn | Phe Thr | Ser Ile | Ser Ala | Ile Gly | | |
| 1205 | | | 1210 | | | 1215 | | | |
| Ser Leu | Pro Ser | Ser Ser | Glu Asn | Ile Ser | Arg Leu | Asn Val | Gly Asn | | |
| 1220 | | | 1225 | | | 1230 | | | |
| Tyr Asn | Tyr Arg | Val Asn | Ala Asn | Lys Asn | Ala Tyr | His Lys | Thr Gln | | |
| 1235 | | | 1240 | | | 1245 | | | |
| Gln Glu | Leu Asn | Leu Lys | Leu Asn | Lys Ile | Val Glu | Val Asn | Ala Pro | | |
| 1250 | | | 1255 | | | 1260 | | | |
| Thr Gly | Asn Asn | Arg Val | Tyr Asn | Arg Val | Ser Thr | Tyr Asn | Leu Thr | | |
| 1265 | | | 1270 | | | 1275 | | | |
| Asn Asp | Glu Ile | Asn Lys | Ile Asn | Lys Gln | Ala Phe | Lys Ala | Ala Asn | | |
| 1280 | | | 1285 | | | 1290 | | | |
| Ser Gly | Leu Asn | Leu Asn | Asp Asn | Asn Asp | Ile Thr | Val Ser | Asn Asn | | |
| 1295 | | | 1300 | | | 1305 | | | |
| Phe Asp | His Arg | Asn Val | Ser Asn | Ser Val | Thr Val | Thr Ile | Arg Lys | | |
| 1310 | | | 1315 | | | 1320 | | | |
| Gly Asp | Leu Ile | Lys Glu | Phe Asn | Ser Ser | Asn Leu | Asn Asn | Met Asn | | |
| 1325 | | | 1330 | | | 1335 | | | |
| Phe Leu | Arg Trp | Val Asn | Ile Asn | Arg Asp | Asp Tyr | Thr Ile | Ser Trp | | |
| 1340 | | | 1345 | | | 1350 | | | |
| Thr Ser | Ser Lys | Ile Gln | Gly Asn | Arg Asn | Thr Asp | Gly Gly | Leu Glu | | |
| 1355 | | | 1360 | | | 1365 | | | |
| Trp Ser | Pro Asp | His Lys | Ser Asn | Leu Ile | Tyr Lys | Tyr Asp | Ala Thr | | |
| 1370 | | | 1375 | | | 1380 | | | |
| Leu Gly | Arg Gln | Ile Asn | Thr Asn | Asn Asp | Val Leu | Thr Leu | Leu Gln | | |
| 1385 | | | 1390 | | | 1395 | | | |
| Ala Thr | Ala Lys | Asn Ser | Asn Asn | Leu Arg | Ser Asn | Ile Asn | Ser Asn | | |
| 1400 | | | 1405 | | | 1410 | | | |
| Glu Lys | Gln Leu | Ala Glu | Arg Asn | Gly Ser | Asn Gly | Tyr Ser | Lys Ser | | |

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| | | | | | | | | | | | | | | | | |
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| Ile | Ile | Arg | Asp | Asp | Gly | Glu | Lys | Ser | Tyr | Leu | Leu | Asn | Ser | Asn | | |
| 1430 | | | | | | 1435 | | | | | 1440 | | | | | |
| Pro | Ile | Gln | Val | Leu | Asp | Leu | Val | Glu | Pro | Asp | Asn | Gly | Tyr | Gly | | |
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| Gly | Arg | Gln | Val | Ser | His | Ser | Asn | Val | Ile | Tyr | Asn | Glu | Lys | Asn | | |
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| Ala | Phe | Asn | Ile | Asp | Lys | Val | Val | Lys | Ala | Asn | Ala | Ala | Asn | Asn | | |
| 1490 | | | | | | 1495 | | | | | 1500 | | | | | |
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| Asn | Thr | Asn | Asn | Val | Ile | Asn | Val | Tyr | Phe | Val | Pro | Ser | Asp | Lys | | |
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| Val | Asn | Pro | Ser | Ile | Thr | Val | Gly | Asn | Tyr | Asp | His | His | Thr | Val | | |
| 1550 | | | | | | 1555 | | | | | 1560 | | | | | |
| Tyr | Ser | Gly | Glu | Thr | Phe | Lys | Asn | Thr | Ile | Asn | Val | Asn | Asp | Asn | | |
| 1565 | | | | | | 1570 | | | | | 1575 | | | | | |
| Tyr | Gly | Leu | Asn | Thr | Val | Ala | Ser | Thr | Ser | Asp | Ser | Ala | Ile | Thr | | |
| 1580 | | | | | | 1585 | | | | | 1590 | | | | | |
| Met | Thr | Arg | Asn | Asn | Asn | Glu | Leu | Val | Gly | Gln | Ala | Pro | Asn | Val | | |
| 1595 | | | | | | 1600 | | | | | 1605 | | | | | |
| Thr | Asn | Ser | Thr | Asn | Lys | Ile | Val | Lys | Val | Lys | Ala | Thr | Asp | Lys | | |
| 1610 | | | | | | 1615 | | | | | 1620 | | | | | |
| Ser | Gly | Asn | Glu | Ser | Ile | Val | Ser | Phe | Thr | Val | Asn | Ile | Lys | Pro | | |
| 1625 | | | | | | 1630 | | | | | 1635 | | | | | |
| Leu | Asn | Glu | Lys | Tyr | Arg | Ile | Thr | Thr | Ser | Ser | Ser | Asn | Gln | Thr | | |
| 1640 | | | | | | 1645 | | | | | 1650 | | | | | |
| Pro | Val | Arg | Ile | Ser | Asn | Ile | Gln | Asn | Asn | Ala | Asn | Leu | Ser | Ile | | |
| 1655 | | | | | | 1660 | | | | | 1665 | | | | | |
| Glu | Asp | Gln | Asn | Arg | Val | Lys | Ser | Ser | Leu | Ser | Met | Thr | Lys | Ile | | |
| 1670 | | | | | | 1675 | | | | | 1680 | | | | | |
| Leu | Gly | Thr | Arg | Asn | Tyr | Val | Asn | Glu | Ser | Asn | Asn | Asp | Val | Arg | | |
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| 1730 | | | | | | 1735 | | | | | 1740 | | | | | |
| Arg | Thr | Thr | Val | Arg | Gly | Gln | Gln | Phe | Pro | Thr | Gly | Lys | Gly | Thr | | |
| 1745 | | | | | | 1750 | | | | | 1755 | | | | | |
| Ser | Pro | Asn | Asp | Phe | Phe | Ser | Leu | Arg | Thr | Gly | Gly | Pro | Val | Asp | | |
| 1760 | | | | | | 1765 | | | | | 1770 | | | | | |
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| 1775 | | | | | | 1780 | | | | | 1785 | | | | | |
| Asn | Gln | Ile | Gly | Arg | Asp | Leu | Thr | Leu | His | Ala | Glu | Ile | Phe | Phe | | |
| 1790 | | | | | | 1795 | | | | | 1800 | | | | | |
| Asp | Gly | Glu | Thr | Thr | Pro | Ile | Arg | Lys | Asp | Thr | Thr | Tyr | Lys | Leu | | |
| 1805 | | | | | | 1810 | | | | | 1815 | | | | | |

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| | | | | | | | | | | | | | | |
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| 1820 | | | | | | 1825 | | | | | | 1830 | | |
| Arg | Phe | Asn | Ser | Ser | Gly | Asp | Ala | Tyr | Pro | Gly | Asn | Phe | Val | Gln |
| 1835 | | | | | | 1840 | | | | | | 1845 | | |
| Ala | Val | Asn | Gln | Tyr | Trp | Pro | Glu | His | Met | Asp | Phe | Arg | Trp | Ala |
| 1850 | | | | | | 1855 | | | | | | 1860 | | |
| Gln | Gly | Ser | Gly | Thr | Pro | Ser | Ser | Arg | Asn | Ala | Gly | Ser | Phe | Thr |
| 1865 | | | | | | 1870 | | | | | | 1875 | | |
| Lys | Thr | Val | Thr | Val | Val | Tyr | Gln | Asn | Gly | Gln | Thr | Glu | Asn | Val |
| 1880 | | | | | | 1885 | | | | | | 1890 | | |
| Asn | Val | Leu | Phe | Lys | Val | Lys | Pro | Asn | Lys | Pro | Val | Ile | Asp | Ser |
| 1895 | | | | | | 1900 | | | | | | 1905 | | |
| Asn | Ser | Val | Ile | Ser | Lys | Gly | Gln | Leu | Asn | Gly | Gln | Gln | Ile | Leu |
| 1910 | | | | | | 1915 | | | | | | 1920 | | |
| Val | Arg | Asn | Val | Pro | Gln | Asn | Ala | Gln | Val | Thr | Leu | Tyr | Gln | Ser |
| 1925 | | | | | | 1930 | | | | | | 1935 | | |
| Asn | Gly | Thr | Val | Ile | Pro | Asn | Thr | Asn | Thr | Thr | Ile | Asp | Ser | Asn |
| 1940 | | | | | | 1945 | | | | | | 1950 | | |
| Gly | Ile | Ala | Thr | Val | Thr | Ile | Gln | Gly | Thr | Leu | Pro | Thr | Gly | Asn |
| 1955 | | | | | | 1960 | | | | | | 1965 | | |
| Ile | Thr | Ala | Lys | Thr | Ser | Met | Thr | Asn | Asn | Val | Thr | Tyr | Thr | Lys |
| 1970 | | | | | | 1975 | | | | | | 1980 | | |
| Gln | Asn | Ser | Ser | Gly | Ile | Ala | Ser | Asn | Thr | Thr | Glu | Asp | Ile | Ser |
| 1985 | | | | | | 1990 | | | | | | 1995 | | |
| Val | Phe | Ser | Glu | Asn | Ser | Asp | Gln | Val | Asn | Val | Thr | Ala | Gly | Met |
| 2000 | | | | | | 2005 | | | | | | 2010 | | |
| Gln | Ala | Lys | Asn | Asp | Gly | Ile | Lys | Ile | Ile | Lys | Gly | Thr | Asn | Tyr |
| 2015 | | | | | | 2020 | | | | | | 2025 | | |
| Asn | Phe | Asn | Asp | Phe | Asn | Ser | Phe | Ile | Ser | Asn | Ile | Pro | Ala | His |
| 2030 | | | | | | 2035 | | | | | | 2040 | | |
| Ser | Thr | Leu | Thr | Trp | Asn | Glu | Glu | Pro | Asn | Ser | Trp | Lys | Asn | Asn |
| 2045 | | | | | | 2050 | | | | | | 2055 | | |
| Ile | Gly | Thr | Thr | Thr | Lys | Thr | Val | Thr | Val | Thr | Leu | Pro | Asn | His |
| 2060 | | | | | | 2065 | | | | | | 2070 | | |
| Gln | Gly | Thr | Arg | Thr | Val | Asp | Ile | Pro | Ile | Thr | Ile | Tyr | Pro | Thr |
| 2075 | | | | | | 2080 | | | | | | 2085 | | |
| Val | Thr | Ala | Lys | Asn | Pro | Val | Arg | Asp | Gln | Lys | Gly | Arg | Asn | Leu |
| 2090 | | | | | | 2095 | | | | | | 2100 | | |
| Thr | Asn | Gly | Thr | Asp | Val | Tyr | Asn | Tyr | Ile | Ile | Phe | Glu | Asn | Asn |
| 2105 | | | | | | 2110 | | | | | | 2115 | | |
| Asn | Arg | Leu | Gly | Gly | Thr | Ala | Ser | Trp | Lys | Asp | Asn | Arg | Gln | Pro |
| 2120 | | | | | | 2125 | | | | | | 2130 | | |
| Asp | Lys | Asn | Ile | Ala | Gly | Val | Gln | Asn | Leu | Ile | Ala | Leu | Val | Asn |
| 2135 | | | | | | 2140 | | | | | | 2145 | | |
| Tyr | Pro | Gly | Ile | Ser | Thr | Pro | Leu | Glu | Val | Pro | Val | Lys | Val | Trp |
| 2150 | | | | | | 2155 | | | | | | 2160 | | |
| Val | Tyr | Asn | Phe | Asp | Phe | Thr | Gln | Pro | Ile | Tyr | Lys | Ile | Gln | Val |
| 2165 | | | | | | 2170 | | | | | | 2175 | | |
| Gly | Asp | Thr | Phe | Pro | Lys | Gly | Thr | Trp | Ala | Gly | Tyr | Tyr | Lys | His |
| 2180 | | | | | | 2185 | | | | | | 2190 | | |
| Leu | Glu | Asn | Gly | Glu | Gly | Leu | Pro | Ile | Asp | Gly | Trp | Lys | Phe | Tyr |
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| | | | | | | | | | | | | | | |
|-----|------|-----|-----|-----|-----|------|-----|-----|-----|-----|------|-----|-----|-----|
| Trp | Asn | Gln | Gln | Ser | Thr | Gly | Thr | Thr | Ser | Asp | Gln | Trp | Gln | Ser |
| | 2210 | | | | | 2215 | | | | | 2220 | | | |
| Leu | Ala | Tyr | Thr | Arg | Thr | Pro | Phe | Val | Lys | Thr | Gly | Thr | Tyr | Asp |
| | 2225 | | | | | 2230 | | | | | 2235 | | | |
| Val | Val | Asn | Pro | Ser | Asn | Trp | Gly | Val | Trp | Gln | Thr | Ser | Gln | Ser |
| | 2240 | | | | | 2245 | | | | | 2250 | | | |
| Ala | Lys | Phe | Ile | Val | Thr | Asn | Ala | Lys | Pro | Asn | Gln | Pro | Thr | Ile |
| | 2255 | | | | | 2260 | | | | | 2265 | | | |
| Thr | Gln | Ser | Lys | Thr | Gly | Asp | Val | Thr | Val | Thr | Pro | Gly | Ala | Val |
| | 2270 | | | | | 2275 | | | | | 2280 | | | |
| Arg | Asn | Ile | Leu | Ile | Ser | Gly | Thr | Asn | Asp | Tyr | Ile | Gln | Ala | Ser |
| | 2285 | | | | | 2290 | | | | | 2295 | | | |
| Ala | Asp | Lys | Ile | Val | Ile | Asn | Lys | Asn | Gly | Asn | Lys | Leu | Thr | Thr |
| | 2300 | | | | | 2305 | | | | | 2310 | | | |
| Phe | Val | Lys | Asn | Asn | Asp | Gly | Arg | Trp | Thr | Val | Glu | Thr | Gly | Ser |
| | 2315 | | | | | 2320 | | | | | 2325 | | | |
| Pro | Asp | Ile | Asn | Gly | Ile | Gly | Pro | Thr | Asn | Asn | Gly | Thr | Ala | Ile |
| | 2330 | | | | | 2335 | | | | | 2340 | | | |
| Ser | Leu | Ser | Arg | Leu | Ala | Val | Arg | Pro | Gly | Asp | Ser | Ile | Glu | Ala |
| | 2345 | | | | | 2350 | | | | | 2355 | | | |
| Ile | Ala | Thr | Glu | Gly | Ser | Gly | Glu | Thr | Ile | Ser | Thr | Ser | Ala | Thr |
| | 2360 | | | | | 2365 | | | | | 2370 | | | |
| Ser | Glu | Ile | Tyr | Ile | Val | Lys | Ala | Pro | Gln | Pro | Glu | Gln | Val | Ala |
| | 2375 | | | | | 2380 | | | | | 2385 | | | |
| Thr | His | Thr | Tyr | Asp | Asn | Gly | Thr | Phe | Asp | Ile | Leu | Pro | Asp | Asn |
| | 2390 | | | | | 2395 | | | | | 2400 | | | |
| Ser | Arg | Asn | Ser | Leu | Asn | Pro | Thr | Glu | Arg | Val | Glu | Ile | Asn | Tyr |
| | 2405 | | | | | 2410 | | | | | 2415 | | | |
| Thr | Glu | Lys | Leu | Asn | Gly | Asn | Glu | Thr | Gln | Lys | Ser | Phe | Thr | Ile |
| | 2420 | | | | | 2425 | | | | | 2430 | | | |
| Thr | Lys | Asn | Asn | Asn | Gly | Lys | Trp | Thr | Ile | Asn | Asn | Lys | Pro | Asn |
| | 2435 | | | | | 2440 | | | | | 2445 | | | |
| Tyr | Val | Glu | Phe | Asn | Gln | Asp | Asn | Gly | Lys | Val | Val | Phe | Ser | Ala |
| | 2450 | | | | | 2455 | | | | | 2460 | | | |
| Asn | Thr | Ile | Lys | Pro | Asn | Ser | Gln | Ile | Thr | Ile | Thr | Pro | Lys | Ala |
| | 2465 | | | | | 2470 | | | | | 2475 | | | |
| Gly | Gln | Gly | Asn | Thr | Glu | Asn | Thr | Asn | Pro | Thr | Val | Ile | Gln | Ala |
| | 2480 | | | | | 2485 | | | | | 2490 | | | |
| Pro | Ala | Gln | His | Thr | Leu | Thr | Ile | Asn | Glu | Ile | Val | Lys | Glu | Gln |
| | 2495 | | | | | 2500 | | | | | 2505 | | | |
| Gly | Gln | Asn | Val | Thr | Asn | Asp | Asp | Ile | Asn | Asn | Ala | Val | Gln | Val |
| | 2510 | | | | | 2515 | | | | | 2520 | | | |
| Pro | Asn | Lys | Asn | Arg | Val | Ala | Ile | Lys | Gln | Gly | Asn | Ala | Leu | Pro |
| | 2525 | | | | | 2530 | | | | | 2535 | | | |
| Thr | Asn | Leu | Ala | Gly | Gly | Ser | Thr | Ser | His | Ile | Pro | Val | Val | Ile |
| | 2540 | | | | | 2545 | | | | | 2550 | | | |
| Tyr | Tyr | Ser | Asp | Gly | Ser | Ser | Glu | Glu | Ala | Thr | Glu | Thr | Val | Arg |
| | 2555 | | | | | 2560 | | | | | 2565 | | | |
| Thr | Lys | Val | Asn | Lys | Thr | Glu | Leu | Ile | Asn | Ala | Arg | Arg | Arg | Leu |
| | 2570 | | | | | 2575 | | | | | 2580 | | | |
| Asp | Glu | Glu | Ile | Ser | Lys | Glu | Asn | Lys | Thr | Pro | Ser | Ser | Ile | Arg |
| | 2585 | | | | | 2590 | | | | | 2595 | | | |
| Asn | Phe | Asp | Gln | Ala | Met | Asn | Arg | Ala | Gln | Ser | Gln | Ile | Asn | Thr |

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|---------------------------------------------------------------------|------|------|
| 2600 | 2605 | 2610 |
| Ala Lys Ser Asp Ala Asp Gln Val Ile Gly Thr Glu Phe Ala Thr 2615 | 2620 | 2625 |
| Pro Gln Gln Val Asn Ser Ala Leu Ser Lys Val Gln Ala Ala Gln 2630 | 2635 | 2640 |
| Asn Lys Ile Asn Glu Ala Lys Ala Leu Leu Gln Asn Lys Ala Asp 2645 | 2650 | 2655 |
| Asn Ser Gln Leu Val Arg Ala Lys Glu Gln Leu Gln Gln Ser Ile 2660 | 2665 | 2670 |
| Gln Pro Ala Ala Ser Thr Asp Gly Met Thr Gln Asp Ser Thr Arg 2675 | 2680 | 2685 |
| Asn Tyr Lys Asn Lys Arg Gln Ala Ala Glu Gln Ala Ile Gln His 2690 | 2695 | 2700 |
| Ala Asn Ser Val Ile Asn Asn Gly Asp Ala Thr Ser Gln Gln Ile 2705 | 2710 | 2715 |
| Asn Asp Ala Lys Asn Thr Val Glu Gln Ala Gln Arg Asp Tyr Val 2720 | 2725 | 2730 |
| Glu Ala Lys Ser Asn Leu Arg Ala Asp Lys Ser Gln Leu Gln Ser 2735 | 2740 | 2745 |
| Ala Tyr Asp Thr Leu Asn Arg Asp Val Leu Thr Asn Asp Lys Lys 2750 | 2755 | 2760 |
| Pro Ala Ser Val Arg Arg Tyr Asn Glu Ala Ile Ser Asn Ile Arg 2765 | 2770 | 2775 |
| Lys Glu Leu Asp Thr Ala Lys Ala Asp Ala Ser Ser Thr Leu Arg 2780 | 2785 | 2790 |
| Asn Thr Asn Pro Ser Val Glu Gln Val Arg Asp Ala Leu Asn Lys 2795 | 2800 | 2805 |
| Ile Asn Thr Val Gln Pro Lys Val Asn Gln Ala Ile Ala Leu Leu 2810 | 2815 | 2820 |
| Gln Pro Lys Glu Asn Asn Ser Glu Leu Val Gln Ala Lys Lys Arg 2825 | 2830 | 2835 |
| Leu Gln Asp Ala Val Asn Asp Ile Pro Gln Thr Gln Gly Met Thr 2840 | 2845 | 2850 |
| Gln Gln Thr Ile Asn Asn Tyr Asn Asp Lys Gln Arg Glu Ala Glu 2855 | 2860 | 2865 |
| Arg Ala Leu Thr Ser Ala Gln Arg Val Ile Asp Asn Gly Asp Ala 2870 | 2875 | 2880 |
| Thr Thr Gln Glu Ile Thr Ser Glu Lys Ser Lys Val Glu Gln Ala 2885 | 2890 | 2895 |
| Met Gln Ala Leu Thr Asn Ala Lys Ser Asn Leu Arg Ala Asp Lys 2900 | 2905 | 2910 |
| Asn Glu Leu Gln Thr Ala Tyr Asn Lys Leu Ile Glu Asn Val Ser 2915 | 2920 | 2925 |
| Thr Asn Gly Lys Lys Pro Ala Ser Ile Arg Gln Tyr Glu Thr Ala 2930 | 2935 | 2940 |
| Lys Ala Arg Ile Gln Asn Gln Ile Asn Asp Ala Lys Asn Glu Ala 2945 | 2950 | 2955 |
| Glu Arg Ile Leu Gly Asn Asp Asn Pro Gln Val Ser Gln Val Thr 2960 | 2965 | 2970 |
| Gln Ala Leu Asn Lys Ile Lys Ala Ile Gln Pro Lys Leu Thr Glu 2975 | 2980 | 2985 |
| Ala Ile Asn Met Leu Gln Asn Lys Glu Asn Asn Thr Glu Leu Val 2990 | 2995 | 3000 |

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|---------|---------------------|---------------------|-------------|
| Asn Ala | Lys Asn Arg Leu Glu | Asn Ala Val Asn Asp | Thr Asp Pro |
| 3005 | 3010 | 3015 | |
| Thr His | Gly Met Thr Gln Glu | Thr Ile Asn Asn Tyr | Asn Ala Lys |
| 3020 | 3025 | 3030 | |
| Lys Arg | Glu Ala Gln Asn Glu | Ile Gln Lys Ala Asn | Met Ile Ile |
| 3035 | 3040 | 3045 | |
| Asn Asn | Gly Asp Ala Thr Ala | Gln Asp Ile Ser Ser | Glu Lys Ser |
| 3050 | 3055 | 3060 | |
| Lys Val | Glu Gln Val Leu Gln | Ala Leu Gln Asn Ala | Lys Asn Asp |
| 3065 | 3070 | 3075 | |
| Leu Arg | Ala Asp Lys Arg Glu | Leu Gln Thr Ala Tyr | Asn Lys Leu |
| 3080 | 3085 | 3090 | |
| Ile Gln | Asn Val Asn Thr Asn | Gly Lys Lys Pro Ser | Ser Ile Gln |
| 3095 | 3100 | 3105 | |
| Asn Tyr | Lys Ser Ala Arg Arg | Asn Ile Glu Asn Gln | Tyr Asn Thr |
| 3110 | 3115 | 3120 | |
| Ala Lys | Asn Glu Ala His Asn | Val Leu Glu Asn Thr | Asn Pro Thr |
| 3125 | 3130 | 3135 | |
| Val Asn | Ala Val Glu Asp Ala | Leu Arg Lys Ile Asn | Ala Ile Gln |
| 3140 | 3145 | 3150 | |
| Pro Glu | Val Thr Lys Ala Ile | Asn Ile Leu Gln Asp | Lys Glu Asp |
| 3155 | 3160 | 3165 | |
| Asn Ser | Glu Leu Val Arg Ala | Lys Glu Lys Leu Asp | Gln Ala Ile |
| 3170 | 3175 | 3180 | |
| Asn Ser | Gln Pro Ser Leu Asn | Gly Met Thr Gln Glu | Ser Ile Asn |
| 3185 | 3190 | 3195 | |
| Asn Tyr | Thr Thr Lys Arg Arg | Glu Ala Gln Asn Ile | Ala Ser Ser |
| 3200 | 3205 | 3210 | |
| Ala Asp | Thr Ile Ile Asn Asn | Gly Asp Ala Ser Ile | Glu Gln Ile |
| 3215 | 3220 | 3225 | |
| Thr Glu | Asn Lys Ile Arg Val | Glu Glu Ala Thr Asn | Ala Leu Asn |
| 3230 | 3235 | 3240 | |
| Glu Ala | Lys Gln His Leu Thr | Ala Asp Thr Thr Ser | Leu Lys Thr |
| 3245 | 3250 | 3255 | |
| Glu Val | Arg Lys Leu Ser Arg | Arg Gly Asp Thr Asn | Asn Lys Lys |
| 3260 | 3265 | 3270 | |
| Pro Ser | Ser Val Ser Ala Tyr | Asn Asn Thr Ile His | Ser Leu Gln |
| 3275 | 3280 | 3285 | |
| Ser Glu | Ile Thr Gln Thr Glu | Asn Arg Ala Asn Thr | Ile Ile Asn |
| 3290 | 3295 | 3300 | |
| Lys Pro | Ile Arg Ser Val Glu | Glu Val Asn Asn Ala | Leu His Glu |
| 3305 | 3310 | 3315 | |
| Val Asn | Gln Leu Asn Gln Arg | Leu Thr Asp Thr Ile | Asn Leu Leu |
| 3320 | 3325 | 3330 | |
| Gln Pro | Leu Ala Asn Lys Glu | Ser Leu Lys Glu Ala | Arg Asn Arg |
| 3335 | 3340 | 3345 | |
| Leu Glu | Ser Lys Ile Asn Glu | Thr Val Gln Thr Asp | Gly Met Thr |
| 3350 | 3355 | 3360 | |
| Gln Gln | Ser Val Glu Asn Tyr | Lys Gln Ala Lys Ile | Lys Ala Gln |
| 3365 | 3370 | 3375 | |
| Asn Glu | Ser Ser Ile Ala Gln | Thr Leu Ile Asn Asn | Gly Asp Ala |
| 3380 | 3385 | 3390 | |

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|------|-----|-----|-----|-----|-----|------|-----|-----|-----|-----|------|-----|-----|-----|
| Ser | Asp | Gln | Glu | Val | Ser | Thr | Glu | Ile | Glu | Lys | Leu | Asn | Gln | Lys |
| 3395 | | | | | | 3400 | | | | | 3405 | | | |
| Leu | Ser | Glu | Leu | Thr | Asn | Ser | Ile | Asn | His | Leu | Thr | Val | Asn | Lys |
| 3410 | | | | | | 3415 | | | | | 3420 | | | |
| Glu | Pro | Leu | Glu | Thr | Ala | Lys | Asn | Gln | Leu | Gln | Ala | Asn | Ile | Asp |
| 3425 | | | | | | 3430 | | | | | 3435 | | | |
| Gln | Lys | Pro | Ser | Thr | Asp | Gly | Met | Thr | Gln | Gln | Ser | Val | Gln | Ser |
| 3440 | | | | | | 3445 | | | | | 3450 | | | |
| Tyr | Glu | Arg | Lys | Leu | Gln | Glu | Ala | Lys | Asp | Lys | Ile | Asn | Ser | Ile |
| 3455 | | | | | | 3460 | | | | | 3465 | | | |
| Asn | Asn | Val | Leu | Ala | Asn | Asn | Pro | Asp | Val | Asn | Ala | Ile | Arg | Thr |
| 3470 | | | | | | 3475 | | | | | 3480 | | | |
| Asn | Lys | Val | Glu | Thr | Glu | Gln | Ile | Asn | Asn | Glu | Leu | Thr | Gln | Ala |
| 3485 | | | | | | 3490 | | | | | 3495 | | | |
| Lys | Gln | Gly | Leu | Thr | Val | Asp | Lys | Gln | Pro | Leu | Ile | Asn | Ala | Lys |
| 3500 | | | | | | 3505 | | | | | 3510 | | | |
| Thr | Ala | Leu | Gln | Gln | Ser | Leu | Asp | Asn | Gln | Pro | Ser | Thr | Thr | Gly |
| 3515 | | | | | | 3520 | | | | | 3525 | | | |
| Met | Thr | Glu | Ala | Thr | Ile | Gln | Asn | Tyr | Asn | Ala | Lys | Arg | Gln | Lys |
| 3530 | | | | | | 3535 | | | | | 3540 | | | |
| Ala | Glu | Gln | Val | Ile | Gln | Asn | Ala | Asn | Lys | Ile | Ile | Glu | Asn | Ala |
| 3545 | | | | | | 3550 | | | | | 3555 | | | |
| Gln | Pro | Ser | Val | Gln | Gln | Val | Ser | Asp | Glu | Lys | Ser | Lys | Val | Glu |
| 3560 | | | | | | 3565 | | | | | 3570 | | | |
| Gln | Ala | Leu | Ser | Glu | Leu | Asn | Asn | Ala | Lys | Ser | Ala | Leu | Arg | Ala |
| 3575 | | | | | | 3580 | | | | | 3585 | | | |
| Asp | Lys | Gln | Glu | Leu | Gln | Gln | Ala | Tyr | Asn | Gln | Leu | Ile | Gln | Pro |
| 3590 | | | | | | 3595 | | | | | 3600 | | | |
| Thr | Asp | Leu | Asn | Asn | Lys | Lys | Pro | Ala | Ser | Ile | Thr | Ala | Tyr | Asn |
| 3605 | | | | | | 3610 | | | | | 3615 | | | |
| Gln | Arg | Tyr | Gln | Gln | Phe | Ser | Asn | Glu | Leu | Asn | Ser | Thr | Lys | Thr |
| 3620 | | | | | | 3625 | | | | | 3630 | | | |
| Asn | Thr | Asp | Arg | Ile | Leu | Lys | Glu | Gln | Asn | Pro | Ser | Val | Ala | Asp |
| 3635 | | | | | | 3640 | | | | | 3645 | | | |
| Val | Asn | Asn | Ala | Leu | Asn | Lys | Val | Arg | Glu | Val | Gln | Gln | Lys | Leu |
| 3650 | | | | | | 3655 | | | | | 3660 | | | |
| Asn | Glu | Ala | Arg | Ala | Leu | Leu | Gln | Asn | Lys | Glu | Asp | Asn | Ser | Ala |
| 3665 | | | | | | 3670 | | | | | 3675 | | | |
| Leu | Val | Arg | Ala | Lys | Glu | Gln | Leu | Gln | Gln | Ala | Val | Asp | Gln | Val |
| 3680 | | | | | | 3685 | | | | | 3690 | | | |
| Pro | Ser | Thr | Glu | Gly | Met | Thr | Gln | Gln | Thr | Lys | Asp | Asp | Tyr | Asn |
| 3695 | | | | | | 3700 | | | | | 3705 | | | |
| Ser | Lys | Gln | Gln | Ala | Ala | Gln | Gln | Glu | Ile | Ser | Lys | Ala | Gln | Gln |
| 3710 | | | | | | 3715 | | | | | 3720 | | | |
| Val | Ile | Asp | Asn | Gly | Asp | Ala | Thr | Thr | Gln | Gln | Ile | Ser | Asn | Ala |
| 3725 | | | | | | 3730 | | | | | 3735 | | | |
| Lys | Thr | Asn | Val | Glu | Arg | Ala | Leu | Glu | Ala | Leu | Asn | Asn | Ala | Lys |
| 3740 | | | | | | 3745 | | | | | 3750 | | | |
| Thr | Gly | Leu | Arg | Ala | Asp | Lys | Glu | Glu | Leu | Gln | Asn | Ala | Tyr | Asn |
| 3755 | | | | | | 3760 | | | | | 3765 | | | |
| Gln | Leu | Thr | Gln | Asn | Ile | Asp | Thr | Ser | Gly | Lys | Thr | Pro | Ala | Ser |
| 3770 | | | | | | 3775 | | | | | 3780 | | | |
| Ile | Arg | Lys | Tyr | Asn | Glu | Ala | Lys | Ser | Arg | Ile | Gln | Thr | Gln | Ile |

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|------|-----|-----|-----|-----|-----|------|-----|-----|-----|-----|-----|------|-----|-----|
| 3785 | | | | | | 3790 | | | | | | 3795 | | |
| Asp | Ser | Ala | Lys | Asn | Glu | Ala | Asn | Ser | Ile | Leu | Thr | Asn | Asp | Asn |
| 3800 | | | | | | 3805 | | | | | | 3810 | | |
| Pro | Gln | Val | Ser | Gln | Val | Thr | Ala | Ala | Leu | Asn | Lys | Ile | Lys | Ala |
| 3815 | | | | | | 3820 | | | | | | 3825 | | |
| Val | Gln | Pro | Glu | Leu | Asp | Lys | Ala | Ile | Ala | Met | Leu | Lys | Asn | Lys |
| 3830 | | | | | | 3835 | | | | | | 3840 | | |
| Glu | Asn | Asn | Asn | Ala | Leu | Val | Gln | Ala | Lys | Gln | Gln | Leu | Gln | Gln |
| 3845 | | | | | | 3850 | | | | | | 3855 | | |
| Ile | Val | Asn | Glu | Val | Asp | Pro | Thr | Gln | Gly | Met | Thr | Thr | Asp | Thr |
| 3860 | | | | | | 3865 | | | | | | 3870 | | |
| Ala | Asn | Asn | Tyr | Lys | Ser | Lys | Lys | Arg | Glu | Ala | Glu | Asp | Glu | Ile |
| 3875 | | | | | | 3880 | | | | | | 3885 | | |
| Gln | Lys | Ala | Gln | Gln | Ile | Ile | Asn | Asn | Gly | Asp | Ala | Thr | Glu | Gln |
| 3890 | | | | | | 3895 | | | | | | 3900 | | |
| Gln | Ile | Thr | Asn | Glu | Thr | Asn | Arg | Val | Asn | Gln | Ala | Ile | Asn | Ala |
| 3905 | | | | | | 3910 | | | | | | 3915 | | |
| Ile | Asn | Lys | Ala | Lys | Asn | Asp | Leu | Arg | Ala | Asp | Lys | Ser | Gln | Leu |
| 3920 | | | | | | 3925 | | | | | | 3930 | | |
| Glu | Asn | Ala | Tyr | Asn | Gln | Leu | Ile | Gln | Asn | Val | Asp | Thr | Asn | Gly |
| 3935 | | | | | | 3940 | | | | | | 3945 | | |
| Lys | Lys | Pro | Ala | Ser | Ile | Gln | Gln | Tyr | Gln | Ala | Ala | Arg | Gln | Ala |
| 3950 | | | | | | 3955 | | | | | | 3960 | | |
| Ile | Glu | Thr | Gln | Tyr | Asn | Asn | Ala | Lys | Ser | Glu | Ala | His | Gln | Ile |
| 3965 | | | | | | 3970 | | | | | | 3975 | | |
| Leu | Glu | Asn | Ser | Asn | Pro | Ser | Val | Asn | Glu | Val | Ala | Gln | Ala | Leu |
| 3980 | | | | | | 3985 | | | | | | 3990 | | |
| Gln | Lys | Val | Glu | Ala | Val | Gln | Leu | Lys | Val | Asn | Asp | Ala | Ile | His |
| 3995 | | | | | | 4000 | | | | | | 4005 | | |
| Ile | Leu | Gln | Asn | Lys | Glu | Asn | Asn | Ser | Ala | Leu | Val | Thr | Ala | Lys |
| 4010 | | | | | | 4015 | | | | | | 4020 | | |
| Asn | Gln | Leu | Gln | Gln | Ser | Val | Asn | Asp | Gln | Pro | Leu | Thr | Thr | Gly |
| 4025 | | | | | | 4030 | | | | | | 4035 | | |
| Met | Thr | Gln | Asp | Ser | Ile | Asn | Asn | Tyr | Glu | Ala | Lys | Arg | Asn | Glu |
| 4040 | | | | | | 4045 | | | | | | 4050 | | |
| Ala | Gln | Ser | Ala | Ile | Arg | Asn | Ala | Glu | Ala | Val | Ile | Asn | Asn | Gly |
| 4055 | | | | | | 4060 | | | | | | 4065 | | |
| Asp | Ala | Thr | Ala | Lys | Gln | Ile | Ser | Asp | Glu | Lys | Ser | Lys | Val | Glu |
| 4070 | | | | | | 4075 | | | | | | 4080 | | |
| Gln | Ala | Leu | Ala | His | Leu | Asn | Asp | Ala | Lys | Gln | Gln | Leu | Thr | Ala |
| 4085 | | | | | | 4090 | | | | | | 4095 | | |
| Asp | Thr | Thr | Glu | Leu | Gln | Thr | Ala | Val | Gln | Gln | Leu | Asn | Arg | Arg |
| 4100 | | | | | | 4105 | | | | | | 4110 | | |
| Gly | Asp | Thr | Asn | Asn | Lys | Lys | Pro | Arg | Ser | Ile | Asn | Ala | Tyr | Asn |
| 4115 | | | | | | 4120 | | | | | | 4125 | | |
| Lys | Ala | Ile | Gln | Ser | Leu | Glu | Thr | Gln | Ile | Thr | Ser | Ala | Lys | Asp |
| 4130 | | | | | | 4135 | | | | | | 4140 | | |
| Asn | Ala | Asn | Ala | Val | Ile | Gln | Lys | Pro | Ile | Arg | Thr | Val | Gln | Glu |
| 4145 | | | | | | 4150 | | | | | | 4155 | | |
| Val | Asn | Asn | Ala | Leu | Gln | Gln | Val | Asn | Gln | Leu | Asn | Gln | Gln | Leu |
| 4160 | | | | | | 4165 | | | | | | 4170 | | |
| Thr | Glu | Ala | Ile | Asn | Gln | Leu | Gln | Pro | Leu | Ser | Asn | Asn | Asp | Ala |
| 4175 | | | | | | 4180 | | | | | | 4185 | | |

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|------|-----|-----|-----|-----|-----|------|-----|-----|-----|-----|------|-----|-----|-----|
| Asn | Gln | Lys | Gln | Gln | Glu | Asn | Tyr | Leu | Asp | Ala | Ser | Asn | Asn | Lys |
| 4580 | | | | | | 4585 | | | | | 4590 | | | |
| Arg | Gln | Asp | Tyr | Asp | Asn | Ala | Val | Asn | Ala | Ala | Lys | Gly | Ile | Leu |
| 4595 | | | | | | 4600 | | | | | 4605 | | | |
| Asn | Gln | Thr | Gln | Ser | Pro | Thr | Met | Ser | Ala | Asp | Val | Ile | Asp | Gln |
| 4610 | | | | | | 4615 | | | | | 4620 | | | |
| Lys | Ala | Glu | Asp | Val | Lys | Arg | Thr | Lys | Thr | Ala | Leu | Asp | Gly | Asn |
| 4625 | | | | | | 4630 | | | | | 4635 | | | |
| Gln | Arg | Leu | Glu | Val | Ala | Lys | Gln | Gln | Ala | Leu | Asn | His | Leu | Asn |
| 4640 | | | | | | 4645 | | | | | 4650 | | | |
| Thr | Leu | Asn | Asp | Leu | Asn | Asp | Ala | Gln | Arg | Gln | Thr | Leu | Thr | Asp |
| 4655 | | | | | | 4660 | | | | | 4665 | | | |
| Thr | Ile | Asn | His | Ser | Pro | Asn | Ile | Asn | Ser | Val | Asn | Gln | Ala | Lys |
| 4670 | | | | | | 4675 | | | | | 4680 | | | |
| Glu | Lys | Ala | Asn | Thr | Val | Asn | Thr | Ala | Met | Thr | Gln | Leu | Lys | Gln |
| 4685 | | | | | | 4690 | | | | | 4695 | | | |
| Thr | Ile | Ala | Asn | Tyr | Asp | Asp | Glu | Leu | His | Asp | Gly | Asn | Tyr | Ile |
| 4700 | | | | | | 4705 | | | | | 4710 | | | |
| Asn | Ala | Asp | Lys | Asp | Lys | Lys | Asp | Ala | Tyr | Asn | Asn | Ala | Val | Asn |
| 4715 | | | | | | 4720 | | | | | 4725 | | | |
| Asn | Ala | Lys | Gln | Leu | Ile | Asn | Gln | Ser | Asp | Ala | Asn | Gln | Ala | Gln |
| 4730 | | | | | | 4735 | | | | | 4740 | | | |
| Leu | Asp | Pro | Ala | Glu | Ile | Asn | Lys | Val | Thr | Gln | Arg | Val | Asn | Thr |
| 4745 | | | | | | 4750 | | | | | 4755 | | | |
| Thr | Lys | Asn | Asp | Leu | Asn | Gly | Asn | Asp | Lys | Leu | Ala | Glu | Ala | Lys |
| 4760 | | | | | | 4765 | | | | | 4770 | | | |
| Arg | Asp | Ala | Asn | Thr | Thr | Ile | Asp | Gly | Leu | Thr | Tyr | Leu | Asn | Glu |
| 4775 | | | | | | 4780 | | | | | 4785 | | | |
| Ala | Gln | Arg | Asn | Lys | Ala | Lys | Glu | Asn | Val | Gly | Lys | Ala | Ser | Thr |
| 4790 | | | | | | 4795 | | | | | 4800 | | | |
| Lys | Thr | Asn | Ile | Thr | Ser | Gln | Leu | Gln | Asp | Tyr | Asn | Gln | Leu | Asn |
| 4805 | | | | | | 4810 | | | | | 4815 | | | |
| Ile | Ala | Met | Gln | Ala | Leu | Arg | Asn | Ser | Val | Asn | Asp | Val | Asn | Asn |
| 4820 | | | | | | 4825 | | | | | 4830 | | | |
| Val | Lys | Ala | Asn | Ser | Asn | Tyr | Ile | Asn | Glu | Asp | Asn | Gly | Pro | Lys |
| 4835 | | | | | | 4840 | | | | | 4845 | | | |
| Glu | Ala | Tyr | Asn | Gln | Ala | Val | Thr | His | Ala | Gln | Thr | Leu | Ile | Asn |
| 4850 | | | | | | 4855 | | | | | 4860 | | | |
| Ala | Gln | Ser | Asn | Pro | Glu | Met | Ser | Arg | Asp | Val | Val | Asn | Gln | Lys |
| 4865 | | | | | | 4870 | | | | | 4875 | | | |
| Thr | Gln | Ala | Val | Asn | Thr | Ala | His | Gln | Asn | Leu | His | Gly | Gln | Gln |
| 4880 | | | | | | 4885 | | | | | 4890 | | | |
| Lys | Leu | Glu | Gln | Ala | Gln | Ser | Ser | Ala | Asn | Thr | Glu | Ile | Gly | Asn |
| 4895 | | | | | | 4900 | | | | | 4905 | | | |
| Leu | Pro | Asn | Leu | Thr | Asn | Thr | Gln | Lys | Ala | Lys | Glu | Lys | Glu | Leu |
| 4910 | | | | | | 4915 | | | | | 4920 | | | |
| Val | Asn | Ser | Lys | Gln | Thr | Arg | Thr | Glu | Val | Gln | Glu | Gln | Leu | Asn |
| 4925 | | | | | | 4930 | | | | | 4935 | | | |
| Gln | Ala | Lys | Ser | Leu | Asp | Ser | Ser | Met | Gly | Thr | Leu | Lys | Ser | Leu |
| 4940 | | | | | | 4945 | | | | | 4950 | | | |
| Val | Ala | Lys | Gln | Pro | Thr | Val | Gln | Lys | Thr | Ser | Val | Tyr | Ile | Asn |
| 4955 | | | | | | 4960 | | | | | 4965 | | | |
| Glu | Asp | Gln | Pro | Glu | Gln | Ser | Ala | Tyr | Asn | Asp | Ser | Ile | Thr | Met |

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| 4970 | | 4975 | | 4980 | | | | | | | | | | |
|------|-----|------|-----|------|-----|------|-----|-----|-----|-----|------|-----|-----|-----|
| Gly | Gln | Thr | Ile | Ile | Asn | Lys | Thr | Ala | Asp | Pro | Val | Leu | Asp | Lys |
| 4985 | | | | | | 4990 | | | | | 4995 | | | |
| Thr | Leu | Val | Asp | Asn | Ala | Ile | Ser | Asn | Ile | Ser | Thr | Lys | Glu | Asn |
| 5000 | | | | | | 5005 | | | | | 5010 | | | |
| Ala | Leu | His | Gly | Glu | Gln | Lys | Leu | Thr | Thr | Ala | Lys | Thr | Glu | Ala |
| 5015 | | | | | | 5020 | | | | | 5025 | | | |
| Ile | Asn | Ala | Leu | Asn | Thr | Leu | Ala | Asp | Leu | Asn | Thr | Pro | Gln | Lys |
| 5030 | | | | | | 5035 | | | | | 5040 | | | |
| Glu | Ala | Ile | Lys | Thr | Ala | Ile | Asn | Thr | Ala | His | Thr | Arg | Thr | Asp |
| 5045 | | | | | | 5050 | | | | | 5055 | | | |
| Val | Thr | Ala | Glu | Gln | Ser | Lys | Ala | Asn | Gln | Ile | Asn | Ser | Ala | Met |
| 5060 | | | | | | 5065 | | | | | 5070 | | | |
| His | Thr | Leu | Arg | Gln | Asn | Ile | Ser | Asp | Asn | Glu | Ser | Val | Thr | Asn |
| 5075 | | | | | | 5080 | | | | | 5085 | | | |
| Glu | Ser | Asn | Tyr | Ile | Asn | Ala | Glu | Pro | Glu | Lys | Gln | His | Ala | Phe |
| 5090 | | | | | | 5095 | | | | | 5100 | | | |
| Thr | Glu | Ala | Leu | Asn | Asn | Ala | Lys | Glu | Ile | Val | Asn | Glu | Gln | Gln |
| 5105 | | | | | | 5110 | | | | | 5115 | | | |
| Ala | Thr | Leu | Asp | Ala | Asn | Ser | Ile | Asn | Gln | Lys | Ala | Gln | Ala | Ile |
| 5120 | | | | | | 5125 | | | | | 5130 | | | |
| Leu | Thr | Thr | Lys | Asn | Ala | Leu | Asp | Gly | Glu | Glu | Gln | Leu | Arg | Arg |
| 5135 | | | | | | 5140 | | | | | 5145 | | | |
| Ala | Lys | Glu | Asn | Ala | Asp | Gln | Glu | Ile | Asn | Thr | Leu | Asn | Gln | Leu |
| 5150 | | | | | | 5155 | | | | | 5160 | | | |
| Thr | Asp | Ala | Gln | Arg | Asn | Ser | Glu | Lys | Gly | Leu | Val | Asn | Ser | Ser |
| 5165 | | | | | | 5170 | | | | | 5175 | | | |
| Gln | Thr | Arg | Thr | Glu | Val | Ala | Ser | Gln | Leu | Ala | Lys | Ala | Lys | Glu |
| 5180 | | | | | | 5185 | | | | | 5190 | | | |
| Leu | Asn | Lys | Val | Met | Glu | Gln | Leu | Asn | His | Leu | Ile | Asn | Gly | Lys |
| 5195 | | | | | | 5200 | | | | | 5205 | | | |
| Asn | Gln | Met | Ile | Asn | Ser | Ser | Lys | Phe | Ile | Asn | Glu | Asp | Ala | Asn |
| 5210 | | | | | | 5215 | | | | | 5220 | | | |
| Gln | Gln | Gln | Ala | Tyr | Ser | Asn | Ala | Ile | Ala | Ser | Ala | Glu | Ala | Leu |
| 5225 | | | | | | 5230 | | | | | 5235 | | | |
| Lys | Asn | Lys | Ser | Gln | Asn | Pro | Glu | Leu | Asp | Lys | Val | Thr | Ile | Glu |
| 5240 | | | | | | 5245 | | | | | 5250 | | | |
| Gln | Ala | Ile | Asn | Asn | Ile | Asn | Ser | Ala | Ile | Asn | Asn | Leu | Asn | Gly |
| 5255 | | | | | | 5260 | | | | | 5265 | | | |
| Glu | Ala | Lys | Leu | Thr | Lys | Ala | Lys | Glu | Asp | Ala | Val | Ala | Ser | Ile |
| 5270 | | | | | | 5275 | | | | | 5280 | | | |
| Asn | Asn | Leu | Ser | Gly | Leu | Thr | Asn | Glu | Gln | Lys | Pro | Lys | Glu | Asn |
| 5285 | | | | | | 5290 | | | | | 5295 | | | |
| Gln | Ala | Val | Asn | Gly | Ala | Gln | Thr | Arg | Asp | Gln | Val | Ala | Asn | Lys |
| 5300 | | | | | | 5305 | | | | | 5310 | | | |
| Leu | Arg | Asp | Ala | Glu | Ala | Leu | Asp | Gln | Ser | Met | Gln | Thr | Leu | Arg |
| 5315 | | | | | | 5320 | | | | | 5325 | | | |
| Asp | Leu | Val | Asn | Asn | Gln | Asn | Ala | Ile | His | Ser | Thr | Ser | Asn | Tyr |
| 5330 | | | | | | 5335 | | | | | 5340 | | | |
| Phe | Asn | Glu | Asp | Ser | Thr | Gln | Lys | Asn | Thr | Tyr | Asp | Asn | Ala | Ile |
| 5345 | | | | | | 5350 | | | | | 5355 | | | |
| Asp | Asn | Gly | Ser | Thr | Tyr | Ile | Thr | Gly | Gln | His | Asn | Pro | Glu | Leu |
| 5360 | | | | | | 5365 | | | | | 5370 | | | |

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|------|-----|-----|-----|-----|-----|------|-----|-----|-----|-----|------|-----|-----|-----|
| Asn | Lys | Ser | Thr | Ile | Asp | Gln | Thr | Ile | Ser | Arg | Ile | Asn | Thr | Ala |
| 5375 | | | | | | 5380 | | | | | 5385 | | | |
| Lys | Asn | Asp | Leu | His | Gly | Val | Glu | Lys | Leu | Gln | Arg | Asp | Lys | Gly |
| 5390 | | | | | | 5395 | | | | | 5400 | | | |
| Thr | Ala | Asn | Gln | Glu | Ile | Gly | Gln | Leu | Gly | Tyr | Leu | Asn | Asp | Pro |
| 5405 | | | | | | 5410 | | | | | 5415 | | | |
| Gln | Lys | Ser | Gly | Glu | Glu | Ser | Leu | Val | Asn | Gly | Ser | Asn | Thr | Arg |
| 5420 | | | | | | 5425 | | | | | 5430 | | | |
| Ser | Glu | Val | Glu | Glu | His | Leu | Asn | Glu | Ala | Lys | Ser | Leu | Asn | Asn |
| 5435 | | | | | | 5440 | | | | | 5445 | | | |
| Ala | Met | Lys | Gln | Leu | Arg | Asp | Lys | Val | Ala | Glu | Lys | Thr | Asn | Val |
| 5450 | | | | | | 5455 | | | | | 5460 | | | |
| Lys | Gln | Ser | Ser | Asp | Tyr | Ile | Asn | Asp | Ser | Thr | Glu | His | Gln | Arg |
| 5465 | | | | | | 5470 | | | | | 5475 | | | |
| Gly | Tyr | Asp | Gln | Ala | Leu | Gln | Glu | Ala | Glu | Asn | Ile | Ile | Asn | Glu |
| 5480 | | | | | | 5485 | | | | | 5490 | | | |
| Ile | Gly | Asn | Pro | Thr | Leu | Asn | Lys | Ser | Glu | Ile | Glu | Gln | Lys | Leu |
| 5495 | | | | | | 5500 | | | | | 5505 | | | |
| Gln | Gln | Leu | Thr | Asp | Ala | Gln | Asn | Ala | Leu | Gln | Gly | Ser | His | Leu |
| 5510 | | | | | | 5515 | | | | | 5520 | | | |
| Leu | Glu | Glu | Ala | Lys | Asn | Asn | Ala | Ile | Thr | Gly | Ile | Asn | Lys | Leu |
| 5525 | | | | | | 5530 | | | | | 5535 | | | |
| Thr | Ala | Leu | Asn | Asp | Ala | Gln | Arg | Gln | Lys | Ala | Ile | Glu | Asn | Val |
| 5540 | | | | | | 5545 | | | | | 5550 | | | |
| Gln | Ala | Gln | Gln | Thr | Ile | Pro | Ala | Val | Asn | Gln | Gln | Leu | Thr | Leu |
| 5555 | | | | | | 5560 | | | | | 5565 | | | |
| Asp | Arg | Glu | Ile | Asn | Thr | Ala | Met | Gln | Ala | Leu | Arg | Asp | Lys | Val |
| 5570 | | | | | | 5575 | | | | | 5580 | | | |
| Gly | Gln | Gln | Asn | Asn | Val | His | Gln | Gln | Ser | Asn | Tyr | Phe | Asn | Glu |
| 5585 | | | | | | 5590 | | | | | 5595 | | | |
| Asp | Glu | Gln | Pro | Lys | His | Asn | Tyr | Asp | Asn | Ser | Val | Gln | Ala | Gly |
| 5600 | | | | | | 5605 | | | | | 5610 | | | |
| Gln | Thr | Ile | Ile | Asp | Lys | Leu | Gln | Asp | Pro | Ile | Met | Asn | Lys | Asn |
| 5615 | | | | | | 5620 | | | | | 5625 | | | |
| Glu | Ile | Glu | Gln | Ala | Ile | Asn | Gln | Ile | Asn | Thr | Thr | Gln | Thr | Ala |
| 5630 | | | | | | 5635 | | | | | 5640 | | | |
| Leu | Ser | Gly | Glu | Asn | Lys | Leu | His | Thr | Asp | Gln | Glu | Ser | Thr | Asn |
| 5645 | | | | | | 5650 | | | | | 5655 | | | |
| Arg | Gln | Ile | Glu | Gly | Leu | Ser | Ser | Leu | Asn | Thr | Ala | Gln | Ile | Asn |
| 5660 | | | | | | 5665 | | | | | 5670 | | | |
| Ala | Glu | Lys | Asp | Leu | Val | Asn | Gln | Ala | Lys | Thr | Arg | Thr | Asp | Val |
| 5675 | | | | | | 5680 | | | | | 5685 | | | |
| Ala | Gln | Lys | Leu | Ala | Ala | Ala | Lys | Glu | Ile | Asn | Ser | Ala | Met | Ser |
| 5690 | | | | | | 5695 | | | | | 5700 | | | |
| Asn | Leu | Arg | Asp | Gly | Ile | Gln | Asn | Lys | Glu | Asp | Ile | Lys | Arg | Ser |
| 5705 | | | | | | 5710 | | | | | 5715 | | | |
| Ser | Ala | Tyr | Ile | Asn | Ala | Asp | Pro | Thr | Lys | Val | Thr | Ala | Tyr | Asp |
| 5720 | | | | | | 5725 | | | | | 5730 | | | |
| Gln | Ala | Leu | Gln | Asn | Ala | Glu | Asn | Ile | Ile | Asn | Ala | Thr | Pro | Asn |
| 5735 | | | | | | 5740 | | | | | 5745 | | | |
| Val | Glu | Leu | Asn | Lys | Ala | Thr | Ile | Glu | Gln | Ala | Leu | Ser | Arg | Val |
| 5750 | | | | | | 5755 | | | | | 5760 | | | |

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|------|-----|-----|-----|-----|-----|------|-----|-----|-----|-----|------|------|-----|-----|
| Gln | Gln | Ala | Gln | Gln | Asp | Leu | Asp | Gly | Val | Gln | Gln | Leu | Ala | Asn |
| 5765 | | | | | | 5770 | | | | | | 5775 | | |
| Ala | Lys | Gln | Gln | Ala | Thr | Gln | Thr | Val | Asn | Gly | Leu | Asn | Ser | Leu |
| 5780 | | | | | | 5785 | | | | | 5790 | | | |
| Asn | Asp | Gly | Gln | Lys | Arg | Glu | Leu | Asn | Leu | Leu | Ile | Asn | Ser | Ala |
| 5795 | | | | | | 5800 | | | | | 5805 | | | |
| Asn | Thr | Arg | Thr | Lys | Val | Gln | Glu | Glu | Leu | Asn | Lys | Ala | Thr | Glu |
| 5810 | | | | | | 5815 | | | | | 5820 | | | |
| Leu | Asn | His | Ala | Met | Glu | Ala | Leu | Arg | Asn | Ser | Val | Gln | Asn | Val |
| 5825 | | | | | | 5830 | | | | | 5835 | | | |
| Asp | Gln | Val | Lys | Gln | Ser | Ser | Asn | Tyr | Val | Asn | Glu | Asp | Gln | Pro |
| 5840 | | | | | | 5845 | | | | | 5850 | | | |
| Glu | Gln | His | Asn | Tyr | Asp | Asn | Ala | Val | Asn | Glu | Ala | Gln | Ala | Thr |
| 5855 | | | | | | 5860 | | | | | 5865 | | | |
| Ile | Asn | Asn | Asn | Ala | Gln | Pro | Val | Leu | Asp | Lys | Leu | Ala | Ile | Glu |
| 5870 | | | | | | 5875 | | | | | 5880 | | | |
| Arg | Leu | Thr | Gln | Thr | Val | Asn | Thr | Thr | Lys | Asp | Ala | Leu | His | Gly |
| 5885 | | | | | | 5890 | | | | | 5895 | | | |
| Ala | Gln | Lys | Leu | Thr | Gln | Asp | Gln | Gln | Ala | Ala | Glu | Thr | Gly | Ile |
| 5900 | | | | | | 5905 | | | | | 5910 | | | |
| Arg | Gly | Leu | Thr | Ser | Leu | Asn | Glu | Pro | Gln | Lys | Asn | Ala | Glu | Val |
| 5915 | | | | | | 5920 | | | | | 5925 | | | |
| Ala | Lys | Val | Thr | Ala | Ala | Thr | Thr | Arg | Asp | Glu | Val | Arg | Asn | Ile |
| 5930 | | | | | | 5935 | | | | | 5940 | | | |
| Arg | Gln | Glu | Ala | Thr | Thr | Leu | Asp | Thr | Ala | Met | Leu | Gly | Leu | Arg |
| 5945 | | | | | | 5950 | | | | | 5955 | | | |
| Lys | Ser | Ile | Lys | Asp | Lys | Asn | Asp | Thr | Lys | Asn | Ser | Ser | Lys | Tyr |
| 5960 | | | | | | 5965 | | | | | 5970 | | | |
| Ile | Asn | Glu | Asp | His | Asp | Gln | Gln | Gln | Ala | Tyr | Asp | Asn | Ala | Val |
| 5975 | | | | | | 5980 | | | | | 5985 | | | |
| Asn | Asn | Ala | Gln | Gln | Val | Ile | Asp | Glu | Thr | Gln | Ala | Thr | Leu | Ser |
| 5990 | | | | | | 5995 | | | | | 6000 | | | |
| Ser | Asp | Thr | Ile | Asn | Gln | Leu | Ala | Asn | Ala | Val | Thr | Gln | Ala | Lys |
| 6005 | | | | | | 6010 | | | | | 6015 | | | |
| Ser | Asn | Leu | His | Gly | Asp | Thr | Lys | Leu | Gln | His | Asp | Lys | Asp | Ser |
| 6020 | | | | | | 6025 | | | | | 6030 | | | |
| Ala | Lys | Gln | Thr | Ile | Ala | Gln | Leu | Gln | Asn | Leu | Asn | Ser | Ala | Gln |
| 6035 | | | | | | 6040 | | | | | 6045 | | | |
| Lys | His | Met | Glu | Asp | Ser | Leu | Ile | Asp | Asn | Glu | Ser | Thr | Arg | Thr |
| 6050 | | | | | | 6055 | | | | | 6060 | | | |
| Gln | Val | Gln | His | Asp | Leu | Thr | Glu | Ala | Gln | Ala | Leu | Asp | Gly | Leu |
| 6065 | | | | | | 6070 | | | | | 6075 | | | |
| Met | Gly | Ala | Leu | Lys | Glu | Ser | Ile | Lys | Asp | Tyr | Thr | Asn | Ile | Val |
| 6080 | | | | | | 6085 | | | | | 6090 | | | |
| Ser | Asn | Gly | Asn | Tyr | Ile | Asn | Ala | Glu | Pro | Ser | Lys | Lys | Gln | Ala |
| 6095 | | | | | | 6100 | | | | | 6105 | | | |
| Tyr | Asp | Ala | Ala | Val | Gln | Asn | Ala | Gln | Asn | Ile | Ile | Asn | Gly | Thr |
| 6110 | | | | | | 6115 | | | | | 6120 | | | |
| Asn | Gln | Pro | Thr | Ile | Asn | Lys | Gly | Asn | Val | Thr | Thr | Ala | Thr | Gln |
| 6125 | | | | | | 6130 | | | | | 6135 | | | |
| Thr | Val | Lys | Asn | Thr | Lys | Asp | Ala | Leu | Asp | Gly | Asp | His | Arg | Leu |
| 6140 | | | | | | 6145 | | | | | 6150 | | | |
| Glu | Glu | Ala | Lys | Asn | Asn | Ala | Asn | Gln | Thr | Ile | Arg | Asn | Leu | Ser |

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|------|-----|-----|-----|-----|-----|------|-----|-----|-----|-----|------|-----|-----|-----|
| Leu | Ile | Asn | Asn | Ala | Asp | Thr | Arg | Asp | Glu | Val | Asn | Lys | Gln | Leu |
| 6560 | | | | | | 6565 | | | | | 6570 | | | |
| Glu | Ile | Ala | Lys | Gln | Leu | Asn | Gly | Asp | Met | Ser | Thr | Leu | His | Lys |
| 6575 | | | | | | 6580 | | | | | 6585 | | | |
| Val | Ile | Asn | Asp | Lys | Asp | Gln | Ile | Gln | His | Leu | Ser | Asn | Tyr | Ile |
| 6590 | | | | | | 6595 | | | | | 6600 | | | |
| Asn | Ala | Asp | Asn | Asp | Lys | Lys | Gln | Asn | Tyr | Asp | Asn | Ala | Ile | Lys |
| 6605 | | | | | | 6610 | | | | | 6615 | | | |
| Glu | Ala | Glu | Asp | Leu | Ile | His | Asn | His | Pro | Asp | Thr | Leu | Asp | His |
| 6620 | | | | | | 6625 | | | | | 6630 | | | |
| Lys | Ala | Leu | Gln | Asp | Leu | Leu | Asn | Lys | Ile | Asp | Gln | Ala | His | Asn |
| 6635 | | | | | | 6640 | | | | | 6645 | | | |
| Glu | Leu | Asn | Gly | Glu | Ser | Arg | Phe | Lys | Gln | Ala | Leu | Asp | Asn | Ala |
| 6650 | | | | | | 6655 | | | | | 6660 | | | |
| Leu | Asn | Asp | Ile | Asp | Ser | Leu | Asn | Ser | Leu | Asn | Val | Pro | Gln | Arg |
| 6665 | | | | | | 6670 | | | | | 6675 | | | |
| Gln | Thr | Val | Lys | Asp | Asn | Ile | Asn | His | Val | Thr | Thr | Leu | Glu | Ser |
| 6680 | | | | | | 6685 | | | | | 6690 | | | |
| Leu | Ala | Gln | Glu | Leu | Gln | Lys | Ala | Lys | Glu | Leu | Asn | Asp | Ala | Met |
| 6695 | | | | | | 6700 | | | | | 6705 | | | |
| Lys | Ala | Met | Arg | Asp | Ser | Ile | Met | Asn | Gln | Glu | Gln | Ile | Arg | Lys |
| 6710 | | | | | | 6715 | | | | | 6720 | | | |
| Asn | Ser | Asn | Tyr | Thr | Asn | Glu | Asp | Leu | Ala | Gln | Gln | Asn | Ala | Tyr |
| 6725 | | | | | | 6730 | | | | | 6735 | | | |
| Asn | His | Ala | Val | Asp | Lys | Ile | Asn | Asn | Ile | Ile | Gly | Glu | Asp | Asn |
| 6740 | | | | | | 6745 | | | | | 6750 | | | |
| Ala | Thr | Met | Asp | Pro | Gln | Ile | Ile | Lys | Gln | Ala | Thr | Gln | Asp | Ile |
| 6755 | | | | | | 6760 | | | | | 6765 | | | |
| Asn | Thr | Ala | Ile | Asn | Gly | Leu | Asn | Gly | Asp | Gln | Lys | Leu | Gln | Asp |
| 6770 | | | | | | 6775 | | | | | 6780 | | | |
| Ala | Lys | Thr | Asp | Ala | Lys | Gln | Gln | Ile | Thr | Asn | Phe | Thr | Gly | Leu |
| 6785 | | | | | | 6790 | | | | | 6795 | | | |
| Thr | Glu | Pro | Gln | Lys | Gln | Ala | Leu | Glu | Asn | Ile | Ile | Asn | Gln | Gln |
| 6800 | | | | | | 6805 | | | | | 6810 | | | |
| Thr | Ser | Arg | Ala | Asn | Val | Ala | Lys | Gln | Leu | Ser | His | Ala | Lys | Phe |
| 6815 | | | | | | 6820 | | | | | 6825 | | | |
| Leu | Asn | Gly | Lys | Met | Glu | Glu | Leu | Lys | Val | Ala | Val | Ala | Lys | Ala |
| 6830 | | | | | | 6835 | | | | | 6840 | | | |
| Ser | Leu | Val | Arg | Gln | Asn | Ser | Asn | Tyr | Ile | Asn | Glu | Asp | Val | Ser |
| 6845 | | | | | | 6850 | | | | | 6855 | | | |
| Glu | Lys | Glu | Ala | Tyr | Glu | Gln | Ala | Ile | Ala | Lys | Gly | Gln | Glu | Ile |
| 6860 | | | | | | 6865 | | | | | 6870 | | | |
| Ile | Asn | Ser | Glu | Asn | Asn | Pro | Thr | Ile | Ser | Ser | Thr | Asp | Ile | Asn |
| 6875 | | | | | | 6880 | | | | | 6885 | | | |
| Arg | Thr | Ile | Gln | Glu | Ile | Asn | Asp | Ala | Glu | Gln | Asn | Leu | His | Gly |
| 6890 | | | | | | 6895 | | | | | 6900 | | | |
| Asp | Asn | Lys | Leu | Arg | Gln | Ala | Gln | Glu | Ile | Ala | Lys | Asn | Glu | Ile |
| 6905 | | | | | | 6910 | | | | | 6915 | | | |
| Gln | Asn | Leu | Asp | Gly | Leu | Asn | Ser | Ala | Gln | Ile | Thr | Lys | Leu | Ile |
| 6920 | | | | | | 6925 | | | | | 6930 | | | |
| Gln | Asp | Ile | Gly | Arg | Thr | Thr | Thr | Lys | Pro | Ala | Val | Thr | Gln | Lys |
| 6935 | | | | | | 6940 | | | | | 6945 | | | |

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|------|-----|-----|-----|-----|-----|------|-----|-----|-----|-----|------|-----|-----|-----|
| Leu | Glu | Glu | Ala | Lys | Ala | Ile | Asn | Gln | Ala | Met | Gln | Gln | Leu | Lys |
| 6950 | | | | | | 6955 | | | | | 6960 | | | |
| Gln | Ser | Ile | Ala | Asp | Lys | Asp | Ala | Thr | Leu | Asn | Ser | Ser | Asn | Tyr |
| 6965 | | | | | | 6970 | | | | | 6975 | | | |
| Leu | Asn | Glu | Asp | Ser | Glu | Lys | Lys | Leu | Ala | Tyr | Asp | Asn | Ala | Val |
| 6980 | | | | | | 6985 | | | | | 6990 | | | |
| Ser | Gln | Ala | Glu | Gln | Leu | Ile | Asn | Gln | Leu | Asn | Asp | Pro | Thr | Met |
| 6995 | | | | | | 7000 | | | | | 7005 | | | |
| Asp | Ile | Ser | Asn | Ile | Gln | Ala | Ile | Thr | Gln | Lys | Val | Ile | Gln | Ala |
| 7010 | | | | | | 7015 | | | | | 7020 | | | |
| Lys | Asp | Ser | Leu | His | Gly | Ala | Asn | Lys | Leu | Ala | Gln | Asn | Gln | Ala |
| 7025 | | | | | | 7030 | | | | | 7035 | | | |
| Asp | Ser | Asn | Leu | Ile | Ile | Asn | Gln | Ser | Thr | Asn | Leu | Asn | Asp | Lys |
| 7040 | | | | | | 7045 | | | | | 7050 | | | |
| Gln | Lys | Gln | Ala | Leu | Asn | Asp | Leu | Ile | Asn | His | Ala | Gln | Thr | Lys |
| 7055 | | | | | | 7060 | | | | | 7065 | | | |
| Gln | Gln | Val | Ala | Glu | Ile | Ile | Ala | Gln | Ala | Asn | Lys | Leu | Asn | Asn |
| 7070 | | | | | | 7075 | | | | | 7080 | | | |
| Glu | Met | Gly | Thr | Leu | Lys | Thr | Leu | Val | Glu | Glu | Gln | Ser | Asn | Val |
| 7085 | | | | | | 7090 | | | | | 7095 | | | |
| His | Gln | Gln | Ser | Lys | Tyr | Ile | Asn | Glu | Asp | Pro | Gln | Val | Gln | Asn |
| 7100 | | | | | | 7105 | | | | | 7110 | | | |
| Ile | Tyr | Asn | Asp | Ser | Ile | Gln | Lys | Gly | Arg | Glu | Ile | Leu | Asn | Gly |
| 7115 | | | | | | 7120 | | | | | 7125 | | | |
| Thr | Thr | Asp | Asp | Val | Leu | Asn | Asn | Asn | Lys | Ile | Ala | Asp | Ala | Ile |
| 7130 | | | | | | 7135 | | | | | 7140 | | | |
| Gln | Asn | Ile | His | Leu | Thr | Lys | Asn | Asp | Leu | His | Gly | Asp | Gln | Lys |
| 7145 | | | | | | 7150 | | | | | 7155 | | | |
| Leu | Gln | Lys | Ala | Gln | Gln | Asp | Ala | Thr | Asn | Glu | Leu | Asn | Tyr | Leu |
| 7160 | | | | | | 7165 | | | | | 7170 | | | |
| Thr | Asn | Leu | Asn | Asn | Ser | Gln | Arg | Gln | Ser | Glu | His | Asp | Glu | Ile |
| 7175 | | | | | | 7180 | | | | | 7185 | | | |
| Asn | Ser | Ala | Pro | Ser | Arg | Thr | Glu | Val | Ser | Asn | Asp | Leu | Asn | His |
| 7190 | | | | | | 7195 | | | | | 7200 | | | |
| Ala | Lys | Ala | Leu | Asn | Glu | Ala | Met | Arg | Gln | Leu | Glu | Asn | Glu | Val |
| 7205 | | | | | | 7210 | | | | | 7215 | | | |
| Ala | Leu | Glu | Asn | Ser | Val | Lys | Lys | Leu | Ser | Asp | Phe | Ile | Asn | Glu |
| 7220 | | | | | | 7225 | | | | | 7230 | | | |
| Asp | Glu | Ala | Ala | Gln | Asn | Glu | Tyr | Ser | Asn | Ala | Leu | Gln | Lys | Ala |
| 7235 | | | | | | 7240 | | | | | 7245 | | | |
| Lys | Asp | Ile | Ile | Asn | Gly | Val | Pro | Ser | Ser | Thr | Leu | Asp | Lys | Ala |
| 7250 | | | | | | 7255 | | | | | 7260 | | | |
| Thr | Ile | Glu | Asp | Ala | Leu | Leu | Glu | Leu | Gln | Asn | Ala | Arg | Glu | Ser |
| 7265 | | | | | | 7270 | | | | | 7275 | | | |
| Leu | His | Gly | Glu | Gln | Lys | Leu | Gln | Glu | Ala | Lys | Asn | Gln | Ala | Val |
| 7280 | | | | | | 7285 | | | | | 7290 | | | |
| Ala | Glu | Ile | Asp | Asn | Leu | Gln | Ala | Leu | Asn | Pro | Gly | Gln | Val | Leu |
| 7295 | | | | | | 7300 | | | | | 7305 | | | |
| Ala | Glu | Lys | Thr | Leu | Val | Asn | Gln | Ala | Ser | Thr | Lys | Pro | Glu | Val |
| 7310 | | | | | | 7315 | | | | | 7320 | | | |
| Gln | Glu | Ala | Leu | Gln | Lys | Ala | Lys | Glu | Leu | Asn | Glu | Ala | Met | Lys |
| 7325 | | | | | | 7330 | | | | | 7335 | | | |
| Ala | Leu | Lys | Thr | Glu | Ile | Asn | Lys | Lys | Glu | Gln | Ile | Lys | Ala | Asp |

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|-------------------------------------|------|-----------------------------------------|
| 7340 | 7345 | 7350 |
| Ser Arg Tyr Val Asn Ala Asp 7355 | 7360 | Ser Gly Leu Gln Ala Asn Tyr Asn 7365 |
| Ser Ala Leu Asn Tyr Gly Ser 7370 | 7375 | Gln Ile Ile Ala Thr Thr Gln Pro 7380 |
| Pro Glu Leu Asn Lys Asp Val 7385 | 7390 | Ile Asn Arg Ala Thr Gln Thr Ile 7395 |
| Lys Thr Ala Glu Asn Asn Leu 7400 | 7405 | Asn Gly Gln Ser Lys Leu Ala Glu 7410 |
| Ala Lys Ser Asp Gly Asn Gln 7415 | 7420 | Ser Ile Glu His Leu Gln Gly Leu 7425 |
| Thr Gln Ser Gln Lys Asp Lys 7430 | 7435 | Gln His Asp Leu Ile Asn Gln Ala 7440 |
| Gln Thr Lys Gln Gln Val Asp 7445 | 7450 | Asp Ile Val Asn Asn Ser Lys Gln 7455 |
| Leu Asp Asn Ser Met Asn Gln 7460 | 7465 | Leu Gln Gln Ile Val Asn Asn Asp 7470 |
| Asn Thr Val Lys Gln Asn Ser 7475 | 7480 | Asp Phe Ile Asn Glu Asp Ser Ser 7485 |
| Gln Gln Asp Ala Tyr Asn His 7490 | 7495 | Ala Ile Gln Ala Ala Lys Asp Leu 7500 |
| Ile Thr Ala His Pro Thr Ile 7505 | 7510 | Met Asp Lys Asn Gln Ile Asp Gln 7515 |
| Ala Ile Glu Asn Ile Lys Gln 7520 | 7525 | Ala Leu Asn Asp Leu His Gly Ser 7530 |
| Asn Lys Leu Ser Glu Asp Lys 7535 | 7540 | Lys Glu Ala Ser Glu Gln Leu Gln 7545 |
| Asn Leu Asn Ser Leu Thr Asn 7550 | 7555 | Gly Gln Lys Asp Thr Ile Leu Asn 7560 |
| His Ile Phe Ser Ala Pro Thr 7565 | 7570 | Arg Ser Gln Val Gly Glu Lys Ile 7575 |
| Ala Ser Ala Lys Gln Leu Asn 7580 | 7585 | Asn Thr Met Lys Ala Leu Arg Asp 7590 |
| Ser Ile Ala Asp Asn Asn Glu 7595 | 7600 | Ile Leu Gln Ser Ser Lys Tyr Phe 7605 |
| Asn Glu Asp Ser Glu Gln Gln 7610 | 7615 | Asn Ala Tyr Asn Gln Ala Val Asn 7620 |
| Lys Ala Lys Asn Ile Ile Asn 7625 | 7630 | Asp Gln Pro Thr Pro Val Met Ala 7635 |
| Asn Asp Glu Ile Gln Ser Val 7640 | 7645 | Leu Asn Glu Val Lys Gln Thr Lys 7650 |
| Asp Asn Leu His Gly Asp Gln 7655 | 7660 | Lys Leu Ala Asn Asp Lys Thr Asp 7665 |
| Ala Gln Ala Thr Leu Asn Ala 7670 | 7675 | Leu Asn Tyr Leu Asn Gln Ala Gln 7680 |
| Arg Gly Asn Leu Glu Thr Lys 7685 | 7690 | Val Gln Asn Ser Asn Ser Arg Pro 7695 |
| Glu Val Gln Lys Val Val Gln 7700 | 7705 | Leu Ala Asn Gln Leu Asn Asp Ala 7710 |
| Met Lys Lys Leu Asp Asp Ala 7715 | 7720 | Leu Thr Gly Asn Asp Ala Ile Lys 7725 |
| Gln Thr Ser Asn Tyr Ile Asn 7730 | 7735 | Glu Asp Thr Ser Gln Gln Val Asn 7740 |

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|------|-----|-----|-----|-----|-----|------|-----|-----|-----|-----|------|-----|-----|-----|
| Phe | Asp | Glu | Tyr | Thr | Asp | Arg | Gly | Lys | Asn | Ile | Val | Ala | Glu | Gln |
| 7745 | | | | | | 7750 | | | | | 7755 | | | |
| Thr | Asn | Pro | Asn | Met | Ser | Pro | Thr | Asn | Ile | Asn | Thr | Ile | Ala | Asp |
| 7760 | | | | | | 7765 | | | | | 7770 | | | |
| Lys | Ile | Thr | Glu | Ala | Lys | Asn | Asp | Leu | His | Gly | Val | Gln | Lys | Leu |
| 7775 | | | | | | 7780 | | | | | 7785 | | | |
| Lys | Gln | Ala | Gln | Gln | Gln | Ser | Ile | Asn | Thr | Ile | Asn | Gln | Met | Thr |
| 7790 | | | | | | 7795 | | | | | 7800 | | | |
| Gly | Leu | Asn | Gln | Ala | Gln | Lys | Glu | Gln | Leu | Asn | Gln | Glu | Ile | Gln |
| 7805 | | | | | | 7810 | | | | | 7815 | | | |
| Gln | Thr | Gln | Thr | Arg | Ser | Glu | Val | His | Gln | Val | Ile | Asn | Lys | Ala |
| 7820 | | | | | | 7825 | | | | | 7830 | | | |
| Gln | Ala | Leu | Asn | Asp | Ser | Met | Asn | Thr | Leu | Arg | Gln | Ser | Ile | Thr |
| 7835 | | | | | | 7840 | | | | | 7845 | | | |
| Asp | Glu | His | Glu | Val | Lys | Gln | Thr | Ser | Asn | Tyr | Ile | Asn | Glu | Thr |
| 7850 | | | | | | 7855 | | | | | 7860 | | | |
| Val | Gly | Asn | Gln | Thr | Ala | Tyr | Asn | Asn | Ala | Val | Asp | Arg | Val | Lys |
| 7865 | | | | | | 7870 | | | | | 7875 | | | |
| Gln | Ile | Ile | Asn | Gln | Thr | Ser | Asn | Pro | Thr | Met | Asn | Pro | Leu | Glu |
| 7880 | | | | | | 7885 | | | | | 7890 | | | |
| Val | Glu | Arg | Ala | Thr | Ser | Asn | Val | Lys | Ile | Ser | Lys | Asp | Ala | Leu |
| 7895 | | | | | | 7900 | | | | | 7905 | | | |
| His | Gly | Glu | Arg | Glu | Leu | Asn | Asp | Asn | Lys | Asn | Ser | Lys | Thr | Phe |
| 7910 | | | | | | 7915 | | | | | 7920 | | | |
| Ala | Val | Asn | His | Leu | Asp | Asn | Leu | Asn | Gln | Ala | Gln | Lys | Glu | Ala |
| 7925 | | | | | | 7930 | | | | | 7935 | | | |
| Leu | Thr | His | Glu | Ile | Glu | Gln | Ala | Thr | Ile | Val | Ser | Gln | Val | Asn |
| 7940 | | | | | | 7945 | | | | | 7950 | | | |
| Asn | Ile | Tyr | Asn | Lys | Ala | Lys | Ala | Leu | Asn | Asn | Asp | Met | Lys | Lys |
| 7955 | | | | | | 7960 | | | | | 7965 | | | |
| Leu | Lys | Asp | Ile | Val | Ala | Gln | Gln | Asp | Asn | Val | Arg | Gln | Ser | Asn |
| 7970 | | | | | | 7975 | | | | | 7980 | | | |
| Asn | Tyr | Ile | Asn | Glu | Asp | Ser | Thr | Pro | Gln | Asn | Met | Tyr | Asn | Asp |
| 7985 | | | | | | 7990 | | | | | 7995 | | | |
| Thr | Ile | Asn | His | Ala | Gln | Ser | Ile | Ile | Asp | Gln | Val | Ala | Asn | Pro |
| 8000 | | | | | | 8005 | | | | | 8010 | | | |
| Thr | Met | Ser | His | Asp | Glu | Ile | Glu | Asn | Ala | Ile | Asn | Asn | Ile | Lys |
| 8015 | | | | | | 8020 | | | | | 8025 | | | |
| His | Ala | Ile | Asn | Ala | Leu | Asp | Gly | Glu | His | Lys | Leu | Gln | Gln | Ala |
| 8030 | | | | | | 8035 | | | | | 8040 | | | |
| Lys | Glu | Asn | Ala | Asn | Leu | Leu | Ile | Asn | Ser | Leu | Asn | Asp | Leu | Asn |
| 8045 | | | | | | 8050 | | | | | 8055 | | | |
| Ala | Pro | Gln | Arg | Asp | Ala | Ile | Asn | Arg | Leu | Val | Asn | Glu | Ala | Gln |
| 8060 | | | | | | 8065 | | | | | 8070 | | | |
| Thr | Arg | Glu | Lys | Val | Ala | Glu | Gln | Leu | Gln | Ser | Ala | Gln | Ala | Leu |
| 8075 | | | | | | 8080 | | | | | 8085 | | | |
| Asn | Asp | Ala | Met | Lys | His | Leu | Arg | Asn | Ser | Ile | Gln | Asn | Gln | Ser |
| 8090 | | | | | | 8095 | | | | | 8100 | | | |
| Ser | Val | Arg | Gln | Glu | Ser | Lys | Tyr | Ile | Asn | Ala | Ser | Asp | Ala | Lys |
| 8105 | | | | | | 8110 | | | | | 8115 | | | |
| Lys | Glu | Gln | Tyr | Asn | His | Ala | Val | Arg | Glu | Val | Glu | Asn | Ile | Ile |
| 8120 | | | | | | 8125 | | | | | 8130 | | | |

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|------|-----|-----|-----|-----|-----|------|-----|-----|-----|-----|------|-----|-----|-----|
| Asn | Glu | Gln | His | Pro | Thr | Leu | Asp | Lys | Glu | Ile | Ile | Lys | Gln | Leu |
| 8135 | | | | | | 8140 | | | | | 8145 | | | |
| Thr | Asp | Gly | Val | Asn | Gln | Ala | Asn | Asn | Asp | Leu | Asn | Gly | Val | Glu |
| 8150 | | | | | | 8155 | | | | | 8160 | | | |
| Leu | Leu | Asp | Ala | Asp | Lys | Gln | Asn | Ala | His | Gln | Ser | Ile | Pro | Thr |
| 8165 | | | | | | 8170 | | | | | 8175 | | | |
| Leu | Met | His | Leu | Asn | Gln | Ala | Gln | Gln | Asn | Ala | Leu | Asn | Glu | Lys |
| 8180 | | | | | | 8185 | | | | | 8190 | | | |
| Ile | Asn | Asn | Ala | Val | Thr | Arg | Thr | Glu | Val | Ala | Ala | Ile | Ile | Gly |
| 8195 | | | | | | 8200 | | | | | 8205 | | | |
| Gln | Ala | Lys | Leu | Leu | Asp | His | Ala | Met | Glu | Asn | Leu | Glu | Glu | Ser |
| 8210 | | | | | | 8215 | | | | | 8220 | | | |
| Ile | Lys | Asp | Lys | Glu | Gln | Val | Lys | Gln | Ser | Ser | Asn | Tyr | Ile | Asn |
| 8225 | | | | | | 8230 | | | | | 8235 | | | |
| Glu | Asp | Ser | Asp | Val | Gln | Glu | Thr | Tyr | Asp | Asn | Ala | Val | Asp | His |
| 8240 | | | | | | 8245 | | | | | 8250 | | | |
| Val | Thr | Glu | Ile | Leu | Asn | Gln | Thr | Val | Asn | Pro | Thr | Leu | Ser | Ile |
| 8255 | | | | | | 8260 | | | | | 8265 | | | |
| Glu | Asp | Ile | Glu | His | Ala | Ile | Asn | Glu | Val | Asn | Gln | Ala | Lys | Lys |
| 8270 | | | | | | 8275 | | | | | 8280 | | | |
| Gln | Leu | Arg | Gly | Lys | Gln | Lys | Leu | Tyr | Gln | Thr | Ile | Asp | Leu | Ala |
| 8285 | | | | | | 8290 | | | | | 8295 | | | |
| Asp | Lys | Glu | Leu | Ser | Lys | Leu | Asp | Asp | Leu | Thr | Ser | Gln | Gln | Ser |
| 8300 | | | | | | 8305 | | | | | 8310 | | | |
| Ser | Ser | Ile | Ser | Asn | Gln | Ile | Tyr | Thr | Ala | Lys | Thr | Arg | Thr | Glu |
| 8315 | | | | | | 8320 | | | | | 8325 | | | |
| Val | Ala | Gln | Ala | Ile | Glu | Lys | Ala | Lys | Ser | Leu | Asn | His | Ala | Met |
| 8330 | | | | | | 8335 | | | | | 8340 | | | |
| Lys | Ala | Leu | Asn | Lys | Val | Tyr | Lys | Asn | Ala | Asp | Lys | Val | Leu | Asp |
| 8345 | | | | | | 8350 | | | | | 8355 | | | |
| Ser | Ser | Arg | Phe | Ile | Asn | Glu | Asp | Gln | Pro | Glu | Lys | Lys | Ala | Tyr |
| 8360 | | | | | | 8365 | | | | | 8370 | | | |
| Gln | Gln | Ala | Ile | Asn | His | Val | Asp | Ser | Ile | Ile | His | Arg | Gln | Thr |
| 8375 | | | | | | 8380 | | | | | 8385 | | | |
| Asn | Pro | Glu | Met | Asp | Pro | Thr | Val | Ile | Asn | Ser | Ile | Thr | His | Glu |
| 8390 | | | | | | 8395 | | | | | 8400 | | | |
| Leu | Glu | Thr | Ala | Gln | Asn | Asn | Leu | His | Gly | Asp | Gln | Lys | Leu | Ala |
| 8405 | | | | | | 8410 | | | | | 8415 | | | |
| His | Ala | Gln | Gln | Asp | Ala | Ala | Asn | Val | Ile | Asn | Gly | Leu | Ile | His |
| 8420 | | | | | | 8425 | | | | | 8430 | | | |
| Leu | Asn | Val | Ala | Gln | Arg | Glu | Val | Met | Ile | Asn | Thr | Asn | Thr | Asn |
| 8435 | | | | | | 8440 | | | | | 8445 | | | |
| Ala | Thr | Thr | Arg | Glu | Lys | Val | Ala | Lys | Asn | Leu | Asp | Asn | Ala | Gln |
| 8450 | | | | | | 8455 | | | | | 8460 | | | |
| Ala | Leu | Asp | Lys | Ala | Met | Glu | Thr | Leu | Gln | Gln | Val | Val | Ala | His |
| 8465 | | | | | | 8470 | | | | | 8475 | | | |
| Lys | Asn | Asn | Ile | Leu | Asn | Asp | Ser | Lys | Tyr | Leu | Asn | Glu | Asp | Ser |
| 8480 | | | | | | 8485 | | | | | 8490 | | | |
| Lys | Tyr | Gln | Gln | Gln | Tyr | Asp | Arg | Val | Ile | Ala | Asp | Ala | Glu | Gln |
| 8495 | | | | | | 8500 | | | | | 8505 | | | |
| Leu | Leu | Asn | Gln | Thr | Thr | Asn | Pro | Thr | Leu | Glu | Pro | Tyr | Lys | Val |
| 8510 | | | | | | 8515 | | | | | 8520 | | | |
| Asp | Ile | Val | Lys | Asp | Asn | Val | Leu | Ala | Asn | Glu | Lys | Ile | Leu | Phe |

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| 8525 | 8530 | 8535 |
| Gly Ala Glu Lys Leu Ser Tyr 8540 | Asp Lys Ser Asn 8545 | Ala Asn Asp Glu 8550 |
| Ile Lys His Met Asn Tyr 8555 | Leu 8560 | Asn Asn Ala Gln Lys Gln Ser Ile 8565 |
| Lys Asp Met Ile Ser His 8570 | Ala Leu Arg Thr 8575 | Glu Val Lys Gln 8580 |
| Leu Leu Gln Gln Ala Lys 8585 | Ile 8590 | Leu Asp Glu Ala Met Lys Ser Leu 8595 |
| Glu Asp Lys Thr Gln Val 8600 | Val 8605 | Ile Thr Asp Thr Thr Leu Pro Asn 8610 |
| Tyr Thr Glu Ala Ser Glu 8615 | Asp 8620 | Lys Lys Glu Lys Val Asp Gln Thr 8625 |
| Val Ser His Ala Gln Ala 8630 | Ile 8635 | Ile Asp Lys Ile Asn Gly Ser Asn 8640 |
| Val Ser Leu Asp Gln Val 8645 | Arg 8650 | Gln Ala Leu Glu Gln Leu Thr Gln 8655 |
| Ala Ser Glu Asn Leu Asp 8660 | Gly 8665 | Asp Gln Arg Val Glu Glu Ala Lys 8670 |
| Val His Ala Asn Gln Thr 8675 | Ile 8680 | Asp Gln Leu Thr His Leu Asn Ser 8685 |
| Leu Gln Gln Gln Thr Ala 8690 | Lys 8695 | Glu Ser Val Lys Asn Ala Thr Lys 8700 |
| Leu Glu Glu Ile Ala Thr 8705 | Val 8710 | Ser Asn Asn Ala Gln Ala Leu Asn 8715 |
| Lys Val Met Gly Lys Leu 8720 | Glu 8725 | Gln Phe Ile Asn His Ala Asp Ser 8730 |
| Val Glu Asn Ser Asp Asn Tyr 8735 | Tyr 8740 | Arg Gln Ala Asp Asp Asp Lys Ile 8745 |
| Ile Ala Tyr Asp Glu Ala 8750 | Leu 8755 | Glu His Gly Gln Asp Ile Gln Lys 8760 |
| Thr Asn Ala Thr Gln Asn 8765 | Glu 8770 | Thr Lys Gln Ala Leu Gln Gln Leu 8775 |
| Ile Tyr Ala Glu Thr Ser 8780 | Leu 8785 | Asn Gly Phe Glu Arg Leu Asn His 8790 |
| Ala Arg Pro Arg Ala Leu 8795 | Glu 8800 | Tyr Ile Lys Ser Leu Glu Lys Ile 8805 |
| Asn Asn Ala Gln Lys Ser 8810 | Ala 8815 | Leu Glu Asp Lys Val Thr Gln Ser 8820 |
| His Asp Leu Leu Glu Leu 8825 | Glu 8830 | His Ile Val Asn Glu Gly Thr Asn 8835 |
| Leu Asn Asp Ile Met Gly 8840 | Glu 8845 | Leu Ala Asn Ala Ile Val Asn Asn 8850 |
| Tyr Ala Pro Thr Lys Ala 8855 | Ser 8860 | Ile Asn Tyr Ile Asn Ala Asp Asn 8865 |
| Leu Arg Lys Asp Asn Phe 8870 | Thr 8875 | Gln Ala Ile Asn Asn Ala Arg Asp 8880 |
| Ala Leu Asn Lys Thr Gln 8885 | Gly 8890 | Gln Asn Leu Asp Phe Asn Ala Ile 8895 |
| Asp Thr Phe Lys Asp Asp 8900 | Ile 8905 | Phe Lys Thr Lys Asp Ala Leu Asn 8910 |
| Gly Ile Glu Arg Leu Thr 8915 | Ala 8920 | Ala Lys Ser Lys Ala Glu Lys Leu 8925 |

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|------|-----|-----|-----|-----|-----|------|-----|-----|-----|-----|------|-----|-----|-----|
| Ile | Asp | Ser | Leu | Lys | Phe | Ile | Asn | Lys | Ala | Gln | Phe | Thr | His | Ala |
| 8930 | | | | | | 8935 | | | | | 8940 | | | |
| Asn | Asp | Glu | Ile | Ile | Asn | Thr | Asn | Ser | Ile | Ala | Gln | Leu | Ser | Arg |
| 8945 | | | | | | 8950 | | | | | 8955 | | | |
| Ile | Val | Asn | Gln | Ala | Phe | Asp | Leu | Asn | Asp | Ala | Met | Lys | Ser | Leu |
| 8960 | | | | | | 8965 | | | | | 8970 | | | |
| Arg | Asp | Glu | Leu | Asn | Asn | Gln | Ala | Phe | Pro | Val | Gln | Ala | Ser | Ser |
| 8975 | | | | | | 8980 | | | | | 8985 | | | |
| Asn | Tyr | Ile | Asn | Ser | Asp | Glu | Asp | Leu | Lys | Gln | Gln | Phe | Asp | His |
| 8990 | | | | | | 8995 | | | | | 9000 | | | |
| Ala | Leu | Ser | Asn | Ala | Arg | Lys | Val | Leu | Ala | Lys | Glu | Asn | Gly | Lys |
| 9005 | | | | | | 9010 | | | | | 9015 | | | |
| Asn | Leu | Asp | Glu | Lys | Gln | Ile | Gln | Gly | Leu | Lys | Gln | Val | Ile | Glu |
| 9020 | | | | | | 9025 | | | | | 9030 | | | |
| Asp | Thr | Lys | Asp | Ala | Leu | Asn | Gly | Ile | Gln | Arg | Leu | Ser | Lys | Ala |
| 9035 | | | | | | 9040 | | | | | 9045 | | | |
| Lys | Ala | Lys | Ala | Ile | Gln | Tyr | Val | Gln | Ser | Leu | Ser | Tyr | Ile | Asn |
| 9050 | | | | | | 9055 | | | | | 9060 | | | |
| Asp | Ala | Gln | Arg | His | Ile | Ala | Glu | Asn | Asn | Ile | His | Asn | Ser | Asp |
| 9065 | | | | | | 9070 | | | | | 9075 | | | |
| Asp | Leu | Ser | Ser | Leu | Ala | Asn | Thr | Leu | Ser | Lys | Ala | Ser | Asp | Leu |
| 9080 | | | | | | 9085 | | | | | 9090 | | | |
| Asp | Asn | Ala | Met | Lys | Asp | Leu | Arg | Asp | Thr | Ile | Glu | Ser | Asn | Ser |
| 9095 | | | | | | 9100 | | | | | 9105 | | | |
| Thr | Ser | Val | Pro | Asn | Ser | Val | Asn | Tyr | Ile | Asn | Ala | Asp | Lys | Asn |
| 9110 | | | | | | 9115 | | | | | 9120 | | | |
| Leu | Gln | Ile | Glu | Phe | Asp | Glu | Ala | Leu | Gln | Gln | Ala | Ser | Ala | Thr |
| 9125 | | | | | | 9130 | | | | | 9135 | | | |
| Ser | Ser | Lys | Thr | Ser | Glu | Asn | Pro | Ala | Thr | Ile | Glu | Glu | Val | Leu |
| 9140 | | | | | | 9145 | | | | | 9150 | | | |
| Gly | Leu | Ser | Gln | Ala | Ile | Tyr | Asp | Thr | Lys | Asn | Ala | Leu | Asn | Gly |
| 9155 | | | | | | 9160 | | | | | 9165 | | | |
| Glu | Gln | Arg | Leu | Ala | Thr | Glu | Lys | Ser | Lys | Asp | Leu | Lys | Leu | Ile |
| 9170 | | | | | | 9175 | | | | | 9180 | | | |
| Lys | Gly | Leu | Lys | Asp | Leu | Asn | Lys | Ala | Gln | Leu | Glu | Asp | Val | Thr |
| 9185 | | | | | | 9190 | | | | | 9195 | | | |
| Asn | Lys | Val | Asn | Ser | Ala | Asn | Thr | Leu | Thr | Glu | Leu | Ser | Gln | Leu |
| 9200 | | | | | | 9205 | | | | | 9210 | | | |
| Thr | Gln | Ser | Thr | Leu | Glu | Leu | Asn | Asp | Lys | Met | Lys | Leu | Leu | Arg |
| 9215 | | | | | | 9220 | | | | | 9225 | | | |
| Asp | Lys | Leu | Lys | Thr | Leu | Val | Asn | Pro | Val | Lys | Ala | Ser | Leu | Asn |
| 9230 | | | | | | 9235 | | | | | 9240 | | | |
| Tyr | Arg | Asn | Ala | Asp | Tyr | Asn | Leu | Lys | Arg | Gln | Phe | Asn | Lys | Ala |
| 9245 | | | | | | 9250 | | | | | 9255 | | | |
| Leu | Lys | Glu | Ala | Lys | Gly | Val | Leu | Asn | Lys | Asn | Ser | Gly | Thr | Asn |
| 9260 | | | | | | 9265 | | | | | 9270 | | | |
| Val | Asn | Ile | Asn | Asp | Ile | Gln | His | Leu | Leu | Thr | Gln | Ile | Asp | Asn |
| 9275 | | | | | | 9280 | | | | | 9285 | | | |
| Ala | Lys | Asp | Gln | Leu | Asn | Gly | Glu | Arg | Arg | Leu | Lys | Glu | His | Gln |
| 9290 | | | | | | 9295 | | | | | 9300 | | | |
| Gln | Lys | Ser | Glu | Val | Phe | Ile | Ile | Lys | Glu | Leu | Asp | Ile | Leu | Asn |
| 9305 | | | | | | 9310 | | | | | 9315 | | | |

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|------|-----|-----|-----|-----|-----|------|-----|-----|-----|-----|------|-----|-----|-----|
| Asn | Ala | Gln | Lys | Ala | Ala | Ile | Ile | Asn | Gln | Ile | Arg | Ala | Ser | Lys |
| 9320 | | | | | | 9325 | | | | | 9330 | | | |
| Asp | Ile | Lys | Ile | Ile | Asn | Gln | Ile | Val | Asp | Asn | Ala | Ile | Glu | Leu |
| 9335 | | | | | | 9340 | | | | | 9345 | | | |
| Asn | Asp | Ala | Met | Gln | Gly | Leu | Lys | Glu | His | Val | Ala | Gln | Leu | Thr |
| 9350 | | | | | | 9355 | | | | | 9360 | | | |
| Ala | Thr | Thr | Lys | Asp | Asn | Ile | Glu | Tyr | Leu | Asn | Ala | Asp | Glu | Asp |
| 9365 | | | | | | 9370 | | | | | 9375 | | | |
| His | Lys | Leu | Gln | Tyr | Asp | Tyr | Ala | Ile | Asn | Leu | Ala | Asn | Asn | Val |
| 9380 | | | | | | 9385 | | | | | 9390 | | | |
| Leu | Asp | Lys | Glu | Asn | Gly | Thr | Asn | Lys | Asp | Ala | Asn | Ile | Ile | Ile |
| 9395 | | | | | | 9400 | | | | | 9405 | | | |
| Gly | Met | Ile | Gln | Asn | Met | Asp | Asp | Ala | Arg | Ala | Leu | Leu | Asn | Gly |
| 9410 | | | | | | 9415 | | | | | 9420 | | | |
| Ile | Glu | Arg | Leu | Lys | Asp | Ala | Gln | Thr | Lys | Ala | His | Asn | Asp | Ile |
| 9425 | | | | | | 9430 | | | | | 9435 | | | |
| Lys | Asp | Thr | Leu | Lys | Arg | Gln | Leu | Asp | Glu | Ile | Glu | His | Ala | Asn |
| 9440 | | | | | | 9445 | | | | | 9450 | | | |
| Ala | Thr | Ser | Asn | Ser | Lys | Ala | Gln | Ala | Lys | Gln | Met | Val | Asn | Glu |
| 9455 | | | | | | 9460 | | | | | 9465 | | | |
| Glu | Ala | Arg | Lys | Ala | Leu | Ser | Asn | Ile | Asn | Asp | Ala | Thr | Ser | Asn |
| 9470 | | | | | | 9475 | | | | | 9480 | | | |
| Asp | Leu | Val | Asn | Gln | Ala | Lys | Asp | Glu | Gly | Gln | Ser | Ala | Ile | Glu |
| 9485 | | | | | | 9490 | | | | | 9495 | | | |
| His | Ile | His | Ala | Asp | Glu | Leu | Pro | Lys | Ala | Lys | Leu | Asp | Ala | Asn |
| 9500 | | | | | | 9505 | | | | | 9510 | | | |
| Gln | Met | Ile | Asp | Gln | Lys | Val | Glu | Asp | Ile | Asn | His | Leu | Ile | Ser |
| 9515 | | | | | | 9520 | | | | | 9525 | | | |
| Gln | Asn | Pro | Asn | Leu | Ser | Asn | Glu | Glu | Lys | Asn | Lys | Leu | Ile | Ser |
| 9530 | | | | | | 9535 | | | | | 9540 | | | |
| Gln | Ile | Asn | Lys | Leu | Val | Asn | Gly | Ile | Lys | Asn | Glu | Ile | Gln | Gln |
| 9545 | | | | | | 9550 | | | | | 9555 | | | |
| Ala | Ile | Asn | Lys | Gln | Gln | Ile | Glu | Asn | Ala | Thr | Thr | Lys | Leu | Asp |
| 9560 | | | | | | 9565 | | | | | 9570 | | | |
| Glu | Val | Ile | Glu | Thr | Thr | Lys | Lys | Leu | Ile | Ile | Ala | Lys | Ala | Glu |
| 9575 | | | | | | 9580 | | | | | 9585 | | | |
| Ala | Lys | Gln | Met | Ile | Lys | Glu | Leu | Ser | Gln | Lys | Lys | Arg | Asp | Ala |
| 9590 | | | | | | 9595 | | | | | 9600 | | | |
| Ile | Asn | Asn | Asn | Thr | Asp | Leu | Thr | Pro | Ser | Gln | Lys | Ala | His | Ala |
| 9605 | | | | | | 9610 | | | | | 9615 | | | |
| Leu | Ala | Asp | Ile | Asp | Lys | Thr | Glu | Lys | Asp | Ala | Leu | Gln | His | Ile |
| 9620 | | | | | | 9625 | | | | | 9630 | | | |
| Glu | Asn | Ser | Asn | Ser | Ile | Asp | Asp | Ile | Asn | Asn | Asn | Lys | Glu | His |
| 9635 | | | | | | 9640 | | | | | 9645 | | | |
| Ala | Phe | Asn | Thr | Leu | Ala | His | Ile | Ile | Ile | Trp | Asp | Thr | Asp | Gln |
| 9650 | | | | | | 9655 | | | | | 9660 | | | |
| Gln | Pro | Leu | Val | Phe | Glu | Leu | Pro | Glu | Leu | Ser | Leu | Gln | Asn | Ala |
| 9665 | | | | | | 9670 | | | | | 9675 | | | |
| Leu | Val | Thr | Ser | Glu | Val | Val | Val | His | Arg | Asp | Glu | Thr | Ile | Ser |
| 9680 | | | | | | 9685 | | | | | 9690 | | | |
| Leu | Glu | Ser | Ile | Ile | Gly | Ala | Met | Thr | Leu | Thr | Asp | Glu | Leu | Lys |
| 9695 | | | | | | 9700 | | | | | 9705 | | | |
| Val | Asn | Ile | Val | Ser | Leu | Pro | Asn | Thr | Asp | Lys | Val | Ala | Asp | His |

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| | | |
|--------------------------------------|------------------------------|----------------------|
| 9710 | 9715 | 9720 |
| Leu Thr Ala Lys Val Lys Val 9725 | Ile Leu Ala Asp Gly 9730 | Ser Tyr Val 9735 |
| Thr Val Asn Val Pro Val Lys 9740 | Val Val Glu Lys Glu 9745 | Leu Gln Ile 9750 |
| Ala Lys Lys Asp Ala Ile Lys 9755 | Thr Ile Asp Val Leu 9760 | Val Lys Gln 9765 |
| Lys Ile Lys Asp Ile Asp Ser 9770 | Asn Asn Glu Leu Thr 9775 | Ser Thr Gln 9780 |
| Arg Glu Asp Ala Lys Ala Glu 9785 | Ile Glu Arg Leu Lys 9790 | Lys Gln Ala 9795 |
| Ile Asp Lys Val Asn His Ser 9800 | Lys Ser Ile Lys Asp 9805 | Ile Glu Thr 9810 |
| Val Lys Arg Thr Asp Phe Glu 9815 | Glu Ile Asp Gln Phe 9820 | Asp Pro Lys 9825 |
| Arg Phe Thr Leu Asn Lys Ala 9830 | Lys Lys Asp Ile Ile 9835 | Thr Asp Val 9840 |
| Asn Thr Gln Ile Gln Asn Gly 9845 | Phe Lys Glu Ile Glu 9850 | Thr Ile Lys 9855 |
| Gly Leu Thr Ser Asn Glu Lys 9860 | Thr Gln Phe Asp Lys 9865 | Gln Leu Thr 9870 |
| Ala Leu Gln Lys Glu Phe Leu 9875 | Glu Lys Val Glu His 9880 | Ala His Asn 9885 |
| Leu Val Glu Leu Asn Gln Leu 9890 | Gln Gln Glu Phe Asn 9895 | Asn Arg Tyr 9900 |
| Lys His Ile Leu Asn Gln Ala 9905 | His Leu Leu Gly Glu 9910 | Lys His Ile 9915 |
| Ala Glu His Lys Leu Gly Tyr 9920 | Val Val Val Asn Lys 9925 | Thr Gln Gln 9930 |
| Ile Leu Asn Asn Gln Ser Ala 9935 | Ser Tyr Phe Ile Lys 9940 | Gln Trp Ala 9945 |
| Leu Asp Arg Ile Lys Gln Ile 9950 | Gln Leu Glu Thr Met 9955 | Asn Ser Ile 9960 |
| Arg Gly Ala His Thr Val Gln 9965 | Asp Val His Lys Ala 9970 | Leu Leu Gln 9975 |
| Gly Ile Glu Gln Ile Leu Lys 9980 | Val Asn Val Ser Ile 9985 | Ile Asn Gln 9990 |
| Ser Phe Asn Asp Ser Leu His 9995 | Asn Phe Asn Tyr Leu 10000 | His Ser Lys 10005 |
| Phe Asp Ala Arg Leu Arg Glu 10010 | Lys Asp Val Ala Asn 10015 | His Ile Val 10020 |
| Gln Thr Glu Thr Phe Lys Glu 10025 | Val Leu Lys Gly Thr 10030 | Gly Val Glu 10035 |
| Pro Gly Lys Ile Asn Lys Glu 10040 | Thr Gln Gln Pro Lys 10045 | Leu His Lys 10050 |
| Asn Asp Asn Asp Ser Leu Phe 10055 | Lys His Leu Val Asp 10060 | Asn Phe Gly 10065 |
| Lys Thr Val Gly Val Ile Thr 10070 | Leu Thr Gly Leu Leu 10075 | Ser Ser Phe 10080 |
| Trp Leu Val Leu Ala Lys Arg 10085 | Arg Lys Lys Glu Glu 10090 | Glu Glu Lys 10095 |
| Gln Ser Ile Lys Asn His His 10100 | Lys Asp Ile Arg Leu 10105 | Ser Asp Thr 10110 |

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Asp Lys Ile Asp Pro Ile Val Ile Thr Lys Arg Lys Ile Asp Lys
 10115 10120 10125
 Glu Glu Gln Ile Gln Asn Asp Asp Lys His Ser Ile Pro Val Ala
 10130 10135 10140
 Lys His Lys Lys Ser Lys Glu Lys Gln Leu Ser Glu Glu Asp Ile
 10145 10150 10155
 His Ser Ile Pro Val Val Lys Arg Lys Gln Asn Ser Asp Asn Lys
 10160 10165 10170
 Asp Thr Lys Gln Lys Lys Val Thr Ser Lys Lys Lys Lys Thr Pro
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 Gln Ser Thr Lys Lys Val Val Lys Thr Lys Lys Arg Ser Lys Lys
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<210> SEQ ID NO 24

<211> LENGTH: 1973

<212> TYPE: PRT

<213> ORGANISM: Staphylococcus epidermidis

<400> SEQUENCE: 24

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

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| 275 | | | | | 280 | | | | | 285 | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Arg | Thr | Thr | Asp | Gly | Trp | Asn | Arg | Asn | Val | Ile | Asp | Ser | Lys | Val | Ala |
| 290 | | | | | | 295 | | | | | 300 | | | | |
| Gly | Ile | Tyr | Gln | Asp | Ile | Asp | Val | Val | Pro | Gly | Ser | Glu | Leu | Asn | Val |
| 305 | | | | 310 | | | | | | 315 | | | | | 320 |
| Asn | Phe | Ile | Ser | Thr | Ser | Pro | Val | Phe | Ser | Asp | Gly | Ala | Ala | Gly | Ala |
| | | | 325 | | | | | | 330 | | | | | 335 | |
| Lys | Leu | Lys | Ile | Ser | Asn | Val | Glu | Gln | Asn | Arg | Val | Leu | Phe | Asp | Ser |
| | | | 340 | | | | | 345 | | | | | 350 | | |
| Arg | Leu | Asn | Gly | Met | Gly | Pro | Tyr | Pro | Thr | Gly | Lys | Leu | Ser | Ala | Met |
| | | 355 | | | | | 360 | | | | | 365 | | | |
| Val | Asn | Ile | Pro | Asn | Asp | Ile | Asn | Arg | Val | Arg | Ile | Ser | Phe | Leu | Pro |
| | 370 | | | | | 375 | | | | | 380 | | | | |
| Val | Ser | Ser | Thr | Gly | Arg | Val | Ser | Val | Gln | Arg | Ser | Ser | Arg | Glu | His |
| 385 | | | | 390 | | | | | | 395 | | | | | 400 |
| Gly | Phe | Gly | Asp | Asn | Ser | Ser | Tyr | Tyr | His | Gly | Gly | Ser | Val | Ser | Asp |
| | | | | 405 | | | | | 410 | | | | | 415 | |
| Val | Arg | Ile | Asn | Ser | Gly | Ser | Tyr | Val | Val | Ser | Lys | Val | Thr | Gln | Arg |
| | | | 420 | | | | | 425 | | | | | 430 | | |
| Glu | Tyr | Thr | Thr | Arg | Pro | Asn | Ser | Ser | Asn | Asp | Thr | Phe | Ala | Arg | Ala |
| | | | 435 | | | | 440 | | | | | 445 | | | |
| Thr | Ile | Asn | Leu | Ser | Val | Glu | Asn | Lys | Gly | His | Asn | Gln | Ser | Lys | Asp |
| | 450 | | | | | 455 | | | | | 460 | | | | |
| Thr | Tyr | Tyr | Glu | Val | Ile | Leu | Pro | Gln | Asn | Ser | Arg | Leu | Ile | Ser | Thr |
| 465 | | | | | 470 | | | | | 475 | | | | | 480 |
| Arg | Gly | Gly | Ser | Gly | Asn | Tyr | Asn | Asn | Ala | Thr | Asn | Lys | Leu | Ser | Ile |
| | | | | 485 | | | | | 490 | | | | | 495 | |
| Arg | Leu | Asp | Asn | Leu | Asn | Pro | Gly | Asp | Arg | Arg | Asp | Ile | Ser | Tyr | Thr |
| | | | 500 | | | | | 505 | | | | | 510 | | |
| Val | Asp | Phe | Glu | Ser | Ser | Ser | Pro | Lys | Leu | Ile | Asn | Leu | Asn | Ala | His |
| | | 515 | | | | | 520 | | | | | 525 | | | |
| Leu | Leu | Tyr | Lys | Thr | Asn | Ala | Thr | Phe | Arg | Gly | Asn | Asp | Gly | Gln | Arg |
| | 530 | | | | 535 | | | | | | 540 | | | | |
| Thr | Gly | Asp | Asn | Ile | Val | Asp | Leu | Gln | Ser | Ile | Ala | Leu | Leu | Met | Asn |
| 545 | | | | | 550 | | | | | 555 | | | | | 560 |
| Lys | Asp | Val | Leu | Glu | Thr | Glu | Leu | Asn | Glu | Ile | Asp | Lys | Phe | Ile | Arg |
| | | | | 565 | | | | | 570 | | | | | 575 | |
| Asp | Leu | Asn | Glu | Ala | Asp | Phe | Thr | Ile | Asp | Ser | Trp | Ser | Ala | Leu | Gln |
| | | | 580 | | | | | 585 | | | | | 590 | | |
| Glu | Lys | Met | Thr | Glu | Gly | Gly | Asn | Ile | Leu | Asn | Glu | Gln | Gln | Asn | Gln |
| | | 595 | | | | | 600 | | | | | 605 | | | |
| Val | Ala | Leu | Glu | Asn | Gln | Ala | Ser | Gln | Glu | Thr | Ile | Asn | Asn | Val | Thr |
| | 610 | | | | | 615 | | | | | 620 | | | | |
| Gln | Ser | Leu | Glu | Ile | Leu | Lys | Asn | Asn | Leu | Lys | Tyr | Lys | Thr | Pro | Ser |
| 625 | | | | | 630 | | | | | 635 | | | | | 640 |
| Gln | Pro | Ile | Ile | Lys | Ser | Asn | Asn | Gln | Ile | Pro | Asn | Ile | Thr | Ile | Ser |
| | | | | 645 | | | | | 650 | | | | | 655 | |
| Pro | Ala | Asp | Lys | Ala | Asp | Lys | Leu | Thr | Ile | Thr | Tyr | Gln | Asn | Thr | Asp |
| | | | 660 | | | | | 665 | | | | | 670 | | |
| Asn | Glu | Ser | Ala | Ser | Ile | Ile | Gly | Asn | Lys | Leu | Asn | Asn | Gln | Trp | Ser |
| | | | 675 | | | | 680 | | | | | 685 | | | |
| Leu | Asn | Asn | Asn | Ile | Pro | Gly | Ile | Glu | Ile | Asp | Met | Gln | Thr | Gly | Leu |
| | 690 | | | | | 695 | | | | | 700 | | | | |

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Val Thr Ile Asp Tyr Lys Ala Val Tyr Pro Glu Ser Val Val Gly Ala
 705 710 715 720
 Asn Asp Lys Thr Gly Asn Ser Asp Ala Ser Glu Ser Arg Ile Thr
 725 730 735
 Met Pro Arg Lys Glu Ala Thr Pro Leu Ser Pro Ile Val Glu Ala Asn
 740 745 750
 Glu Glu Arg Val Asn Val Val Ile Ala Pro Asn Gly Glu Ala Thr Gln
 755 760 765
 Ile Ala Ile Lys Tyr Arg Thr Pro Asp Gly Gln Glu Ala Thr Leu Val
 770 775 780
 Ala Ser Lys Asn Gly Ser Ser Trp Thr Leu Asn Lys Gln Ile Asp Tyr
 785 790 795 800
 Val Asn Ile Glu Glu Asn Ser Gly Lys Val Thr Ile Gly Tyr Gln Ala
 805 810 815
 Val Gln Pro Glu Ser Glu Val Ile Ala Thr Glu Thr Lys Gly Asn Ser
 820 825 830
 Asp Glu Ser Ala Glu Ser Arg Val Thr Met Pro Arg Lys Glu Ala Thr
 835 840 845
 Pro His Ser Pro Ile Val Glu Ala Asn Glu Glu His Val Asn Val Thr
 850 855 860
 Ile Ala Pro Asn Gly Glu Ala Thr Gln Ile Ala Ile Lys Tyr Arg Thr
 865 870 875 880
 Pro Asp Gly Gln Glu Thr Thr Leu Ile Ala Ser Lys Asn Gly Ser Ser
 885 890 895
 Trp Thr Leu Asn Lys Gln Ile Asp Tyr Val Asn Ile Glu Glu Asn Ser
 900 905 910
 Gly Lys Val Thr Ile Gly Tyr Gln Ala Val Gln Leu Glu Ser Glu Val
 915 920 925
 Ile Ala Thr Glu Thr Lys Gly Asn Ser Asp Ala Ser Ala Glu Ser Arg
 930 935 940
 Ile Thr Met Leu Arg Lys Glu Ala Thr Pro His Ser Pro Ile Val Glu
 945 950 955 960
 Ala Asn Glu Glu His Val Asn Val Thr Ile Ala Pro Asn Gly Glu Ala
 965 970 975
 Thr Gln Ile Ala Ile Lys Tyr Arg Thr Pro Asp Gly Gln Glu Ala Thr
 980 985 990
 Leu Val Ala Ser Lys Asn Glu Ser Ser Trp Thr Leu Asn Lys Gln Ile
 995 1000 1005
 Asp His Val Asn Ile Asp Glu Asn Ser Gly Lys Val Thr Ile Gly
 1010 1015 1020
 Tyr Gln Ala Val Gln Pro Glu Ser Glu Ile Ile Ala Thr Glu Thr
 1025 1030 1035
 Lys Gly Asn Ser Asp Ala Ser Ala Glu Ser Arg Ile Thr Met Pro
 1040 1045 1050
 Arg Lys Glu Ala Thr Pro Ile Pro Pro Thr Leu Glu Ala Ser Val
 1055 1060 1065
 Gln Glu Ala Ser Val Thr Val Thr Pro Asn Glu Asn Ala Thr Lys
 1070 1075 1080
 Val Phe Ile Lys Tyr Leu Asp Ile Asn Asp Glu Ile Ser Thr Ile
 1085 1090 1095
 Ile Ala Ser Lys Ile Asn Gln Gln Trp Thr Leu Asn Lys Asp Asn
 1100 1105 1110

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| | | | | | | | | | |
|---------|---------|---------|------|---------|---------|------|---------|-----|--|
| Phe Gly | Ile Lys | Ile Asn | Pro | Leu Thr | Gly Lys | Val | Ile Ile | Ser | |
| 1115 | | | 1120 | | | 1125 | | | |
| Tyr Val | Ala Val | Gln Pro | Glu | Ser Asp | Val Ile | Ala | Ile Glu | Ser | |
| 1130 | | | 1135 | | | 1140 | | | |
| Gln Gly | Asn Ser | Asp Leu | Ser | Glu Glu | Ser Arg | Ile | Ile Met | Pro | |
| 1145 | | | 1150 | | | 1155 | | | |
| Thr Lys | Glu Glu | Pro Pro | Glu | Pro Pro | Ile Leu | Glu | Ser Asp | Ser | |
| 1160 | | | 1165 | | | 1170 | | | |
| Ile Glu | Ala Lys | Val Asn | Ile | Phe Pro | Asn Asp | Glu | Ala Thr | Arg | |
| 1175 | | | 1180 | | | 1185 | | | |
| Ile Val | Ile Met | Tyr Thr | Ser | Leu Glu | Gly Gln | Glu | Ala Thr | Leu | |
| 1190 | | | 1195 | | | 1200 | | | |
| Val Ala | Ser Lys | Asn Glu | Ser | Ser Trp | Thr Leu | Asn | Lys Gln | Ile | |
| 1205 | | | 1210 | | | 1215 | | | |
| Asp His | Val Asn | Ile Asp | Glu | Asn Ser | Gly Lys | Val | Thr Ile | Gly | |
| 1220 | | | 1225 | | | 1230 | | | |
| Tyr Gln | Ala Val | Gln Pro | Glu | Ser Glu | Val Ile | Ala | Thr Glu | Thr | |
| 1235 | | | 1240 | | | 1245 | | | |
| Lys Gly | Asn Ser | Asp Ala | Ser | Ala Glu | Ser Arg | Val | Thr Met | Pro | |
| 1250 | | | 1255 | | | 1260 | | | |
| Arg Lys | Glu Ala | Thr Pro | His | Ser Pro | Ile Val | Glu | Thr Asn | Glu | |
| 1265 | | | 1270 | | | 1275 | | | |
| Glu Arg | Val Asn | Val Val | Ile | Ala Pro | Asn Gly | Glu | Ala Thr | Gln | |
| 1280 | | | 1285 | | | 1290 | | | |
| Ile Ala | Ile Lys | Tyr Arg | Thr | Pro Asp | Gly Gln | Glu | Thr Thr | Leu | |
| 1295 | | | 1300 | | | 1305 | | | |
| Ile Ala | Ser Lys | Asn Gly | Ser | Ser Trp | Thr Leu | Asn | Lys Gln | Ile | |
| 1310 | | | 1315 | | | 1320 | | | |
| Asp His | Val Asn | Ile Asp | Glu | Asn Ser | Gly Lys | Val | Thr Ile | Gly | |
| 1325 | | | 1330 | | | 1335 | | | |
| Tyr Gln | Ala Val | Gln Pro | Glu | Ser Glu | Ile Ile | Ala | Thr Glu | Thr | |
| 1340 | | | 1345 | | | 1350 | | | |
| Lys Gly | Asn Ser | Asp Ala | Ser | Ala Glu | Ser Arg | Ile | Thr Met | Pro | |
| 1355 | | | 1360 | | | 1365 | | | |
| Arg Lys | Glu Ala | Ile Pro | His | Ser Pro | Ile Val | Glu | Ala Asn | Glu | |
| 1370 | | | 1375 | | | 1380 | | | |
| Glu His | Val Asn | Val Thr | Ile | Ala Pro | Asn Gly | Glu | Thr Thr | Gln | |
| 1385 | | | 1390 | | | 1395 | | | |
| Ile Ala | Val Lys | Tyr Arg | Thr | Pro Asp | Gly Gln | Glu | Ala Thr | Leu | |
| 1400 | | | 1405 | | | 1410 | | | |
| Ile Ala | Ser Lys | Asn Glu | Ser | Ser Trp | Thr Leu | Asn | Lys Gln | Ile | |
| 1415 | | | 1420 | | | 1425 | | | |
| Asp His | Val Asn | Ile Asp | Glu | Asn Ser | Gly Lys | Val | Thr Ile | Gly | |
| 1430 | | | 1435 | | | 1440 | | | |
| Tyr Gln | Ala Val | Gln Pro | Glu | Ser Glu | Val Ile | Ala | Thr Glu | Thr | |
| 1445 | | | 1450 | | | 1455 | | | |
| Lys Gly | Asn Ser | Asp Ala | Ser | Ala Glu | Ser Arg | Ile | Thr Met | Pro | |
| 1460 | | | 1465 | | | 1470 | | | |
| Val Lys | Glu Lys | Thr Pro | Ala | Pro Pro | Ile Ser | Ile | Ile Asn | Glu | |
| 1475 | | | 1480 | | | 1485 | | | |
| Ser Asn | Ala Ser | Val Glu | Ile | Ile Pro | Gln Val | Asn | Val Thr | Gln | |
| 1490 | | | 1495 | | | 1500 | | | |
| Leu Ser | Leu Gln | Tyr Ile | Asp | Ala Lys | Gly Gln | Gln | Gln Asn | Leu | |

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| | | |
|-------------------------------------|-----------------------------|-------------------------|
| 1505 | 1510 | 1515 |
| Ile Ala Thr Leu Asn Gln Asn 1520 | Gln Trp Thr Leu Asn 1525 | Lys Asn Val 1530 |
| Ser His Ile Thr Val Asp Lys 1535 | Asn Thr Gly Lys 1540 | Val Leu Ile Asn 1545 |
| Tyr Gln Ala Val Tyr Pro Glu 1550 | Ser Glu Val Ile 1555 | Ala Arg Glu Ser 1560 |
| Lys Gly Asn Ser Asp Ser Ser 1565 | Asn Val Ser Met 1570 | Val Ile Met Pro 1575 |
| Arg Lys Thr Ala Thr Pro Lys 1580 | Pro Pro Ile Ile 1585 | Lys Val Asp Glu 1590 |
| Met Asn Ala Ser Leu Ala Ile 1595 | Ile Pro Tyr Lys 1600 | Asn Asn Thr Ala 1605 |
| Ile Asn Ile His Tyr Ile Asp 1610 | Lys Lys Gly Ile 1615 | Lys Ser Met Val 1620 |
| Thr Ala Ile Lys Asn Asn Asp 1625 | Gln Trp Gln Leu 1630 | Asp Glu Lys Ile 1635 |
| Lys Tyr Val Lys Ile Asp Ala 1640 | Lys Thr Gly Thr 1645 | Val Ile Ile Asn 1650 |
| Tyr Gln Ile Val Gln Glu Asn 1655 | Ser Glu Ile Ile 1660 | Ala Thr Ala Ile 1665 |
| Asn Gly Asn Ser Asp Lys Ser 1670 | Glu Glu Val Lys 1675 | Val Leu Met Pro 1680 |
| Ile Lys Glu Phe Thr Pro Leu 1685 | Ala Pro Leu Leu 1690 | Glu Thr Asn Tyr 1695 |
| Lys Lys Ala Thr Val Ser Ile 1700 | Leu Pro Gln Ser 1705 | Asn Ala Thr Lys 1710 |
| Leu Asp Phe Lys Tyr Arg Asp 1715 | Lys Lys Gly Asp 1720 | Ser Lys Ile Ile 1725 |
| Ile Val Lys Arg Phe Lys Asn 1730 | Ile Trp Lys Ala 1735 | Asn Glu Gln Ile 1740 |
| Ser Gly Val Thr Ile Asn Pro 1745 | Glu Phe Gly Gln 1750 | Val Val Ile Asn 1755 |
| Tyr Gln Ala Val Tyr Pro Glu 1760 | Ser Asp Ile Leu 1765 | Ala Ala Gln Tyr 1770 |
| Val Gly Asn Ser Asp Ala Ser 1775 | Glu Trp Ala Lys 1780 | Val Lys Met Pro 1785 |
| Lys Lys Glu Leu Ala Pro His 1790 | Ser Pro Ser Leu 1795 | Ile Tyr Asp Asn 1800 |
| Arg Asn Asn Lys Ile Leu Ile 1805 | Ala Pro Asn Ser 1810 | Asn Ala Thr Glu 1815 |
| Met Glu Leu Ser Tyr Val Asp 1820 | Lys Asn Asn Gln 1825 | Ser Leu Lys Val 1830 |
| Lys Ala Leu Lys Ile Asn Asn 1835 | Arg Trp Lys Phe 1840 | Asp Ser Ser Val 1845 |
| Ser Asn Ile Ser Ile Asn Pro 1850 | Asn Thr Gly Lys 1855 | Ile Val Leu Gln 1860 |
| Pro Gln Phe Leu Leu Thr Asn 1865 | Ser Lys Ile Ile 1870 | Val Phe Ala Lys 1875 |
| Lys Gly Asn Ser Asp Ala Ser 1880 | Ile Ser Val Ser 1885 | Leu Arg Val Pro 1890 |
| Ala Val Lys Lys Ile Glu Leu 1895 | Glu Pro Met Phe 1900 | Asn Val Pro Val 1905 |

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Leu Val Ser Leu Asn Lys Lys Arg Ile Gln Phe Asp Asp Cys Ser
1910 1915 1920

Gly Val Lys Asn Cys Leu Asn Lys Gln Ile Ser Lys Thr Gln Leu
1925 1930 1935

Pro Asp Thr Gly Tyr Ser Asp Lys Ala Ser Lys Ser Asn Ile Leu
1940 1945 1950

Ser Val Leu Leu Leu Gly Phe Gly Phe Leu Ser Tyr Ser Arg Lys
1955 1960 1965

Arg Lys Glu Lys Gln
1970

<210> SEQ ID NO 25

<211> LENGTH: 10

<212> TYPE: PRT

<213> ORGANISM: Enterococcus faecalis

<220> FEATURE:

<221> NAME/KEY: misc_feature

<222> LOCATION: (3)..(3)

<223> OTHER INFORMATION: Xaa can be any naturally occurring amino acid

<400> SEQUENCE: 25

Leu Pro Xaa Thr Ser Ala Gly Ala Asn Ser
1 5 10

<210> SEQ ID NO 26

<211> LENGTH: 30

<212> TYPE: DNA

<213> ORGANISM: Enterococcus faecalis

<400> SEQUENCE: 26

ccgcatgccca agagcaaaca gcaaaagaag 30

<210> SEQ ID NO 27

<211> LENGTH: 33

<212> TYPE: DNA

<213> ORGANISM: Enterococcus faecalis

<400> SEQUENCE: 27

ccgtcgactt aagtaccaga agtgggtggtt ttc 33

<210> SEQ ID NO 28

<211> LENGTH: 30

<212> TYPE: DNA

<213> ORGANISM: Enterococcus faecalis

<400> SEQUENCE: 28

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<212> TYPE: DNA

<213> ORGANISM: Enterococcus faecalis

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<210> SEQ ID NO 30

<211> LENGTH: 33

<212> TYPE: DNA

<213> ORGANISM: Enterococcus faecalis

<400> SEQUENCE: 30

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| ccggatccgc agctaataaa gaagaatttt tag | 33 |
| <210> SEQ ID NO 31 <211> LENGTH: 33 <212> TYPE: DNA <213> ORGANISM: Enterococcus faecalis | |
| <400> SEQUENCE: 31 | |
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| <210> SEQ ID NO 32 <211> LENGTH: 28 <212> TYPE: DNA <213> ORGANISM: Enterococcus faecalis | |
| <400> SEQUENCE: 32 | |
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| <400> SEQUENCE: 33 | |
| ccctgcagtt acccaccaaa tgtgataacc c | 31 |
| <210> SEQ ID NO 34 <211> LENGTH: 33 <212> TYPE: DNA <213> ORGANISM: Enterococcus faecalis | |
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| ccggatccga agaaataact gatttatfff tac | 33 |
| <210> SEQ ID NO 35 <211> LENGTH: 36 <212> TYPE: DNA <213> ORGANISM: Enterococcus faecalis | |
| <400> SEQUENCE: 35 | |
| ccgagctctt attgttctcg aattaatttt tctaac | 36 |
| <210> SEQ ID NO 36 <211> LENGTH: 27 <212> TYPE: DNA <213> ORGANISM: Enterococcus faecalis | |
| <400> SEQUENCE: 36 | |
| ccgcatgctc gcaagcaagc gttcaag | 27 |
| <210> SEQ ID NO 37 <211> LENGTH: 33 <212> TYPE: DNA <213> ORGANISM: Enterococcus faecalis | |
| <400> SEQUENCE: 37 | |
| ccctgcagtt agaagcctga ctcttttact ttt | 33 |
| <210> SEQ ID NO 38 <211> LENGTH: 30 <212> TYPE: DNA <213> ORGANISM: Enterococcus faecalis | |
| <400> SEQUENCE: 38 | |

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ccggatccca agaagtaaca agtgatgctg 30

<210> SEQ ID NO 39
 <211> LENGTH: 32
 <212> TYPE: DNA
 <213> ORGANISM: Enterococcus faecalis

<400> SEQUENCE: 39

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<400> SEQUENCE: 40

ccggatccga aacaggatat gcgcaaac 28

<210> SEQ ID NO 41
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ccgagctctt attccttatt acgaatcgcc tg 32

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<400> SEQUENCE: 42

gcgggatccg aagaaaatgg ggagagcgc 29

<210> SEQ ID NO 43
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<400> SEQUENCE: 43

gcggagctct taggtacctt tgtgtttggt tgg 33

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<400> SEQUENCE: 44

gaattgagca aaagttcaat cg 22

<210> SEQ ID NO 45
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<400> SEQUENCE: 45

caagtaaaaa agccggtaca gc 22

<210> SEQ ID NO 46
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<400> SEQUENCE: 46
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<210> SEQ ID NO 47
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<400> SEQUENCE: 47
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<210> SEQ ID NO 48
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<400> SEQUENCE: 49
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 <212> TYPE: DNA
 <213> ORGANISM: Enterococcus faecalis

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<400> SEQUENCE: 51
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<210> SEQ ID NO 52
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<400> SEQUENCE: 52
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<210> SEQ ID NO 53
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<400> SEQUENCE: 53
 aagcctgact cttttacttt tttattg 27

<210> SEQ ID NO 54
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<400> SEQUENCE: 54

ggtaaccttg tgtttgttg gtac 24

<210> SEQ ID NO 55

<211> LENGTH: 33

<212> TYPE: DNA

<213> ORGANISM: Enterococcus faecalis

<400> SEQUENCE: 55

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<210> SEQ ID NO 56

<211> LENGTH: 35

<212> TYPE: DNA

<213> ORGANISM: Enterococcus faecalis

<400> SEQUENCE: 56

tctgcagttc aattgactac tttcaatata ctgtc 35

<210> SEQ ID NO 57

<211> LENGTH: 36

<212> TYPE: DNA

<213> ORGANISM: Enterococcus faecalis

<400> SEQUENCE: 57

cccaagcttt cagaatgctt gaccttgatt attgta 36

What is claimed is:

1. A method of raising an immune response in a human or animal patient comprising administering to the human or animal patient an effective amount of an isolated protein comprising the amino acid sequence of SEQ ID NO: 9.

2. A method of eliciting an immune response in a human or animal comprising administering to said human or animal an immunologically effective amount of an isolated protein comprising the amino acid sequence of SEQ ID NO: 9.

3. A method of raising an immune response in a human or animal patient comprising administering to the human or

animal patient an effective amount of an isolated peptide comprising the amino acid sequence of amino acids 63-1067 of SEQ ID NO: 9.

4. A method of eliciting an immune response in a human or animal comprising administering to said human or animal an immunologically effective amount of an isolated peptide comprising the amino acid sequence of amino acids 63-1067 of SEQ ID NO: 9.

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