



(86) Date de dépôt PCT/PCT Filing Date: 2008/01/10  
 (87) Date publication PCT/PCT Publication Date: 2008/07/17  
 (85) Entrée phase nationale/National Entry: 2009/07/13  
 (86) N° demande PCT/PCT Application No.: EP 2008/050227  
 (87) N° publication PCT/PCT Publication No.: 2008/084072  
 (30) Priorité/Priority: 2007/01/12 (EP07000602.8)

(51) Cl.Int./Int.Cl. *A61K 38/00* (2006.01)  
 (71) Demandeur/Applicant:  
 INTERCELL AG, AT  
 (72) Inventeurs/Inventors:  
 SENN, BEATRICE, AT;  
 NAGY, ESZTER, AT;  
 MEINKE, ANDREAS, AT;  
 VON GABAIN, ALEXANDER, AT;  
 MAIERHOFER, BARBARA, AT;  
 STIERSCHNEIDER, ULRIKE, AT;  
 BERGER, MANFRED, AT;  
 ...  
 (74) Agent: BERESKIN & PARR LLP/S.E.N.C.R.L.,S.R.L.

(54) Titre : PROTEINES PROTECTRICES DE S. AGALACTIA, LEURS COMBINAISONS ET LEURS PROCEDES D'UTILISATION  
 (54) Title: PROTECTIVE PROTEINS OF S. AGALACTIAE, COMBINATIONS THEREOF AND METHODS OF USING THE SAME

(57) **Abrégé/Abstract:**

The invention relates to a composition comprising at least two protective proteins against *Streptococcus agalactiae* (*S. agalactiae*) or functionally active variant thereof; a protective peptide against *S. agalactiae*; one or more nucleic acid(s) encoding the at least two proteins and/or the protective peptide; a method of producing the composition; a pharmaceutical composition, especially a vaccine, comprising the composition and/or at least one protective peptide; methods for producing antibodies; a mixture of antibodies against the at least two proteins of the composition; the use of the composition and/or at least one protective peptide and/or one or more nucleic acid(s) for the manufacture of a medicament for the immunization or treatment of a subject; methods of diagnosing a *S. agalactiae* infection; a method for identifying a ligand capable of binding the composition and/or at least one protective peptide; and the use of the composition and/or at least one protective peptide for the isolation and/or purification and/or identification of an interaction partner of the composition and/or peptide.



(72) Inventeurs(suite)/Inventors(continued): NEUBAUER, CHRISTINA, AT; COHEN, KATHERINE, AT

(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization  
International Bureau(43) International Publication Date  
17 July 2008 (17.07.2008)

PCT

(10) International Publication Number  
**WO 2008/084072 A2**(51) International Patent Classification:  
A61K 38/00 (2006.01)Wiener Neustadt (AT). **COHEN, Katherine** [AT/AT];  
Hermannngasse 6/4/44, A-1070 Vienna (AT).

(21) International Application Number:

PCT/EP2008/050227

(74) Agent: **COHEN, Katherine**; Intercell AG, Corporate De-  
velopment & Intellectual Property, Campus Vienna Biocen-  
ter 6, A-1030 Vienna (AT).

(22) International Filing Date: 10 January 2008 (10.01.2008)

(81) Designated States (*unless otherwise indicated, for every  
kind of national protection available*): AE, AG, AL, AM,  
AO, AT, AU, AZ, BA, BB, BG, BH, BR, BW, BY, BZ, CA,  
CH, CN, CO, CR, CU, CZ, DE, DK, DM, DO, DZ, EC, EE,  
EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID,  
IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC,  
LK, LR, LS, LT, LU, LY, MA, MD, ME, MG, MK, MN,  
MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH,  
PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV,  
SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN,  
ZA, ZM, ZW.

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

07000602.8 12 January 2007 (12.01.2007) EP

(71) Applicant (*for all designated States except US*): **INTER-  
CELL AG** [AT/AT]; Campus Vienna Biocenter 6, A-1030  
Vienna (AT).

(72) Inventors; and

(75) Inventors/Applicants (*for US only*): **SENN, Beatrice**  
[CH/AT]; Gumpendorferstrasse 135/41, A-1060 Vienna  
(AT). **NAGY, Eszter** [HU/AT]; Klimschgasse 30/II/21,  
A-1030 Vienna (AT). **MEINKE, Andreas** [DE/AT]; Piet-  
teggasse 26/1, A-3013 Pressbaum (AT). **VON GABAIN,  
Alexander** [AT/AT]; Hockegasse 77, A-1180 Vienna  
(AT). **MAIERHOFER, Barbara** [AT/AT]; Bachzeile  
322, A-2272 Niederabsdorf (AT). **STIERSCHNEIDER,  
Ulrike** [AT/AT]; Kaiserstrasse 66/2/14, A-1070 Vienna  
(AT). **BERGER, Manfred** [AT/AT]; Francesco Solimena  
Weg 18, A-2700 Wiener Neustadt (AT). **NEUBAUER,  
Christina** [AT/AT]; Ausstellungsgasse 6/3/11, A-2700(84) Designated States (*unless otherwise indicated, for every  
kind of regional protection available*): ARIPO (BW, GH,  
GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM,  
ZW), Eurasian (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM),  
European (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI,  
FR, GB, GR, HR, HU, IE, IS, IT, LT, LU, LV, MC, MT, NL,  
NO, PL, PT, RO, SE, SI, SK, TR), OAPI (BF, BJ, CF, CG,  
CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

Published:

- without international search report and to be republished  
upon receipt of that report
- with sequence listing part of description published sepa-  
rately in electronic form and available upon request from  
the International Bureau

(54) Title: PROTECTIVE PROTEINS OF S. AGALACTIAE, COMBINATIONS THEREOF AND METHODS OF USING THE  
SAME

(57) Abstract: The invention relates to a composition comprising at least two protective proteins against Streptococcus agalactiae (S. agalactiae) or functionally active variant thereof; a protective peptide against S. agalactiae; one or more nucleic acid(s) encoding the at least two proteins and/or the protective peptide; a method of producing the composition; a pharmaceutical composition, especially a vaccine, comprising the composition and/or at least one protective peptide; methods for producing antibodies; a mixture of antibodies against the at least two proteins of the composition; the use of the composition and/or at least one protective peptide and/or one or more nucleic acid(s) for the manufacture of a medicament for the immunization or treatment of a subject; methods of diagnosing a S. agalactiae infection; a method for identifying a ligand capable of binding the composition and/or at least one protective peptide; and the use of the composition and/or at least one protective peptide for the isolation and/or purification and/or identification of an interaction partner of the composition and/or peptide.

WO 2008/084072 A2



**Protective proteins of *S. agalactiae*, combinations thereof  
and methods of using the same**

5

The invention relates to a composition comprising at least two protective proteins against *Streptococcus agalactiae* (*S. agalactiae*) or functionally active variants thereof; a protective peptide against *S. agalactiae*; one or more nucleic acid(s) encoding the at least two proteins and/or the protective peptide; a method of producing the composition; a pharmaceutical composition, especially a vaccine, comprising the composition and/or at least one protective peptide; methods for producing antibodies; a mixture of antibodies against the at least two proteins of the composition; the use of the composition and/or at least one protective peptide and/or one or more nucleic acid(s) for the manufacture of a medicament for the immunization or treatment of a subject; methods of diagnosing a *S. agalactiae* infection; a method for identifying a ligand capable of binding the composition and/or at least one protective peptide; and the use of the composition and/or at least one protective peptide for the isolation and/or purification and/or identification of an interaction partner of the composition and/or peptide.

20 *S. agalactiae* is an encapsulated gram-positive bacterium, which belongs to the Group B Streptococci (GBS) based on its haemolysis pattern on blood agar. Capsules form the basis for classifying GBS into nine distinct serotypes. Most of them have been shown to cause serious diseases, and the two most common serotypes - type III and V - are estimated to account for the majority (~80%) of invasive diseases worldwide. The ranking and serotype prevalence differs by age group and geographic area.

25 *Streptococcus agalactiae* is a frequent cause of infections in neonates, pregnant women and in chronically ill and elderly patients. In newborns Group B Streptococcus even represents the predominant pathogen in the United States causing life threatening diseases, such as sepsis, pneumonia and meningitis. GBS diseases are associated with a high mortality rate (~5%) and a large percentage (~20%) of children surviving GBS infections becomes permanently handicapped with hearing, learning and visual disabilities.

30



- 2 -

Newborns usually acquire the pathogen during delivery from their GBS-colonized mothers. Twenty-five to 40% of pregnant women are colonized with GBS, but are asymptomatic. Due to vertical transmission during birth, 50-70% of neonates born to colonized women - that is approximately 10-25% of all newborns - become colonized by  
5 GBS during delivery which is a prerequisite for infection and disease. In the United States, GBS infections affect 1-5 newborns/1,000 live births. Pre-term infants are at the highest risk for invasive disease due to their immature immune system and the low level of maternal antibody transfer before the 34<sup>th</sup> pregnancy week.

10 GBS disease occurs throughout the world. The highest prevalence of invasive disease in newborns occurs in Western countries, due to the elimination and reduction of other infectious agents and also due to the increased survival of very immature newborns. Before prevention by intrapartum antibiotic treatment was introduced, about 17,000 cases of invasive GBS diseases (sepsis, pneumonia and/or meningitis) were reported in the US  
15 annually. The rates of serious GBS infections are higher among newborns than among any other age group. Nonetheless, serious Group B streptococcal infections occur in other age groups in both men and women. Among non-pregnant adults, rates of serious disease range from 4.1 to 7.2 cases per 100,000 and increase with age. The average death rate for invasive infections is 8-10% for adults between ages 18-64 and 15-25% for adults > 65  
20 years of age. Serious disease is most common among elderly, bedridden patients and people suffering from severe medical conditions including diabetes mellitus, liver disease, history of stroke, history of cancer or bedsores.

Currently, disease management fully relies on antibiotics, mainly Penicillin G. In order to  
25 prevent invasive disease in newborns, pregnant women are screened for carriage of GBS at 35<sup>th</sup> to 37<sup>th</sup> weeks of gestation. Colonized mothers are then treated with high dose antibiotics during delivery to prevent neonatal GBS disease.

Current standard treatment of GBS infections is also based on antibiotics. Route, dosage,  
30 schedule and duration of therapy depend on the severity of the illness. Ten days of treatment is recommended for bacteraemia, pneumonia and soft tissue infections, while 2-3 weeks is recommended for meningitis and 3-4 weeks for osteomyelitis.

- 3 -

Invasive GBS diseases are associated with 5% mortality and 20% permanent damage in spite of effective antibiotic therapy, due to a very rapid and dramatic clinical course. Before prevention direct medical costs of neonatal disease were ~\$300 million annually in the US; and GBS still poses a considerable economic burden.

5

Although intrapartum prophylaxis has decreased the incidence of early-onset GBS disease, currently available strategies are not ideal as they can neither prevent late-onset infections nor disease in premature babies which are at highest risk for invasive disease.

10 Currently, no effective preventive vaccine is available. There are efforts focusing on using capsular polysaccharides (with or without protein-conjugation) as immunogens, but several arguments militate against that approach. Polysaccharides induce IgG2 antibodies, which cross the placenta less efficiently than IgG1 or IgG3 antibodies. This especially poses a problem for the most susceptible early-born neonates, since placental antibody transfer is  
15 low before the 34<sup>th</sup> pregnancy week and about 10% of deliveries occur before that time. An additional disadvantage of polysaccharide vaccines is the incomplete vaccine coverage among GBS serotypes. Given adequate ecological pressure, replacement disease by non-vaccine serotypes remains a real threat, particularly in areas with high disease burden.

20 Taking these insufficiencies into account, new generation immune interventions against GBS disease are needed. Given the very recent acceptance of the use of a cervical cancer-preventing vaccine in teenage girls, a new approach would be the use of combinations of proteins as a prophylactic GBS vaccine in order to provide protection against more than one *S. agalactiae* strain or serotype.

25

Accordingly, one problem underlying the present invention was to provide alternative means for the development of medicaments such as vaccines against *S. agalactiae* infection. More particularly, one problem was to provide combinations of protective proteins, particularly more effective combinations, derived from *S. agalactiae* that can be  
30 used for the manufacture of said medicaments.

Surprisingly, this object has been solved by combinations of protective proteins/peptides comprising or consisting of the amino acid sequences as defined in SEQ ID NOS: 1 to 6 or functionally active variants thereof.



Accordingly, a first subject of the present invention relates to a composition comprising at least two proteins selected from the group consisting of

- 5 i) a protective protein comprising or consisting of the protective peptide of SEQ ID NO: 1 (gbs0233p) or functionally active variant thereof;
- ii) a protective protein comprising or consisting of the protective peptide of SEQ ID NO: 2 (gbs1087p) or functionally active variant thereof;
- iii) a protective protein comprising or consisting of the protective peptide of SEQ ID NO: 3 (gbs1309p) or functionally active variant thereof;
- 10 iv) a protective protein comprising or consisting of the protective peptide of SEQ ID NO: 4 (gbs1477p) or functionally active variant thereof;
- v) a protective protein comprising or consisting of the protective peptide of SEQ ID NO: 5 (gbs1478p) or functionally active variant thereof; and
- 15 vi) a protective protein comprising or consisting of the protective peptide of SEQ ID NO: 6 (gbs2018p) or functionally active variant thereof.

Surprisingly, it was found that combinations of the above protective proteins provide a better protection against *S. agalactiae* than a protective protein when used alone. A better protection in the context of the present invention may refer to a situation in which  
20 protection provided by the combination is improved quantitatively in comparison to the single components of the composition. For example, the combination may provide protection against at least one serotype of *S. agalactiae* against which at least one of the protective proteins present in the composition does not provide protection. Accordingly, the number of serotypes against which the combination provides protection is increased.  
25 Additionally or alternatively, protection provided by the combination is improved qualitatively in comparison to the single components of the composition. For example, the survival of mice challenged with GBS strains may be improved when a composition of protective proteins is used in comparison to the single components of the composition. Both, quantitatively and qualitatively sufficient protection, are important for successful  
30 prevention and/or treatment, since it is the goal striven for to provide protection which is as high as possible and which protects against as many serotypes as possible.

Additionally, combinations of different protective proteins are in general advantageous in comparison to single protective proteins, since in the case of vaccines employing different



- 5 -

protective proteins/antibodies the probability of a serotype switch of the pathogen in question leading to reduced effectiveness of the vaccine is strongly diminished. This is due to the fact that more than one mutation in *S. agalactiae* proteins at defined sites would be required in order to render the respective *S. agalactiae* strain unsusceptible to the vaccine.

5

The protective protein consisting of the amino acid sequence of SEQ ID NO: 1 is derived from *S. agalactiae* strain 12403 and has been denoted by gbs0233p (partial gbs0233) in accordance with the genome of NEM316 (ATCC12403). The DNA sequence encoding the full length protein gbs0233 (consisting of 308 amino acids; SEQ ID NO: 229) from which  
10 the protective protein consisting of the amino acid sequence of the SEQ ID NO: 1 is derived is disclosed at GenBank<sup>®</sup> accession number AL732656 (complete genome of *Streptococcus agalactiae* NEM316) and the amino acid sequence of the full length protein is disclosed in WO2004/099242 (see SEQ ID NO: 475). The amino acid sequence of SEQ ID NO: 1 is disclosed in the Examples as well as in the attached Sequence listing. The  
15 protective protein comprising or consisting of the protective peptide of SEQ ID NO: 1 or a functionally active variant thereof are referred to as (protective) proteins of subgroup i).

The protective protein consisting of the amino acid sequence of SEQ ID NO: 2 is derived from *S. agalactiae* strain 6313 and has been denoted by gbs1087p (partial gbs1087) in  
20 accordance with the genome of NEM316 (ATCC12403). The amino acid and encoding DNA sequences of the full length protein gbs1087 (also referred to as FbsA and consisting of 442 amino acids; SEQ ID NO: 230) from which the protective protein consisting of the amino acid sequence of the SEQ ID NO: 2 is derived is disclosed in WO2004/035618 (see Fig. 1 and SEQ ID NO: 11). The amino acid sequence of SEQ ID NO: 2 is disclosed in the  
25 Examples as well as in the attached Sequence listing. The protective protein comprising or consisting of the protective peptide of SEQ ID NO: 2 or a functionally active variant thereof are referred to as (protective) proteins of subgroup ii).

The protective protein consisting of the amino acid sequence of SEQ ID NO: 3 is derived  
30 from *S. agalactiae* strain 12403 and has been denoted by gbs1309p (partial gbs1309) in accordance with the genome of NEM316 (ATCC12403). The DNA sequence encoding the full length protein gbs1309 (consisting of 403 amino acids; SEQ ID NO: 231) from which the protective protein consisting of the amino acid sequence of the SEQ ID NO: 3 is derived is disclosed in GenBank<sup>®</sup> accession number AL732656 (complete genome of

- 6 -

*Streptococcus agalactiae* NEM316) and the amino acid sequence of the full length protein is disclosed in WO2004/099242 (see SEQ ID NO: 307). The amino acid sequence of SEQ ID NO: 3 is disclosed in the Examples as well as in the attached Sequence listing. The protective protein comprising or consisting of the protective peptide of SEQ ID NO: 3 or a functionally active variant thereof are referred to as (protective) proteins of subgroup iii).

The protective protein consisting of the amino acid sequence of SEQ ID NO: 4 is derived from *S. agalactiae* strain 6313 and has been denoted by gbs1477p (partial gbs1477) in accordance with the genome of NEM316 (ATCC12403). The amino acid and encoding DNA sequences of the full length protein gbs1477 (also referred to as PabB and consisting of 674 amino acids; SEQ ID NO: 232) from which the protective protein consisting of the amino acid sequence of the SEQ ID NO: 4 is derived is disclosed in WO2004/035618 (see Fig. 16 and SEQ ID NO: 18). The amino acid sequence of SEQ ID NO: 4 is disclosed in the Examples as well as in the attached Sequence listing. The protective protein comprising or consisting of the protective peptide of SEQ ID NO: 4 or a functionally active variant thereof are referred to as (protective) proteins of subgroup iv).

The protective protein consisting of the amino acid sequence of SEQ ID NO: 5 is derived from *S. agalactiae* strain 6313 and has been denoted by gbs1478p (partial gbs1478) in accordance with the genome of NEM316 (ATCC12403). The amino acid and encoding DNA sequences of the full length protein gbs1478 (also referred to as PabA and consisting of 901 amino acids; SEQ ID NO: 233) from which the protective protein consisting of the amino acid sequence of the SEQ ID NO: 5 is derived is disclosed in WO2004/035618 (see Fig. 16 and SEQ ID NO: 17). The amino acid sequence of SEQ ID NO: 5 is disclosed in the Examples as well as in the attached Sequence listing. The protective protein comprising or consisting of the protective peptide of SEQ ID NO: 5 or a functionally active variant thereof are referred to as (protective) proteins of subgroup v).

The protective protein consisting of the amino acid sequence of SEQ ID NO: 6 is derived from *S. agalactiae* strain 12403 and has been denoted by gbs2018p (partial gbs2018) in accordance with the genome of NEM316 (ATCC12403). The DNA sequence encoding the full length protein gbs2018 (also referred to as BibA (Santi *et al.*, 2007, Mol. Microbiol. 63:754-767) and consisting of 643 amino acids; SEQ ID NO: 234) from which the protective protein consisting of the amino acid sequence of the SEQ ID NO: 6 is derived is



- 7 -

disclosed at GenBank<sup>®</sup> accession number AL732656 (complete genome of *Streptococcus agalactiae* NEM316) and the amino acid sequence of the full length protein is disclosed in WO2004/099242 (see SEQ ID NO: 364). The amino acid sequence of SEQ ID NO: 6 is disclosed in the Examples as well as in the attached Sequence listing. The protective protein comprising or consisting of the protective peptide of SEQ ID NO: 6 or a functionally active variant thereof are referred to as (protective) proteins of subgroup vi).

The combinations of the protective proteins of the sequences of SEQ ID NO: 1 to 6 have been shown to induce a protective immune response against different serotypes and/or to show increased protection against *S. agalactiae* in an animal model (see Examples). Functionally active variants may be obtained by changing the sequence of at least one of the protective proteins of SEQ ID NO: 1 to 6 and are characterized by having a biological activity similar to that displayed by the respective protective protein of the sequence of SEQ ID NO: 1 to 6 from which the variant is derived, including the ability to induce protective immune responses and/or to show protection against *S. agalactiae* e.g. in an animal model, wherein any variant may be tested in any of the tests described in the Examples. The functionally active variant of a protective protein may be obtained by sequence alterations in the protective protein, wherein the protein with the sequence alterations essentially retains a function of the unaltered protective protein, e.g. having a biological activity similar to that displayed by the unaltered protective protein (see above) including the ability to induce protective immune responses and/or to show protection against *S. agalactiae*. Such sequence alterations can include, but are not limited to, (conservative) substitutions, deletions, mutations and insertions.

In a preferred embodiment of the invention the composition comprises at least three proteins selected from the group consisting of subgroup i) to vi). In an even more preferred embodiment of the invention the composition comprises at least four proteins selected from the group consisting of subgroup i) to vi).

In a preferred embodiment of the invention the at least two, three or four proteins of the composition of the invention are selected from different subgroups i) to vi). Alternatively or additionally, at least two of the proteins of the composition of the invention are selected from one of the subgroups i) to vi).



- 8 -

Examples of combinations of the first alternative (selection of protective proteins from different groups) are compositions comprising:

- one protein of subgroup i) and one protein of subgroup ii);
- one protein of subgroup i) and one protein of subgroup iii);
- 5 - one protein of subgroup i) and one protein of subgroup iv);
- one protein of subgroup i) and one protein of subgroup v);
- one protein of subgroup i) and one protein of subgroup vi);
- one protein of subgroup ii) and one protein of subgroup iii);
- one protein of subgroup ii) and one protein of subgroup iv);
- 10 - one protein of subgroup ii) and one protein of subgroup v);
- one protein of subgroup ii) and one protein of subgroup vi);
- one protein of subgroup iii) and one protein of subgroup iv);
- one protein of subgroup iii) and one protein of subgroup v);
- one protein of subgroup iii) and one protein of subgroup vi);
- 15 - one protein of subgroup iv) and one protein of subgroup v);
- one protein of subgroup iv) and one protein of subgroup vi);
- one protein of subgroup v) and one protein of subgroup vi);
- one protein of subgroup i) and one protein of subgroup ii) and one protein selected from any of the subgroups iii) to vi);
- 20 - one protein of subgroup i) and one protein of subgroup iii) and one protein selected from any of the subgroups ii) or iv) to vi);
- one protein of subgroup i) and one protein of subgroup iv) and one protein selected from any of the subgroups ii), iii), v) or vi);
- one protein of subgroup i) and one protein of subgroup v) and one protein selected from any of the subgroups ii) to iv) or vi);
- 25 - one protein of subgroup i) and one protein of subgroup vi) and one protein selected from any of the subgroups ii) to v);
- one protein of subgroup ii) and one protein of subgroup iii) and one protein selected from any of the subgroups i) or iv) to vi);
- 30 - one protein of subgroup ii) and one protein of subgroup iv) and one protein selected from any of the subgroups i), iii), v) or vi);
- one protein of subgroup ii) and one protein of subgroup v) and one protein selected from any of the subgroups i), iii), iv) or vi);

- 9 -

- one protein of subgroup ii) and one protein of subgroup vi) and one protein selected from any of the subgroups i) or iii) to v);
- one protein of subgroup iii) and one protein of subgroup iv) and one protein selected from any of the subgroups i), ii), v) or vi);
- 5 - one protein of subgroup iii) and one protein of subgroup v) and one protein selected from any of the subgroups i), ii), iv) or vi);
- one protein of subgroup iii) and one protein of subgroup vi) and one protein selected from any of the subgroups i), ii), iv) or v);
- one protein of subgroup iv) and one protein of subgroup v) and one protein selected from any of the subgroups i) to iii) or vi);
- 10 - one protein of subgroup iv) and one protein of subgroup vi) and one protein selected from any of the subgroups i) to iii) or v); or
- one protein of subgroup v) and one protein of subgroup vi) and one protein selected from any of the subgroups i) to iv).

15

Preferred examples are:

- one protein of subgroup iv) and one protein of subgroup vi);
- one protein of subgroup iv), one protein of subgroup vi) and one protein of subgroup ii);
- 20 - one protein of subgroup iv), one protein of subgroup vi), one protein of subgroup ii) and one protein of subgroup v);
- one protein of subgroup iv), one protein of subgroup vi), one protein of subgroup ii) one protein of subgroup v) and one protein of subgroup i);
- one protein of subgroup iv), one protein of subgroup vi), one protein of subgroup ii), one protein of subgroup v) and one protein of subgroup iii); or
- 25 - one protein of subgroup iv), one protein of subgroup vi), one protein of subgroup ii), one protein of subgroup v), one protein of subgroup i) and one protein of subgroup iii).

30 In an alternative preferred embodiment of the invention at least two of the proteins of the composition of the invention may be selected from one of the subgroups i) to vi). The at least two proteins may be selected in order to cover different strains or serotypes of *S. agalactiae* and, accordingly, to provide protection against e.g. different strains or serotypes of *S. agalactiae*. The complete genome of *Streptococcus agalactiae* NEM316 (strain



- 10 -

12403) is available at GenBank<sup>®</sup> accession number AL732656. Furthermore, the complete or incomplete genomic sequences of the following strains of *Streptococcus agalactiae* are available at GenBank<sup>®</sup> (NIH genetic sequence database; <http://www.ncbi.nlm.nih.gov/>) or NCBI (National Center for Biotechnology Information, Bethesda, MD, USA; <http://www.ncbi.nlm.nih.gov/>) using the indicated accession numbers:

Strain	Serotype	Source
515	Ia	NCBI: NZ_AAJP00000000 GenBank <sup>®</sup> : AAJP00000000
A909	Ia/c	NCBI: NC_007432 GenBank <sup>®</sup> : CP000114
H36B	Ib	NCBI: NZ_AAJS00000000 GenBank <sup>®</sup> : AAJS00000000
18RS21	II	NCBI: NZ_AAJO00000000 GenBank <sup>®</sup> : AAJO00000000
COH1	III	NCBI: NZ_AAJR00000000 GenBank <sup>®</sup> : AAJR00000000
ATCC12403 (NEM316)	III	NCBI: NC_004368 GenBank <sup>®</sup> : AL732656
2603V/R	V	NCBI: NC_004116 GenBank <sup>®</sup> : AE009948
CJB111	V	NCBI: NZ_AAJQ00000000 GenBank <sup>®</sup> : AAJQ00000000

Using the sequences of SEQ ID NO: 1 to 6 as specified in the present description and knowing the sequences of other *S. agalactiae* strains (e.g. *vide supra*) the skilled person is able to identify the corresponding sequences of *S. agalactiae* strains other than 12403 (for SEQ ID NO: 1, 3 and 6) or 6313 (for SEQ ID NO: 2, 4 and 5) without undue burden. The corresponding sequences may be identified using e.g. the tools and sequences provided by “The Comprehensive Microbial Resource (CMR)” (see <http://cmr.tigr.org/>). However, it should be understood that the above strains are listed as examples of different *S. agalactiae* strains and that the present invention is not to be limited to those strains.



- 11 -

Additionally, examples of sequences of the proteins corresponding to SEQ ID NO: 1 to 6 and derived from other serotypes are published or disclosed in:

<b>Protein or Analogue</b>	<b>NCBI Accession Number (strain [serotype]) or SEQ ID NO</b>
gbs0233	Any sequence of SEQ ID NO: 55 to 60 (see Table 7), 229 and 235 to 286.
gbs1087	CAD12883 (6313); CAD27183 (706 S2 [Ia]); CAD27181 (SS1169 [V]); CAD27186 (O90R); CAD27182 (O176 H4A [II]) or any sequence of SEQ ID NO: 67 to 72 (see Table 8), 230 and 287 to 316.
gbs1309	Any sequence of SEQ ID NO: 79 to 84 (see Table 9), 231 and 317 to 359.
gbs1477	Any sequence of SEQ ID NO: 91 to 132 (see Table 10), 223 to 228 (see Fig. 5), 232 and 360 to 362.
gbs1478	Any sequence of SEQ ID NO: 185 to 203 (see Table 11), 233 and 363 to 378.
gbs2018	CAJ66802 (CCH57); CAJ66794 (CCH180); CAJ66788 (NEM1002); CAJ 66790 (NEM1560) or any sequence of SEQ ID NO: 175 to 179 (see Table 12), 234 and 379 to 425.

- 5 However, it should be understood that the present invention is not limited to the variants and corresponding proteins described above. Other naturally occurring proteins corresponding to those of SEQ ID NO: 1 to 6 may be identified as described above and used in order to carry out the present invention.
- 10 Examples of combinations of the second alternative (selection of protective proteins from one group only) are compositions comprising:
- at least two different proteins of subgroup i), preferably selected from the group consisting of proteins comprising or consisting of SEQ ID NO: 1 or a naturally occurring variant thereof;

- 12 -

- at least two different proteins of subgroup ii), preferably selected from the group consisting of proteins comprising or consisting of SEQ ID NO: 2 or a naturally occurring variant thereof;
- at least two different proteins of subgroup iii), preferably selected from the group consisting of proteins comprising or consisting of SEQ ID NO: 3 or a naturally occurring variant thereof;
- at least two different proteins of subgroup iv), preferably selected from the group consisting of proteins comprising or consisting of SEQ ID NO: 4 or a naturally occurring variant thereof;
- at least two different proteins of subgroup v), preferably selected from the group consisting of proteins comprising or consisting of SEQ ID NO: 5 or a naturally occurring variant thereof;
- at least two different proteins of subgroup vi), preferably selected from the group consisting of proteins comprising or consisting of SEQ ID NO: 6 or a naturally occurring variant thereof;
- preferably at least two protective proteins each comprising or consisting of a sequence selected from SEQ ID NO: 55 to 60 (see Table 7), 235 to 286, and optionally 229;
- preferably at least two protective proteins each comprising or consisting of a sequence selected from SEQ ID NO: 67 to 72 (see Table 8), 287 to 316, and optionally 230;
- preferably at least two protective proteins each comprising or consisting of a sequence selected from SEQ ID NO: 79 to 84 (see Table 9), 317 to 359, and optionally 231;
- preferably at least two protective proteins each comprising or consisting of a sequence selected from SEQ ID NO: 185 to 203 (see Table 11), 363 to 378, and optionally 233;
- preferably at least two protective proteins each comprising or consisting of a sequence selected from SEQ ID NO: 175 to 179 (see Table 12), 379 to 425 and optionally 234; or
- more preferably at least two protective proteins each comprising or consisting of a sequence selected from SEQ ID NO: 91 to 132 (see Table 10), 360 to 362, or 223 to 228 (see Fig. 5) and optionally 232.



- 13 -

In a preferred embodiment the naturally occurring variants are those derived from *S. agalactiae* strains selected from the group consisting of IC97, IC98, IC105, IC108, IC216, IC244, IC245, IC246, IC247, IC250, IC251, IC252, IC253, IC254, IC255, IC287, IC288, IC289, IC290, IC291, IC304, IC305, IC306, IC361, IC363, IC364, IC365, IC366, IC367, 5 IC368, IC377, IC379, IC432, IC434, IC455, IC457, IC458, IC459, IC460, IC461, IC462, IC463, IC469, IC470, 126H4A, 5095S2, 6313, 12351, 12403 (NEM316), 12401, COH1, BAA23, 0176H4A, A909, C388/90, BAA22, 2603V/R, 49447, BAA611, 515, H36B, 18RS21, CJB111, and those disclosed in Tables 7 to 13.

10 In another preferred embodiment of the present invention the composition of the invention comprises

- at least one protein of subgroup iv);
- at least one protein of subgroup vi);
- at least one protein of subgroup iv) and at least one protein of subgroup vi);
- 15 - at least one protein of subgroup iv), at least one protein of subgroup vi) and at least one protein of subgroup ii); or
- at least one protein of subgroup iv), at least one protein of subgroup vi), at least one protein of subgroup ii) and at least one protein of subgroup v).

20 In a preferred embodiment of the invention one of the at least two proteins comprises or consists of the protective peptide of SEQ ID NO: 4 (gbs1477p) or a functionally active variant thereof, preferably protective peptide of SEQ ID NO: 4 (gbs1477p) or a naturally occurring functionally active variant thereof, more preferably a naturally occurring functionally active variant as listed in Tables 10, 13, and in the Sequence listing.

25

In another preferred embodiment of the invention one of the at least two proteins comprises or consists of the protective peptide of SEQ ID NO: 6 (gbs2018p) or a functionally active variant thereof, preferably protective peptide of SEQ ID NO: 6 (gbs2018p) or a naturally occurring functionally active variant thereof, more preferably a 30 naturally occurring functionally active variant as listed in Tables 12, 13, and in the Sequence listing.

In another preferred embodiment of the invention one of the at least two proteins comprises or consists of the protective peptide of SEQ ID NO: 2 (gbs1087p) or a



- 14 -

functionally active variant thereof, preferably protective peptide of SEQ ID NO: 2 (gbs1087p) or a naturally occurring functionally active variant thereof, more preferably a naturally occurring functionally active variant as listed in Tables 8, 13, and in the Sequence listing.

5

In another preferred embodiment of the invention one of the at least two proteins comprises or consists of the protective peptide of SEQ ID NO: 5 (gbs1478p) or a functionally active variant thereof, preferably protective peptide of SEQ ID NO: 5 (gbs1478p) or a naturally occurring functionally active variant thereof, more preferably a naturally occurring functionally active variant as listed in Tables 11, 13, and in the Sequence listing.

10

In another preferred embodiment of the invention one of the at least two proteins comprises or consists of the protective peptide of SEQ ID NO: 1 (gbs0233p) or a functionally active variant thereof, preferably protective peptide of SEQ ID NO: 1 (gbs0233p) or a naturally occurring functionally active variant thereof, more preferably a naturally occurring functionally active variant as listed in Tables 7, 13, and in the Sequence listing.

15

In another preferred embodiment of the invention one of the at least two proteins comprises or consists of the protective peptide of SEQ ID NO: 3 (gbs1309p) or a functionally active variant thereof, preferably protective peptide of SEQ ID NO: 3 (gbs1309p) or a naturally occurring functionally active variant thereof, more preferably a naturally occurring functionally active variant as listed in Tables 9, 13, and in the Sequence listing.

20

In a more referred embodiment of the invention the at least two proteins of the composition of the invention encompass:

- the protective peptide of SEQ ID NO: 4 (gbs1477p) and the protective peptide of SEQ ID NO: 6 (gbs2018p);
- the protective peptide of SEQ ID NO: 4 (gbs1477p) and the protective peptide of SEQ ID NO: 6 (gbs2018p) and the protective peptide of SEQ ID NO: 2 (gbs1087p); or

30

- 15 -

– the protective peptide of SEQ ID NO: 4 (gbs1477p) and the protective peptide of SEQ ID NO: 6 (gbs2018p) and the protective peptide of SEQ ID NO: 2 (gbs1087p) and the protective peptide of SEQ ID NO: 5 (gbs1478p).

5 In a further preferred embodiment of the invention a naturally occurring functionally active variant of any of the protective peptides of SEQ ID NO: 1 to 6 of the above list of compositions may be used. Examples of the resulting combinations are:

- I. the protective peptide of SEQ ID NO: 4 (gbs1477p) and a naturally occurring  
10 functionally active variant of SEQ ID NO: 6 (gbs2018p);
- II. a naturally occurring functionally active variant of SEQ ID NO: 4 (gbs1477p) and the protective peptide of SEQ ID NO: 6 (gbs2018p);
- III. a naturally occurring functionally active variant of SEQ ID NO: 4 (gbs1477p) and a naturally occurring functionally active variant of SEQ ID NO: 6 (gbs2018p);
- 15 IV. the protective peptide of SEQ ID NO: 4 (gbs1477p) and the protective peptide of SEQ ID NO: 6 (gbs2018p) and a naturally occurring functionally active variant of SEQ ID NO: 2 (gbs1087p);
- V. the protective peptide of SEQ ID NO: 4 (gbs1477p) and a naturally occurring functionally active variant of SEQ ID NO: 6 (gbs2018p) and the protective peptide of  
20 SEQ ID NO: 2 (gbs1087p);
- VI. a naturally occurring functionally active variant of SEQ ID NO: 4 (gbs1477p) and the protective peptide of SEQ ID NO: 6 (gbs2018p) and the protective peptide of SEQ ID NO: 2 (gbs1087p);
- VII. a naturally occurring functionally active variant of SEQ ID NO: 4 (gbs1477p) and a  
25 naturally occurring functionally active variant of SEQ ID NO: 6 (gbs2018p) and the protective peptide of SEQ ID NO: 2 (gbs1087p);
- VIII. the protective peptide of SEQ ID NO: 4 (gbs1477p) and a naturally occurring functionally active variant of SEQ ID NO: 6 (gbs2018p) and a naturally occurring functionally active variant of SEQ ID NO: 2 (gbs1087p);
- 30 IX. a naturally occurring functionally active variant of SEQ ID NO: 4 (gbs1477p) and the protective peptide of SEQ ID NO: 6 (gbs2018p) and a naturally occurring functionally active variant of SEQ ID NO: 2 (gbs1087p);



- 16 -

- X. a naturally occurring functionally active variant of SEQ ID NO: 4 (gbs1477p) and a naturally occurring functionally active variant of SEQ ID NO: 6 (gbs2018p) and a naturally occurring functionally active variant of SEQ ID NO: 2 (gbs1087p);
- XI. any of the compositions of I to X in combination with the protective peptide of SEQ ID NO: 5 (gbs1478p); or
- XII. any of the compositions of I to X in combination with a naturally occurring functionally active variant of SEQ ID NO: 5 (gbs1478p).

wherein the naturally occurring functionally active variant is selected from those listed in Tables 7 to 13 and Fig. 5 and those of SEQ ID NO: 229 to 234 and 235 to 425.

Preferred *S. agalactiae* strains from which the naturally occurring functionally active variant may be derived include IC97, IC98, IC105, IC108, IC216, IC244, IC245, IC246, IC247, IC250, IC251, IC252, IC253, IC254, IC255, IC287, IC288, IC289, IC290, IC291, IC304, IC305, IC306, IC361, IC363, IC364, IC365, IC366, IC367, IC368, IC377, IC379, IC432, IC434, IC455, IC457, IC458, IC459, IC460, IC461, IC462, IC463, IC469, IC470, 126H4A, 5095S2, 6313, 12351, 12403 (NEM316), 12401, COH1, BAA23, 0176H4A, A909, C388/90, BAA22, 2603V/R, 49447, BAA611, 515, H36B, 18RS21, CJB111, and those disclosed in Tables 7 to 13.

20

In one embodiment of the invention two or more proteins of the at least two proteins of the composition of the invention may be combined into at least one fusion protein. The resulting fusion protein may encompass two or more of the proteins of subgroups i), ii), iii), iv), v) and/or vi) as defined above. Any of the specific combinations mentioned above may be combined into at least one fusion protein. The fusion protein may encompass e.g. the same protein components as described in the section "examples of combinations of the first alternative (selection of protective proteins from different groups)" for the components of the composition, wherein the proteins can be arranged in the fusion protein in any suitable manner.

30

The fusion protein may comprise or consist of two or more proteins as defined above. Additionally, the fusion protein may encompass a linker, such as a protein linker, to connect the two or more proteins or additional C- or N-terminal sequences, such as a tag in order to purify the fusion protein. Additional sequences may also result from genetic

- 17 -

engineering and the use of suitable restriction sites when preparing the nucleic acid sequences underlying the fusion protein.

The proteins of subgroup i), ii), iii), iv), v) and/or vi) combined in a fusion protein may be directly joined to each other or may be combined over a linker. The linker may be e.g. a short amino acid sequence. The linker may result from the genetic engineering of a suitable fusion protein or may be introduced in order to allow the single proteins to operate effectively.

10 In a preferred embodiment the functionally active variant

a) is a functionally active fragment of the protective peptide, the functionally active fragment comprising at least 50% of the sequence of the protective peptide, preferably at least 70%, more preferably at least 80%, still more preferably at least 90%, even more preferably at least 95% and most preferably at least 97%, 98% or 15 99%;

b) is derived from the protective peptide by at least one amino acid substitution, addition and/or deletion, wherein the functionally active variant has a sequence identity to the protective peptide or to the functionally active fragment as defined in a) of at least 40%, preferably at least 60%, more preferably at least 75%, still more preferably at least 90%, even more preferably at least 95% and most preferably at 20 least 97%, 98% or 99%; and/or

c) consists of the protective peptide or a functionally active variant thereof and additionally at least one amino acid heterologous to the protective peptide, preferably wherein the functionally active variant is derived from or identical to any of the 25 naturally occurring variants of any of the sequences of SEQ ID NO: 55 to 60, 67 to 72, 79 to 84, 91 to 132, 175 to 179, 185 to 203, 223 to 234, and 235 to 425.

The combinations of protective peptides of SEQ ID NO: 1 to 6 have been shown to induce a protective immune response against different serotypes and/or to show protection against 30 *S. agalactiae* in a sepsis model (see Examples). Functionally active variants may be obtained by changing the sequence of the protective peptide as defined above and are characterized by having a biological activity similar to that displayed by the respective protective peptide of the sequence of SEQ ID NO: 1 to 6 from which the variant is derived, including the ability to induce protective immune responses and/or to show protection



- 18 -

against *S. agalactiae* e.g. in a sepsis model, wherein any variant may be tested in any of the tests described in the Examples within a composition of at least two proteins as defined above. The functionally active variant of a protective peptide may be obtained by sequence alterations in the protective peptide, wherein the peptide with the sequence alterations  
5 retains a function of the unaltered protective peptide, e.g. having a biological activity similar to that displayed by the unaltered protective peptide (see above), when used in combination of the invention. Such sequence alterations can include, but are not limited to, (conservative) substitutions, additions, deletions, mutations and insertions.

10 The variant of the protective peptide is functionally active in the context of the present invention, if the activity of the composition of the invention including the variant (but not the original protein) amounts to at least 10%, preferably at least 25%, more preferably at least 50%, even more preferably at least 70%, still more preferably at least 80%, especially at least 90%, particularly at least 95%, most preferably at least 99% of the activity of the  
15 composition of the invention including the protective peptide without sequence alteration (i.e. the original protein). The activity of the composition including the variant may be determined or measured as described in the Examples and then compared to that obtained for the composition including the respective protective peptide of the amino acid sequence of SEQ ID NO: 1 to 6 instead of the variant.

20

The functionally active fragment of the protective peptide is characterized by being derived from the protective peptide of SEQ ID NO: 1 to 6 by one or more deletions resulting in a peptide comprising at least 50% of the sequence of the protective peptide, preferably at least 70%, more preferably at least 80%, still more preferably at least 90%, even more  
25 preferably at least 95% and most preferably at least 97%, 98% or 99%. Sequence identity may be determined as described below. The deletion(s) may be C-terminally, N-terminally and/or internally. Preferably the fragment is obtained by 1, 2, 3, 4, 5, 6, 7, 8, 9 or 10, more preferably 1, 2, 3, 4 or 5, even more preferably 1, 2 or 3, still more preferably 1 or 2, most preferably 1 deletion(s).

30

Alternatively or additionally, the variant may be obtained from the protective peptide by at least one amino acid substitution, addition and/or deletion, wherein the functionally active variant has a sequence identity to the protective peptide or to the functionally active fragment as defined in a) of at least 40%, preferably at least 60%, more preferably at least

- 19 -

75%, still more preferably at least 90%, even more preferably at least 95% and most preferably at least 97%, 98% or 99%. Sequence identity may be determined as described below. The substitution(s), addition(s) and/or deletion(s) may be C-terminally, N-terminally and/or internally. Preferably, the functionally active variant is obtained from the protective peptide or the fragment, preferably the protective peptide, by 1, 2, 3, 4, 5, 6, 7, 8, 9 or 10, more preferably 1, 2, 3, 4 or 5, even more preferably 1, 2 or 3, still more preferably 1 or 2, most preferably 1 amino acid substitution(s), addition(s) and/or deletion(s).

Furthermore, the variant may consist of the protective peptide or the functionally active variant thereof, preferably the variant of a) and/or b), and at least one amino acid residue heterologous to the protective peptide or variant thereof, such as a marker protein. The feature "heterologous amino acid" or "amino acid heterologous to the protective peptide or variant thereof" refers to any amino acid which is different from that amino acid located adjacent to the protective protein in any naturally occurring protein of *S. agalactiae*, particularly from that of strain 12403 (for SEQ ID NO: 1, 3 and 6) or 6313 (for SEQ ID NO: 2, 4 and 5), especially the sequence made reference to above. The one or more additional amino acids may be C-terminally, N-terminally or C- and N-terminally to the protective peptide or variant thereof.

20

The substituted or additional sequence or amino acid residue(s) as defined above consists of (an) amino acid residue(s), which may be any amino acid, which may be either an L- and/or a D-amino acid, naturally occurring and otherwise. Preferably the amino acid is any naturally occurring amino acid such as alanine, cysteine, aspartic acid, glutamic acid, phenylalanine, glycine, histidine, isoleucine, lysine, leucine, methionine, asparagine, proline, glutamine, arginine, serine, threonine, valine, tryptophan or tyrosine.

However, the amino acid may also be a modified or an unusual amino acid. Examples of those are 2-aminoadipic acid, 3-aminoadipic acid, beta-alanine, 2-aminobutyric acid, 4-aminobutyric acid, 6-aminocaproic acid, 2-aminoheptanoic acid, 2-aminoisobutyric acid, 3-aminoisobutyric acid, 2-aminopimelic acid, 2,4-diaminobutyric acid, desmosine, 2,2'-diaminopimelic acid, 2,3-diaminopropionic acid, N-ethylglycine, N-ethylasparagine, hydroxylysine, allo-hydroxylysine, 3-hydroxyproline, 4-hydroxyproline, isodesmosine, allo-isoleucine, N-methylglycine, N-methylisoleucine, 6-N-methyllysine, N-methylvaline,



- 20 -

norvaline, norleucine or ornithine. Additionally, the amino acid may be subject to modifications such as posttranslational modifications. Examples of modifications include acetylation, amidation, blocking, formylation, gamma-carboxyglutamic acid hydroxylation, glycosylation, methylation, phosphorylation and sulfatation. If more than one substituted or additional heterologous amino acid residue is present in the peptide, the amino acid residues may be the same or different from one another.

In one preferred embodiment of the invention, the functionally active variant of the peptide of the invention is essentially identical to the protective peptide of subgroups i) to vi), but differs from the peptide of the SEQ ID NO: 1 to 6, respectively, in that it is derived from a homologous sequence of a different strain or even serotype of *S. agalactiae*. As detailed above different strains and serotypes of *S. agalactiae* have been identified so far. Accordingly, any of these serotypes may be the basis for the functionally active variant. These are referred to as naturally occurring variants (see also above). Preferably, these naturally occurring variants are derived from *S. agalactiae* strains selected from the group consisting of IC97, IC98, IC105, IC108, IC216, IC244, IC245, IC246, IC247, IC250, IC251, IC252, IC253, IC254, IC255, IC287, IC288, IC289, IC290, IC291, IC304, IC305, IC306, IC361, IC363, IC364, IC365, IC366, IC367, IC368, IC377, IC379, IC432, IC434, IC455, IC457, IC458, IC459, IC460, IC461, IC462, IC463, IC469, IC470, 126H4A, 5095S2, 6313, 12351, 12403 (NEM316), 12401, COH1, BAA23, 0176H4A, A909, C388/90, BAA22, 2603V/R, 49447, BAA611, 515, H36B, 18RS21, CJB111, and those disclosed in Tables 7 to 13.

However, the term "functionally active variant" includes naturally occurring allelic variants, as well as mutants or any other non-naturally occurring variants. As is known in the art, an allelic variant is an alternate form of a (poly)peptide that is characterized as having a substitution, deletion, or addition of one or more amino acids that does essentially not alter the biological function of the polypeptide. By "biological function" is meant a function of the peptide in the cell it naturally occurs in, even if the function is not necessary for the growth or survival of the cells. For example, the biological function of a porin is to allow the entry into cells of compounds present in the extracellular medium. The biological function is distinct from the antigenic function. A polypeptide can have more than one biological function.

- 21 -

Accordingly, the present invention also relates to compositions comprising protective peptides including functionally active variants thereof of different *S. agalactiae* isolates. Such homologues may easily be identified and isolated based on the nucleic acid and amino acid sequences disclosed herein as discussed above. A homologous protective peptide of a different strain or even serotype may be identified by e.g. sequence alignment. The homologous sequence may vary from any of the protective peptides of subgroups i) to vi), by one or more amino acid substitutions, deletions and/or additions.

Percentage of sequence identity can be determined e.g. by sequence alignment. Methods of alignment of sequences for comparison are well known in the art. Various programs and alignment algorithms have been described e.g. in Smith and Waterman, Adv. Appl. Math. 2: 482, 1981 or Pearson and Lipman, Proc. Natl. Acad. Sci. U.S.A. 85: 2444-2448, 1988.

The NCBI Basic Local Alignment Search Tool (NCBI BLAST) (Altschul et al., J. Mol. Biol. 215: 403-410, 1990) is available from several sources, including the National Center for Biotechnology Information (NCBI, Bethesda, MD) and on the Internet, for use in connection with the sequence analysis programs blastp, blastn, blastx, tblastn and tblastx. Variants, e.g. of any protective peptide of the sequences of SEQ ID NO: 1 to 6, are typically characterized using the NCBI Blast 2.0, gapped blastp set to default parameters. For comparisons of amino acid sequences of e.g. at least 85 amino acids, the "Blast 2 sequences" function may be employed using the default BLOSUM62 matrix set to default parameters, (gap existence cost of 11, and a per residue gap cost of 1).

In a preferred embodiment, the functionally active variant derived from the peptide as defined above by amino acid exchanges, deletions or insertions may also conserve, or more preferably improve, the activity (as defined above). Furthermore, these peptides may also cover epitopes, which trigger the same or preferably an improved T cell response. These epitopes are referred to as "heteroclitic". They have a similar or preferably greater affinity to MHC/HLA molecules, and the ability to stimulate the T cell receptors (TCR) directed to the original epitope in a similar or preferably stronger manner. Heteroclitic epitopes can be obtained by rational design i. e. taking into account the contribution of individual residues to binding to MHC/HLA as for instance described by (Rammensee, H. et al., 1999, Immunogenetics. 50: 213-219), combined with a systematic exchange of residues potentially interacting with the TCR and testing the resulting sequences with T cells



- 22 -

directed against the original epitope. Such a design is possible for a skilled man in the art without much experimentation.

Conservative substitutions are those that take place within a family of amino acids that are related in their side chains and chemical properties. Examples of such families are amino acids with basic side chains, with acidic side chains, with non-polar aliphatic side chains, with non-polar aromatic side chains, with uncharged polar side chains, with small side chains, with large side chains etc.. In one embodiment, one conservative substitution is included in the peptide. In another embodiment, two conservative substitutions or less are included in the peptide. In a further embodiment, three conservative substitutions or less are included in the peptide.

Examples of conservative amino acid substitutions include, but are not limited to, those listed below:

15

<u>Original Residue</u>	<u>Conservative Substitutions</u>
Ala	Ser
Arg	Lys
Asn	Gln; His
20 Asp	Glu
Cys	Ser
Gln	Asn
Glu	Asp
His	Asn; Gln
25 Ile	Leu; Val
Leu	Ile; Val
Lys	Arg; Gln; Asn
Met	Leu; Ile
Phe	Met; Leu; Tyr
30 Ser	Thr
Thr	Ser
Trp	Tyr
Tyr	Trp; Phe
Val	Ile; Leu

In another embodiment of the invention the peptide as defined above may be modified by a variety of chemical techniques to produce derivatives having essentially the same activity (as defined above for fragments and variants) as the modified peptides, and optionally  
5 having other desirable properties. For example, carboxylic acid groups of the protein, whether C-terminal or side chain, may be provided in the form of a salt of a pharmaceutically-acceptable cation or esterified to form an ester, or converted to an amide. Amino groups of the peptide, whether amino-terminal or side chain, may be in the form of a pharmaceutically-acceptable acid addition salt, such as the HCl, HBr, acetic, benzoic,  
10 toluene sulfonic, maleic, tartaric and other organic salts, or may be converted to an amide. Hydroxyl groups of the peptide side chains may be converted to alkoxy or to an ester using well recognized techniques. Phenyl and phenolic rings of the peptide side chains may be substituted with one or more halogen atoms, such as fluorine, chlorine, bromine or iodine, or with alkyl, alkoxy, carboxylic acids and esters thereof, or amides of such carboxylic  
15 acids. Thiols can be protected with any one of a number of well recognized protecting groups, such as acetamide groups.

Peptides of this invention may be in combination with outer surface proteins or other proteins or antigens of other proteins. In such combination, the peptide may be in the form  
20 of a fusion protein. The peptides/proteins of the composition of the invention may be optionally fused to a selected peptide or protein derived from other microorganisms. For example, a peptide or protein may be fused at its N-terminus or C-terminus to a polypeptide from another pathogen or to more than one polypeptide in sequence. Peptides which may be useful for this purpose include polypeptides identified by the prior art.

25

In a preferred embodiment of the invention a protein/peptide of the composition of the invention is fused to an epitope tag which provides an epitope to which an anti-tag substance can selectively bind. The epitope tag is generally placed at the N- or C-terminus of the peptide but may be incorporated as an internal insertion or substitution as the  
30 biological activity permits. The presence of such epitope-tagged forms of a peptide can be detected using a substance such as an antibody against the tagged peptide. Also, provision of the epitope tag enables the peptide to be readily purified by affinity purification using an anti-tag antibody or another type of affinity matrix that binds to the epitope tag. Various tag polypeptides and their respective antibodies are well known in the art. Examples



- 24 -

include a poly-histidine (poly-his) tag, e.g. a hexa-histidine tag as described in the Examples, a poly-histidine-glycine (poly-his-gly) tag, the HA tag polypeptide, the c-myc tag, the Strep tag and the FLAG tag.

5 Fusions also may include the peptides/proteins of the composition of this invention fused or coupled to moieties other than amino acids, including lipids and carbohydrates. Further, peptides/proteins/compositions of this invention may be employed in combination with other vaccinal agents described by the prior art, as well as with other types of vaccinal agents derived from other microorganisms. Such peptides/proteins are useful in the  
10 prevention, treatment and diagnosis of diseases caused by a wide spectrum of *Streptococcus* isolates.

These fusion proteins are constructed for use in the methods and compositions of this invention. These fusion proteins or multimeric proteins may be produced recombinantly, or  
15 may be synthesized chemically.

The peptides and proteins described herein may be prepared by any of a number of conventional techniques. Desired peptides may be chemically synthesized. An alternative approach involves generating the fragments of known peptides by enzymatic digestion,  
20 e.g., by treating the protein with an enzyme known to cleave proteins at sites defined by particular amino acid residues, or by digesting the DNA with suitable restriction enzymes, expressing the digested DNA and isolating the desired fragment. Yet another suitable technique involves isolating and amplifying a DNA fragment encoding a desired peptide  
25 termini of the DNA fragment are employed as the 5' and 3' primers in the PCR. Techniques for making mutations, such as deletions, insertions and substitutions, at predetermined sites in DNA, and therefore in proteins having a known sequence are well known. One of skill in the art using conventional techniques, such as PCR, may readily use the peptides, proteins and compositions provided herein to identify and isolate other similar proteins.  
30 Such methods are routine and not considered to require undue experimentation, given the information provided herein. For example, variations can be made using oligonucleotide-mediated site-directed mutagenesis (Carter et al., Nucl. Acids Res., 13:4431 (1985); Zoller et al., Nucl. Acids Res. 10:6487 (1987)), cassette mutagenesis (Wells et al., Gene, 34:315 (1985)), restriction selection mutagenesis (Wells et al., Philos. Trans. R. Soc. London

- 25 -

SerA, 317:415 (1986)), PCR mutagenesis, or other known techniques can be performed on the cloned DNA to produce the peptide or composition of the invention.

Another subject of the invention relates to a protective peptide consisting of the amino acid  
5 sequence selected from the group consisting of SEQ ID NO: 1, SEQ ID NO: 2, SEQ ID  
NO: 3, SEQ ID NO: 4, SEQ ID NO: 5, SEQ ID NO: 6, SEQ ID NO: 55 to 57, 59, 60, 68,  
69, 71, 72, 79 to 84, 91 to 132, 175 to 179, 185 to 203, 223 to 234, and 235 to 425 which  
have been shown to provide protection against *S. agalactiae* (see Examples).

10 Another subject of the invention relates to one or more nucleic acid(s) encoding the at least  
two proteins comprised in the composition according to the invention and/or any of the  
protective peptides according to the invention.

Nucleic acid molecules of the present invention may be in the form of RNA, such as  
15 mRNA or cRNA, or in the form of DNA, including, for instance, cDNA and genomic  
DNA e.g. obtained by cloning or produced by chemical synthetic techniques or by a  
combination thereof. The DNA may be triple-stranded, double-stranded or single-stranded.  
Single-stranded DNA may be the coding strand, also known as the sense strand, or it may  
be the non-coding strand, also referred to as the anti-sense strand. Nucleic acid molecule as  
20 used herein also refers to, among others, single- and double-stranded DNA, DNA that is a  
mixture of single- and double-stranded RNA, and RNA that is a mixture of single- and  
double-stranded regions, hybrid molecules comprising DNA and RNA that may be single-  
stranded or, more typically, double-stranded, or triple-stranded, or a mixture of single- and  
double-stranded regions. In addition, nucleic acid molecule as used herein refers to triple-  
25 stranded regions comprising RNA or DNA or both RNA and DNA.

The nucleic acid may be a fragment of a nucleic acid occurring naturally in *S. agalactiae*.  
The nucleic acid also includes sequences that are a result of the degeneration of the genetic  
code. There are 20 natural amino acids, most of which are specified by more than one  
30 codon. Therefore, all nucleotide sequences are included in the invention which result in the  
peptide as defined above.

Preferred examples of the nucleic acid(s) encoding the at least two proteins comprised in  
the composition according to the invention and/or any of the protective peptides according



- 26 -

to the invention are those comprising or consisting of at least one nucleic acid sequence selected from the group consisting of SEQ ID NO: 7, SEQ ID NO: 8, SEQ ID NO: 9, SEQ ID NO: 10, SEQ ID NO: 11, SEQ ID NO: 12, SEQ ID NO: 61 to 66, SEQ ID NO: 73 to 78, SEQ ID NO: 85 to 90, SEQ ID NO: 133 to 174, SEQ ID NO: 180 to 184 and SEQ ID  
5 NO: 204 to 222. The above sequences are indicated in the Examples, Tables 7 to 12 and the attached Sequence listing.

Additionally, the nucleic acid may contain one or more modified bases. Such nucleic acids may also contain modifications e.g. in the ribose-phosphate backbone to increase stability  
10 and half life of such molecules in physiological environments. Thus, DNAs or RNAs with backbones modified for stability or for other reasons are "nucleic acid molecules" as that feature is intended herein. Moreover, DNAs or RNAs comprising unusual bases, such as inosine, or modified bases, such as tritylated bases, to name just two examples, are nucleic acid molecules within the context of the present invention. It will be appreciated that a  
15 great variety of modifications have been made to DNA and RNA that serve many useful purposes known to those of skill in the art. The term nucleic acid molecule as it is employed herein embraces such chemically, enzymatically or metabolically modified forms of nucleic acid molecule, as well as the chemical forms of DNA and RNA characteristic of viruses and cells, including simple and complex cells, *inter alia*. For  
20 example, nucleotide substitutions can be made which do not affect the peptide or protein or composition of the invention encoded by the nucleic acid, and thus any nucleic acid molecule which encodes an antigenic peptide or functionally active variant thereof or a composition of the invention as defined above is encompassed by the present invention.

25 Furthermore, any of the nucleic acid molecules encoding a peptide or composition of the invention can be functionally linked, using standard techniques such as standard cloning techniques, to any desired regulatory sequences, whether a *S. agalactiae* regulatory sequence or a heterologous regulatory sequence, heterologous leader sequence, heterologous marker sequence or a heterologous coding sequence to create a fusion protein.

30

The nucleic acid of the invention may be originally formed *in vitro* or in a cell in culture, in general, by the manipulation of nucleic acids by endonucleases and/or exonucleases and/or polymerases and/or ligases and/or recombinases or other methods known to the skilled practitioner to produce the nucleic acids.

In one embodiment of the invention, the nucleic acid(s) according to the invention is/are located in a vector or a cell other than *S. agalactiae*.

5 A vector may further include nucleic acid sequences that permit it to replicate in the host cell, such as an origin of replication, one or more therapeutic genes and/or selectable marker genes and other genetic elements known in the art such as regulatory elements directing transcription, translation and/or secretion of the encoded peptide or protein. The vector may be used to transduce, transform or infect a cell, thereby causing the cell to  
10 express nucleic acids and/or proteins other than those native to the cell. The vector optionally includes materials to aid in achieving entry of the nucleic acid into the cell, such as a viral particle, liposome, protein coating or the like. Numerous types of appropriate expression vectors for protein expression are known in the art, which may be used in standard molecular biology techniques. Such vectors are selected from among  
15 conventional vector types including insects, e.g., baculovirus expression, or yeast, fungal, bacterial or viral expression systems. Other appropriate expression vectors, of which numerous types are known in the art, can also be used for this purpose. Methods for obtaining such expression vectors are well-known (see, e.g. Sambrook et al., Molecular Cloning. A Laboratory Manual, 2nd edition, Cold Spring Harbor Laboratory, New York  
20 (1989)). In one embodiment, the vector is a viral vector. Viral vectors include, but are not limited to, retroviral and adenoviral vectors.

Suitable host cells or cell lines for transfection by this method include bacterial cells. For example, the various strains of *E. coli* are well-known as host cells in the field of  
25 biotechnology. Various strains of *B. subtilis*, *Pseudomonas*, *Streptomyces*, and other bacilli and the like may also be employed in this method. Many strains of yeast cells known to those skilled in the art are also available as host cells for expression of the peptides of the present invention. Other fungal cells or insect cells such as *Spodoptera frugiperda* (Sf9) cells may also be employed as expression systems. Alternatively, mammalian cells, such as  
30 human 293 cells, Chinese hamster ovary cells (CHO), the monkey COS-1 cell line or murine 3T3 cells derived from Swiss, BALB/c or NIH mice may be used. Still other suitable host cells, as well as methods for transfection, culture, amplification, screening, production, and purification are known in the art.



- 28 -

A further subject of the invention relates to a method of producing the composition according to the invention or the protective peptide according to the invention, comprising

- (a) introducing the one or more nucleic acids into a host cell;
- (b) expressing the protein(s) and/or peptide(s) encoded by the nucleic acid by culturing  
5 the host cell under conditions conducive to the expression of the protein(s) and/or peptide(s); and
- (c) collecting and/or isolating the expressed protein(s) and/or peptide(s) of step (b).

A peptide or composition of the invention or component thereof may be produced by  
10 expressing a nucleic acid of the invention in a suitable host cell. The nucleic acid encoding the peptide/protein can be introduced into a host cell by any conventional technique. The host cells can e.g. be transfected, e.g. by conventional means such as electroporation with at least one expression vector containing a nucleic acid of the invention under the control of a transcriptional regulatory sequence. The transfected or transformed host cell is then  
15 cultured under conditions that allow expression of the protein. The expressed protein is recovered, isolated, and optionally purified from the cell (or from the culture medium, if expressed extracellularly) by appropriate means known to one of skill in the art. For example, the proteins are isolated in soluble form following cell lysis, or extracted using known techniques, e.g. in guanidine chloride. If desired, the peptides or fragments of the  
20 invention are produced as a fusion protein. Such fusion proteins are those described above. Alternatively, for example, it may be desirable to produce fusion proteins to enhance expression of the protein in a selected host cell or to improve purification. The molecules comprising the peptides and compositions of this invention may be further purified using any of a variety of conventional methods including, but not limited to: liquid  
25 chromatography such as normal or reversed phase, using HPLC, FPLC and the like; affinity chromatography (such as with inorganic ligands or monoclonal antibodies); size exclusion chromatography; immobilized metal chelate chromatography; gel electrophoresis; and the like. One of skill in the art may select the most appropriate isolation and purification techniques without departing from the scope of this invention.  
30 Such purification provides the peptide/protein/composition in a form substantially free from other proteinaceous and non-proteinaceous materials of the microorganism.

Still another subject of the invention relates to a pharmaceutical composition, especially a vaccine, comprising

- 29 -

- (i) the composition according to the invention and/or at least one protective peptide according to the invention; and
- (ii) optionally a pharmaceutically acceptable carrier or excipient.

5 A peptide or composition of the invention may be used for methods for immunizing or treating humans and/or animals with the disease caused by infection with *S. agalactiae*. Therefore, the peptide or composition may be used within a pharmaceutical composition. The pharmaceutical composition of the present invention may further encompass pharmaceutically acceptable carriers and/or excipients. The pharmaceutically acceptable  
10 carriers and/or excipients useful in this invention are conventional and may include buffers, stabilizers, diluents, preservatives, and solubilizers. Remington's Pharmaceutical Sciences, by E. W. Martin, Mack Publishing Co., Easton, PA, 15th Edition (1975), describes compositions and formulations suitable for pharmaceutical delivery of the (poly)peptides/proteins herein disclosed.

15

If the pharmaceutical composition comprises at least two protective proteins as defined above, the proteins of subgroup i) to vi) may be formulated into one or more pharmaceutical composition(s). Additionally, the two or more pharmaceutical compositions may be administered together, simultaneously or consecutively.

20

In general, the nature of the carrier or excipients will depend on the particular mode of administration being employed. For instance, parenteral formulations usually comprise injectable fluids that include pharmaceutically and physiologically acceptable fluids such as water, physiological saline, balanced salt solutions, aqueous dextrose, glycerol or the  
25 like as a vehicle. For solid compositions (e. g. powder, pill, tablet, or capsule forms), conventional non-toxic solid carriers can include, for example, pharmaceutical grades of mannitol, lactose, starch, or magnesium stearate. In addition to biologically neutral carriers, pharmaceutical compositions to be administered can contain minor amounts of non-toxic auxiliary substances, such as wetting or emulsifying agents, preservatives, and  
30 pH buffering agents and the like, for example sodium acetate or sorbitan monolaurate.

In a preferred embodiment the pharmaceutical composition further comprises an immunostimulatory substance such as an adjuvant. The adjuvant can be selected based on the method of administration and may include mineral oil-based adjuvants such as Freund's



- 30 -

complete and incomplete adjuvant, Montanide incomplete Seppic adjuvant such as ISA or ISA206 (SEPPIC, Paris, France), oil in water emulsion adjuvants such as the Ribi adjuvant system, syntax adjuvant formulation containing muramyl dipeptide, IC31<sup>®</sup> (Intercell; a synthetic adjuvant comprising the peptide motif KLK [WO 02/32451] and an oligonucleotide [WO 01/93905]), or aluminum salt adjuvants, preferably aluminum hydroxide or aluminum phosphate.

In a more preferred embodiment the immunostimulatory substance is selected from the group comprising polycationic polymers, especially polycationic peptides such as polyarginine, immunostimulatory deoxynucleotides (ODNs), especially Oligo(dIdC)<sub>13</sub>, peptides containing at least two LysLeuLys motifs, especially KLKLLLLLKLK, neuroactive compounds, especially human growth hormone, alum, adjuvants and combinations thereof. Preferably the combination is either a polycationic polymer and immunostimulatory deoxynucleotides or of a peptide containing at least two LysLeuLys motifs and immunostimulatory deoxynucleotides. In a still more preferred embodiment the polycationic polymer is a polycationic peptide.

The term "Oligo(dIdC)<sub>13</sub>" as used in the present invention means a phosphodiester backbone single-stranded DNA molecule containing 13 deoxy (inosine-cytosine) motifs, also defined by the term [oligo-d(IC)<sub>13</sub>]. The exact sequence is 5'-dIdCdIdCdIdCdIdCdIdCdIdCdIdCdIdCdIdCdIdCdIdCdIdC-3'. Oligo(dIdC)<sub>13</sub> can also be defined by the terms (oligo-dIC<sub>26</sub>); oligo-dIC<sub>26-mer</sub>; oligo-deoxy IC, 26-mer; or oligo-dIC, 26-mer, as specified for example in WO 01/93903 and WO 01/93905.

In an even more preferred embodiment of the invention the immunostimulatory substance is at least one immunostimulatory nucleic acid. Immunostimulatory nucleic acids are e.g. natural or artificial CpG containing nucleic acids, short stretches of nucleic acids derived from non-vertebrates or in form of short oligonucleotides (ODNs) containing non-methylated cytosine-guanine dinucleotides (CpG) in a defined base context (e.g. as described in WO 96/02555). Alternatively, also nucleic acids based on inosine and cytidine as e.g. described in WO 01/93903, or deoxynucleic acids containing deoxy-inosine and/or deoxyuridine residues (described in WO 01/93905 and WO 02/095027) may preferably be used as immunostimulatory nucleic acids in the present invention. Preferably, mixtures of different immunostimulatory nucleic acids are used in the present invention. Additionally,

- 31 -

the aforementioned polycationic compounds may be combined with any of the immunostimulatory nucleic acids as aforementioned. Preferably, such combinations are according to the ones described in WO 01/93905, WO 02/32451, WO 01/54720, WO 01/93903, WO 02/13857, WO 02/095027 and WO 03/047602.

5

In addition or alternatively, such a vaccine composition may comprise a neuroactive compound. Preferably, the neuroactive compound is human growth factor, e.g. described in WO 01/24822. Also preferably, the neuroactive compound is combined with any of the polycationic compounds and/or immunostimulatory nucleic acids as defined above.

10

In a highly preferred embodiment of the invention, the adjuvants are those used in the Examples, e.g. Complete Freund's adjuvant, aluminum hydroxide or/and an adjuvant comprising the KLKLLLLLKLK peptide and [dIdC]<sub>13</sub> phosphodiester ssDNA, such as IC31<sup>®</sup> (Intercell AG, Vienna, Austria; described above).

15

The composition may be used e.g. for immunization or treatment of a subject. The pharmaceutical composition encompasses at least one peptide or composition of the invention; however, it may also contain a cocktail (i.e., a simple mixture) containing different peptides and/or compositions of the invention, optionally mixed with different antigenic peptides or proteins of other pathogens. Such mixtures of these peptides, polypeptides, proteins or fragments or variants thereof are useful e.g. in the generation of desired antibodies to a wide spectrum of *S. agalactiae* isolates. The (poly)peptide(s)/composition(s) of the present invention may also be used in the form of a pharmaceutically acceptable salt. Suitable acids and bases which are capable of forming salts with the peptides of the present invention are well known to those of skill in the art, and include inorganic and organic acids and bases.

25

Still another subject of the invention relates to a pharmaceutical composition comprising

- (i) the one or more nucleic acid(s) according to the invention or one or more nucleic acid(s) complementary thereto, and
- (ii) optionally a pharmaceutically acceptable carrier or excipient.

30

The nucleic acid sequences, alone or in combination with other nucleic acid sequences encoding peptides/proteins/compositions or antibodies or directed to other pathogenic



- 32 -

microorganisms, may further be used as components of a pharmaceutical composition. The composition may be used for immunizing or treating humans and/or animals with the disease caused by infection with *S. agalactiae*.

5 The pharmaceutically acceptable carrier or excipient may be as defined above.

In another embodiment, the nucleic acid sequences of this invention, alone or in combination with nucleic acid sequences encoding other antigens or antibodies from other pathogenic microorganisms, may further be used in compositions directed to actively  
10 induce a protective immune response in a subject to the pathogen. These components of the present invention are useful in methods for inducing a protective immune response in humans and/or animals against infection with *S. agalactiae*.

For use in the preparation of the therapeutic or vaccine compositions, nucleic acid delivery  
15 compositions and methods are useful, which are known to those of skill in the art. The nucleic acids of the present invention or one or more nucleic acid(s) complementary thereto may be employed in the methods of this invention or in the compositions described herein as DNA sequences, either administered as naked DNA, or associated with a pharmaceutically acceptable carrier and provide for *in vivo* expression of the antigen,  
20 peptide or polypeptide. So-called "naked DNA" may be used to express the peptide or composition of the invention *in vivo* in a patient. (See, e.g., J. Cohen, Science, 259:1691-1692, which describes similar uses of "naked DNA"). For example, "naked DNA" associated with regulatory sequences may be administered therapeutically or as part of the vaccine composition e.g., by injection.

25

Alternatively, a nucleic acid encoding a peptide or composition of the invention or a nucleic acid complementary thereto may be used within a pharmaceutical composition, e.g. in order to express the peptide or composition of the invention *in vivo*, e.g., to induce antibodies.

30

A preferred embodiment of the invention relates to a pharmaceutical composition, wherein the nucleic acid is comprised in a vector and/or a cell other than *S. agalactiae*. Vectors and cells suitable in the context of the present invention are described above. Vectors are particularly employed for a DNA vaccine. An appropriate vector for delivery may be

- 33 -

readily selected by one of skill in the art. Exemplary vectors for *in vivo* gene delivery are readily available from a variety of academic and commercial sources, and include, e.g., adeno-associated virus (International patent application No. PCT/US91/03440), adenovirus vectors (M. Kay et al., Proc. Natl. Acad. Sci. USA, 91:2353 (1994); S. Ishibashi et al., J. Clin. Invest., 92:883 (1993)), or other viral vectors, e.g., various poxviruses, vaccinia, etc.. Recombinant viral vectors, such as retroviruses or adenoviruses, are preferred for integrating the exogenous DNA into the chromosome of the cell.

Another subject of the invention relates to a method for producing antibodies, characterized by the following steps:

- (a) administering an effective amount of the composition according to the invention and/or at least one protective peptide according to the invention to an animal; and
- (b) isolating the antibodies produced by the animal in response to the administration of step (a) from the animal.

15

A further subject of the invention relates to a method for producing antibodies, characterized by the following steps:

- (a) contacting a B cell with an effective amount of the composition according to the invention and/or at least one protective peptide according to the invention;
- (b) fusing the B cell of step (a) with a myeloma cell to obtain a hybridoma cell; and
- (c) isolating the antibodies produced by the cultivated hybridoma cell.

Also included in the scope of the invention is the production of antibodies against a peptide or composition according to the invention. This includes, for example, monoclonal and polyclonal antibodies, chimeric, single chain, and humanized antibodies, as well as Fab fragments, or the product of a Fab expression library, which are able to specifically bind to the peptide or composition according to the invention.

In a preferred embodiment the antibody is a monoclonal, polyclonal, chimeric or humanized antibody or functionally active fragment thereof. In another preferred embodiment the functionally active fragment comprises a Fab fragment.

Antibodies generated against the peptide or composition according to the invention can be obtained by direct injection of the peptide or composition according to the invention into



- 34 -

an animal or administering of the peptide or composition according to the invention to an animal, preferably a non-human. The antibody so obtained will then bind the peptide or composition according to the invention. Such antibodies can then be used to isolate reactive antigens, peptide or proteins from a tissue expressing those.

5

For preparation of monoclonal antibodies, any technique known in the art, which provides antibodies produced by continuous cell line cultures, e.g. a hybridoma cell line, can be used.

10

Techniques described for the production of single chain antibodies (U. S. Patent No. 4,946,778) can be adapted to produce single chain antibodies to the antigenic peptides or compositions according to the invention. Also, transgenic mice or other organisms such as other mammals may be used to express humanized antibodies to the antigenic peptides or compositions according to the invention.

15

Antibodies may be also produced using a hybridoma cell line. Hybridoma cell lines expressing desirable monoclonal antibodies are generated by well-known conventional techniques. The hybridoma cell can be generated by fusing a normal-activated, antibody-producing B cell with a myeloma cell. In the context of the present invention the hybridoma cell is able to produce an antibody specifically binding to the antigenic peptide or composition according to the invention.

20

Similarly, desirable high titer antibodies are generated by applying known recombinant techniques to the monoclonal or polyclonal antibodies developed to these peptides/proteins/compositions (see, e.g., PCT Patent Application No. PCT/GB85/00392; British Patent Application Publication No. GB2188638A; Amit et al., Science, 233:747-753 (1986); Queen et al., Proc. Natl. Acad. Sci. USA, 86:10029-10033 (1989); PCT Patent Application No. WO90/07861; Riechmann et al., Nature, 332:323-327 (1988); Huse et al., Science, 246:1275-1281 (1988)).

25

30

Particularly, the antibody may be produced by initiating an immune response in a non-human animal by administering a peptide or composition of the invention to an animal, removing an antibody-containing body fluid from said animal, and producing the antibodies by subjecting said antibodies containing body fluid to further purification steps.

Alternatively, the antibody may be produced by initiating an immune response in a non-human animal by administering a peptide or composition, as defined in the present invention, to said animal, removing the spleen or spleen cells from said animal and/or  
5 producing hybridoma cells of said spleen or spleen cells, selecting and cloning hybridoma cells specific for the peptide or composition according to the invention and producing the antibody by cultivation of said cloned hybridoma cells.

Alternatively, the antibody may be produced employing a phage display antibody library.  
10 The method is based on the selective binding of one or more members of a phage display antibody library to a surface-bound antigen. The method may e.g. be carried out as follows: an antigen of choice is immobilized to a solid surface, such as nitrocellulose, magnetic beads, a column matrix or, the most widely used, plastic surfaces as polystyrene tubes or 96-well plates. The antibody phages are incubated with the surface-bound antigen,  
15 followed by thorough washing to remove the excess nonbinders. The bound antibody phage can subsequently be eluted and e.g. amplified by infection of *Escherichia coli*. This method allows the detection of a single antibody phage and as it can be selected by e.g. its resistance marker, it can give rise to a bacterial colony after elution. The isolation of antibodies using phage display antibody libraries has been described in more details by  
20 Mancini et al., New Microbiol. 2004 Oct;27(4):315-328 and Pini et al., Curr Protein Pept Sci. 2004 Dec;5(6):487-496.

In a preferred embodiment the antibodies produced according to a method of the invention are additionally purified. Methods of purification are known to the skilled artisan.  
25

The antibody may be used in methods for treating an infection. Accordingly, still another subject of the invention relates to a pharmaceutical composition, especially a vaccine, comprising the antibody produced according to the invention. The pharmaceutical composition may encompass further components as detailed above. The composition may  
30 further encompass substances increasing their capacity to stimulate T cells. These include T helper cell epitopes, lipids or liposomes or preferred modifications as described in WO01/78767. Another way to increase the T cell stimulating capacity of epitopes is their formulation with immune stimulating substances for instance cytokines or chemokines like



- 36 -

interleukin-2, -7, -12, -18, class I and II interferons (IFN), especially IFN-gamma, GM-CSF, TNF-alpha, flt3-ligand and others.

Another subject of the invention relates to a mixture of antibodies against the at least two  
5 proteins of the composition according to the invention and/or against the at least one protective peptide according to the invention. The mixture of antibodies may be further characterized and produced as described above.

Methods of producing antibodies, mixtures of antibodies, as well as the use of antibodies  
10 are also described in Examples 4 and 5, and Figures 4, 8, and 10 to 13.

Another subject of the invention relates to the use of the composition according to the invention and/or at least one protective peptide according to the invention and/or one or more of the nucleic acid(s) according to the invention for the manufacture of a medicament  
15 for the immunization or treatment of a subject, preferably against *S. agalactiae*, more preferably against pneumonia, septicemia, meningitis, fever, vomiting, poor feeding, irritability, urinary tract infection and/or vaginal infection caused by *S. agalactiae*.

The peptides, proteins, compositions or the nucleic acids of the invention are generally  
20 useful for inducing an immune response in a subject. The vaccine used for immunization may be administered to a subject susceptible to infection by *S. agalactiae*, preferably mammals, and still more preferably humans, in any conventional manner, including oral, topical, intranasal, intramuscular, intra-lymph node, intradermal, intraperitoneal, subcutaneous, and combinations thereof, but most preferably through intramuscular  
25 injection. The volume of the dose for intramuscular administration is preferably up to about 5 ml, still more preferably between 0.5 ml and 3 ml, and most preferably about 1 to 2 ml. The volume of the dose when subcutaneous injection is the selected administration route is preferably up to about 5 ml, still more preferably between 0.5 ml and 3 ml, and most preferably about 1 to 2 ml. The amount of substance in each dose should be enough  
30 to confer effective immunity against and decrease the risk of developing clinical signs resulting from *S. agalactiae* infection to a subject receiving a vaccination therewith. Preferably, the unit dose of protein should be up to about 5 µg protein/kg body weight, more preferably between about 0.2 to 3 µg, still more preferably between about 0.3 to 1.5 µg, more preferably between about 0.4 to 0.8 µg, and still more preferably about 0.6 µg.

- 37 -

Alternative preferred unit doses of protein could be up to about 6 µg protein/kg body weight, more preferably between about 0.05 to 5 µg, still more preferably between about 0.1 to 4 µg. The dose is preferably administered 1 to 3 times, e.g. with an interval of 1 to 4 weeks. Preferred amounts of protein per dose are from approximately 1 µg to  
5 approximately 1 mg, more preferably from approximately 5 µg to approximately 500 µg, still more preferably from approximately 10 µg to approximately 250 µg and most preferably from approximately 25 µg to approximately 100 µg.

In still another aspect of the invention the mixture of antibodies or the antibody produced  
10 according to the invention or functional fragment thereof is used for the manufacture of a medicament for the treatment of an infection, preferably a *S. agalactiae* infection. The treatment involves administering an effective amount of the antibody to a subject, preferably a mammal, more preferably a human. Thus, antibodies against the peptides or the composition of the present invention may be employed to inhibit and/or treat  
15 infections, particularly bacterial infections and especially infections arising from *S. agalactiae*.

An "effective amount" of peptides, proteins, compositions or the nucleic acids of the invention or an antibody produced according to the invention may be calculated as that  
20 amount capable of exhibiting an *in vivo* effect, e.g. preventing or ameliorating a sign or symptom of infection, particularly *S. agalactiae* infection. Such amounts may be determined by one of skill in the art. Such a substance may be administered in any conventional manner, including oral, topical, intranasal, intramuscular, intra-lymph node, intradermal, intraperitoneal, subcutaneous, and combinations thereof, but preferably  
25 intramuscularly or subcutaneously. However, it may also be formulated to be administered by any other suitable route, including orally or topically. The selection of the route of delivery and dosage of such therapeutic compositions is within the skill of the art.

Treatment in the context of the present invention refers to both therapeutic treatment and  
30 prophylactic or preventive measures, wherein the object is to prevent or slow down (lessen) the targeted pathologic condition or disorder. Those in need of treatment include those already with the disorder as well as those prone to have the disorder or those in whom the disorder is to be prevented.



- 38 -

Another subject of the invention relates to a method of diagnosing a *S. agalactiae* infection comprising the steps of:

- (a) contacting a sample obtained from a subject with the composition according to the invention and/or at least one protective peptide according to the invention; and
- 5 (b) detecting the presence of an antibody against the protective peptide, the functionally active variant and/or the composition in the sample,

wherein the presence of the antibody is indicative for the *S. agalactiae* infection.

Another subject of the invention relates to a method of diagnosing a *S. agalactiae* infection comprising the steps of:

- (a) contacting a sample obtained from a subject with the mixture of antibodies according to the invention; and
- (b) detecting the presence of the at least two proteins of the composition according to the invention and/or of the at least one protective peptide according to the invention in  
15 the sample,

wherein the presence of the at least two proteins and/or of the at least one protective peptide is indicative for the *S. agalactiae* infection.

The protective peptides or compositions of the invention or alternatively a mixture of  
20 antibodies may be used for the detection of *S. agalactiae*. Preferably such detection is for diagnosis, more preferably for the diagnosis of a disease, most preferably for the diagnosis of a *S. agalactiae* infection. The protective peptides or compositions may be used to detect the presence of a *S. agalactiae*-specific antibody or fragment thereof e.g. in a sample obtained from a subject. Alternatively, the mixture of antibodies may be used to detect the  
25 presence of *S. agalactiae* proteins, e.g. in a sample obtained from a subject. The sample may be e.g. a blood sample.

The present invention also relates to diagnostic assays such as quantitative and diagnostic assays for detecting levels of the proteins, compositions and/or mixtures of antibodies of  
30 the present invention in cells and tissues or body fluids, including determination of normal and abnormal levels. Assay techniques that can be used to determine levels of a peptide, a composition or an antibody, in a sample derived from a host are well known to those of skill in the art. Such assay methods include radioimmunoassays, competitive-binding assays, Western Blot analysis and ELISAs. Among these, ELISAs frequently are preferred.

- 39 -

An ELISA initially comprises preparing an antibody or antibodies specific to the peptide or composition, preferably a monoclonal antibody. In addition, a reporter antibody generally is prepared which binds to the monoclonal antibody. The reporter antibody is attached to a detectable reagent such as radioactive, fluorescent or enzymatic reagent, such as  
5 horseradish peroxidase enzyme.

The peptides or compositions of the present invention may also be used for the purpose of or in connection with an array. More particularly, at least one of the peptides or compositions of the present invention may be immobilized on a support. Said support  
10 typically comprises a variety of peptides/proteins whereby the variety may be created by using one or several of the peptides or compositions of the present invention. The characterizing feature of such array as well as of any array in general is the fact that at a distinct or predefined region or position on said support or a surface thereof, a distinct polypeptide is immobilized. Because of this any activity at a distinct position or region of  
15 an array can be correlated with a specific polypeptide. The number of different peptides or antibodies of the present invention immobilized on a support may range from as little as 10 to several 1000 different peptides or compositions of the present invention. Alternatively, antibodies produced according to the present invention may be used to detect peptides or compositions of the invention.

20

The manufacture of such arrays is known to the one skilled in the art and, for example, described in US patent 5,744,309. The array preferably comprises a planar, porous or non-porous solid support having at least a first surface. Preferred support materials are, among others, glass or cellulose. It is also within the present invention that the array is used for  
25 any of the diagnostic applications described herein. Apart from the peptides or antibodies of the present invention also the nucleic acid molecules according to the present invention may be used for the generation of an array as described above.

Another subject of the invention relates to a method for diagnosing an infection with *S. agalactiae* comprising the steps of:  
30

- (a) contacting a sample obtained from a subject with a primer and/or a probe specific for the one or more nucleic acid(s) according to the invention; and
- (b) detecting the presence of one or more nucleic acid(s) according to the invention in the sample,



- 40 -

wherein the presence of the one or more nucleic acid(s) is indicative for the *S. agalactiae* infection.

5 A series of methods for detecting nucleic acids in samples by using specific primers and/or probes is known in the art. In general, these methods are based on the specific binding of a primer or probe to the nucleic acid in question. The methods may involve amplification of the nucleic acid, e.g. RNA or DNA, before the actual detection step. Therefore, primers may be used to specifically induce transcription and/or amplification of RNA or DNA in order to generate a detectable amount of nucleic acid. Suitable well known techniques may be PCR and RT-PCR. Suitable primers and probes for the method of the invention may be produced based on sequence information provided in the present application. Guidelines and computer-assisted programs (e.g. Primer Express<sup>®</sup>, Applied Biosystems, Foster City, CA, USA) for designing primers and probes to a specific nucleic acid are known to the person skilled in the art.

15

After the amplification step the amplified nucleic acid, in general DNA, may be detected e.g. by its size (e.g. involving agarose gel electrophoresis) or using labeled probes which specifically bind to the amplified nucleic acid. The probes may be labeled with a dye, radioactive marker, a fluorescent marker, an enzyme-linked marker or any other marker.

20

For example, FRET (Förster resonance energy transfer) may be used for the detection of the nucleic acid of the invention. In FRET, a donor fluorophore molecule absorbs excitation energy and delivers this via dipole-dipole interaction to a nearby acceptor fluorophore molecule. This process only occurs when the donor and acceptor molecules are sufficiently close to one another. Several different strategies for determining the optimal physical arrangement of the donor and acceptor moieties are known to the skilled practitioner. For this, a fluorescent donor is excited at its specific fluorescence excitation wavelength. By a long-range dipole-dipole coupling mechanism, this excited state is then nonradiatively transferred to a second molecule, the acceptor. The donor returns to the electronic ground state. The described energy transfer mechanism is termed "Förster resonance energy transfer" (FRET). The process involves measuring fluorescence as FRET donor and acceptor moieties are brought together as a result of DNA hybridization. For examples two probes each labeled with a suitable marker hybridize to the nucleic acid of the invention within a distance which allows FRET to occur. Suitable markers include

25  
30

- 41 -

Cyan 500, Cy5, Cy3, SYBR Green I, fluorescein, HEX, Red 610 and Red 640, wherein the two marker involved have to be selected based on there excitation and emission spectrums as known by the skilled person. A suitable system for the detection of nucleic acids is the LightCycler<sup>®</sup> (Roche Diagnostics).

5

Another subject of the invention relates to a method for identifying a ligand capable of binding the composition according to the invention and/or at least one protective peptide according to the invention comprising:

- (a) providing a test system comprising the peptide and/or composition,
- 10 (b) contacting the test system with a test compound, and
- (c) detecting a signal generated in response to the binding of the test compound to the peptide and/or composition.

More particularly, the method may be carried out by contacting an isolated or immobilized protective peptide or composition according to the invention with a candidate ligand under  
15 conditions to permit binding of the candidate ligand to the peptide, wherein the test system comprises a component capable of providing a detectable signal in response to the binding of the candidate ligand to said peptide; and detecting the presence or absence of a signal generated in response to the binding of the ligand to the peptide. The ligand may be an agonist or an antagonist.

20

Test systems for detection binding of a ligand are known to the skilled artisan and include e.g. binding assays with labeled ligand such as radioligands, fluorescence-labeled ligands or enzyme-labeled ligands.

25 The test compound can be any test compound either naturally occurring or chemically synthesized. Naturally occurring test compounds include in particular antibodies, preferably those showing similarity to the antibodies of the invention. In one preferred embodiment of the invention the test compound is provided in the form of a chemical compound library. Chemical compound libraries include a plurality of chemical  
30 compounds and have been assembled from any of multiple sources, including chemically synthesized molecules and natural products, or have been generated by combinatorial chemistry techniques. They are especially suitable for high throughput screening. They may be comprised of chemical compounds of a particular structure or compounds of a particular creature such as a plant.



A further subject of the invention relates to the use of the composition according to the invention and/or at least one protective peptide according to the invention for the isolation and/or purification and/or identification of an interaction partner of the composition and/or peptide. The isolation and/or purification and/or identification of the ligand may be carried out as detailed above or as known to the person skilled in the art. In a preferred embodiment of the invention an affinity device may be used. The affinity device may comprise at least a support material and any antigenic peptide or composition according to the present invention, which is attached to the support material. Because of the specificity of the protective peptides and/or compositions according to the present invention for their target cells or target molecules or their interaction partners, the peptides and/or compositions allow a selective removal of their interaction partner(s) from any kind of sample applied to the support material provided that the conditions for binding are met. The sample may be a biological or medical sample, including but not limited to, fermentation broth, cell debris, cell preparation, tissue preparation, organ preparation, blood, urine, lymph liquid, liquor and the like. The peptide or composition may be attached to the matrix in a covalent or non-covalent manner. Suitable support material is known to the one skilled in the art and can be selected from the group comprising cellulose, silicon, glass, aluminium, paramagnetic beads, starch and dextrane.

20

The present invention is further illustrated by the following Figures, Examples and the Sequence listing, from which further features, embodiments and advantages may be taken. It is to be understood that the present examples are given by way of illustration only and not by way of limitation of the disclosure.

25

## FIGURE LEGENDS

**Figure 1** shows the protection achieved by passive immunization with selected hyper-immune rabbit sera generated by immunization of rabbits with *S. agalactiae* antigens in a mouse lethality model. CD-1 mice (10 mice per group) were immunized intraperitoneally with 150 µl hyper-immune rabbit sera 1 – 3 hours before they were intraperitoneally challenged. (A) gbs1309p, gbs1478p, gbs2018p, Sip and PBS-induced hyperimmune sera,

30

- 43 -

challenge with  $1 \times 10^7$  cfu C388/90 (serotype Ia/c). **(B)** gbs1478p, gbs2018p, Sip and PBS-induced hyperimmune sera, challenge with  $5 \times 10^6$  cfu ATCC12401 (serotype Ib). **(C)** gbs0233p, gbs1087p, gbs1477p, Sip and PBS-induced hyperimmune sera, challenge with  $1 \times 10^8$  cfu ATCC12403 (serotype III). **(D)** gbs0233p, gbs1087p, gbs2018p, Sip and PBS-induced hyperimmune sera, challenge with  $1 \times 10^8$  cfu ATCC49447 (serotype V). Survival was monitored for 14 days post-challenge. Numbers of surviving mice are plotted as percentage of total mice.

**Figure 2** shows the protection achieved with a combination of *S. agalactiae* antigen-specific hyperimmune rabbit sera in a mouse lethality model. CD-1 mice (10 mice per group) were immunized intraperitoneally with different combinations of hyperimmune rabbit sera (150  $\mu$ l per hyper-immune rabbit sera) 1 – 3 hours before an intraperitoneal challenge. Sip- and PBS-induced sera were used as positive and negative controls, respectively. Mice were immunized with combinations of sera induced by gbs1087p, gbs1477p, gbs1478p and gbs2018p; challenge with **(A)**  $5 \times 10^6$  cfu ATCC12401 (serotype Ib); **(B)**  $1 \times 10^8$  cfu ATCC12403 (serotype III) and **(C)**  $1 \times 10^8$  cfu ATCC49447 (serotype V). Survival was monitored for 11 days post-challenge. Numbers of surviving mice are plotted as percentage of total mice.

**Figure 3** shows the surface staining of the serotype III GBS strain ATCC12403. The results for the hyperimmune rabbit sera (black) are shown in comparison to those for the respective preimmune sera (white).

**Figure 4** shows an opsonophagocytic killing assay with hyperimmune rabbit sera and different GBS strains. **(A)** Serotype II GBS strain AC3912, not suitable for animal testing. **(B)** Serotype V GBS strain BAA23, used for animal testing. PI, preimmune sera; HI, hyperimmune sera. GBS cells in the exponential phase were opsonised with 200-fold diluted sera in the presence of 5% guinea pig complement for 60 minutes. Phagocytic cells (RAW264.7) were added to opsonised bacteria and incubated for an additional 60 minutes at 37°C. Surviving bacteria were counted on agar plates after overnight incubation at 37°C. Percentage of killing was calculated based on CFU obtained after incubation with the different hyperimmune sera relative to CFU obtained with preimmune sera at 0 min and after 60 min of incubation as described under experimental procedure.



- 44 -

**Figure 5 A-D** show the sequence alignment of protein sequences homologous to gbs1477 from genomic and sequenced strains. Alignment of sequences was performed using the software from Vector NTI (Suite 7.1; Invitrogen, Austria). The name on the left of the sequence indicates the strain name. Amino acids in bold, residue identical in at least 50% of sequences. \*, indicates position of STOP codon.

**Figure 6** shows active protection of adult mice by immunizing with a combination of GBS antigens. CD-1 mice (10 mice per group) were immunized with different combinations of the recombinant proteins (gbs1087p, gbs1477p, gbs1478p and gbs2018p; 25 µg each). As positive control 25 µg Sip protein (open circle) was used. For the negative control, PBS (open square) was used with ALUM 1%. One week after the last booster immunization, mice were challenged with **(A)**  $3.5 \times 10^6$  cfu 12401; **(B)**  $8.8 \times 10^7$  cfu 12403 or with **(C)**  $1.1 \times 10^8$  cfu 49447. Numbers of surviving mice are plotted as a percentage of the total number of mice.

**Figure 7** shows active protection of adult mice by immunizing with single GBS antigens. CD-1 mice (10 mice per group) were immunized with different recombinant proteins (gbs0233p, gbs1087p, gbs1309p, gbs1477p, gbs1478p and gbs2018p; 50 µg each). As positive control 50 µg Sip protein (open circle) was used. For the negative control, PBS (open square) was used with ALUM 1%. One week after the last booster immunization, mice were challenged with **(A)**  $3.5 \times 10^6$  cfu 12401; **(B)**  $8.8 \times 10^7$  cfu 12403 or with **(C)**  $1.1 \times 10^8$  cfu 49447. Numbers of surviving mice are plotted as a percentage of the total number of mice.

**Figure 8** shows the protection of neonatal mice by immunization of mothers passively with rabbit sera generated with a combination of recombinant GBS antigens. Pregnant CD-1 mice were immunized at day 18 post gestation with 500 µl of combinations of gbs1087p, gbs1477p, gbs1478p and gbs2018p-induced sera or PBS-induced control sera. Neonates were challenged within 24 – 38 hours after birth with lethal challenge doses of **(A)**  $1.2 \times 10^7$  cfu C388/90; **(B)**  $1.3 \times 10^6$  cfu 12403; **(C)**  $5.7 \times 10^6$  cfu BAA23 or **(D)**  $1.8 \times 10^8$  cfu 2603V/R. Numbers of surviving neonates are plotted as a percentage of the total number of challenged neonates.

- 45 -

**Figure 9** shows active protection of adult mice by immunizing with different clade gbs1477p proteins. CD-1 mice (10 mice per group) were immunized with different clade proteins of gbs1477p (gbs1477p-2603V/R, gbs1477p-49447, gbs1477p-C388/90, gbs1477p-126H4A and gbs1477p-NEM316; 50 µg each). As negative control, PBS (open square) was used with ALUM 1%. One week after the last booster immunization, mice were challenged with **(A)**  $1.4 \times 10^7$  cfu C388/90; **(B)**  $1.2 \times 10^8$  cfu 12403 or with **(C)**  $1.6 \times 10^8$  cfu BAA23. Numbers of surviving mice are plotted as a percentage of the total number of mice.

**Figure 10** shows the protection of adult mice by immunization with mouse mAbs against different recombinant GBS antigens. CD-1 mice were immunized intraperitoneally with 50 µg of the respective mouse mAb. 1 to 3 hours later, mice were challenged intraperitoneally with **(A)**  $1.2 \times 10^8$  cfu 12403; **(B)**  $1.5 \times 10^8$  cfu 12403 or **(C)**  $1.1 \times 10^8$  cfu 49447. Numbers of surviving mice are plotted as a percentage of the total number of challenged mice.

15

**Figure 11** shows the protection of adult mice by immunization with a combination of five mouse mAbs against different recombinant GBS antigens. CD-1 mice were immunized intraperitoneally with 25 µg of each mouse mAb. 1 to 3 hours later, mice were challenged intraperitoneally with **(A)**  $1.2 \times 10^7$  cfu C388/90; **(B)**  $9.6 \times 10^7$  cfu 12403 or **(C)**  $1.7 \times 10^8$  cfu BAA23. Numbers of surviving mice are plotted as a percentage of the total number of challenged mice.

25

**Figure 12** shows surface staining of the serotype III GBS strain ATCC12403. The results for the monoclonal antibodies (black) are shown with the buffer control (white).

**Figure 13** shows opsonophagocytic killing assay with hyperimmune rabbit sera and different GBS strains. Rabbit sera were tested in the opsonophagocytic killing assay at a serum dilution of 1:1,000, % killing was calculated in relation to the respective pre-immune sera. The mouse monoclonal antibodies were tested with two different amounts added to the opsonophagocytic killing assay, 1.0 and 0.1 µg. The % killing for the monoclonal antibodies was calculated in relation to the complement control.

30

## EXAMPLES



**Example 1: Group B streptococcal antigens and combinations thereof inducing protective immune responses against lethal sepsis in an i.p. challenge model.**

5 ***Experimental procedures***

*Cloning and expression of recombinant Group B streptococcal proteins*

Cloning of genes: The gene of interest was amplified from genomic DNA of *S. agalactiae*  
 10 ATCC12403 (serotype III) by PCR using gene specific primers. Apart from the gene  
 specific part, the primers had restriction sites that aided in a directional cloning of the  
 amplified PCR product. The gene annealing (specific) part of the primer ranged between  
 15 15-30 bases in length. The PCR products obtained were digested with the appropriate  
 restriction enzymes and cloned into the pET28b (+) vector (Novagen) for His-tagged  
 proteins. Once the recombinant plasmid was confirmed to contain the gene of interest, *E.*  
*coli* BL21 star<sup>®</sup> cells (Invitrogen) that served as expression host were transformed. Cloning  
 of the gbs1087, gbs1477 and gbs1478 genes has been performed using genomic DNA from  
 strain *S. agalactiae* 6313 (serotype III) in the vector pET28a (+). The origin of the gene and  
 position within the full length gene of the selected antigens are listed in Table 1. The  
 20 amino acid and nucleic acid sequences are as follows:

Amino acid sequences:

**SEQ ID NO: 1**

25 **Construct 1: gbs0233p**

LCLALLTISGCQLTDTKKPGHTTIKVAQAQSSSTESSIMANIVTELIHHELGYNTTLISNLGS  
 STVTHQALLRGDADIAATRYTGTDITGTLGLKAVKDTKEASKIVKTEFQKRYNQTWYPTYG  
 FSDTYAFMVTKEFARQNKIKITKISDLKKLSTTMKAGVDSWMMNREGDGYTDFAKTYGFESH  
 IYPMQIGLVYDAVESNKMQSVLGYSTDGRISYDLEILRDDKKFFPPYEASMVVNSIIKK  
 30 DPKLKKLLHRLDGKINLKTMQNLNYMVDDKLEPSVVAKQFLEKNHYFRGD

**SEQ ID NO: 2**

**Construct 2: gbs1087p**

MDSVGNQSQGNVLERRQRDAENRSQGNVLERRQRDVENKSQGNVLERRQRDAENKSQGNVL  
 35 ERRQRDAENRSQGNVLERRQRDAENRSQGNVLERRQRDAENRSQGNVLERRQRDAENRSQGNV  
 NVLERRQRDAENRSQGNVLERRQRDVENKSQGNVLERRQRDAENKSQGNVLERRQRDAENR  
 SQGNVLERRQRDAENRSQGNVLERRQRDAENRSQGNVLERRQRDAENRSQGNVLERRQRDA  
 ENRSQGNVLERRQRDAENRSQGNVLERRQRDAENRSQGNVLERRQRDAENRSQGNVLERRQ  
 RDAENKSQVGQLIG

40

**SEQ ID NO: 3**

- 47 -

**Construct 3: gbs1309p**

SVTYSQSERTVVFSFGEITFSRSRWTNGFETRI PVDEWLGLEKYKRY SIEFLYHVAKLATM  
 MPYRQVCKVIDSTLQTIITKDCVLKAVKFVEKLLKEKERYRFYLEEPERKKVKKLYVEGD  
 GVMIKSTDSREERRYLDLTHFVIHTGSKKVSTKRYELQDKHEILQLNYDKAKYNLLDYIYN  
 5 NYEVDDDTILITNSDMGKGYTSRVFKELGKALKVKKHEHFWDIYHVKEKLS SYLRKYPIEL  
 TDFALDAVKKYNSDKLELVFDTVESLICDELEDQEFQKFKKKVLNNFKYIKPAHLRNLSNR  
 GIGIMESQHRKITYRMKRRGMYWSKWGI STMANMI ILERANGLRELF FGSWRKVYSEYKEG  
 SFSAGRLFKKTDELDFSKPLLKNRGRKWSITGIK

10 **SEQ ID NO: 4****Construct 4: gbs1477p**

DDVTTDTVTLHKIVMPQAAFDNFTEGTGKKNDS DYVGKQINDLKS YFGSTDAKEIKGAFFV  
 FKNETGTFITENGKEVD TLEAKDAEGGAVLSGLTKDTGFAFNTAKLKGTYQIVELKEKSN  
 YDNNGSILADSKAVPVKITLPLVNNQGVV KDAHIYPKNTETKPQVDKNFADKDLDYTDNRK  
 15 DKGVVSATVGDKKEYIVGTKILKGS DYKKLVWTD SMTKGLTFNNNVKVTLDGKDFPVLNYK  
 LVTDDQGFRLALNATGLAAVAAA AKDKDVEIKITYSATVNGSTTVEVPETNDVKLDYGNNP  
 TEESEPQEGTPANQEIKVIKDWAVDGTITDVNVAVKAI FTLQEKQTDGTWVNVASHEATKP  
 SRFEHTFTGLDNTKTYRVVERVSGYTP EYVSFKNGVVTIKNNKNSNDPTPINPSEPKVVTY  
 GRKFVKTNQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAKLALDEAVKAYNDL  
 20 TKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKKLISNAGGQFEITG  
 LDKGTYSLEETQAPAGYATLSGDVNF EVTATSYSKGATTDIAYDKGSVKKDAQQVQNKKVT  
 IPQTGGIGT

**SEQ ID NO: 5**25 **Construct 5: gbs1478p**

ESTVPENGA KGLVVKKTDDQNKPLSKATFVLKTTAH PESKIEKVTAELTGEATFDNLI PG  
 DYT LSEETAPEGYKKTNQ TWQVKVESNGKTTIQNSGDKNSTIGQNHEELDKQYPPTGIYED  
 TKESYKLEHVKG SVPNGKSEAKAVNPYSSEGEHIREIPEGTLSKRISEVGD LAHNKYKIEL  
 TVSGKTIVKPV DKQKPLDVVFLDNSNSMNN DGNFQRHNKAKKAAEALGTAVKDILGANS  
 30 DNRVALVTYGS DIFDGRSVDVVKGFKEDDKYYGLQTKFTIQTENYSHKQLTNNAEEI IKRI  
 PTEAPRAK WGSTTNGLTPEQQKQYYLSKVGETFTMKAFMEADDILSQVDRNSQKIIVHITD  
 GVPTRSYAINNF KLGASYESQFEQMKN GYLNKSNFLLDKPEDIKNGESYFLFPLDSYQ  
 TQIISGNLQKLHYLDLNLNYPKGTIYRNGPVREHGTPTKLYINSLKQKNYDIFNFGIDISA  
 FRQVYNEDYKKNQDGT FQKLKEEAFELSDGEITELMKSFS SKPEYYTPIVTS SDASNNEIL  
 35 SKIQQQFEKVLTKENSIVNGTIEDPMGDKINLQLGNGQTLQPSDYTLQGNDGSIMKDSIAT  
 GGPNNDDGGILKGVKLEYIKNKLYVRGLNLGEGQKVTLT YDVKLDDSFISNKFYDTNGRTTL  
 NPKSEDPNTLRDFPIPKIRDVREYPTITIKNEKKLGEIEFTKVDKDNNKLLLKGATFELQE  
 FNEDYKLYLP IKNNSKVVTGENGKISYKDLKDGKYQLIEAVSPKDYQKITNKPILT FEVV  
 KGSIQNI IAVNKQISEYHEEGDKHLITNTHIPP KGIIPMTGGK

40

**SEQ ID NO: 6****Construct 6: gbs2018p**

DTSSGISASIPHKKQVNLGAVTLKNLISKYRGNDKAI AILLSRVNDFNRSQDTLPQLINS  
 TEAEIRNILYQGQIGKQNKPSVTT HAKVSDQELGKQSRRSQDI IKS LGFLSSDQKDILVKS  
 45 ISSSKDSQLILKFVTQATQLNNAESTKAKQMAQNDVALIKNISPEVLEEYKEKIQRASTKS  
 QVDEFVAEAKKVNSNKETLVNQANGKKQEIAKLENLSNDEMLRYNTAIDNVVKQYNEGKL  
 NITAAMNALNSIKQAAQEVAQKNLQKQYAKKIERISSKGLALS KKAKEIYEKHK SILPTPG  
 YYADSVGTYLNRFRDKQTFGNRSVWTGQSGLDEAKKMLDEVKLLKELQDLTRGTKEDKKP  
 DVKPEAKPEAKPNIQVPKQAPTEAAKPALSPEALTRLTTWYNQAKDLLKDDQVKDKYVDIL  
 50 AVQKAVDQAYDHVEEGKFITTDQANQLANKLRDALQSLELKD KKVAKPEAKPEAKPEAKPE  
 AKPEAKPEAKPEAKPEAKPDVKPEAKPDVKPEAKPEAKPEAKSEAKPEAKLEAKPEAKPAT



- 48 -

KKSVENTSGNLAACKAIENKKYSKKLPST

Nucleic acid sequences:5 **SEQ ID NO: 7****Construct 1: gbs0233p**

CTTTGCTTAGCCCTCTTAACGATTTCTGGTTGTCAATTAACCGATACTAAAAACCTGGTC  
ATACCACAATTAAGGTTGCTGCCCCAAAGTTCTACAGAGTCTAGTATCATGGCAAATATTGT  
CACCGAATTAATTCATCACGAATTAGGATACAACAACCTTTAATAAGCAATCTTGGTTCC  
10 TCTACGGTTACTCACCAAGCTTTGCTCCGTGGTGATGCTGACATTGCTGCCACACGTTATA  
CAGGAACAGACATCACAGGAACCTTTGGCTTAAAAGCTGTTAAAGACACTAAAGAAGCTTC  
TAAGATTGTAAAAACTGAATTCCAAAAACGCTACAATCAAACCTGGTATCCTACTTATGGT  
TTTTCTGATACTTATGCATTCATGGTTACTAAAGAGTTTGCCAGACAGAATAAAATCACCA  
AGATCTCTGATCTCAAAAAGTTATCAACAACCTATGAAGGCAGGGGTTGATAGTTCATGGAT  
15 GAATCGCGAGGGAGATGGATACACTGATTTTCGCTAAAACATACGGTTTTGAATTTTCACAT  
ATTTACCCTATGCAAATTGGCTTAGTCTATGATGCAGTTGAAAGTAACAAAATGCAATCTG  
TATTAGGCTACTCCACTGACGGTCGTATTTTCGAGCTATGATTTAGAAATTTTAAGGGATGA  
TAAAAAATTCTTTCCTCCTTATGAAGCCTCTATGGTTGTCAACAATTCTATCAAAAAA  
GATCCTAAACTAAAAAATTACTCCATCGACTCGATGGTAAAATCAATTTAAAAACGATGC  
20 AAAACCTTAATTATATGGTAGATGATAAACTTTTAGAACCTTCAGTTGTTGCCAAACAATT  
TTTAGAAAAAACCATTATTTTAGAGGAGAT

**SEQ ID NO: 8****Construct 2: gbs1087p**

ATGGATAGTGTTGGAAATCAAAGTCAGGGCAATGTTTTAGAGCGTCGTCAACGTGATGCAG  
AAAACAGAAGCCAAGGCAATGTTCTAGAGCGTCGTCAACGCGATGTTGAGAATAAGAGCCA  
AGGCAATGTTTTAGAGCGTCGTCAACGTGATGCGGAAAACAAGAGCCAAGGCAATGTTTTA  
GAGCGTCGTCAACGTGATGCAGAAAACAGAAGCCAAGGCAATGTTCTAGAGCGTCGTCAAC  
GTGATGCAGAAAACAGAAGCCAAGGCAATGTTCTAGAGCGTCGTCAACGCGATGCAGAAAA  
30 CAGAAGCCAAGGTAATGTTCTAGAGCGTCGTCAACGTGATGCAGAAAACAGAAGCCAAGGT  
AATGTTCTAGAGCGTCGTCAACGTGATGCAGAAAACAGAAGCCAAGGTAATGTTCTAGAGC  
GTCGTCAACGCGATGTTGAGAATAAGAGCCAAGGCAATGTTTTAGAGCGTCGTCAACGTGA  
TGCGGAAAACAAGAGCCAAGGCAATGTTTTAGAGCGTCGTCAACGTGATGCAGAAAACAGA  
AGCCAAGGCAATGTTTTAGAGCGTCGTCAACGTGATGCAGAAAACAGAAGCCAAGGCAATG  
35 TTCTAGAGCGTCGTCAACGTGATGCAGAAAACAGAAGCCAAGGCAATGTTCTAGAGCGTCG  
TCAACGTGATGCAGAAAACAGAAGCCAAGGCAATGTTCTAGAGCGTCGTCAACGCGATGCA  
GAAAACAGAAGCCAAGGTAATGTTCTAGAGCGTCGTCAACGTGATGCAGAAAACAGAAGCC  
AAGGCAATGTTTTAGAGCGTCGTCAACGTGATGCAGAAAACAGAAGCCAAGGCAATGTTTT  
AGAGCGTCGTCAACGTGATGCAGAAAACAGAAGCCAAGGCAATGTTTTAGAGCGTCGTCAA  
40 CGTGATGCGGAAAACAAGAGCCAAGTAGGTCAACTTATAGGG

**SEQ ID NO: 9****Construct 3: gbs1309p**

AGTGTAACCTATTCACAGTCTGAACGTACGGTTGTTTTCTCTTTTGGAGAAATAACATTTA  
45 GTAGGAGTCGCTGGACAAATGGCTTTGAAACTAGAATACCAGTAGATGAGTGGTTAGGTCT  
TGAAAAATATAAGAGATATTCAATAGAATTCTTATATCATGTTGCAAATTTGGCTACAATG  
ATGCCTTATCGTCAAGTTTGCAAAGTAATAGATAGCACTTTGCAAACAATCATAACAAAAG  
ACTGTGTTTTTAAAAGCAGTAAAATTTGTAGAAAAATTGTTAAAAGAAAAAGAACGCTATCG  
TTTTTATTTGGAAGAGCCACCCGAACGTAAAAAAGTGAAAAAAGTGTATGTTGAGGGTGAT  
50 GGAGTCATGATTAAAAGCACAGATTCTAGAGAGGAAAGAAGGTATTTAGATTTAACACATT  
TTGTTATTCATACAGGCTCAAAAAAAGTTTCTACTAAAAGATATGAATTGCAGGACAAGCA



- 49 -

CGAAATATTACAGCTTAATTATGATAAAGCTAAATATAATCTTTTAGATTATATTTATAAT  
AACTATGAAGTAGATGACGATACTATTTTAATCACTAACTCTGATATGGGTAAAGGCTATA  
CTAGTAGAGTTTTTAAGGAATTAGGAAAAGCACTTAAGGTAAAGAAACATGAGCATTTTTG  
GGATATCTATCATGTTAAAGAAAAGTTAAGTTCATACCTTAGAAAATATCCAATTGAATTA  
5 ACCGATTTTGCTTTAGATGCGGTAAAAAATATAATTCTGATAAGCTTGAATTAGTTTTTG  
ATACTGTTGAATCACTGATTTGTGATGAACTTGAAGATCAAGAATTTTCAGAAGTTTAAGAA  
AAAAGTATTAAATAATTTCAAATATATAAAACCAGCTCATCTTAGAAATCTTTCAAATCGT  
GGTATTGGTATCATGGAATCACAAACACAGAAAGATAACGTATAGAATGAAGCGACGTGGCA  
TGTATTGGTCAAAGTGGGGAATCTCCACAATGGCAAATATGATTATACTTGAAAGAGCTAA  
10 CGGTTTACGAGAATTATTTTTTCGGTTCCTTGGAGAAAGGTATACAGTGAGTATAAAGAAGGT  
TCATTTAGTGCAGGGCGACTTTTTTAAAAAGACAGATGAATTAGATAAAATTTTCTAAGCCCC  
TTCTAAAAAATGGCAGAAAATGGAGTATAACAGGAATCAAA

**SEQ ID NO: 10****Construct 4: gbs1477p**

GACGACGTAACAACCTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCATTTG  
ATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAA  
TGACCTTAAATCTTATTTTTGGCTCAACCGATGCTAAAGAAATTAAGGGTGCTTTCTTTGTT  
TTCAAAAATGAAACTGGTACAAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGG  
20 AAGCTAAAGATGCTGAAGGTGGTGTCTTTCAGGGTTAACAAAAGACACTGGTTTTTGC  
TTTTAACACTGCTAAGTTAAAAGGAACCTACCAAATCGTTGAATTGAAAGAAAAATCAAAC  
TACGATAACAACGGTTCATCTTGGCTGATTCAAAGCAGTTCCAGTTAAAATCACTCTGC  
CATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAAC  
AAAACCACAAGTAGATAAGAACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAA  
25 GACAAAGGTGTTGTCTCAGCGACAGTTGGTGACAAAAAAGAATACATAGTTGGAACAAAAA  
TTCTTAAAGGCTCAGACTATAAGAACTGGTTTTGGACTGATAGCATGACTAAAGGTTTTGAC  
GTTCAACAACAACGTTAAAGTAACATTGGATGGTAAAGATTTTCCTGTTTTAACTACAAA  
CTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACAGGTCTTGCAGCAGTAG  
CAGCTGCTGCAAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTGAACGG  
30 CTCCACTACTGTTGAAGTTCCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCA  
ACGGAAGAAAGTGAACCACAAGAAGGTAAGTACTCCAGCTAACCAAGAAATTAAGTCATTAAAG  
ACTGGGCAGTAGATGGTACAATTACTGATGTTAATGTTGCAGTTAAAGCTATCTTTACCTT  
GCAAGAAAAACAACGGATGGTACATGGGTGAACGTTGCTTCCACACGAAGCAACAAAACCA  
TCACGCTTTGAACATACTTTCACAGGTTTGGATAATACTAAAACCTTACCGCGTTGTGGAAC  
35 GTGTTAGCGGCTACACTCCAGAATATGTATCATTTAAAAATGGTGTGTTGTGACTATCAAGAA  
CAACAAAAACTCAAATGATCCAACCTCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCC  
TTGTTAAGAAAGAAGGAAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAA  
GGCAGCTGTAAAAACTGCTAAACTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTG  
40 ACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAAACAGCATTGGCTACTGTTGATCAAAAAC  
AAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATATGAATGGGTGTCAGATAA  
AAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATTACTGGT  
TTGGATAAAGGCACTTATAGCTTGGAAAGAACTCAAGCACCAGCAGGTTATGCGACATTGT  
CAGGTGATGTAAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACTGACAT  
45 CGCATATGATAAAGGATCTGTAAAAAAGATGCCCAACAAGTTCAAAACAAAAAAGTAACC  
ATCCACAAACAGGTGGTATTGGTACA

**SEQ ID NO: 11****Construct 5: gbs1478p**

GAAAGTACCGTACCGGAAAATGGTGCTAAAGGAAAGTTAGTTGTTAAAAAGACAGATGACC  
50 AGAACAAACCACTTTCAAAGCTACCTTTGTTTTAAAACTACTGCTCATCCAGAAAGTAA



- 50 -

AATAGAAAAAGTAACTGCTGAGCTAACAGGTGAAGCTACTTTTGATAATCTCATACCTGGA  
GATTATACTTTATCAGAAGAAACAGCGCCCGAAGGTTATAAAAAGACTAACCAGACTTGGC  
AAGTTAAGGTTGAGAGTAATGGAAAACTACGATACAAAATAGTGGTGATAAAAATTCCAC  
AATTGGACAAAATCACGAAGAAGTAGATAAGCAGTATCCCCCACAGGAATTTATGAAGAT  
5 ACAAAGGAATCTTATAAACTTGAGCATGTTAAAGGTTTCAGTTCCAAATGGAAAGTCAGAGG  
CAAAAGCAGTTAACCCATATTCAAGTGAAGGTGAGCATATAAGAGAAATTCAGAGGGAAC  
ATTATCTAAACGTATTTTCAGAAGTAGGTGATTTAGCTCATAATAAATATAAAATTGAGTTA  
ACTGTCAGTGGAAAAACCATAGTAAAACCAGTGGACAAAACAAAAGCCGTTAGATGTTGTCT  
TCGTA CTGATAATTCTAACTCAATGAATAACGATGGCCCAAATTTTCAAAGGCATAATAA  
10 AGCCAAGAAAGCTGCCGAAGCTCTTGGGACCGCAGTAAAAGATATTTTAGGAGCAAACAGT  
GATAATAGGGTTGCATTAGTTACCTATGGTTCAGATATTTTTGATGGTAGGAGTGTAGATG  
TCGTAAAAGGATTTAAAGAAGATGATAAATATTATGGCCTTCAAAC TAAGTTCACAATTCA  
GACAGAGAATTATAGTCATAAACAATTAACAAAATAATGCTGAAGAGATTATAAAAAGGATT  
CCTACAGAAGCTCCTAGAGCTAAATGGGGATCAACTACAAACGGACTTACTCCAGAGCAAC  
15 AAAAGCAGTACTATCTTAGTAAAGTAGGGGAAACATTTACTATGAAAGCCTTCATGGAGGC  
AGATGATATTTTTGAGTCAAGTAGATCGAAATAGTCAAAAAATTATTGTTTCATATAACTGAT  
GGTGTTCACAACAAGATCATATGCTATTAATAATTTTAAATTGGGTGCATCATATGAAAGCC  
AATTTGAACAAATGAAAAAAAATGGATATCTAAATAAAAAGTAATTTTCTACTTACTGATAA  
GCCCGAGGATATAAAAAGGAAATGGGGAGAGTTACTTTTTGTTTCCCTTAGATAGTTATCAA  
20 ACACAGATAATCTCTGGAACTTACAAAACTTCATTATTTAGATTTAAATCTTAATTACC  
CTAAAGGTACAATTTATCGAAATGGACCAGTAAGAGAACATGGAACACCAACCAAACCTTTA  
TATAAATAGTTTAAAACAGAAAAATTATGACATCTTTAATTTTGGTATAGATATATCTGCT  
TTTAGACAAGTTTATAATGAGGATTATAAGAAAAATCAAGATGGTACTTTTCAA AAATTGA  
AAGAGGAAGCTTTTGAAC TTTTCAGATGGGGAAATAACAGA ACTAATGAAGTCATTCTCTTC  
25 TAAACCTGAGTATTATAACCCCGATAGTAACTTCATCCGATGCATCTAACAATGAAATTTTA  
TCTAAAATTCAGCAACAATTTGAAAAGGTTTTAACAAAAGAAA ACTCAATTGTTAATGGAA  
CTATAGAAGATCCTATGGGTGACAAAATCAATTTACAGCTTGGCAACGGACAAACATTGCA  
ACCAAGTGATTATACTTTACAGGGAAATGATGGAAGTATAATGAAAGATAGCATTGCAACT  
GGTGGGCCTAATAATGATGGTGGAACTTAAAGGGGTTAAATTAGAATACATCAA AAATA  
30 AACTCTACGTTAGAGGTTTGAAC TTAGGGGAGGGACAAAAGTAACACTCACATATGATGT  
GAAACTAGATGACAGTTTTATAAGTAACAAATTTCTATGACACTAATGGTAGAAC AACATTG  
AATCCTAAATCAGAGGATCCTAATACACTTAGAGATTTTCCAATCCCTAAAATTCGTGATG  
TGAGAGAATATCCTACAATAACGATTAAAACGAGAAGAAGTTAGGTGAAATTGAATTTAC  
AAAAGTTGATAAAGATAATAATAAGTTGCTTCTCAAAGGAGCTACGTTTGAAC TCAAGAA  
35 TTTAATGAAGATTATAAACTTTATTTACCAATAAAAAATAATAATTCAAAGTAGTGACGG  
GAGAAAACGGCAAAATTTCTTACAAAGATTTGAAAGATGGCAAAATATCAGTTAATAGAAGC  
AGTTTCGCCGAAGGATTATCAA AAAATTAATAATAACCAATTTTAACTTTTGAAGTTGTT  
AAAGGATCGATACAAAATATAATAGCTGTTAATAAACAGATTTCTGAATATCATGAGGAAG  
GTGACAAGCATTTAATTACCAACACGCATATTCCACCAAAGGAATTATTCCGATGACAGG  
40 TGGGAAA

**SEQ ID NO: 12****Construct 6: gbs2018p**

GATACTAGTTCAGGAATATCGGCTTCAATTCCTCATAAGAAACAAGTTAATTTAGGGGCGG  
45 TTACTCTGAAGAATTTGATTTCTAAATATCGTGGTAATGACAAAGCTATTGCTATACTTTT  
AAGTAGAGTAAATGATTTTAAATAGAGCATCACAGGATACACTTCCACAATTAATTAATAGT  
ACTGAAGCAGAAATTAGAAATATTTTATATCAAGGACAAATTGGTAAGCAAATAAACCAA  
GTGTA ACTACACATGCTAAAGTTAGTGATCAAGA ACTAGGTAAGCAGTCAAGACGTTCTCA  
AGATATCATTAAGTCATTAGGTTTCCTTTTCATCAGACCAA AAAAGATATTTTAGTTAAATCT  
50 ATTAGCTCTTCAAAGATTCGCAACTTATTCTTAAATTTGTA ACTCAAGCCACGCAACTGA  
ATAATGCTGAATCAACAAAAGCTAAGCAAATGGCTCAA AATGACGTGGCCTTAATAAAAAA



- 51 -

TATAAGCCCCGAAGTCTTAGAAGAATATAAAGAAAAAATTCAAAGAGCTAGCACTAAGAGT  
 CAAGTTGATGAGTTTGTAGCAGAAGCTAAAAAAGTTGTTAATTCCAATAAAGAAACGTTGG  
 TAAATCAGGCCAATGGTAAAAAGCAAGAAATTGCTAAGTTAGAAAATTTATCTAACGATGA  
 AATGTTGAGATATAATACTGCAATTGATAATGTAGTGAAACAGTATAATGAAGGTAAGCTC  
 5 AATATTACTGCTGCAATGAATGCTTTAAATAGTATTAAGCAAGCAGCACAGGAAGTTGCCC  
 AGAAAACTTACAAAAGCAGTATGCTAAAAAAATTGAAAGAATAAGTTCAAAGGATTAGC  
 GTTATCTAAAAAGGCTAAAGAAATTTATGAAAAGCATAAAAGTATTTTGCCTACACCTGGA  
 TATTATGCAGACTCTGTGGGAACTTATTTGAATAGGTTTAGAGATAAACAAACTTTTCGGAA  
 ATAGGAGTGTTTGGACTGGTCAAAGTGGACTTGATGAAGCAAAAAAAATGCTTGATGAAGT  
 10 CAAAAAGCTTTTAAAAGAACTTCAAGACCTTACCAGAGGTACTAAAGAAGATAAAAAACCA  
 GACGTTAAGCCAGAAGCCAAACCAGAGGCCAAACCAAATATTCAAGTACCTAAACAAGCAC  
 CTACAGAAGCTGCAAAAACCAGCTTTGTCCACCAGAAGCCTTGACAAGATTGACTACATGGTA  
 TAATCAAGCTAAAGATCTGCTTAAAGATGATCAAGTAAAGGACAAATACGTAGATATACTT  
 GCAGTTCAAAAAGCTGTTGACCAAGCTTATGATCATGTGGAAGAGGGAAAATTTATTACCA  
 15 CTGATCAAGCAAATCAATTAGCTAACAAGCTACGTGATGCTTTACAAAGTTTAGAATTAAA  
 AGATAAAAAAGTAGCCAAACCAGAAGCCAAACCAGAGGCCAAACCAGAAGCTAAGCCAGAA  
 GCTAAGCCAGAAGCTAAGCCAGAAGCTAAGCCAGAGGCCAAACCAGAAGCTAAGCCAGACG  
 TTAAGCCAGAAGCTAAACCAGACGTTAAACCAGAGGCTAAGCCAGAAGCTAAACCAGAGGC  
 TAAGTCAGAAGCTAAACCAGAGGCTAAGCTAGAAGCTAAACCAGAGGCCAAACCAGCAACC  
 20 AAAAAATCGGTTAATACTAGCGGAACTTGGCGGCTAAAAAAGCTATTGAAAACAAAAAGT  
 ATAGTAAAAAATTACCATCAACG

Expression and purification of proteins: *E. coli* BL21 star<sup>®</sup> cells harboring the recombinant  
 25 plasmid were grown into log phase in the required culture volume. Once an OD<sub>600nm</sub> of 0.6  
 was reached the culture was induced with 0.5 mM IPTG for 3 hours at 37°C. The cells  
 were harvested by centrifugation, lysed by a combination of the freeze-thaw method  
 followed by disruption of cells with 'Bug-buster<sup>®</sup>', (Novagen). The lysate was separated  
 by centrifugation into soluble (supernatant) and insoluble (pellet) fractions. Depending on  
 30 the location of the protein different purification strategies were applied. A) If the His-  
 tagged protein was in the soluble fraction, protein purification was done by binding the  
 supernatant to Ni-Sepharose beads (Ni-Sepharose<sup>™</sup> 6 Fast Flow, GE Healthcare). Due to  
 the presence of the hexa Histidine (6xHIS) at the C terminus of the expressed protein, it  
 bound to the Ni-Sepharose while the other contaminating proteins were washed from the  
 35 column by wash buffer. The protein was eluted by 500 mM Imidazole in 20 mM NaH<sub>2</sub>PO<sub>4</sub>,  
 0.5 mM NaCl buffer at pH 7.4. The eluate was concentrated, assayed by Bradford for  
 protein concentration and checked by SDS-PAGE and Western blot. B) If the protein was  
 present in the insoluble fraction, the pellet was solubilized in suitable buffer containing 8  
 M urea and applied onto the Ni-NTA column under denaturing conditions (in buffer  
 40 containing 8 M urea) using the same materials and procedure as mentioned above.  
 Contaminating proteins were washed from the column by wash buffer without urea.



- 52 -

Refolding of the His-tagged protein was performed while the protein was immobilized on the Ni-NTA matrix. After renaturation, proteins were eluted by the addition of 500 mM Imidazole. The eluate was dialyzed to remove traces of urea and concentrated if the volume was large, checked by SDS-PAGE and measured by the Bradford method.

5

#### *Animal protection studies*

Animals: CD-1 female mice (6-8 weeks) were used for these studies.

Active immunization, generation of hyper-immune mouse sera: 50 µg of recombinant protein was injected subcutaneously into CD-1 mice, adjuvanted with Complete Freund's adjuvant (CFA). Animals were boosted twice with the same amount of protein and Incomplete Freund's adjuvant (IFA) at days 14 and 28. The published protective Sip (gbs0031) protein antigen (Brodeur et al., Infect Immun. 68(10):5610-5618 (2000)) was used as a positive control, while mice immunized with adjuvant only served as negative controls. Antibody titres were measured at day 35 by ELISA using the respective recombinant proteins. In case of hyper-immune sera generation mice were terminally bled at day 35.

Generation of hyperimmune rabbit sera: Polyclonal rabbit sera were generated for gbs0031, gbs0233p, gbs1087p, gbs1309p, gbs1477p, gbs1478p and gbs2018p at Charles River Laboratories, Kislegg, Germany. 250 µg of recombinant protein was injected into New Zealand White rabbits, adjuvanted with Complete Freund's adjuvant (CFA). Animals were boosted three times with the same amount of protein, but with Incomplete Freund's adjuvant (IFA) at days 28, 42 and 56. Antibody titers were measured at day 38 and 52 by ELISA using the respective recombinant proteins. Rabbits were terminally bled at day 70.

Passive immunization: CD-1 mice were immunized intraperitoneally 1 to 3 hours before the bacterial challenge with 150 µl mouse or rabbit hyperimmune sera.

Bacterial challenge: Freshly grown *S. agalactiae* strains C388/90 (serotype Ia/c), A909 (serotype Ia/c), ATCC12401 (serotype Ib), ATCC12403 (serotype III), COH1 (serotype III), BAA22 (serotype III), 2603V/R (serotype V), ATCC49447 (serotype V), BAA23 (serotype V) were used for animal challenge studies. In order to determine the viable cell numbers present in the bacterial inoculum, cfus were determined via plating on blood agar

- 53 -

plates.  $10^6$  -  $10^8$  cfus were applied intraperitoneally into mice. Protection by immunization was measured by a lethal sepsis model, where survival rates were followed for 1 to 2 weeks post-challenge and survival was expressed as percentage of the total number of animals (10 mice / group).

5

### **Results**

By using a genomic scale antigen identification method we selected Group B streptococcal antigens based on immunogenicity in humans (WO2004/099242) and pre-selected vaccine candidates based on *in vitro* assays. Here we show immune protection by six Group B streptococcal antigens in animal models. The first screening model was set up using adult mice and the mouse-adapted *S. agalactiae* ATCC12403 serotype III strain that was also used for the genomic library construction and cloning of some of the vaccine candidates. We set up the method with CD-1 mice and defined the LD<sub>90</sub> - LD<sub>100</sub> dose. The model set up was further optimized by using positive and negative control sera. Protection was estimated by reduced lethality of mice immunized with Sip or anti-Sip immune sera relative to animals immunized with adjuvant alone or treated with control sera. Based on these data, CD-1 mice and a challenge dose between  $5 \times 10^7$  to  $1 \times 10^8$  cfu was used for further studies. Mice were immunized first with the recombinant antigens adjuvanted with CFA/IFA and in subsequent experiments with hyper-immune mouse sera transferred to naïve animals before challenge with *S. agalactiae* ATCC12403 (serotype III). In the active, as well as in the passive model, several protective antigens were identified that showed variable protection levels, ranging from higher, equal or lower survival relative to Sip. Since several different Group B Streptococcus serotypes are able to cause severe disease in humans, it is important to test cross-protection of vaccine candidates against all major serotypes in animal experiments. Moreover, it has been firmly demonstrated that protective antigens show strain-dependent variations not only in their primary sequences and expression, but also in their protective capacity. For that reason, we have set up the screening model with several different *S. agalactiae* strains representing the major serotypes, Ia, Ib, III and V. Strain-dependent protection within one serotype was also addressed by using 2-3 different strains of the most common serotypes Ia, III and V. In order to perform this large number of experiments with the minimal animal sacrifice and good comparability, we generated hyper-immune rabbit sera for all *in vitro* selected recombinant antigens. Three rabbits were immunized with each individual antigen

10  
15  
20  
25  
30



- 54 -

adjuvanted with CFA/IFA using a standard protocol. Animals were pre-screened for pre-existing GBS-specific antibodies by testing their sera with ELISA and only animals without a significant reaction were included in immunization studies. The individual hyper-immune sera were then analyzed for antigen-specific antibody levels and used in pools for further analyses. Thus, the very same immune sera were used for passive protection studies with nine different GBS strains that we found useful for animal studies. As a result of these experiments we could identify six novel vaccine candidates – gbs0233p, gbs1087p, gbs1309p, gbs1477p, gbs1478p and gbs2018p – that showed protection against at least one serotype when used as a sole antigen (Figure 1).

10

In order to examine benefits of combinations of different antigenic components, we performed passive protection studies by combining rabbit sera with different antigen specificities. With different combinations using these six protective vaccine candidates, we could demonstrate increased protection compared to the single proteins against all the tested GBS serotypes. The combination of gbs1477p + gbs2018p provided a significantly increased level of protection against many serotypes. The best protection seen so far was achieved with a combination of gbs1087p + gbs1477p + gbs1478p + gbs2018p that protected most of the mice against all nine tested GBS strains (Figure 2).

20

## **Example 2: Surface exposure and induction of functional antibodies by Group B streptococcal antigens.**

### ***Experimental procedures***

25

#### ***FACS analysis:***

The *S. agalactiae* strain to be tested was inoculated from a glycerol stock into 5 ml THB medium and incubated over night at 37°C. The overnight culture was reinoculated by adding 200 µl into 10 ml fresh THB medium and incubated until an OD<sub>600nm</sub> of approximately 1 was reached (~5 x 10<sup>8</sup> cells/ml). The bacterial cells were pelleted by centrifugation at 4,000 rpm for 5 min and washed twice with 2 ml HBSS. The final pellet was resuspended in HBSS with 1% BSA to give a cell density of 5 x 10<sup>6</sup> cells/ml. To 100 µl bacterial suspension 1 µl serum was added and incubated for 45 min on ice. Bacteria

30

- 55 -

were pelleted by centrifugation at 1,000 g for 4 min and washed once with 150  $\mu$ l HBSS with 1% BSA and resuspended in 100  $\mu$ l HBSS with 1% BSA. To the opsonised bacteria 1  $\mu$ l of the secondary antibody (goat F(ab)2 fragment anti rabbit IgG coupled with PE) was added and incubated for 45 min on ice in the dark. The cells were washed twice with 150  $\mu$ l HBSS as described above and dissolved in 250  $\mu$ l HBSS, the cells were fixed by the addition of 250  $\mu$ l 4% para-formaldehyde. The fluorescent staining of the bacteria was measured by FACS analysis.

### *Opsonophagocytic killing assay*

10

Preparation of bacterial cells: The *S. agalactiae* strain to be tested was inoculated from a glycerol stock into 5 ml THB medium and incubated overnight at 37°C. The over night culture was reinoculated by adding 200  $\mu$ l into 10 ml fresh THB medium and incubated until an OD<sub>600nm</sub> of approximately 1 was reached. The bacteria were pelleted by centrifugation at 4,000 rpm for 5 min and washed twice with 2 ml HBSS. The final pellet was resuspended in HBSS with 0.125% BSA to give a final concentration of  $5 \times 10^4$  cells/85  $\mu$ l.

Preparation of RAW264.7 cells: Cells were cultivated in T175 flasks with 25 ml DMEM high glucose medium at 37°C with 5% CO<sub>2</sub>. Cells were detached from the flasks by scraping and collected by low speed centrifugation at 1,000 rpm for 10 min and washed twice with 50 ml HBSS with 10 mM glucose and resuspended in HBSS with 10 mM glucose to give a cell concentration of  $1 \times 10^7$  cells/ml.

Opsonophagocytic killing assay: Bacteria (85  $\mu$ l) were mixed with 10  $\mu$ l guinea pig complement and 5  $\mu$ l prediluted serum and incubated for 60 min at 6°C with shaking (500 rpm). To the opsonised bacteria 100  $\mu$ l ( $1 \times 10^6$  cells) RAW264.7 cells were added. Three aliquots of 10  $\mu$ l were taken and each added to 1.5 ml water after 5 min incubation, 100  $\mu$ l were plated on blood agar plates to determine the initial bacterial count, T<sub>0</sub>. The suspensions with opsonised bacteria and RAW264.7 cells were incubated for one hour at 37°C with shaking (500 rpm). After 60 min incubation three aliquots of 10  $\mu$ l were removed and each diluted in 1.5 ml, after 5 min incubation, 100  $\mu$ l were plated on blood agar plates to determine T<sub>60</sub>. After overnight cultivation cfus were determined with a colony counter.



Evaluation: For each sample the relationship between the cfu at T<sub>0</sub> and T<sub>60</sub> was determined. The percentage killing of each test serum was related to the respective preimmune serum using the relationship between T<sub>0</sub> and T<sub>60</sub> with the formula 100-  
5 100x(test serum/preimmune serum).

### **Results**

The analyses of surface expression of gbs0233, gbs1087, gbs1309, gbs1477, gbs1478 and  
10 gbs2018 have been performed by FACS analysis using the very same pooled rabbit hyperimmune sera that were tested for protection in animal studies. These six protective antigens were detected on the surface of Group B streptococcal strains. Four of the antigens (gbs1087, gbs1477, gbs1478, gbs2018) were most consistently detected (Figure 3), gbs0233 was not expressed *in vitro* by all strains and gbs1309 was mainly detected in  
15 the bacterial supernatant. The *in vitro* expression experiments have been performed with nine different strains from the serotypes Ia, Ib, II, III, IV and V; the most comprehensive studies have been performed with the serotype III strain ATCC12403.

Based on the passive protection data, it is firmly established that protection by the selected  
20 six vaccine candidates is mainly mediated by antibodies. The ability to measure functional antibodies in *in vitro* assays is essential for the development of both a prophylactic vaccine and an antibody-based therapy or prevention. Nine different *S. agalactiae* strains representing six serotypes (Ia, Ib, II, III, IV and V) were used to evaluate gbs0233p, gbs1087p, gbs1309p, gbs1477p, gbs1478p and gbs2018p for induction of functional  
25 antibodies. Included in the opsonophagocytic killing assays were two GBS strains representing serotypes II and IV that were not suitable for animal testing. As an example of the *in vitro* assays, results with two strains are presented in Figure 4, the serotype II strain AC3912 (Figure 4A) and the serotype V strain BAA23 (Figure 4B). Simultaneously with the opsonophagocytic killing assay cells were tested for *in vitro* expression of the tested  
30 antigens by Western blot and FACS analysis. At a serum dilution of 1:200, only gbs1478p showed more than 50% killing of the serotype II strain AC3912 (Figure 4A), both gbs1477p and gbs1478p showed more than 50% killing of the serotype V strain BAA23 (Figure 4B). The remaining antigens showed less than 50% killing of the strains tested in

- 57 -

Figure 4, which in most cases can be explained by poor *in vitro* expression of the antigens in these strains.

### 5 **Example 3: Sequence conservation of protective Group B streptococcal antigens**

#### *Experimental procedures*

Sequence analyses of *S. agalactiae* genes: In order to determine the sequence of an antigen  
10 from diverse *S. agalactiae* strains, PCR was performed with primers specific for the gene  
of interest. *S. agalactiae* strains used for these analyses are shown in Tables 2 and 13.  
Oligonucleotide sequences as primers for PCR were designed for the selected antigens in  
order to be able to amplify the full gene. Sequencing was performed with dedicated  
primers using the PCR products as templates. The sequences of the oligonucleotides are  
15 listed in Table 3. Genomic DNA of all *S. agalactiae* strains was prepared as described in  
WO2004/099242. PCR was performed in a reaction volume of 25 µl using Taq polymerase  
(1 U), 200 nM dNTPs, 10 pMol of each oligonucleotide and the kit according to the  
manufacturer's instructions (Invitrogen, The Netherlands). As standard, 30 cycles (1x: 5  
min. 95°C, 30x: 30 sec. 95°C, 30 sec. 56°C, 30 sec. 72°C, 1x: 4 min. 72°C) were  
20 performed, unless conditions had to be adapted for individual primer pairs. PCR samples  
were sequenced with the oligonucleotides as listed in Table 3.

#### *Results*

25 The genomic sequence of eight individual strains of *S. agalactiae* (Tables 2 and 4) has  
been published and was compared for the six antigens shown to be protective under  
Example 1. The comparison showed that the proteins gbs0233 and gbs1087 are highly  
conserved (more than 99 and 91% identity, respectively; Tables 2 and 4), although  
gbs1087 displayed various numbers of repeats in the different GBS strains (see also  
30 WO2004/035618). This high degree of protein sequence identity (gbs0233: >99%;  
gbs1087: >86%) could also be observed for the strains that were subjected to DNA  
sequence analyses as listed in Tables 5, 7, 8, and Table 13 and in the Sequence listing. The  
gbs0233 protein from any of the analyzed strains showed at least 98.7% amino acid  
sequence identity to gbs0233 from *S. agalactiae* NEM316, with only 6 amino acid position



- 58 -

showing a change. The sequences of the gbs1087 proteins from the analyzed strains were also highly conserved, yet the different strains harboured between a single and up to 29 repeats of a highly conserved 17 amino acid long sequence. The sequences of proteins gbs1309 and gbs2018 showed high sequence conservation in 7 genomic strains (more than 5 87 and 77% identity, respectively), while protein sequences diverged more significantly in strain COH1 (69.9 and 47.7%, respectively; Table 4). The gbs1309 protein showed a similar high degree of amino acid sequence identity (89.6%) in the sequenced GBS strains (Table 5, 9, 13 and Sequence listing), while the gbs2018 protein can be classified in two clades, with 95% of strains belonging to one clade with at least 60.8% sequence identity and 3 strains COH1(III), BAA22(III) and 49447(V) belonging to the second clade. The 10 protein gbs1478 is highly conserved in 6 genomic strains (more than 87% identity), yet the strains COH1 and A909 show a lower amino acid sequence identity of approximately 43% (Table 4). Protein gbs1478 is conserved in most analyzed GBS strains as shown in Table 5, 11, 13 and the Sequence listing, but exists as 2 distinct clades with an amino acid sequence 15 identity of more than 80% in the dominant clade (approx. 80% of analyzed strains) and more than 99% in the second clade. The protein gbs1477 shows the highest degree of amino acid sequence variability, with six distinct clades that can be characterized. Strains COH1 and A909 do not encode a homologous protein with significant amino acid sequence identity (Table 4). The sequence analyses of the gbs1477 gene from further 20 distinct GBS strains revealed that all selected strains encode a protein homologous to gbs1477 and that all six clades were covered by these sequences (Table 5, 6, 10, 13, Sequence listing and Figure 5). The prototype sequences for the 6 clades of gbs1477 are: strain 12401 (clade 1; SEQ ID NO: 93), strain IC254 (clade 2; SEQ ID NO: 110), strain 126H4A (clade 3; SEQ ID NO: 94), strain 49447 (clade 4; SEQ ID NO: 95), strain 25 C388/90 (clade 5; SEQ ID NO: 100) and strain NEM316 (clade 6; SEQ ID NO: 223 and SEQ ID NO: 361); (for all sequences, see Sequence listing). Within any single clade the level of amino acid sequence identity reaches at least 98%.

**Example 4: Group B streptococcal antigens and combinations thereof as well as 30 mouse monoclonal antibodies, generated against these antigens, induce protective immune responses against lethal sepsis in an i.p. challenge model.**

#### *Experimental procedures*

*Cloning and expression of recombinant Group B streptococcal proteins*

Cloning of genes: The gene of interest was amplified from genomic DNA of *S. agalactiae* ATCC12403 (serotype III) by PCR using gene specific primers. Apart from the gene specific part, the primers had restriction sites that aided in a directional cloning of the amplified PCR product. The gene annealing (specific) part of the primer ranged between 15-30 bases in length. The PCR products obtained were digested with the appropriate restriction enzymes and cloned into the pET28b (+) vector (Novagen) for His-tagged proteins. Once the recombinant plasmid was confirmed to contain the gene of interest, *E. coli* BL21 star<sup>®</sup> cells (Invitrogen) that served as expression host were transformed. Cloning of the gbs1087, gbs1477 and gbs1478 genes has been performed using genomic DNA from strain *S. agalactiae* 6313 (serotype III) in the vector pET28a (+). The constructs of the selected antigens are listed in Table 1.

15 Expression and purification of proteins:

*E. coli* BL21 star<sup>®</sup> cells harboring the recombinant plasmid were grown into log phase in the required culture volume. Once an OD<sub>600nm</sub> of 0.6 was reached the culture was induced with 0.5 mM IPTG for 3 hours at 37°C. The cells were harvested by centrifugation, lysed by a combination of the freeze-thaw method followed by disruption of cells with 'Bug-buster<sup>®</sup>' (Novagen). The lysate was separated by centrifugation into soluble (supernatant) and insoluble (pellet) fractions. Depending on the location of the protein different purification strategies were applied. A) If the His-tagged protein was in the soluble fraction, protein purification was done by binding the supernatant to Ni-Sepharose beads (Ni-Sepharose<sup>™</sup> 6 Fast Flow, GE Healthcare). Due to the presence of the hexa Histidine (6xHIS) at the C terminus of the expressed protein, it bound to the Ni-Sepharose while the other contaminating proteins were washed from the column by wash buffer. The protein was eluted by 500 mM Imidazole in 20 mM NaH<sub>2</sub>PO<sub>4</sub>, 0.5 mM NaCl buffer at pH 7.4. The eluate was concentrated, assayed by Bradford for protein concentration and checked by SDS-PAGE and Western blot. B) If the protein was present in the insoluble fraction, the pellet was solubilized in suitable buffer containing 8 M Urea and applied onto the Ni-NTA column under denaturing conditions (in buffer containing 8 M Urea) using the same materials and procedure as mentioned above. Contaminating proteins were washed from the column by wash buffer without urea. Refolding of the His-tagged protein was performed while the protein was immobilized on the Ni-NTA matrix. After renaturation,



- 60 -

proteins were eluted by the addition of 500 mM Imidazole. The eluate was dialyzed to remove traces of urea and concentrated if the volume was large, checked by SDS-PAGE and measured by the Bradford method.

5 *Animal protection studies*

Animals: CD-1 female mice (6-8 weeks) were used for these studies.

Active immunization: 25 µg of recombinant protein was injected subcutaneously into CD-1 mice, adjuvanted with ALUM 1%. Animals were boosted twice with the same amount of protein and ALUM 1% at days 14 and 28. The published protective Sip (gbs0031) protein  
10 antigen was used as a positive control, while mice immunized with adjuvant only served as negative controls. Antibody titres were measured at day 35 by ELISA using the respective recombinant proteins.

Generation of hyperimmune rabbit sera: Polyclonal rabbit sera were generated for  
15 gbs0031, gbs0233p, gbs1087p, gbs1309p, gbs1477p, gbs1478p and gbs2018p at Charles River Laboratories, Kislegg, Germany. 250 µg of recombinant protein was injected into New Zealand White rabbits, adjuvanted with Complete Freund adjuvant (CFA). Animals were boosted three times with the same amount of protein, but with Incomplete Freund adjuvant (IFA) at days 28, 42 and 56. Antibody titers were measured at day 38 and 52 by  
20 ELISA using the respective recombinant proteins. Rabbits were terminally bled at day 70.

Generation of mouse monoclonal antibodies: Monoclonal mouse antibodies were generated against gbs0233p, gbs1087p, gbs1477p, gbs1478p and gbs2018p at Abgent, San Diego, USA. 100 µg of recombinant protein was injected into Balb/c mice, adjuvanted  
25 with Complete Freund adjuvant (CFA). Animals were boosted with 50 µg protein and CFA at week 2; at week 3 animals were boosted with the same amount of protein, but with Incomplete Freund adjuvant (IFA) and at week 4 and 5 animals were boosted with 50 µg protein in PBS (without adjuvant). Antibody titers were measured in week 5 by ELISA and Western blotting using the respective recombinant proteins. Spleen cells from mouse with  
30 the best titer were fused with myeloma cell F0 using PEG protocol. Subsequently growing fused hybridoma clones were screened against the respective antigen for test of their specificity and sensitivity. ELISA positive clones were tested also by Western blot. Selected clones from this test were subcloned at least two times and antibodies were purified by protein G affinity chromatography from culture medium.

Passive immunization (neonates): Pregnant CD-1 mice were given 0.5 ml undiluted rabbit hyper-immune sera by intraperitoneal injection 2 to 4 days before delivery. Within 48h after birth, pups were challenged intraperitoneally.

5

Bacterial challenge: Freshly grown *S. agalactiae* strains C388/90 (serotype Ia/c), A909 (serotype Ia/c), ATCC12401 (serotype Ib), ATCC12403 (serotype III), COH1 (serotype III), ATCCBAA22 (serotype III), 2603V/R (serotype V), ATCC49447 (serotype V), ATCCBAA23 (serotype V) were used for animal challenge studies. In order to determine the viable cell numbers present in the bacterial inoculum, cfus were determined via plating on blood agar plates.  $10^6$  -  $10^8$  cfus were applied intraperitoneally into mice. Protection by immunization was measured by a lethal sepsis model, where survival rates were followed for 1 to 2 weeks post-challenge and survival was expressed as percentage of the total number of animals (10 mice / group for active immunization; for neonatal challenge number of animals depends on the litter size).

10  
15

### **Results**

By using a genomic scale antigen identification method we selected Group B antigens based on immunogenicity in humans (WO04/099242) and pre-selected vaccine candidates based on *in vitro* assays. We have shown previously immune protection by six Group B streptococcal antigens in animal models. Additionally, with different combinations using these six protective vaccine candidates, we demonstrated increased protection compared to the single proteins against all the tested GBS serotypes. The combination of gbs1477p + gbs2018p provided a significantly increased level of protection against many serotypes. The best protection seen so far was achieved with a combination of gbs1087p + gbs1477p + gbs1478p + gbs2018p that protected most of the mice against all nine tested GBS strains (see Example 1 and Figure 2). So far these experiments were obtained in serum transfer experiments. We now further substantiated these results by active immunization of mice with two, three or four recombinant proteins using ALUM as adjuvant (Figure 6). Immunization with single proteins verified the data already obtained with the hyper-immune rabbit sera (Figure 7).

20  
25  
30

Since GBS sepsis affects mainly newborns, we have also developed a model that can demonstrate protection in neonatal mice. We established a murine model with passive



- 62 -

immunization of pregnant mice with hyperimmune rabbit sera (500  $\mu$ l i.p.) 2-4 days before delivery and challenging their babies with *S. agalactiae* 24-48h after birth. We observed excellent protection of newborn mice born to mothers immunized with the combination of gbs1087p + gbs1477p + gbs1478p + gbs2018p specific immune sera (4 x 125  $\mu$ l) (Figure 8). Hyperimmune sera against the individuals antigens were also effective in this, but overall was lower than that obtained with the combination of four (data not shown). These findings are very significant, since the models with the different GBS strains were very stringent, resulting in death of infected pups within 24 hours.

Since the protein gbs1477 has the highest sequence variability and exists in different clades (Table 5 & 6, Figure 5) the protection was analyzed using the adult sepsis/lethality model. Mice were immunized with fragments (corresponding to gbs1477p of strain 6313) of six different gbs1477 proteins, originated from distinct clades. Protection was measured against the homologous as well as against the heterologous clade (Figure 9). The best protection was always obtained when immunization and challenge is done with the homologous clade. The more variable the sequences of the different clades are the lower the protection obtained in the sepsis model.

This invention includes also protection data by mouse monoclonal antibodies. mAbs were generated against gbs0233p, gbs1087p, gbs1477p, gbs1478p and gbs2018p. Selection of hybridoma supernatants were performed using antigen-specific ELISA and/or FACS analysis. Per antigen two mAbs were selected and tested in the passive transfer model using 50  $\mu$ g purified mAb. We demonstrate in this invention that we obtain protection with a single mAb against at least one serotype (Figure 10). In order to examine benefits of combinations of different mAb components, we performed passive protection studies by combining mAbs with different antigen specificities. We could demonstrate increased protection compared to the single mAbs against all the tested GBS serotypes. The best protection seen so far was achieved with a combination of mAbs against gbs0233p + gbs1087p + gbs1477p + gbs1478p + gbs2018p that protected most of the mice against all nine tested GBS strains (Figure 11).

**Example 5: Group B streptococcal antigens and mouse monoclonal antibodies, generated against these antigens, induce functional antibodies against group B streptococcus.**

## *Experimental procedures*

### *FACS analysis*

The *Streptococcus agalactiae* strain to be tested was inoculated from a glycerol stock into 5 ml THY medium and incubated over night at 37°C. The overnight culture was re-inoculated by adding 200 µl into 10 ml fresh THY medium and incubated until an OD<sub>600nm</sub> of approximately 1 was reach (~5 x 10<sup>8</sup> cells/ml). The bacteria were pelleted by centrifugation at 4,000 rpm for 5 min and washed twice with 2 ml HBSS. The final pellet was resuspended in HBSS with 1% BSA to give a cell density of 5 x 10<sup>6</sup> cells/ml. To 100 µl bacteria, 1 µl immune serum was added and incubated for 45 min on ice. Bacteria were pelleted by centrifugation at 1,000 g for 4 min and washed once with 150 µl HBSS with 1% BSA and resuspended in 100 µl HBSS with 1% BSA. To the opsonised bacteria, 1 µl of the secondary antibody (goat F(ab)<sub>2</sub> fragment anti rabbit IgG coupled with PE) was added and incubated for 45 min on ice in dark. The cells were washed twice with 150 µl HBSS as described above and dissolved in 250 µl HBSS, the cells were fixed by addition of 250 µl 4% para-formaldehyde. The fluorescent staining of the bacteria was measured by flow cytometry.

### *Opsonophagocytic killing assay*

Preparation of bacterial cells: The *Streptococcus agalactiae* strain to be tested was inoculated from a glycerol stock into 5 ml THY medium and incubated overnight at 37°C. The over night culture was re-inoculated by adding 200 µl into 10 ml fresh THY medium and incubated until an OD<sub>600nm</sub> of approximately 1 was reached. The bacteria were pelleted by centrifugation at 4,000 rpm for 5 min and washed twice with 2 ml HBSS. The final pellet was re-suspended in HBSS with 0.125% BSA to give a final concentration of 5 x 10<sup>4</sup> cells/85 µl.

Preparation of RAW264.7 cells: Cells were cultivated in T175 flasks with 25 ml DMEM high glucose medium at 37°C with 5% CO<sub>2</sub>. Cells were detached from the plates by scraping and collected by low speed centrifugation at 1,000 rpm for 10 min and washed twice with 50 ml HBSS with 10 mM glucose and re-suspended in HBSS with 10 mM glucose to give a cell concentration of 1 x 10<sup>7</sup> cells/ml.

Opsonophagocytic killing assay conditions: Bacterial cells (85 µl) were mixed with 10 µl guinea pig complement and 5 µl pre-diluted serum and incubated for 60 min at 6°C with



- 64 -

shaking (500 rpm). To the opsonised bacteria, 100  $\mu$ l ( $1 \times 10^6$  cells) of RAW264.7 cells were added. Three aliquots of 10  $\mu$ l were taken out and added to 1.5 ml water after 5 min incubation, 100  $\mu$ l were plated out on blood agar plates. This CFU determination served as the initial bacterial count,  $T_0$ . The suspension of opsonised bacteria and RAW264.7 cells  
5 was further incubated at 37°C with shaking (500 rpm) for 60 min and then the  $T_{60}$  was determined as described for the  $T_0$ . Blood agar plates were incubated overnight and the CFUs determined on the next day using a colony counter.

Evaluation: For each serum, the relationship between the CFUs at  $T_0$  and  $T_{60}$  was determined for the pre-immune and the immune serum. The percentage of killing of each  
10 immune serum was determined by the following formula:  $100 - 100 \times (\text{immune serum} / \text{preimmune serum})$ . A reaction without sera was included in each assay as negative (complement) control.

### **Results**

15

Based on the passive protection data, it is firmly established that protection by the selected six vaccine candidates is mainly mediated by antibodies. The ability to measure functional antibodies in *in vitro* assays is essential for the development of both a prophylactic vaccine and an antibody-based therapy or prevention. The same opsonophagocytic killing assay  
20 that was developed for the *in vitro* validation and used for selection of vaccine candidates was employed to analyze the hyperimmune rabbit sera for the presence of functional antibodies. Seven mouse monoclonal antibodies and four rabbit sera representing four antigens were tested in the opsonophagocytic killing assay for functional antibodies and staining in flow cytometry of the serotype III GBS strain ATCC12403 (Figure 12 and 13).  
25 All monoclonal antibodies bound to the serotype III GBS strain ATCC12403 as measured by flow cytometry (Figure 12). The opsonophagocytic killing assay with the rabbit and mouse monoclonal antibodies are shown in Figure 13. At a 1:1,000 dilution of the rabbit sera only gbs1477#28 and gbs1478#4 showed a high killing activity and gbs1087#12 showed low killing activity. The mouse monoclonal antibodies generated against gbs1477p  
30 and gbs1478p showed high killing activity and those against gbs1087p and gbs2018p showed only a killing activity at the higher concentration.

**Table 1: List of genes selected for expression.**

The nomenclature of the proteins is derived from the genome of NEM316 (ATCC12403). The strain and position (start/stop) of the amplicon within the full length gene/protein are indicated.

5

Construct	Gene	Full length	Strain	Vector	nt (start/stop)	SEQ ID NO	aa (start/stop)	SEQ ID NO
1	gbs0233	308	12403	pET28b	37-921	7	13-307	1
2	gbs1087	442	6313	pET28a	106-1062	8	36-354	2
3	gbs1309	403	12403	pET28b	4-1203	9	2-401	3
4	gbs1477	674	6313	pET28a	88-1944	10	30-648	4
5	gbs1478	901	6313	pET28a	88-2595	11	30-865	5
6	gbs2018	643	12403	pET28b	106-1836	12	36-612	6

**Table 2:** List of *S. agalactiae* strains which were used for the first run of sequencing of the most of the genes of interest, and for which the genome has been published. (A complete list of strains used for the first and second run can be found in Table 13.)

10

<b>GBS strains used for sequencing</b>	
<i>S. agalactiae</i> strain	Serotype
IC105	IV
IC458	Ia
0176H4A	II
ATCC12401	Ib
BAA23	V
COH1	III
ATCC12403	III
6313	III
<b>Published genomic GBS strains (Maione <i>et al.</i>, 2005, Science 309(5731): 148-50)</b>	
<i>S. agalactiae</i> strain	Serotype
H36B	Ib
COH1	III
CJB111	V
A909	Ia/c
515	Ia
2603V/R	V
NEM316 (ATCC12403)	III
18RS21	II



**Table 3:** Oligonucleotides used for PCR and sequencing

ORF-protein	PCR	Sequencing Primer	Name	SEQ ID NO:
gbs0233	ICC5455 & ICC5456	GGGGGCACAATTCCTGTTAT	ICC 5455	13
		AAAAAGTGGTGGATAAATTGTTCT	ICC 5456	14
gbs1087	ICC5489 & ICC5494	CATTGTAAATCTTAATGTTAGTATGA	ICC 5489	15
		TGACTTTGATTTCCAACACTATCC	ICC 5490	16
		GGTTTTAGAACTTGGAAATCAGGA	ICC 5491	17
		GGTCTATTAGCTACATTAGTAACCTG	ICC 5492	18
		AGAGAAAATAATCACTCTAGTCAAGG	ICC 5493	19
		AAAAAGTCACCCTAACCAACC	ICC 5494	20
gbs1309	ICC5465 & ICC5468	AATCATCGGTGAAGTGACGA	ICC 5465	21
		CGGTTAATTCAATTGGATATTTTCT	ICC 5466	22
		ACTCTGATATGGGTAAAGGCTAT	ICC 5467	23
		CTTGAATTATTCTTAAAAAGACCAAAA	ICC 5468	24
	ICC5469 & ICC5470	CCAGTAGATGAGTGGTTAGGTCTTG	ICC 5469	25
		AAGATGAGCTGGTTTTATATATTG	ICC 5470	26
gbs1477	ICC5471 & ICC5479	TTGCAGGTGGAATTTATATTTGG	ICC 5471	27
		TTCTTATCTACTTGTGGTTTTGTTTCA	ICC 5477	28
		TCTTGGCTGATTCAAAAGCA	ICC 5478	29
		GGTTCTGATGGGTTGATTGG	ICC 5479	30
	ICC5480 & ICC5472	AATGGCTCTTGCTTATGATCT	ICC 5472	31
		TGTTAGCGGCTACACTCCAG	ICC 5480	32
	ICC5471 & ICC5475	TTGCAGGTGGAATTTATATTTGG	ICC 5471	33
		CAACTTTTGGTTCAGTTGG	ICC 5475	34
		CCCATTGTCAAACCATT	ICC 5473	35
		GCTACTGCTGAAATCGGTMA	ICC 5474	36
	ICC5476 & ICC5472	CATACATGATCTCAGAACGT	ICC 5476	37
		AATGGCTCTTGCTTATGATCT	ICC 5472	38
gbs1478	ICC5481 & ICC5484	TCTAGGATATTCTGTATCTGATCTTAG	ICC 5481	39
		CCATCAAAAATATCTGAACCA	ICC 5482	40
		GAGGGAACATTATCTAAACGTATTTCA	ICC 5483	41

- 67 -

		TTCAATTTTTGAAAAGTACCATCTTG	ICC 5484	42
	ICC5485 & ICC5488	GAACATGGAACACCAACCAA	ICC 5485	43
		TCAATTTACCTAACTTCTTCTCG	ICC 5486	44
		TTTTCCAATCCCTAAAATTCG	ICC 5487	45
		TTTTCATTTCTATCTCCTTCTTATTC	ICC 5488	46
gbs2018	ICC5457 & ICC5460	AAAAGGCAAAGTTCTGATGAGG	ICC 5457	47
		AAAAATGCTTGATGAAGTCAAAA	ICC 5458	48
		GTTTGGCTTCTGGCTTAACG	ICC 5459	49
		TGATCAAGAAGTACTAGGTAAGCAGTCA	ICC 5460	50
	ICC5461 & ICC5462	CAAATTTAAGAATAAGTTGCGAATC	ICC 5461	51
		AGAGTAAATGATTTTAATAGAGCATCA	ICC 5462	52
	ICC5463 & ICC5464	AAAATATTTCTAATTTCTGCTTCAGT	ICC 5463	53
		AATTAATAAATAACGTGGTCCTATCC	ICC 5464	54



**Table 4: Sequence identity of proteins in published genomes of 8 GBS strains.**

ORF indicates the name of the respective gene/protein in the genomic GBS strain. % Id, amino acid sequence identity in percentage. All comparisons are performed to the respective protein of GBS strain NEM316.

ORF name (NEM316)	<i>S. agalactiae</i> H36B		<i>S. agalactiae</i> COHI		<i>S. agalactiae</i> CJB111		<i>S. agalactiae</i> A909		<i>S. agalactiae</i> 515		<i>S. agalactiae</i> 2603V/R		<i>S. agalactiae</i> 18RS21	
	ORF	% Id	ORF	% Id	ORF	% Id	ORF	% Id	ORF	% Id	ORF	% Id	ORF	% Id
gbs0233	SAI_0243	99.0	SAN_0021	99.7	SAM_0244	99.0	SAK_0301	99.0	SAL_0280	99.7	SAG0242	99.0	SAJ_0320	99.0
gbs1087	SAI_2325	93.9	SAN_1174	93.3	SAM_1069	91.5	SAK_1142	92.8	SAL_1159	92.0	SAG1052	100.0	SAJ_1090	100.0
gbs1309	SAI_1330	99.4	SAN_1370	69.9	SAM_1259	100.0	SAK_1321	99.7	SAL_1364	87.7	SAG1237	99.7	SAJ_2108	90.1
gbs1477	SAI_1511	47.0	none		SAM_1372	98.4	none		SAL_1486	75.9	SAG1407	47.3	SAJ_1416	47.3
gbs1478	SAI_1512	87.7	SAN_0702	43.2	SAM_1373	99.1	SAK_0780	43.3	SAL_1487	97.0	SAG1408	97.3	SAJ_1417	97.3
gbs2018	SAI_2103	78.8	SAN_2207	47.7	SAM_1974	77.6	SAK_1999	92.3	SAL_2118	100.0	SAG2063	87.5	SAJ_1966	87.5



**Table 5: Sequence identity of proteins as determined by the first run of sequencing.** % Id, amino acid sequence identity of the respective protein in percentage as determined by DNA sequencing. All comparisons are performed to the respective protein of GBS strain NEM316.

5

	<i>S. agalactiae</i> IC105	<i>S. agalactiae</i> IC458	<i>S. agalactiae</i> 12401	<i>S. agalactiae</i> BAA23	<i>S. agalactiae</i> COH1	<i>S. agalactiae</i> 0176H4A
ORF name (NEM316)	% Id	% Id	% Id	% Id	% Id	% Id
gbs0233	99.0	100	99.0	99.0	99.7	99.7
gbs1087	86.0	95.2	93.9	91.5	92.7	98.6
gbs1309	100	100	99.3	100	99.5	99.5
gbs1477	48.9	49.3	48.8	98.4	Not determined	67.8
gbs1478	88.2	88.1	87.7	99.1	Not determined	Not determined
gbs2018	77.6	99.7	77.2	78.5	Not determined	80.2

**Table 6: Sequence identity of gbs1477 proteins as determined by the first run of sequencing.** % Id, amino acid sequence identity of the respective protein in percentage as determined by DNA sequencing. Pairwise comparisons are performed for the gbs1477 protein among the sequenced GBS strains.

10

	<i>S. agalactiae</i> IC105	<i>S. agalactiae</i> IC458	<i>S. agalactiae</i> 12401	<i>S. agalactiae</i> BAA23	<i>S. agalactiae</i> 0176H4A
gbs1477	% Id	% Id	% Id	% Id	% Id
<i>S. agalactiae</i> NEM316	48.9	49.3	48.8	98.4	67.8
<i>S. agalactiae</i> IC105	100	58.6	99.9	49.4	48.0
<i>S. agalactiae</i> IC458		100	58.6	49.5	53.2
<i>S. agalactiae</i> 12401			100	49.4	47.9
<i>S. agalactiae</i> BAA23				100	68.5
<i>S. agalactiae</i> 0176H4A					100



**Table 7: Amino acid and encoding DNA sequences of gbs0233 proteins derived from different strains of *S. agalactiae***

	<b>Strain 0176H4A</b>
	ORF DNA sequence (SEQ ID NO: 61)
5	ATGCTTAAAAAATCGCACTTTTTACAGATATTTACACTTTGCTTAGCCCTCTTAACGATTTCTGGTTGTCAA
	TTAACCGATACTAAAAACCTGGTCATACCACAATTAAGGTTGCTGCACAAAGTTCTACAGAGTCTAGTATC
	ATGGCAAATATTGTCACCGAATTAATTCATCACGAATTAGGATACAACACAACCTTAATAAGCAATCTTGGT
	TCCTCTACGGTTACTACCAAGCTTTGCTCCGTGGTGATGCTGACATTGCTGCCACACGTTATACAGGAACA
	GACATCACAGGAACTCTTGGCTTAAAAGCTGTTAAAGACCCCTAAAGAAGCTTCTAAGATTGTAAAAACTGAA
10	TTCCAAAACGCTACAATCAAACCTGGTATCCTACTTATGGTTTTTCTGATACTTATGCATTCATGGTTACT
	AAAGAGTTTGCCAGACAGAATAAAAATCACCAAGATCTCTGATCTCAAAAAATTATCAACAACCTATGAAGGCA
	GGGGTTGATAGTTCATGGATGAATCGCGAGGGAGATGGATACACTGATTTTCGCTAAAACATACGGTTTTGAA
	TTTTACATATTTACCTATGCAAATTGGCTTAGTCTATGATGCGGTTGAAAGTAACAAAATGCAATCTGTA
	TTAGGCTACTCCACTGACGGTCGTATTTTCGAGCTATGATTTAGAAATTTTAAGGGATGATAAAAAATTCTTT
15	CCTCCTTATGAAGCCTCTATGGTTGTCAACAATTCTATCATCAAAAAAGATCCTAAACTAAAAAAATTACTC
	CATCGACTCGATGGTAAAATCAATTTAAAAACGATGCAAAACCTTAATTATATGGTAGATGATAAACTTTTA
	GAACCTTCAGTTGTTGCCAAACAATTTTTAGAAAAAAACCATTATTTTAGAGGAGATAAATAA
	ORF amino acid sequence (SEQ ID NO: 55):
20	MLKKSHFLQIFTLCLALLTISGCQLDTPKPGHTTIKVAAQSSTESSIMANIVTELIHHELGYNTTLISNLG
	SSTVTHQALLRGDADIAATRYTGTDITGTLGLKAVKDPKEASKIVKTEFQKRYNQTWYPTYGFSPTYAFMVT
	KEFARQNKITKISDLKLLSTTMKAGVDSWVNREGDGYTDFAKTYGFESHYPMQIGLVYDAVESNKMQSV
	LGYSTDGRISSYDLEILRDDKKFFPPYEASMVVNSIIKKDPKLLKLLHRLDGKINLKTMQNLNMYMVDKLL
	EPSVVAQKFLEKNHYFRGDK
25	<b>Strain 12401</b>
	ORF DNA sequence (SEQ ID NO: 62):
	ATGCTTAAAAAATCGCACTTTTTACAGATATTTACACTTTGCTTAGCCCTCTTAACGATTTCTGGTTGTCAA
	TTAACCGATACTAAAAAGTCTGGTCATACCACAATTAAGGTTGCTGCCCAAAGTTCTACAGAGTCTAGTATT
30	ATGGCAAATATTATCACCGAATTAATTCATCACGAATTAGGATACAACACAACCTTAATAAGCAATCTTGGT
	TCCTCTACGGTTACTACCAAGCTTTGCTCCGTGGTGATGCTGACATTGCTGCCACACGTTATACAGGAACA
	GACATCACAGGGACTCTTGGTTTTAAAAGCTGTTAAAGACCCCTAAAGAAGCTTCTAAGATTGTAAAAACTGAA
	TTCCAAAACGCTACAATCAAACCTGGTATCCTACTTATGGTTTTTCTGATACTTATGCATTCATGGTTACT
	AAAGAGTTTGCCAGACAgAATAAAAATCACTAAGATCTCTGATCTTAAAAAATTATCAACAACCTATGAAGGCA
35	GGGGTTGATAGTTCATGGATGAATCGCGAGGGAGATGGATACACTGATTTTCGCTAAAACATACGGTTTTGAA
	TTTTACATATTTACCTATGCAAATTGGCTTAGTCTATGATGCGGTTGAAAGTAACAAAATGCAATCTGTA
	TTAGGCTACTCCACTGACGGTCGTATTTTCGAGCTATGATTTAGAAATTTTAAGGGATGATAAAAAATTCTTT
	CCTCCTTATGAAGCCTCTATGGTTGTCAACAATTCTATCATCAAAAAAGATCCTAAACTAAAAAAATTACTC
	CATCGACTCGATGGTAAAATCAATTTAAAAACGATGCAAAACCTTAATTATATGGTAGATGATAAACTTTTA
	GAACCTTCAGTTGTTGCCAAACAATTTTTAGAAAAAAACCATTATTTTAGAGGAGATAAATAA
40	ORF amino acid sequence (SEQ ID NO: 56):
	MLKKSHFLQIFTLCLALLTISGCQLDTPKSGHTTIKVAAQSSTESSIMANIIITELIHHELGYNTTLISNLG
	SSTVTHQALLRGDADIAATRYTGTDITGTLGLKAVKDPKEASKIVKTEFQKRYNQTWYPTYGFSPTYAFMVT
	KEFARQNKITKISDLKLLSTTMKAGVDSWVNREGDGYTDFAKTYGFESHYPMQIGLVYDAVESNKMQSV
	LGYSTDGRISSYDLEILRDDKKFFPPYEASMVVNSIIKKDPKLLKLLHRLDGKINLKTMQNLNMYMVDKLL
45	EPSVVAQKFLEKNHYFRGDK
	<b>Strain BAA23</b>
	ORF DNA sequence (SEQ ID NO: 63):
50	ATGCTTAAAAAATCGCACTTTTTACAGATATTTACACTTTGCTTAGCCCTCTTAACGATTTCTGGTTGTCAA
	TTAACCGATACTAAAAAGTCTGGTCATACCACAATTAAGGTTGCTGCCCAAAGTTCTACAGAGTCTAGTATT
	ATGGCAAATATTATCACCGAATTAATTCATCACGAATTAGGATACAACACAACCTTAATAAGCAATCTTGGT
	TCCTCTACGGTTACTACCAAGCTTTGCTCCGTGGTGATGCTGACATTGCTGCCACACGTTATACAGGAACA
	GACATCACAGGGACTCTTGGTTTTAAAAGCTGTTAAAGACCCCTAAAGAAGCTTCTAAGATTGTAAAAACTGAA
	TTCCAAAACGCTACAATCAAACCTGGTATCCTACTTATGGTTTTTCTGATACTTATGCATTCATGGTTACT
55	AAAGAGTTTGCCAGACAGAATAAAAATCACTAAGATCTCTGATCTTAAAAAATTATCAACAACCTATGAAGGCA
	GGGGTTGATAGTTCATGGATGAATCGCGAGGGAGATGGATACACTGATTTTCGCTAAAACATACGGTTTTGAA
	TTTTACATATTTACCTATGCAAATTGGCTTAGTCTATGATGCGGTTGAAAGTAACAAAATGCAATCTGTA
	TTAGGCTACTCCACTGACGGTCGTATTTTCGAGCTATGATTTAGAAATTTTAAGGGATGATAAAAAATTCTTT
	CCTCCTTATGAAGCCTCTATGGTTGTCAACAATTCTATCATCAAAAAAGATCCTAAACTAAAAAAATTACTC



- 71 -

CATCGACTCGATGGTAAAATCAATTTAAAAACGATGCAAAACCTTAATTATATGGTAGATGATAAACTTTTA  
GAACCTTCAGTTGTTGCCAAACAATTTTATAGAAAAAACCATTTATTTTAGAGGAGATAAATAA

ORF amino acid sequence (SEQ ID NO: 57):

5 MLKKSFLQIFTLCLALLTISGCQLDTPKSGHTTIKVAAQSSTESSIMANIITELIHHELGYNTTLISNLG  
SSTVTHQALLRGDADIAATRYTGTDITGTLGLKAVKDPKEASKIVKTEFQKRYNQTWYPTYGFSPTYAFMVT  
KEFARQNKITKISDLKLLSTTMKAGVDSSWMNREGDGYTDFAKTYGFEFSHIYPMQIGLVYDAVESNKMOSV  
LGYSTDGRISSYDLEILRDDKKFFPPYEASMVVNSIIKKDPKLLKLLHRLDGKINLKTMQNLNMYMVDKLL  
EPSVVAQKFLEKNHYFRGDK

#### 10 Strain COH1

ORF DNA sequence (SEQ ID NO: 64):

15 ATGCTTAAAAAATCGCACTTTTTACAGATATTTACACTTTGCTTAGCCCTCTTAACGATTTCTGGTTGTCAA  
TTAACCGATACTAAAAACCTGGTCATAACCACAATTAAGGTTGCTGCCCAAAGTTCTACAGAGTCTAGTATC  
ATGGCAAATATTGTCACCGAATTAATTCATCACGAATTAGGATACAACACAACCTTTAATAAGCAATCTTGGT  
20 TCCTCTACGGTTACTACCAAGCTTTGCTCCGTGGTGATGCTGACATTGCTGCCACACGTTATACAGGAACA  
GACATCACAGGAACCTTTGGCTTAAAAGCTGTTAAAGACCCCTAAAGAAGCTTCTAAGATTGTAAAAACTGAA  
TTCCAAAAACGCTACAATCAAACCTGGTATCCTACTTATGGTTTTTCTGATACTTATGCATTCATGGTTACT  
AAAGAGTTTGCCAGACAGAATAAAAATCACCAAGATCTCTGATCTCAAAAAGTTATCAACAACATGAAGGCA  
GGGGTTGATAGTTCATGGATGAATCGCGAGGGAGATGGATACACTGATTTTCGCTAAAACATACGGTTTTGAA  
25 TTTTCACATATTTACCTATGCAAATTGGCTTAGTCTATGATGCGGTTGAAAGTAACAAAATGCAATCTGTA  
TTAGGCTACTCCACTGACGGTCGTATTTTCGAGCTATGATTTAGAAATTTTAAGGGATGATAAAAAATTCTTT  
CCTCCTTATGAAGCCTCTATGGTTGTCAACAATTCATCATCAAAAAGATCCTAAACTAAAAAAATTACTC  
CATCGACTCGATGGTAAAATCAATTTAAAAACGATGCAAAACCTTAATTATATGGTAGATGATAAACTTTTA  
GAACCTTCAGTTGTTGCCAAACAATTTTATAGAAAAAACCATTTATTTTAGAGGAGATAAATAA

25 ORF amino acid sequence (SEQ ID NO: 58):

30 MLKKSFLQIFTLCLALLTISGCQLDTPKSGHTTIKVAAQSSTESSIMANIVTELIHHELGYNTTLISNLG  
SSTVTHQALLRGDADIAATRYTGTDITGTLGLKAVKDPKEASKIVKTEFQKRYNQTWYPTYGFSPTYAFMVT  
KEFARQNKITKISDLKLLSTTMKAGVDSSWMNREGDGYTDFAKTYGFEFSHIYPMQIGLVYDAVESNKMOSV  
LGYSTDGRISSYDLEILRDDKKFFPPYEASMVVNSIIKKDPKLLKLLHRLDGKINLKTMQNLNMYMVDKLL  
EPSVVAQKFLEKNHYFRGDK

#### Strain IC105

ORF DNA sequence (SEQ ID NO: 65):

35 ATGCTTAAAAAATCGCACTTTTTACAGATATTTACACTTTGCTTAGCCCTCTTAACGATTTCTGGTTGTCAA  
TTAACCGATACTAAAAAGTCTGGTCATAACCACAATTAAGGTTGCTGCCCAAAGTTCTACAGAGTCTAGTATT  
ATGGCAAATATTATCACCGAATTAATTCATCACGAATTAGGATACAACACAACCTTTAATAAGCAATCTTGGT  
40 TCCTCTACGGTTACTACCAAGCTTTGCTCCGTGGTGATGCTGACATTGCTGCCACACGTTATACAGGAACA  
GACATCACAGGGACTCTTGGTTTTAAAAGCTGTTAAAGACCCCTAAAGAAGCTTCTAAGATTGTAAAAACTGAA  
TTCCAAAAACGCTACAATCAAACCTGGTATCCTACTTATGGTTTTTCTGATACTTATGCATTCATGGTTACT  
AAAGAGTTTGCCAGACAGAATAAAAATCACTAAGATCTCTGATCTTAAAAAATTATCAACAACATGAAGGCA  
GGGGTTGATAGTTCATGGATGAATCGCGAGGGAGATGGATACACTGATTTTCGCTAAAACATACGGTTTTGAA  
45 TTTTCACATATTTACCTATGCAAATTGGCTTAGTCTATGATGCGGTTGAAAGTAACAAAATGCAATCTGTA  
TTAGGCTACTCCACTGACGGTCGTATTTTCGAGCTATGATTTAGAAATTTTAAGGGATGATAAAAAATTCTTT  
CCTCCTTATGAAGCCTCTATGGTTGTCAACAATTCATCATCAAAAAGATCCTAAACTAAAAAAATTACTC  
CATCGACTCGATGGTAAAATCAATTTAAAAACGATGCAAAACCTTAATTATATGGTAGATGATAAACTTTTA  
GAACCTTCAGTTGTTGCCAAACAATTTTATAGAAAAAACCATTTATTTTAGAGGAGATAAATAA

ORF amino acid sequence (SEQ ID NO: 59):

50 MLKKSFLQIFTLCLALLTISGCQLDTPKSGHTTIKVAAQSSTESSIMANIITELIHHELGYNTTLISNLG  
SSTVTHQALLRGDADIAATRYTGTDITGTLGLKAVKDPKEASKIVKTEFQKRYNQTWYPTYGFSPTYAFMVT  
KEFARQNKITKISDLKLLSTTMKAGVDSSWMNREGDGYTDFAKTYGFEFSHIYPMQIGLVYDAVESNKMOSV  
LGYSTDGRISSYDLEILRDDKKFFPPYEASMVVNSIIKKDPKLLKLLHRLDGKINLKTMQNLNMYMVDKLL  
EPSVVAQKFLEKNHYFRGDK

#### Strain IC458

ORF DNA sequence (SEQ ID NO: 66):

55 ATGCTTAAAAAATCGCACTTTTTACAGATATTTACACTTTGCTTAGCCCTCTTAACGATTTCTGGTTGTCAA  
TTAACCGATACTAAAAACCTGGTCATAACCACAATTAAGGTTGCTGCCCAAAGTTCTACAGAGTCTAGTATC  
ATGGCAAATATTGTCACCGAATTAATTCATCACGAATTA<sub>g</sub>GATACAACACAACCTTTAATAAGCAATCTTGGT  
60 TCCTCTACGGTTACTACCAAGCTTTGCTCCGTGGTGATGCTGACATTGCTGCCACACGTTATACAGGAACA  
GACATCAcAGGAACCTTTGGCTTAAAAGCTGTTAAAGACACTAAAGAAGCTTCTAAGATTGTAAAAACTGAA



TTCCAAAAACGCTACAATCAAACCTGGTATCCTACTTATGGTTTTTCTGATACTTATGCATTCATGGTTACT  
 AAAGTTTGGCCAGACAGAAATAAATCACCAAGATCTCTGATCTCAAAAAGTTATCAACAACTATGAAGGCA  
 GGGTTGATAGTTCATGGATGAATCGCGAGGGAGATGGATACACTGATTTTCGCTAAAACATACGGTTTTGAA  
 TTTTCACATATTTACCCTATGCAAATTGGCTTAGTCTATGATGCAGTTGAAAGTAACAAAATGCAATCTGTA  
 5 TTAGGCTACTCCACTGACGGTCGTATTTTCGAGCTATGATTTAGAAATTTTAAGGGATGATAAAAAATTCTTT  
 CCTCCTTATGAAGCCTCTATGGTTGTCAACAATTCTATCATCAAAAAGATCCTAAACTAAAAAATTACTC  
 CATCGACTCGATGGTAAAATCAATTTAAAAACGATGCAAAACCTTAATTATATGGTAGATGATAAACTTTTA  
 GAACCTTCAGTTGTTGCCAAACAATTTTATAGAAAAAACCATTTATTTTAGAGGAGATAAATAA

ORF amino acid sequence (SEQ ID NO: 60):

10 MLKKSFLQIFTLCLALLTISGCQLDTPKPGHTTIKVAQSSSTESSIMANIVTELIHHELGYNTTLISNLG  
 SSTVTHQALLRGDADIAATRYTGTDTITGLGLKAVKDTKEASKIVKTEFQKRYNQWYPTYGFSPTYAFMVT  
 KEFARQNKITKISDLKLLSTTMKAGVDSWWMNREGDGYTDFAKTYGFEFSHIYPMQIGLVYDAVESNKMOSV  
 LGYSTDGRISSYDLEILRDDKKFFPPYEASMVVNSIIKKDPKLLKLLHRLDGLKINLKTMLNLMVDDKLL  
 EPSVVAQFLEKNHYFRGDK

**Table 8: Amino acid and encoding DNA sequences of gbs1087 proteins derived from different strains of *S. agalactiae***

**Strain 0176H4A**

ORF DNA sequence (SEQ ID NO: 73):

20 TTGTTCAATAAAAATAGGTTTTAGAACTTGGAAATCAGGAAAGCTTTGGCTTTATATGGGAGTGCTAGGATCA  
 ACTATTATTTTAGGATCAAGTCCTGTATCTGCTATGGATAGTGTGGAAATCAAAGTCAGGGCAATGTTTTA  
 GAGCGTCGTCAACGCGATGCAGAAAACAGAAGCCAAGGTAATGTTCTAGAGCGTCGTCAACGCGATGCAGAA  
 AACAGAAGCCAAGGTAATGTTCTAGAGCGTCGTCAACGTGATGCGGAAAACAAGAGCCAAGTAGGTCAACTT  
 25 ATAGGGAAAAATCCACTTCTTTCAAAGTCAATTATATCTAGAGAAAATAATCACTCTAGTCAAGGTGACTCT  
 AACAAACAGTCATTCTCTAAAAAAGTATCTCAGGTTACTAATGTAGCTAATAGACCGATGTTAACTAATAAT  
 TCTAGAACAATTTTCAGTGATAAATAAATTACCTAAAACAGGTGATGATCAAAATGTCATTTTTTAACTTGTA  
 GGTTTTGGTTTAAATTTTGTTAACAAGTCGCTGCGGTTTGAGACGCAATGAAAATTAA

ORF amino acid sequence (SEQ ID NO: 67):

30 LFNKIGFRTWKSGKLWLYMGVLGSTIILGSSPVSAMDSVGNQSQGNVLERRQRDAENRSQGNVLERRQRDAE  
 NRSQGNVLERRQRDAENKSQVGQLIGKNPLLSKSIISRENNHSSQGDSNKQSFSSKVSQVTNVANRPLMTNN  
 SRTISVINKLPKTGDDQNVIFKLVGFGLILLTSRCGLLRNEN

**Strain 12401**

ORF DNA sequence (SEQ ID NO: 74):

35 TTGTTCAATAAAAATAGGTTTTAGAACTTGGAAATCAGGAAAGCTTTGGCTTTATATGGGAGTGCTAGGATCA  
 ACTATTATTTTAGGATCAAGTCCTGTATCTGCTATGGATAGTGTGGAAATCAAAGTCAAGGTAATGTTCTA  
 GAGCGTCGTCAACGTGATGCGGATAACAAGAGCCAAGGTAATGTTCTAGAACGTCGTCAACGCGATGTGGAA  
 AACAAAAGTCAGGGCAATGTTCTAGAACGTCGTCAACGTGATGTTGAGAATAAGAGCCAAGGCAATGTTCTA  
 40 GAGCGTCGCCAACGTGATGCAGAAAACAAAAGTCAGGGTAATGTTCTAGAGCGTCGTCAACGCGATGCAGAT  
 AACAGAGCCAAGGTAATGTTCTAGAACGTCGTCAACGCGATGTGGAAAACAAAAGTCAGGGCAATGTTCTA  
 GAGCGTCGCCAACGTGATGTTGAGAACAAGAGCCAAGTAGGTCAACTTATAGGGAAAAATCCACTTCTTTCA  
 AAGTCAACTATATCTAGAGAAAATAATCACTCTAGTCAAGGTGACTCTAACAAACAGTCATTCTCTAAAAA  
 GTATCTCAGGTTACTAATGTAGCTAATAGACCAATGTTAACTAATAAATTTCTAGAACAATTTTCAGTGATAAAT  
 45 AAATTACCTAAAACAGGTGATGATCAAAATGTCATTTTTTAACTTGTAGGTTTTGGTTTAAATTTTGTTAACA  
 AGTCGCTGCGGTTTGAGACGCAATGAAAATTAA

ORF amino acid sequence (SEQ ID NO: 68):

50 LFNKIGFRTWKSGKLWLYMGVLGSTIILGSSPVSAMDSVGNQSQGNVLERRQRDADNKSQGNVLERRQRDVE  
 NKSQGNVLERRQRDVENKSQGNVLERRQRDAENKSQGNVLERRQRDADNKSQGNVLERRQRDVENKSQGNV  
 LERQRDVENKSQVGQLIGKNPLLSKSTISRENNHSSQGDSNKQSFSSKVSQVTNVANRPLMTNNSRTISVIN  
 KLPKTGDDQNVIFKLVGFGLILLTSRCGLLRNEN

**Strain BAA23**

ORF DNA sequence (SEQ ID NO: 75):

55 TTGTTCAATAAAAATAGGTTTTAGAACTTGGAAATCAGGAAAGCTTTGGCTTTATATGGGAGTGCTAGGATCA  
 ACTATTATTTTAGGATCAAGTCCTGTATCTGCTATGGATAGTGTGGAAATCAAAGTCAAGGGCAATGTTTTA  
 GAGCGTCGTCAACGTGATGCAGAAAACAGAAGCCAAGGTAATGTTCTAGAGCGTCGTCAACGTGATGCAGAA



AACAGAAGCCAAGGTAATGTTCTAGAGCGTCGTCAACGCGATGCAGAAAACAGAAGCCAAGGTAATGTTCTA  
 GAGCGTCGTCAACGTGATGTTGAGAATAAGAGCCAAGGCAATGTTTTAGAGCGTCGTCAACGCGATGTTGAG  
 AATAAGAGCCAAGGTAATGTTCTAGAGCGTCGTCAACGTGATGCGGAAAACAAGAGCCAAGGCAATGTTCTA  
 GAGCGTCGTCAACGTGATGCGGAAAACAAGAGCCAAGGCAATGTTCTAGAGCGTCGTCAACGCGATGCAGAA  
 5 AACAGAAGCCAAGGTAATGTTTTAGAACGTCGTCAACGCGATGTTGAGAACAAGAGCCAAGGTAACGTTCTA  
 GAGCGTCGCCAACGTGACGTTGAGAACAAGAGCCAAGGTAATGTTTTAGAGCGTCGCCAACGCGATGCGGAT  
 AACAAAAGTCAGGGCAATGTTTTAGAGCGTCGCCAACGTGATGTTGAGAACAAGAGCCAAGGTAATGTTCTA  
 GAGCGTCGCCAAAATAATGTCCTTATTAAGAGTCAAGATAATGTTCTAGAGCGCCGCCAACGTGATGCGGAT  
 10 AACAGAGCCAGGGTAACGTTCTAGAGCGTCGTCAACGCGATGTTGAGAATAAGAGCCAAGGTAATGTTTTA  
 GAGCGTCGCCAACATGATGTTGAGAATAAGAGTCAAGTAGGTCAACTTATAGGGAAAAATCCACTTTTTTCA  
 AAGTCAACTGTATCTAGAGAAAATAATCACTCTAGTCAAGGTGACTCTAACAAACAGTCATTCTCTAAAAAA  
 GTATCTCAGGTTACTAATGTAGCTAATAGACCGATGTTAACTAATAATTCTAGAACAATTTTCAGTGATAAAT  
 AAATTACCTAAAACAGGTGATGATCAAAATGTCATTTTTAACTTGTAGGTTTTGGTTTAAATTTTATTAACA  
 AGTCTCTGCGGTTTGAGACGCAATGAAAATTA

15 ORF amino acid sequence (SEQ ID NO: 69):

LFNKIGFRTWKSGKLWLYMGVLGSTIILGSSPVSAMDSVGNQSQGNVLERRQRDAENRSQGNVLERRQRDAE  
 NRSQGNVLERRQRDAENRSQGNVLERRQRDVENKSQGNVLERRQRDVENKSQGNVLERRQRDAENKSQGNV  
 20 ERRQRDAENKSQGNVLERRQRDAENRSQGNVLERRQRDVENKSQGNVLERRQRDVENKSQGNVLERRQRDAD  
 NKSQGNVLERRQRDVENKSQGNVLERRQNNVLIKSQDNVLERRQRDADNKSQGNVLERRQRDVENKSQGNV  
 ERRQHDVENKSQVQQLIGKNPLFSKSTVSRENNHSSQGDSNKQSFSSKVSQVTNVANRPMLTNNRSRTISVIN  
 KLPKTGDDQNVIFKLVGFGLILLTSLCGLRRNEN

#### Strain COH1

25 ORF DNA sequence (SEQ ID NO: 76):

TTGTTCAATAAAATAGGTTTTAGAACTTGAAATCAGGAAAGCTTTGGCTTTATATGGGAGTGCTAGGATCA  
 ACTATTATTTTAGGATCAAGTCCTGTATCTGCTATGGATAGTGTGGAAATCAAAGTCAAGGTAATGTTTTA  
 GAGCGTCGCCAACGTGATGCGGAAAACAAGTCAAGGTAATGTTTTAGAGCGTCGCCAACGTGATGCGGAA  
 30 AACAGAGCCAAGGCAATGTTTTAGAGCGTCGTCAACGCGATGTTGAGAATAAGAGCCAAGGCAATGTTTTA  
 GAGCGTCGTCAACGTGATGCGGAAAACAAGTCAAGGCAATGTTCTAGAGCGCCGCCAACGTGATGCGGAT  
 AACAGAGCCAAGTAGGTCAACTTATAGGGAAAAATCCACTTTTTTCAAAGCCAAGTGTATCTAGAGAAAAT  
 AATCACTCTAGTCAAGGTGACTCTAACAAACAGTCATTCTCTAAAAAAGTATCTCAGGTTACTAATGTAGCT  
 AATAGACCGATGTTAACTAATAATTCTAGAACAATTTTCAGTGATAAATAAATTACCTAAAACAGGTGGTGAT  
 CAAAATGTCATTTTTAACTTGTAGGTTTTGGTTTAAATTTTGTAAACAAGTCGCTGCGGTTTGAGACGCAAT  
 GAAAATTA

35 ORF amino acid sequence (SEQ ID NO: 70):

LFNKIGFRTWKSGKLWLYMGVLGSTIILGSSPVSAMDSVGNQSQGNVLERRQRDAENKSQGNVLERRQRDAE  
 NKSQGNVLERRQRDVENKSQGNVLERRQRDAENKSQGNVLERRQRDADNKSQVQQLIGKNPLFSKPTVSREN  
 40 NHSSQGDSNKQSFSSKVSQVTNVANRPMLTNNRSRTISVINKLPKTGGDQNVIFKLVGFGLILLTSLCGLRRN  
 EN

#### Strain IC105

45 ORF DNA sequence (SEQ ID NO: 77):

TTGTTCAATAAAATAGGTTTTAGAACTTGAAATCAGGAAAGCTTTGGCTTTATATGGGAGTGCTAGGATCa  
 actattatthtttaggatcaagtcctgtttctgCTATGGATAGTGTGGAAATCAAAGTCAAGGCAATGTTTTA  
 50 GAGCGTCGTCAACGTGATGCGGAAAACAAGAGCCAAGGCAATGTTCTAGAGCGTCGTCAACGCGATGCAGAA  
 AACAGAAGCCAAGGTAATGATCTAGAGCGTCGCCAACGTGACGTTGAGAACAAGAGCCAAGGTAACGTTCTA  
 GAGCGTCGTCAACGTGATGCAGATAACAAGAGCCAGGGCAATGTTTTAGAGCGTCGCCAACGTGATGTTGAG  
 AACAGAGCCAAGGTAATGTTCTAGAGCGTCGCCAAAATAATGTCCTTATTAAGAGTCAAGATAATGATCTA  
 GAGCGCCGCCAACGTGATGCGGATAACAAGAGCCAGGGTAACGTTCTAGAGCGTCGCCAACGTGATGTTGAG  
 55 AACAGAGCCAAGGTAATGTTCTAGAGCGCCGCCAACGTGATGCGGATAACAAGAGCCAGGGTAACGTTCTA  
 GAGTGTGCGCAACGTGATGTTGAGAACAAGAGCCAAGGTAATGTTCTAGAGCGTCGTCAACGTGATGCGGAA  
 AACAGAGCCAAGGCAATGTTCTAGAGCGTCGTCAACGCGATGCAGAAAACAAGAGCCAAGGTAATGTTCTA  
 GAGCGTCGCCAACGTGACGTTGAGAACAAGAGCCAAGGTAACGTTCTAGAGCGTCGTCAACGTGATGCAGAT  
 AACAGAGCCAGGGCAATGTTTTAGAGCGTCGCCAACGTGATGTTGAGAACAAGAGCCAAGGTAATGTTCTA  
 GAGCGTCGCCAAAATAATGTCCTTATTAAGAGTCAAGATAATGTTCTAGAGCGCCGCCAACGTGATGCGGAT  
 AACAGAGCCAGGGTAACGTTCTAGAGCGTCGCCAACGTGATGTTGAGAACAAGAGCCAAGGCAATGTTTTA  
 GAGCGTCGTCAACGCGATGTTGAGAATAAGAGTCAAGTAGGTCAACTTATAGGGAAAAATCCACTTTTTTCA  
 AAGTCAACTGTATCTAGAGAAAATAATCACTCTAGTCAAGGTGACTCTAACAAACAGTCATTCTCTAAAAAA  
 GTATCTCAGGTTACTAATGTAGCTAATAGACCGATGTTAACTAATAATTCTAGAACAATTTTCAGTGATAAAT



AAATTACCTAAAACAGGTGATGATCAAAATGTCATTTTTAAACTTGTAGGTTTTGGTTTAAATTTTATTAACA  
AGTCTCTGCGGTTTGAGACGCAATGAAAATTAA

ORF amino acid sequence (SEQ ID NO: 71):

LFNKIGFRTWKSGKLWLYMGVLGSTIILGSSPVSAMDSQGNVLERRQRDAENKSQGNVLERRQRDAE  
5 NRSQGNLERRQRDVENKSQGNVLERRQRDADNKSQGNVLERRQRDVENKSQGNVLERRQNNVLIKSQDNDL  
ERRQRDADNKSQGNVLERRQRDVENKSQGNVLERRQRDADNKSQGNVLECRQRDVENKSQGNVLERRQRDAE  
NKSQGNVLERRQRDAENRSQGNVLERRQRDVENKSQGNVLERRQRDADNKSQGNVLERRQRDVENKSQGNV  
10 ERRQNNVLIKSQDNLERRQRDADNKSQGNVLERRQRDVENKSQGNVLERRQRDVENKSQVQQLIGKNPLFS  
KSTVSRENNHSSQGDSNKQSFSSKVSQVTNVANRPLMTNNSRTISVINKLPKTGDDQNVIFKLVGFGLILLT  
SLCGLRRNEN

#### Strain IC458

ORF DNA sequence (SEQ ID NO: 78):

TTGTTCAATAAAAATAGGTTTTAGAACTTGGAATCAGGAAAGCTTTGGCTTTATATGGGAGTGCTAGGATCA  
15 ACTATTATTTTAGGATCAAGTCCTGTATCTGCTATGGATAGTGTGGAAATCAAAGTCAGGGCAATGTTTTA  
GAGCGTCGTCAACGCGATGTTGAGAATAAGAGCCAAGGTAATGTTCTAGAGCGTCGTCAACGTGATGCGGAA  
ACAAGAGCCAAGGCAATGTTTTAGAGCGTCGTCAACGTGATGCAGAAAACAGAAGCCAAGGCAATGTTTTA  
GAGCGTCGTCAACGTGATGCAGAAAACAGAAGCCAAGGCAATGTTTTAGAGCGTCGTCAACGCGATGTTGAG  
20 AATAAGAGCCAAGGTAATGTTCTAGAGCGTCGTCAACGCGATGTTGAGAATAAGAGCCAAGGTAATGTTCTA  
GAGCGTCGTCAACGTGATGCGGAAAACAAGAGCCAAGGCAATGTTTTAGAGCGTCGTCAACGTGATGCAGAA  
AACAGAAGCCAAGGCAATGTTTTAGAGCGTCGTCAACGTGATGCAGAAAACAGAAGCCAAGGCAATGTTTTA  
GAGCGTCGTCAACGCGATGTTGAGAATAAGAGCCAAGGCAATGTTTTAGAGCGTCGTCAACGTGATGCAGAA  
AACAGAAGCCAAGGCAATGTTTTAGAGCGTCGTCAACGTGATGCAGAAAACAGAAGCCAAGTAGGTCAACTT  
25 ATAGGGAAAAATCCACTTCTTTCAAAGTCAATTATATCTAGAGAAAATAATCACTCTAGTCAAGGTGACTCT  
AACAAACAGTCATTCTCTAAAAAAGTATCTCAGGTTACTAATGTAGCTAATAGACCGATGTTAACTAATAAT  
TCTAGAACAATTTTCAGTGATAAATAAATTACCTAAAACAGGTGATGATCAAAATGTCATTTTTAACTTGTA  
GGTTTTGGTTTAAATTTTGTTAACAAGTCGCTGCGGTTTGAGACGCAATGAAAATTAA

ORF amino acid sequence (SEQ ID NO: 72):

LFNKIGFRTWKSGKLWLYMGVLGSTIILGSSPVSAMDSVGNQSQGNVLERRQRDVENKSQGNVLERRQRDAE  
30 NKSQGNVLERRQRDAENRSQGNVLERRQRDAENRSQGNVLERRQRDVENKSQGNVLERRQRDVENKSQGNV  
ERRQRDAENKSQGNVLERRQRDAENRSQGNVLERRQRDAENRSQGNVLERRQRDVENKSQGNVLERRQRDAE  
NRSQGNVLERRQRDAENRSQVQQLIGKNPLLSKSIISRENNHSSQGDSNKQSFSSKVSQVTNVANRPLMTN  
SRTISVINKLPKTGDDQNVIFKLVGFGLILLTSRCGLRRNEN

35

**Table 9: Amino acid and encoding DNA sequences of gbs1309 proteins derived from different strains of *S. agalactiae***

#### Strain 0176H4A

ORF DNA sequence (SEQ ID NO: 85):

TTTAGTGTAACCTATTCACAGTCTGAACGTACGGTTGTTTTCTTTTTGGAGAAATAACATTTAGTAGGAGT  
40 CGCTGGACAAATGGCTTTGAAACTAGAATACCAGTAGATGAGTGGTTAGGTCTTGAAAAATATAAGAGATAT  
TCAATAGAATTCTTATATCATGTTGCAAAATTGGCTACAATGATGCCTTATCGTCAAGTTTGCAAAGTAATA  
GATAGCACTTTGCAAACAATCATAACAAAAGACTGTGTTTTAAAAGCAGTAAAATTTGTAGAAAAATTGTTA  
AAAGAAAAAGAACGCTATCGTTTTTTATTTGGAAGAGCCACCCGAACGTAAAAAAGTAAAAAACTGTATGTT  
45 GAGGGTGATGGAGTCATGATTAAAAGCACAGATTCTAGAGAGGAAAGAAGGTATTTAGATTTAACACATTTT  
GTTATTCATACAGGCTCAAAAAAAGTTTCTACTAAAAGATATGAATTGCAGGACAAGCACGAAATATTACAG  
CTTAATTATGATAAAGCTAAATATAATCTTTTAGATTATATTTATAAATAACTATGAAGTAGATGACGATACT  
ATTTTAATCACTAACTCTGATATGGGTAAAGGCTATACTAGTAGAGTTTTTAAGGAATTAGGAAAAGCACTT  
AAGGTAAAGAAACATGAGCATTTTTTGGGATATCTATCATGTTAAAGAAAAGTTAAGTTCATACCTTAGAAAA  
50 TATCCAATTGAATTAACCGATTTTACTTTAGATGCGGTAAAAAATATAATTCTGATAAGCTTGAATTAGTT  
TTTGATACTGTTGAATCACTGATTTGTGATGAACCTGAAGATCAAGAATTTGAGAAGTTAAGAAAAAAGTA  
TTAAATAATTTCAAATATATAAAACCAGCTCATCTTAGAAATCTTTCAAATCGTGGTATTGGTATCATGGAA  
TCACAACACAGAAAGATAACGTATAGAATGAAGCGACGTGGCATGTATTGGTCAAAGTGGGGAATCTCCACA  
ATGGCAAATATGATTATATTTGAAAGAGCTAACGGTTTACGAGAATTATTTTTCGGTTCTTGGAGAAAGGTA  
55 TACAGTGAGTATAAAGAAGGTTTCAATTTAGTGCAGGGCGACTTTTTAAAAGACAGATGAATTAGATAAATTT  
TCTAAGCCCCTTCTAAAAAATGGCAGAAAATGGAGTATAACAGGAATCAAACAAAATAG

ORF amino acid sequence (SEQ ID NO: 79):



FSVTYSQSERTVVFSFGEITFSRSRWTNGFETRI PVDEWLGLEKYKRY SIEFLYHVAKLATMMPYRQVCKVI  
 DSTLQTIITKDCVLKAVK FVEKLLKEKERYRFYLEEPPERKKVKKLYVEGDGVMIKSTDSREERRYLDLTHF  
 VIHTGSKKVSTKRYELQDKHEILQLNYDKAKYNLLDYIYNNYEVD DDTILITNSDMGKGYTSRVFKELGKAL  
 KVKKHEHFWDIYHVKEKLSSYL RKYPIELTDFTLDAVKKYN SDKLELVFDTVESLICDELEDQEFQKFKKKV  
 5 LNNFKYIKPAHLRNLSNRGIGIMESQHRKITYRMKRRGMYWSKWGI STMANMIIFERANGLRELF FGSWRKV  
 YSEYKEGSFSAGRLF KKTDELDFSKPLLKNGR KWSITGIKTK

**Strain 12401**

ORF DNA sequence (SEQ ID NO: 86):

10 TTTAGTGTAACCTATTCACAGTCTGAACGTACGGTTGTTTTCTCTTTTGGAGAAATAACATTTAGTAGGAGT  
 CGCTGGACAAATGGCTTTGAACTAGAATACCAGTAGATGAGTGGTTAGGTCTTGAAAAATATAAGAGATAT  
 TCAATAGAATTCTTATATCATGTTGCAAATTTGGCTACAATGATGCCTTATtGTCAAGTTTGCAAAGTAATA  
 GATAGCACTTTGCAAACAATCATAACAAAAGACTGTGTTTTAAAAGCAGTAAAATTTGTAGAAAAATTGTTA  
 AAAGAAAAAGAACGCTATCGTTTTTATTTGGAAGAGCCACCCGAACgTAAAAAAGtGAAAAAAGTGTATGTT  
 15 GaGGGTGATGGAGTCATGATTAAGCACAGATTCTAGAGAGGAAAGAAGGTATTTAGATTTAACACATTTT  
 GTTATTCATACAGGCTCAAAAAAAGTTTCTACTAAAAGATATGAATTGCAGGACAAGCACGAAATATTACAG  
 CTTAATTATGATAAAGCTAAATATAATCTTTTAGATTATATTTATAATAACTATGAAGTAGATGACGATACT  
 ATTTTAATCACTAACTCTGATATGGGTAAAGGCTATACTAGTAGAGTTTTTAAGGAATTAGGAAAAGCACTT  
 AAGGTAAAGAAACATGAGCATTTTTGGGATATCTATCATGTtAAAGAAAAGTTAAGTTCATACCTTAGAAAA  
 20 TATCCAATTGAATTAACCGATTTTGCTTTAGATGCGGTAAAAAATATAAATCTGATAAGCTTGAATTAGTT  
 TTTGATACTGTTGAATCACTGATTTGTGATGAACTTGAAGATCAAGAATTT CAGAAGTTTAAGAAAAAAGTA  
 TTAAATAATTTCAAATATATAAAAACCAGCTCATCTTAGAAATCTTTCAAATCGTGGTATTGGTATCATGGAA  
 TCACAACACAGAAAGATAACGTATAGAATGAAGCGACGTGGCATGTATTGGTCAAAGTGGGGAATCTCCACA  
 ATGGCAAATATGATTATATTTGAAAGAGCTAACGGTTTACGAGAATTATTTTTTCGGTTCCTGGAGAAAGGTA  
 25 TACAGTGAGTATAAAGAAGGTTTCAATTTAGTGCAGGGCGACTTTTTAAAAGACAGATGAATTATATAAATTT  
 TCTAAGCCCCTTCTAAAAAATGGCAGAAAATGGAGTATAACAGGAATCAAACAAAATAG

ORF amino acid sequence (SEQ ID NO: 80):

30 FSVTYSQSERTVVFSFGEITFSRSRWTNGFETRI PVDEWLGLEKYKRY SIEFLYHVAKLATMMPYCQVCKVI  
 DSTLQTIITKDCVLKAVK FVEKLLKEKERYRFYLEEPPERKKVKKLYVEGDGVMIKSTDSREERRYLDLTHF  
 VIHTGSKKVSTKRYELQDKHEILQLNYDKAKYNLLDYIYNNYEVD DDTILITNSDMGKGYTSRVFKELGKAL  
 KVKKHEHFWDIYHVKEKLSSYL RKYPIELTDFALDAVKKYN SDKLELVFDTVESLICDELEDQEFQKFKKKV  
 LNNFKYIKPAHLRNLSNRGIGIMESQHRKITYRMKRRGMYWSKWGI STMANMIIFERANGLRELF FGSWRKV  
 YSEYKEGSFSAGRLF KKTDELYKFSKPLLKNGR KWSITGIKTK

**Strain BAA23**

ORF DNA sequence (SEQ ID NO: 87):

35 TTTAGTGTAACCTATTCACAGTCTGAACGTACGGTTGTTTTCTCTTTTGGAGAAATAACATTTAGTAGGAGT  
 CGCTGGACAAATGGCTTTGAACTAGAATACCAGTAGATGAGTGGTTAGGTCTTGAAAAATATAAGAGATAT  
 TCAATAGAATTCTTATATCATGTTGCAAATTTGGCTACAATGATGCCTTATCGTCAAGTTTGCAAAGTAATA  
 40 GATAGCACTTTGCAAACAATCATAACAAAAGACTGTGTTTTAAAAGCAGTAAAATTTGTAGAAAAATTGTTA  
 AAAGAAAAAGAACGCTATCGTTTTTATTTGGAAGAGCCACCCGAACGTAAAAAAGTAAAAAAGTGTATGTT  
 GAGGGTGTGATGGAGTCATGATTAAGCACAGATTCTAGAGAGGAAAGAAGGTATTTAGATTTAACACATTTT  
 GTTATTCATACAGGCTCAAAAAAAGTTTCTACTAAAAGATATGAATTGCAGGACAAGCACGAAATATTACAG  
 CTTAATTATGATAAAGCTAAATATAATCTTTTAGATTATATTTATAATAACTATGAAGTAGATGACGATACT  
 45 ATTTTAATCACTAACTCTGATATGGGTAAAGGCTATACTAGTAGAGTTTTTAAGGAATTAGGAAAAGCACTT  
 AAGGTAAAGAAACATGAGCATTTTTGGGATATCTATCATGTAAAGAAAAGTTAAGTTCATACCTTAGAAAA  
 TATCCAATTGAATTAACCGATTTTGCTTTAGATGCGGTAAAAAATATAAATCTGATAAGCTTGAATTAGTT  
 TTTGATACTGTTGAATCACTGATTTGTGATGAACTTGAAGATCAAGAATTT CAGAAGTTTAAGAAAAAAGTA  
 TTAAATAATTTCAAATATATAAAAACCAGCTCATCTTAGAAATCTTTCAAATCGTGGTATTGGTATCATGGAA  
 50 TCACAACACAGAAAGATAACGTATAGAATGAAGCGACGTGGCATGTATTGGTCAAAGTGGGGAATCTCCACA  
 ATGGCAAATATGATTATACTTGAAGAGCTAACGGTTTACGAGAATTATTTTTTCGGTTCCTGGAGAAAGGTA  
 TACAGTGAGTATAAAGAAGGTTTCAATTTAGTGCAGGGCGACTTTTTAAAAGACAGATGAATTAGATAAATTT  
 TCTAAGCCCCTTCTAAAAAATGGCAGAAAATGGAGTATAACAGGAATCAAACAAAATAG

ORF amino acid sequence (SEQ ID NO: 81):

55 FSVTYSQSERTVVFSFGEITFSRSRWTNGFETRI PVDEWLGLEKYKRY SIEFLYHVAKLATMMPYRQVCKVI  
 DSTLQTIITKDCVLKAVK FVEKLLKEKERYRFYLEEPPERKKVKKLYVEGDGVMIKSTDSREERRYLDLTHF  
 VIHTGSKKVSTKRYELQDKHEILQLNYDKAKYNLLDYIYNNYEVD DDTILITNSDMGKGYTSRVFKELGKAL  
 KVKKHEHFWDIYHVKEKLSSYL RKYPIELTDFALDAVKKYN SDKLELVFDTVESLICDELEDQEFQKFKKKV  
 LNNFKYIKPAHLRNLSNRGIGIMESQHRKITYRMKRRGMYWSKWGI STMANMIIFERANGLRELF FGSWRKV  
 60 YSEYKEGSFSAGRLF KKTDELDFSKPLLKNGR KWSITGIKTK



**Strain COH1**

ORF DNA sequence (SEQ ID NO: 88):

5 ATGGAAGTTAAAAATTCTCGGAAAAAGATTTTGTAAATGAAATAAATAAAAACAGAAACAATTTT  
 AGTCAAATTGAACAGTATGAAAGCTATATCGCTCCTCAAATGAGAACGAAAGGCTATAAGAGGATCAATCAG  
 TCTGAACGTACGGTTGTTTTCTCTTTTGGAGAAATAACATTTAGTAGGAGTCGCTGGACAAATGGCTTTGAA  
 ACTAGAATACCAGTAGATGAGTGGTTAGGTCTTGAAAAATATAAGAGATATTCAATAGAATTCTTATATCAT  
 GTTGCAAAATTGGCTACAATGATGCCTTATCGTCAAGTTTGCAAAGTAATAGATAGCACTTTGCAAACAATC  
 10 ATAACAAAAGACTGTGTTTTAAAAGCAGTAAAATTTGTAGAAAAATTGTTAAAAGAAAAAGAACGCTATCGT  
 TTTTATTTGGAAGAGCCACCCGAACGTAAAAAAGTGAAAAAACTGTATGTTGAGGGTGATGGAGTCATGATT  
 AAAAGCACAGATTTAGAGAGGAAAGAAGGTATTTAGATTTAACACATTTTGTATTTCATACAGGCTCAAAA  
 AAAGTTTCTACTAAAAGATATGAATTGCGGGACAAGCACGAAATATTACAGCTTAATTATGATAAAGCTAAA  
 TATAATCTTTTAGATTATATTTATAATAACTATGAAGTAGATGACGATACTATTTTAATCACTAACTCTGAT  
 ATGGGTAAAGGCTATACTAGTAGAGTTTTTAAGGAATTAGGAAAAGCACTTAAGGTAAAGAAACATGAGCAT  
 15 TTTTGGGATATCTATCATGTTAAAGAAAAGTTAAGTTCATACCTTAGAAAATATCCAATTGAATTAACCGAT  
 TTTGCTTTAGATGCGGTAAAAAATATAATTCTGATAAGCTTGAATTAGTTTTTGATACTGTTGAATCACTG  
 ATTTGTGATGAACCTTGAAGATCAAGAATTTGAGAAGTTTAAGAAAAAAGTATTAATAAATTTCAAATATATA  
 AAACCAGCTCATCTTAGAAATCTTTCAAATCGTGGTATTGGTATCATGGAATCACAACACAGAAAGATAACG  
 TATAGAATGAAGCGACGTGGCATGTATTGGTCAAAGTGGGGAATCTCCACAATGGCAAATATGATTATACTT  
 20 GAAAGAGCTAACGGTTTACGAGAATTATTTTTCGGTCTTGGAGAAAGGTATACAGTGAGTATAAAGAAGGT  
 TCATTTAGTGCAGGGCGACTTTTTTAAAAGACAGATGAATTAGATAAATTTTCTAAGCCCCTTCTAAAAAAT  
 GGCAGAAAATGGAGTATAACAGGAATCAAAACAAAATAG

ORF amino acid sequence (SEQ ID NO: 82):

25 MEVKKFSEKDFVNEINKIKQKQFLSQIEQYESYIAPQMRTKGYKRINQSERTVVFSFGEITFSRSRWTNGFE  
 TRIPVDEWLGLEKYKRYSEFLYHVAKLATMMPYRQVCKVIDSTLQTIITKDCVLKAVKFVEKLLKEKERYR  
 FYLEEPPERKKVKKLYVEGDGVMIKSTDSREERRYLDLTHFVIHTGSKKVSTKRYELQDKHEILQLNYDKAK  
 YNLLDYIYNNYEVDLDDTILITNSDMGKGYTSRVFKELGKALKVKKHEHFWDIYHVKEKLSSYLKYP IELTD  
 FALDAVKKYNSDKLELVFDTVESLICDELEDQEFQKFKKKVLNFKYIKPAHLRNLNLRGIGIMESQHRKIT  
 30 YRMKRRGMYWSKWI STMANMI I LERANGLRELFFGSRKVYSEYKEGSFSAGRLFKKTDELDFSKP L LKN  
 GRKWSITGIKTK

**Strain IC105**

ORF DNA sequence (SEQ ID NO: 89):

35 TTTAGTGTAACCTATTCACAGTCTGAACGTACGGTTGTTTTCTCTTTTGGAGAAATAACATTTAGTAGGAGT  
 CGCTGGACAAATGGCTTTGAAACTAGAATACCAGTAGATGAGTGGTTAGGTCTTGAAAAATATAAGAGATAT  
 TCAATAGAATTCTTATATCATGTTGCAAAATTGGCTACAATGATGCCTTATCGTCAAGTTTGCAAAGTAATA  
 GATAGCACTTTGCAAACAATCATAACAAAAGACTGTGTTTTAAAAGCAGTAAAATTTGTAGAAAAATTGTTA  
 AAAGAAAAAGAACGCTATCGTTTTTATTTGGAAGAGCCACCCGAACGTAAAAAAGTGAAAAAACTGTATGTT  
 GAGGGTGATGGAGTCATGATTAAGCACAGATTTCTAGAGAGGAAAGAAGGTATTTAGATTTAACACATTTT  
 40 GTTATTTCATACAGGCTCAAAAAAAGTTTTCTACTAAAAGATATGAATTGCAGGACAAGCACGAAATATTACAG  
 CTTAATTATGATAAAGCTAAATATAATCTTTTAGATTATATTTATAATAACTATGAAGTAGATGACGATACT  
 ATTTTAATCACTAACTCTGATATGGGTAAAGGCTATACTAGTAGAGTTTTTAAGGAATTAGGAAAAGCACTT  
 AAGGTAAAGAAACATGAGCATTTTTGGGATATCTATCATGTTAAAGAAAAGTTAAGTTCATACCTTAGAAAA  
 TATCCAATTGAATTAACCGATTTTGTCTTTAGATGCGGTAAAAAATATAATTCTGATAAGCTTGAATTAGTT  
 45 TTTGATACTGTTGAATCACTGATTTGTGATGAACCTTGAAGATCAAGAATTTGAGAAGTTTAAGAAAAAAGTA  
 TTAAATAATTTCAAATATATAAAACCAGCTCATCTTAGAAATCTTTCAAATCGTGGTATTGGTATCATGGAA  
 TCACAACACAGAAAGATAACGTATAGAATGAAGCGACGTGGCATGTATTGGTCAAAGTGGGGAATCTCCACA  
 ATGGCAAATATGATTATACTTGAAGAGCTAACGGTTTACGAGAATTATTTTTCGGTCTTGGAGAAAGGTA  
 TACAGTGAGTATAAAGAAGGTTTCAATTTAGTGCAGGGCGACTTTTTTAAAAGACAGATGAATTAGATAAATTT  
 50 TCTAAGCCCCTTCTAAAAAATGGCAGAAAATGGAGTATAACAGGAATCAAAACAAAATAG

ORF amino acid sequence (SEQ ID NO: 83):

55 FSVTYSQSERTVVFSFGEITFSRSRWTNGFETRI PVDEWLGLEKYKRYSEFLYHVAKLATMMPYRQVCKVI  
 DSTLQTIITKDCVLKAVKFVEKLLKEKERYRFYLEEPPERKKVKKLYVEGDGVMIKSTDSREERRYLDLTHF  
 VIHTGSKKVSTKRYELQDKHEILQLNYDKAKYNLLDYIYNNYEVDLDDTILITNSDMGKGYTSRVFKELGKAL  
 KVKKHEHFWDIYHVKEKLSSYLKYP IELTDFALDAVKKYNSDKLELVFDTVESLICDELEDQEFQKFKKKV  
 LNNFKYIKPAHLRNLNLRGIGIMESQHRKIT YRMKRRGMYWSKWI STMANMI I LERANGLRELFFGSRKV  
 YSEYKEGSFSAGRLFKKTDELDFSKP L LKNGRKWSITGIKTK

**Strain IC458**

ORF DNA sequence (SEQ ID NO: 90):

60



TTTAGTGTAACCTATTCACAGTCTGAACGTACGGTGTGTTTTCTCTTTTGGAGAAATAACATTTAGTAGGAGT  
 CGCTGGACAAATGGCTTTGAAACTAGAATACCAGTAGATGAGTGGTTAGGTCTTGAAAAATATAAGAGATAT  
 TCAATAGAATTCTTATATCATGTTGCAAAATTGGCTACAATGATGCCTTATCGTCAAGTTTGCAAAGTAATA  
 5 GATAGCACTTTGCAAACAATCATAACAAAAGACTGTGTTTTAAAAGCAGTAAAATTTGTAGAAAAATTGTTA  
 AAAGAAAAAGAACGCTATCGTTTTTTATTTGGAAGAGCCACCCGAACGTAAAAAAGTGAAAAAAGTGTATGTT  
 GAGGGTGATGGAGTCATGATTAAGACACAGATTCTAGAGAGGAAAGAAGGTATTTAGATTTAACACATTTT  
 GTTATTCATACAGGCTCAAAAAAAGTTTCTACTAAAAGATATGAATTGCAGGACAAGCACGAAATATTACAG  
 CTTAATTATGATAAAGCTAAATATAATCTTTTAGATTATATTTATAATAACTATGAAGTAGATGACGATACT  
 ATTTTAATCACTAACTCTGATATGGGTAAAGGCTATACTAGTAGAGTTTTTAAGGAATTAGGAAAAGCACTT  
 10 AAGGTAAAGAAACATGAGCATTGTTGGGATATCTATCATGTTAAAGAAAAGTTAAGTTCATACCTTAGAAAA  
 TATCCAATTGAATTAACCGATTTTGCCTTAGATGCGGTAAAAAATATAATTTCTGATAAGCTTGAATTAGTT  
 TTTGATACTGTTGAATCACTGATTTGTGATGAACTTGAAGATCAAGAATTTTCAAGAGTTTAAAGAAAAAGTA  
 TTAAATAATTTCAAATATATAAAAACCAGCTCATCTTAGAAATCTTCAAATCGTGGTATTGGTATCATGGAA  
 TCACAACACAGAAAGATAACGTATAGAATGAAGCGACGTGGCATGTATTGGTCAAAGTGGGGAATCTCCACA  
 15 ATGGCAAATATGATTATACTTGAAGAGCTAACGGTTTACGAGAATTATTTTTCGGTTCTTGGAGAAAGGTA  
 TACAGTGAGTATAAAGAAGGTTTCAATTTAGTGCAGGGCGACTTTTTAAAAGACAGATGAATTAGATAAATTT  
 TCTAAGCCCCTTCTAAAAAATGGCAGAAAATGGAGTATAACAGGAATCAAACAAAATAG  
 ORF amino acid sequence (SEQ ID NO: 84):  
 FSVTYSQSERTVVFSFGEITFSRSRWTNGFETRI PVDEWLGLEKYKRY SIEFLYHVAKLATMMPYRQVCKVI  
 20 DSTLQTIITKDCVLKAVKFVEKLLKEKERYFYLEPPERKVKKLYVEGDGVMIKSTDSREERRYLDLTHF  
 VIHTGSKKVSTKRYELQDKHEILQLNYDKAKYNLLDYIYNNYEVDDDTILITNSDMGKGYTSRVFKELGKAL  
 KVKKHEHFWDIYHVKEKLSSYLKYPIELTDFALDAVKKYN SDKLELVFDTVESLICDELEDQEFQKFKKKV  
 LNNFKYIKPAHLRNL SNRGIGIMESQHRKITYRMKRRGMYWSK WGI STMANMI I LERANGLRELFFGSRKV  
 YSEYKEGSFSAGRLF KKTDELDFSKPLLNKGRKWSITGIKTK

25

**Table 10: Amino acid and encoding DNA sequences of gbs1477 proteins derived from different strains of *S. agalactiae***

**Strain 0176H4A**  
 30 ORF DNA sequence (SEQ ID NO: 133):  
 ATGAAAAAATCAACAAATTTTTGTGGCGTTCTCAGCGTTGTTACTGATTTTAACGTCATTGCTCTCAGTT  
 GCACCAGCGTTTGC GGAAAAAGAAAAACAAC T GAGACTGTTACTTTGCATAAAATTTTACAACTGATACA  
 AACCTTAAGAATAGTGCTTTCCCTGGTACAAAAGGGCTAGATGGAAC T GAATATGACGGGAAAGCTATTGAT  
 35 AAATTGGATAGCTACTTTGGCAATGACTCAAAAGATATTGGTGGGGCTTACTTTATATTGGCAAATAGCAAG  
 GGTGAATATATCAAAGCTAATGATAAAAATAAATTAAAGCCTGAGTTTAGTGGGAACACTCCGAAAACGACC  
 CTCAATATTAGTGAAGCTGTAGGTGGTTTTGACAGAAGAAAACGCAGGTATTAAGTTTGA AACCACTGGTTTA  
 AGAGGGGATTTCCAGATTATTGAATTGAAAGACAAGTCAACTTACAATAATGGTGGGGCCATCTTGGCTGAT  
 TCAAAGCGGTTCCAGTGAAAATCACTCTTCCATTGATAAACAAGGATGGTGTGTTAAAGATGCACACGTC  
 40 TATCCAAAGAACACTGAAACAAAACCGCAAATTGACAAGA ACTTTGCTGATAAAAATCTTGATTATATTAAC  
 AACCAAAAAGACAAAGGTACTATATCAGCAACTGTTGGTGTATGTTAAAAAATATACTGTTGGGACAAAATC  
 CTTAAAGGATCTGACTATAAAAATTAGTTTTGGACCGATAGCATGACGAAAGGATTGACGTTTAAACAACGAT  
 GTTACTGTAACATTGGATGGTGCAAATTTTGAACAATCAAATTACACCTTAGTAGCTGATGACCAAGGTTTC  
 CGTCTTGTCTTGAATGCAACAGGTCTTTCTAAAGTAGCAGAAGCTGCAAAAACAAAAGATGTTGAAATCAA  
 45 ATCAACTATTCAGCTACAGTAAACGGTTCTACTGTGCTTGA AAAGTCAGAAAATAATGATGTCAAAC TAGAT  
 TATGGTAACAACCAACAAC T GAAAACGAACCACAAC T GGTAATCCAGTTAACAAAGAAATCACAGTTCGA  
 AAGACTTGGGCAGTGGATGGTAATGAAGTGAATAAGGGAGATGAAAAGTTGACGCTGTCTTCACGTTGCAA  
 GTTAAAGATAGTGACAAATGGGTGAATGTGCGATT CAGCAACAGCAACAGCAGCAACTGACTTCAAATACACT  
 TTCAAAAAC T TGGATAATGCCAAAAC T TACCGTGTGTTAGTGAACGTTGTTAGCGGCTACGCTCCAGCCTACGTT  
 50 TCATTTGTGGGTGGAGTTGTGACTATTAAGAATAACAAAAC TCAAATGACCCAAC TCCAATCAATCCATCA  
 GAACCAAAAGTTGTGACTTATGGACGTAAATTTGTGAAAACAAATCAAGATGGCTCTGAACGTTCTAGCAGGA  
 GCTACTTTCTTGTTAAGA ACTCACAAAGTCAACTTGGCACGTAAATCAGGTGTTGCAACTAATGAAGCT  
 CACAAAGCAGTAACAGATGCTAAAGTACAAC TGGATGAAGCTGTTAAAGCTTATAACAAATTGACTAAAGAA  
 CAACAAGAAAGTCAAGATGGTAAAGCAGCATTGAATCTTATTGATGAAAACAAACAGCTTACAATGAAGCT  
 TTTGCTAAAGCAAAC TACTCATATGAATGGGTGTGATGATAAAAACGCTGCAAACGTTGTTAAATTGATTTCT  
 55 AATACAGCTGGTAAATTTGAAATTACAGGTTTGAATGCAGGCGAGTATAGTTTGGAAAGAGACTCAAGCACCA  
 ACAGGTTATGCTAAATTTGTCAAGTGTATCATT TAAAGTAAATGATACATCGTATAGCGAAGGGGCTTCA  
 AATGATATTGCATACGATAAAGACTCCGGTAAAACAGATGCACAAAAGTTGTCAACAAAAGTAACAATC



CCACAAACAGGTGGTATTGGTACAATTCTTTTCACAATTATTGGTTTAAGCATTATGCTTGGAGCGGTAGTT  
ATCATGAAAAGACGTCAATCAGAGGAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 91):

5 MKKINKFFVAFSALLLILTSLLSVAPAFAEKEKTTETVTLHKILQTDNLKNSAFPGTKGLDGTEYDGKAID  
KLDYFNGDSKDIGGAYFILANSKGEYIKANDKNLKPFEFSGNTPKTTLNISEAVGGLTEENAGIKFETTGL  
RGDFQI IELKDKSTYNNGGAILADSKAVPVKITLPLINKDGVVKDAHVYPKNTETKPQIDKNFADKNLDYIN  
10 NQKDKGTISATVGDVKKYTVGTKILKGSYKLVWTD SMTKGLTFNNDVTVTLDGANFEQSNYTLVADDQGF  
RLVLNATGLSKVAEAAKTKDVEIKINYSATVNGSTVVEKSENNDVKLDYGNNPTTENEPQTGNPNVKEITVR  
KTWAVDGNVKNKGDEKVDVFTLQVKDSDKWNVD SATATAATDFKYTFKNLDNAKTYRVVERVSGYAPAYV  
SFVGGVVTIKNNKNSNDPTPINPSEPKVVITYGRKFVKTNDG SERLAGATFLVKNSQSQYLARKSGVATNEA  
15 HKAVTDAKVQLDEAVKAYNKLTKEQQESQDGKALNLI DEKQTAYNEAFKANYSEWVVDKNAANVVKLIS  
NTAGKFEITGLNAGEYSLEETQAPTGYAKLSSDVSFKVNDTSYSEGASNDIAYDKDSGKTDAQKVVNKKVTI  
PQTGGIGTILFTIIGLSIMLGAVVIMKRRQSEEA

### 15 Strain 12351

ORF DNA sequence (SEQ ID NO: 134):

ATGAAAAAATCAACAAATATTTTGCAGTCTTCTCGGCATTGCTACTGACCGTAACATCATTGTTCTCAGTT  
GCACCAGTGTGGCGGAAGAAGCAAAAAC TACTGACACAGTGACCTTGCACAAGATTGTCATGCCCTCGAACT  
GCATTTGACGGTTTTACTGCTGGTACAAAGGTAAGGATAATACTGACTACGTTGGTAAACAAATCGAAGAC  
20 CTTAAACTTACTTTGGCTCAGGCGAAGCGAAAAGAAATCGCAGGTGCTTACTTTGCTTTCAAAAATGAAGCT  
GGTACTAAATACATCACTGAAAATGGTGAAGAAGTTGATACTTTGGATAACAACAGATGCCAAAGGTGGTGCT  
GTTCTTAAAGTTTTAACAACAGACAATGGTTTTCAAATTTAACACTTCTAAATTAACAGGAACTTACCAAATC  
GTTGAATTGAAAGAAAATCTACATAACAACGATGGTCTATCTTGGCTGATTCAAAGCAGTTCCAGTT  
AAAATCACTCTTCCATTGGTAAACGACAATGGTGTGTTAAAGACGCTCACGTTTATCCAAAGAACACTGAA  
25 ACAAACCACAAGTAGATAAGA ACTTCGCAGATAAAGA ACTTGATTATGCGAACAACAAAAAGACAAAGGG  
ACTGTCTCAGCATCTGTTGGTGATGTTAAAAATATCATGTTGGAACAAAAATCCTTAAAGGTTTCAGACTAT  
AAGAAATTAATCTGGACCGATAGCATGACCAAAGGTTTGACTTTCAACAACGATATTGCTGTAACATTGGAT  
GGTGAACCTCTTGATGCTACAAATTACAAACTTGTAGCAGATGACCAAGGTTTCCGCCTTGTCTTGACTGAC  
AAAGGTCTTGAAGCAGTGGCAAAAGCCGCAAAAACAAAAGATGTTGAAATCAAGATCACTTACTCAGCTACT  
30 TTGAACGGTCTGCTGCTGTTGAAGTTCTAGAAACCAATGATGTTAAATTGGACTACGGCAACAACCCAACA  
ATTGAAATGAACCAAAGAAGGTTTCCAGTTGATAAGAAAATCACTGTTAAACAAAACATGGGCAGTAGAT  
GGCAATGAAGTGAATAAAGCAGATGAAACAGTTGATGCTGTCTTACCTTGCAAGTTAAAGATGGTGACAAA  
TGGGTGAATGTTGATTGAGTAAAGCAACAGCTGCAACTAGCTTCAAACACACTTTTGAAAACCTGGATAAT  
GCTAAAACCTACCGCGTTATCGAACGTTAGCGGCTACGCTCCAGAATACGCTCATTGTTAAATGGCGTT  
35 GTAACCATCAAGAACAACAAGACTCAAATGAGCCAAC TCCAATCAACCCATCAGAACCAAAGTGGTGACT  
TATGGACGTAAATTTGTGAAAACAAATAAAGATGGAAAAGAACGCTTGGCAGGAGCTACCTTCCTTGTAAAG  
AAAGATGGCAAGTACTTGGCACGTAAATCAGGTGTTGCAACAGATGCAGAAAAGCTGCTGTAGATTCAACT  
AAATCAGCATTGGATGCTGCTGTTAAAGCTTACAATGATTTGACTAAAGAAAACAAGAAGGTCAAGATGGT  
AAATCAGCATTGGCTACCGTTAGTGAAAACAAAAGCTTACATTGATGCCTTTGTTAAAGCTAACTACTCA  
40 TACGAATGGGTTGAAGATAAAAATGCTAAGAATGTTGTTAAATTGATTTCTAACGATAAAGGTCAATTTGAA  
ATTACTGGCTTGACTGAAGGTCAATACTCATTGGAAGAAACACAAGCACC AACTGGTTATGCTAAATTATCA  
GGTGATGTTTCGTTTAATGTTAATGCTACTTCATACAGTAAAGGTTCTGCTCAAGATATTGAGTATACCCAA  
GGTTCTAAAAC TAAAGATGCACAACAAGTTATCAATAAGAAGGTTACTATTCCACAACAGGTGGTATTGGT  
ACAATTTTTTTT CACAATTATTGGATTAAGTATTATGCTTGGAGCGGTAGTTATCATGAAAAGACGTCAATCA  
45 GAGGAAGTTTAA

ORF amino acid sequence (SEQ ID NO: 92):

50 MKKINKYFAVFSALLLTVTSLFSVAPVF AEEAKTTDTVTLHKIVMPRTAFDGF TAGTKGKDNTDYVGKQIED  
LKTYFGSGEAKEIAGAYFAFKNEAGTKYITENGE EVDLTDTTDAKGGAVLKGLTTDNGFKFN TSKLTGTYQI  
VELKEKSTYNNDSILADSKAVPVKITLPLVNDNGVVKDAHVYPKNTETKPQVDKNFADKELDYANNKKDKG  
TVSASVGDVKKYHVGTKILKGSYKLIWTD SMTKGLTFNNDIAVTLDGATLDATNYKLVADDQGFRLVLTD  
KGLEAVAKAAKTKDVEIKITYSATLNGSAVVEVLETNDVKLDYGNNPTIENEPKEGIPVDKKITVNKTWAVD  
GNEVNKADETVDAVFTLQVKDGDKWNVD SAKATAATSFKHTFENLDNAKTYRVI ERVSGYAPEYVSFVNGV  
VTIKNNKDSNEPTPINPSEPKVVITYGRKFVKTNDGKERLAGATFLVKKDGKYLARKSGVATDAEKA AVDST  
KSAALDAVKAYNDLTKEKQEGQDGKSALATVSEKQKAYIDAFVKANYSYEWVEDKNAKNVVKLISNDKGQFE  
55 ITGLTEGQYSLEETQAPTGYAKLSGDVSFNVNATSYSKGS AQDIEYTQGSKTKDAQQVINKKVTI PQTGGIG  
TIFFTIIGLSIMLGAVVIMKRRQSEEV

### Strain 12401

ORF DNA sequence (SEQ ID NO: 135):



ATGAAAAGAATCAACAAATATTTTGCATGTTCTCGGCATTGTTATTAATTTTAAACATCGTTGTTATCGGTA  
 GCTCCGGTATTTGCTGCTGAGATGGGAAATATCACTAAAACAGTAACCTTACACAAAATTGTTCAAACATCC  
 GATAATTTGGCTAAGCCAAATTTCCCAGGAATAAATGGATTGAATGGAACGAAGTATATGGGTCAAAAACCTT  
 ACTGACATTTTCAGGATATTTTGGGCAAGGTTCTAAAGAAATCGCCGGTGCTTTCTTTGCGGTTATGAATGAA  
 5 AGTCAGACAAAATATATCACAGAAAGTGGTACTGAAGTAGAAAGTATCGATGCAGCAGGTGTCTTAAAGGT  
 TTGACAACTGAAAACGGCATTACATTTAATACTGCAAACCTAAAAGGAACATAACCAAATCGTTGAGTTGCTT  
 GACAAATCTAATTATAAAAATGGTGACAAAGTTCTTGCTGACTCAAAAAGCTGTCCCAGTGAAAATCACTCTT  
 CCTTTGTATAACGAAGAAGGAATTGTCGTGGACGCTGAAGTGTATCCAAAGAATACAGAAGAAGCACCACAA  
 ATCGACAAAACCTTTGCTAAAGCAAATAAATTGTTGAATGACAGTGATAATTCAGCTATTGCAGGTGGGGCA  
 10 GACTACGACAAATATCAGGCAGAAAAGCAAAGCTACTGCTGAAATCGGTCAAGAAATCCCTTACGAAGTT  
 AAAACAAAATCCAAAAGGGTCTAAATACAAAACCTTGCTTGGGTGCGATACCATGTCAAATGGTTTGACA  
 ATGGGTAACACTGTAACTTAGAAGCATCGTCAGGCTCTTTTGTAGAAGGTACAGATTACAATGTTGAACGT  
 GATGACCGTGGTTTCACTTTGAAATTCACAGATACAGGTTTGACTAAGCTACAAAAGAAGCGGAAACACAA  
 GCTGTTGAATTCACATTGACATATAGCGCAACAGTTAACGGTGCAGGCTATTGATGACAAGCCAGAAAGCAAT  
 15 GATATCAAACCTTCAATACGGTAACAAACCAGGTAAAAAGTAAAAGAAATCCCAGTAACACCGTCAAATGGC  
 GAAATCACTGTTAGCAAACCTTGGGACAAAGGTTTCAAGTTTAGAGAATGCGAATGTTGTTTATACCCTTAAA  
 GATGGTGAACAGCTGTTGCCTCAGTTTCATTGACAAAACAACACCAAATGGCGAAATCAACTTAGGTAAT  
 GGTATTAATTTACAGTTACTGGAGCGTTTGCTGGTAAATTCAGTGGTCTGACTGATAGTAAAACATACATG  
 ATCTCAGAACGTATCGCTGGTTATGGTAATACAATCACTACTGGTGCCTGGTAGTGCAGCTATCACCAATACT  
 20 CCAGATTCAGACAACCCAACACCCTTAATCCAACTGAACCAAAGTTGTGACACACGGTAAAAAATTCGTC  
 AAAACAAGTTCGACTGAAACAGAACGCTTGCAAGGTGCACAGTTCGTTGTTAAAGATTCAGCTGGTAAATAC  
 CTTGCATTGAAATCATCTGCGACAATATCAGCTCAAACAACAGCTTACACAAATGCTAAAACCTGCTCTTGAC  
 GCTAAAATCGCAGCTTACAACAACTTTCAGCAGACGATCAAAAAGGTACTAAAGGTGAAACAGCTAAAGCA  
 GAAATCAAACTGCTCAAGACGCTTACAATGCAGCCTTCATCGTAGCTCGTACAGCTTACGAGTGGGTAAC  
 25 AATAAAGAAGATGCTAACGTTGTTAAAGTGACTTCAAACGCTGACGGTCAATTTGAAGTTAGCGGTCTTGCA  
 ACTGGTGATTATAAACTTGAAGAAACACAAGCTCCAGCTGGTTACGCTAAATTAGCAGGTGATGTTGATTTT  
 AAAGTTGGAAACAGCTCAAAGCAGACGACTCAGGTAACATTGATTACACTGCTAGCAGCAATAAAAAAGAC  
 GCTCAACGCATAGAAAACAAAAAGTGACTATTCCACAAACAGGTGGTATTGGTACAATTCTTTTCACAATT  
 30 ATTGTTTTAAGCATTATGCTTGGAGCGGTAATTATCATGAAAAGACGTCAATCAGAGGAAGCTTAA  
 ORF amino acid sequence (SEQ ID NO: 93):  
 MKRINKYFAMFSALLLILTSLLSVAPVFAEMGNITKTVTLHKIVQTSNDLAKPNFPINGLNGTKYMGQKL  
 TDISGYFGQGSKEIAGAFFAVMNESQTKYITESGTEVESIDAAGVLKGLTTENGI TFNTANLKGTYQIVELL  
 DKSNYKNGDKVLADSKAVPVKITLPLYNEEGIVVDAEVYPKNTEEAPQIDKNFAKANKLLNDSNSAIAGGA  
 DYDKYQAEKAKATAEIGQEI PYEVKTKIQKGSKYKNLAWVDTMSNGLTMGNTVNLEASSGSFVEGTDYNVER  
 35 DDRGFTLKFDTGLTKLQKEAETQAVEFTLTYSATVNGAAIDDKPESNDIKLQYGNKPGKKVKEIPVTPSNG  
 EITVSKTWDKGSDELNANVVYTLKDGGTAVASVSLTKTTPNGEINLNGIKFTVTGAFAGKFSGLTDSKTYM  
 ISERIAGYGNTITTTGAGSAAITNTPDSDNPTPLNPTEPKVVTHGKKFVKTSSTETERLQGAQFVVKDSAGKY  
 LALKSSATISAQTTAYTNAKTALDAKIAAYNKLSADDQKGTGETAKAEIKTAQDAYNAAFIVARTAYEWT  
 NKEDANVVKVTSNADGQFEVSGLATGDYKLEETQAPAGYAKLAGDVDFKVGNSKADDSGNI DYTASSNKKD  
 40 AQRIENKQVTIPQTGGIGTILFTIIGLSIMLGAVIIMKRRQSEEA

**Strain 126H4A**

ORF DNA sequence (SEQ ID NO: 136):

ATGAAAAAATCAACAAATATTTTGCAGTCTTCTCGGCCTTGCTACTGACCGTAACATCATTGCTCTCAGTT  
 45 GCACCAGCGTTTGCAGGACGAAGCAACAATAACAGTGACTTTGCACAAGATCTTGCAAACCTGAATCAAAT  
 CTTAATAAAAGTAACTTCCCAGGAACCTACAGGCCTAACGGAGATGACTATAAAGGTGAATCTATTTCTGAC  
 CTTGCTGAATACTTTGGATCAGGTTCTAAAGAAATGACGGTGCTTTCTTTGCTTTGGCTTTAGAAGAGGAA  
 AAAGATGGTGTGCTACAATATGTTAAGGCAAAGCAAATGACAAATTAACACCAGACTTAATTACTAAAGGT  
 ACACCTGCAACAACAACAAAAGTTGAAGAAGCTGTAGGTGGTTTGACAACCTGGTACGGGTATTGTTTTCAAT  
 50 ACAGCTGGTTTGAAAGGTAATTTCAAATTTATTGAATTGAAAGACAAATCAACTTACAACAATAATGGTTCC  
 CTCTTAGCAGCTTCAAAGCAGTTCGCGTGAAAATCACTCTTCCATTGGTAAGCAAAGATGGTGTGTTAAA  
 GATGCACACGTTTATCCAAAGAACACTGAAACAAAACCAGAAGTAGACAAGAATTCGCTAAAACAAACGAT  
 TTGACAGCTCTCAAAGACGCTACTCTTCTTAAGGCTGGTGCAGACTACAAAACCTATTCAGCGACTAAAGCT  
 ACTGTAACAGCTGAAATCGGTAAAGTTATCCCTTACGAAGTTAAAACAAAAGTTCTTAAAGGTTCTAAATAC  
 55 GAAAACCTGGTTTGGACCGATACCATGTCAAATGGTTTGACAATGGGTGATGATGTTAACCTTGCAGTTTCA  
 GGGACTACAACAACCTTTCATTAAAGATATAGATTACTCTTAGCATTGATGACCGTGGTTTACATTGAAA  
 TTCAAAGCTACTGGATTGGACAAATTTGGAAGAAGCAGCTAAAGCATCTGATGTTGAATTTACATTGACTTAT  
 AAAGCTACTGTTAATGGCCAAGCAATTATTGACAACCCAGAAGTCAATGACATCAAATTTGGACTATGGTAAT  
 AAACCTGGTACAGATTTATCAGAACAACCTGTGACACCTGAAGATGGTGAAGTTAAAGTCACTAAAACATGG  
 60 GCAGCAGGTGCTAATAAAGCAGACGCTAAAGTTGTCTACACACTTAAAATGCTACTAAACAAGTCGTAGCT



- 80 -

TCTGTTCGCATTGACCGCAGCTGATACAAAAGGTACGATTAATCTTGGTAAAGGCATGACCTTTGAAATCACA  
 GGAGCTTTCTCAGGTACATTCAAAGGCCTTCAAATAAAGCTTACACTGTTTCTGAACGTGTTGCAGGTTAT  
 ACTAATGCTATTAATGTTACTGGTAATGCTGTTGCTATCACCAATACACCAGACAGTGACAATCCAACGCCA  
 CTTAACCCAACTCAACCAAAAAGTTGAAACACATGGTAAGAAATTTGTCAAAGTTGGCGATGCAGATGCCCGC  
 5 TTAGCTGGTGCACAATTCGTTGTGAAAAATTCAGCTGGTAAATTCCTTGTCTTAAAGAAGATGCAGCTGTA  
 TCAGGAGCTCAAACCTGAATTGGCAACTGCTAAAACAGACTTGGATAATGCCATCAAAGCTTACAACGGTTTG  
 ACAAAGCGCAGCAAGAAGGTGCTGATGGTACATCAGCAAAAAGAACTTATCAACACTAAACAGTCAGCTTAC  
 GACGCAGCCTTCATCAAAGCACGTACAGCTTATATATGGGTAGATGAAAAAACTAAAGCTATTACCTTCACT  
 TCAAATAATCAAGGTCAATTTGAAGTTACTGGTCTTGAAGTAGGTTCTTACAAACTTGAAGAACTCTTGCA  
 10 CCAGCAGGTTATGCTAAATTGTCAGGCGACATTGAGTTTACAGTTGGACACGATTCTTACACAAGTGGTGAC  
 ATCAAGTACAAGACAGATGATGCTAGCAACAATGCACAAAAAGTTTTCAATAAAAAAGTAACCATCCCACAA  
 ACAGGTGGTATTGGTACAATTCTTTTCACAATTATTGGTTTAAGCATTATGCTTGGAGCGGTAGTTATCATG  
 AAAAGACGTCAATCAGAGGAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 94):

15 MKKINKYFAVFSALLLTVTSLLSVAPAFADAEATTNTVTLHKILQTESNLNKS NFPGTTGLNGDDYKGESISD  
 LAEYFGSGSKEIDGAFFALALEEEKDGVVQYVKAKANDKLTPLDITKGT PATTTKVEEAVGGLTTGTGIVFN  
 TAGLKGNFKIIELKDKSTYNNNGSLLAASKAVPVKITLPLVSKDGVVKDAHVYPKNTETKPEVDKNFAKTND  
 LTALKDATLLKAGADYKNYSATKATVTAEIGKVI PYEVKTKVLKGSKYEKLWTDTMSNGLTMGDDVNLAVS  
 20 GTTTTFIKDIDYTLSIDDRGFTLKF KATGLDKLEEAAKASDVEFTLTYKATVNGQAIIDNPEVNDIKLDYGN  
 KPGTDLSEQPVTPEEDGEVKVTKTWAAGANKADAKVVYTLKNATKQVVASVALTAADTKGTINLGKGMTFEIT  
 GAFSGTFKGLQNKAYTVSERVAGYTNAINVTGNAVAITNTPDSDNPTPLNPTQPKVETHGKKFVKVDADAR  
 LAGAQFVVKNSAGKFLALKEDA AVSQAQTE LATAKTDLDNAIKAYNGLTKAQQEGADGTSAKELINTKQSAY  
 DAAFIKARTAYI WVEKTKAITFTSNNQGFVETGLEVGSYKLEETLAPAGYAKLSGDIEFTVGHDSYTSGD  
 IKYKTDDASNNAQKVFNKKVTIPQTGGIGTILFTIIGLSIMLGAVVIMKRRQSEEA

### Strain 49447

ORF DNA sequence (SEQ ID NO: 137):

ATGAAAAAATCAACAAATTTTTTGTGGCGTTCTCAGCGTTGTTACTGATTTTAACGTCATTGCTCTCAGTT  
 GCACCAGCGTTTGC GGAAAAAGAAAAACAACCTGAGACTGTTACTTTGCATAAAATTTTACAACTGATACA  
 30 AACCTTAAGAATAGTGCTTTCCCTGGTACAAAAGGGCTAGATGGAACCTGAATATGACGGGAAAGCTATTGAT  
 AAATTGGATAGCTACTTTGGCAATGACTCAAAGATATTGGTGGGGCTTACTTTATATTGGCAAATAGCAAG  
 GGTGAATATATCAAAGCTAATGATAAAAAATAAATTAAAGCCTGAGTTTAGTGGGAACACTCCGAAAACGACC  
 CTCAATATTAGTGAAGCTGTAGGTGGTTTGACAGAAGAAAACGCAGGTATTAAGTTTGAAACCACTGGTTTA  
 AGAGGGGATTTCCAGATTATTGAATTGAAAGACAAGTCAACTTACAATAATGGTGGGGCCATCTTGGCTGAT  
 35 TCAAAGCGGTTCCAGTGAAAATCACTCTTCCATTGATAAACAAGGATGGTGTGTTAAAGATGCACACGTC  
 TATCCAAAGAACACTGAAACAAAACCGCAAATTGACAAGAACTTTGCTGATAAAAAATCTTGATTATATTAAC  
 AACCAAAAAGACAAAGGTACTATATCAGCAACTGTTGGTGTATGTTAAAAAATATACTGTTGGGACAAAAATC  
 CTTAAAGGATCTGACTATAAAAAATTAGTTTGGACCGATAGCATGACGAAAGGATTGACGTTTAAACAACGAT  
 GTTACTGTAACATTGGATGGTGCAAATTTTGAACAATCAAATTACACCTTAGTAGCTGATGACCAAGGTTTC  
 40 CGTCTTGTCTTGAATGCAACAGGTCTTTCTAAAGTAGCAGAAGCTGCAAAAACAAAAGATGTTGAAATCAA  
 ATCAACTATTCAGCTACAGTAAACGGTTCTACTGTCGTTGAAAAGTCAGAAAATAATGATGTCAAACCTAGAT  
 TATGGTAACAACCCAACAACCTGAAAACGAACCACAAACTGGTAATCCAGTTAACAAAGAAATCACAGTTCGA  
 AAGACTTGGGCAGTGGATGGTAATGAAGTGAATAAGGGAGATGAAAAGTTGACGCTGTCTTCACGTTGCAA  
 GTTAAAGATAGTGACAAATGGGTGAATGTTCGATTGACGCAACAGCAACAGCAGCAACTGACTTCAAATACACT  
 45 TTCAAAAACCTGGATAATGCCAAAACCTACCGTGTGTTAGTAAACGTTAGCGGCTACGCTCCAGCCTACGTT  
 TCATTTGTGGGTGGAGTTGTGACTATTAAGAATAACAAAACCTCAAATGACCCAACCTCAAATCAATCCATCA  
 GAACCAAAAAGTTGTGACTTATGGACGTAAATTTGTGAAAACAAATCAAGATGGCTCTGAACGTCTAGCAGGA  
 GCTACTTTCTTGTTAAGAACTCACAAAGTCAACTTGGCACGTAAATCAGGTGTTGCAACTAATGAAGCT  
 CACAAAGCAGTAACAGATGCTAAAGTACAACCTGGATGAAGCTGTTAAAGCTTATAACAAATTGACTAAAGAA  
 50 CAACAAGAAAGTCAAGATGGTAAAGCAGCATTGAATCTTATTGATGAAAACAAACAGCTTACAATGAAGCT  
 TTTGCTAAAGCAAACCTACTCATATGAATGGGTGTAGATAAAAACGCTGCAAACGTTGTTAAATTGATTTCT  
 AATACAGCTGGTAAATTTGAAATTACAGGTTTGAATGCAGGCGAGTATAGTTTGGAAAGAGACTCAAGCACCA  
 ACAGGTTATGCTAAATTGTCAAGTGTATCATTTAAAGTAAATGATACATCGTATAGCGAAGGGGCTTCA  
 AATGATATTGCATACGATAAAGACTCCGGTAAAACAGATGCACAAAAAGTTGTCAACAAAAAAGTAACAATC  
 55 CCACAAACAGGTGGTATTGGTACAATTCTTTTCACAATTATTGGTTTAAGCATTATGCTTGGAGCGGTAGTT  
 ATCATGAAAAGACGTCAATCAGAGGAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 95):

60 MKKINKFFVAFSALLLILTSLLSVAPAFAEKEKTTETVTLHKILQTDNLNKS AAFPGTKGLDGTEYDGAID  
 KLDSYFGNDSKDIGGAYFILANSKGEYIKANDKNKLKPEFSGNTPKTTLNISEAVGGLTEENAGIKFETTGL  
 RGDFQIIELKDKSTYNNGGAILADSKAVPVKITLPLINKDGVVKDAHVYPKNTETKPKQIDKNFADKNLDYIN



NQKDKGTISATVGDVKKYTVGTKILKGSYKLVWTD SMTKGLTFNNDVTVTL DGANFEQSNYTLVADDQGF  
 RLVLNATGLSKVAEAAKTKDVEIKINYSATVNGSTVVEKSENNDVKLDYGNPPTTENEPQTGNPVNKEITVR  
 KTWAVDGVNEVNGDEKVDVFTLQVKDSKWNVD SATATAATDFKYTFKNLDNAKTYRVVERVSGYAPAYV  
 SFVGGVVTIKNNKNSNDPTPINPSEPKVVITYGRKFVKTNDG SERLAGATFLVKNSQSQYLARKSGVATNEA  
 5 HKAVTDAKVQLDEAVKAYNKLTKEQQESQDGKAA LNLI DEKQTAYNEAFKANYSYEWVVDKNAANVVKLIS  
 NTAGKFEITGLNAGEYSLEETQAPTGYAKLSSDVSFKVNDTSYSEGASNDIAYDKDSGKTDAQKVVNKKVTI  
 PQTGGIGTILFTIIGLSIMLGAVVIMKRRQSEEA

**Strain 5095S2**

10 ORF DNA sequence (SEQ ID NO: 138):  
 ATGAAAAGAATCAACAAATATTTTGC AATGTTCTCGGCATTGTTATTAATTTTAAACATCGTTGTTATCGGTA  
 GCTCCGGTATTTGCTGCTGAGATGGGAAATATCACTAAAACAGTAACCTTACACAAAATTTGTTCAAACATCC  
 GATAATTTGGCTAAGCCAAATTTCCCAGGAATAAATGGATTGAATGGAACGAAGTATATGGGTCAAAAACCTT  
 ACTGACATTTTCAGGATATTTTGGGCAAGGTTCTAAAGAAATCGCCGGTGCCTTTCTTTGCGGTTATGAATGAA  
 15 AGTCAGACAAAATATATCACAGAAAGTGGTACTGAAGTAGAAAGTATCGATGCAGCAGGTGTCCTTAAAGGT  
 TTGACAAC TGA AACGGCATTACATTTAATACTGCAAAC TAAAAGGAACATACCAAATCGTTGAGTTGCTT  
 GACAAATCTAATTATAAAAATGGTGACAAAGTTCTTGCTGACTCAAAGCTGTCCCAGTGAAAATCACTCTT  
 CCTTTGTATAACGAAGAAGGAATTATCGTGGACGCTGAAGTGTATCCAAAGAATACAGAAGAAGCACCACAA  
 ATCGACAAAAC TTTGCTAAAGCAAATAAATTGTTGAATGACAGTGATAATTCAGCTATTGCAGGTGGGGCA  
 20 GACTACGACAAATATCAGGCAGAAAAAGCAAAGCTACTGCTGAAATCGGTCAAGAAATCCCTTACGAAGTT  
 AAAACAAAATCCAAAAGGGTCTAAATACAAAACCTTGCTTGGGTGATACCATGTCAAATGGTTTGACA  
 ATGGGTAACACTGTTAACTTAGAAGCATCGTCAGGCTCTTTTGTAGAAGGTACAGATTACAATGTTGAACGT  
 GATGACCGTGGTTTCACTTTGAAATTCACAGATACAGGTTTGACTAAGCTACAAAAGAAGCGGAAACACAC  
 GCTGTTGAATTCACATTGACATATAGCGCAACAGTTAACGGTGC GGCTATTGATGACAAGCCAGAAAGCAAT  
 25 GATATCAAAC TTTCAATACGGTAACAAACCAGGTAAAAAAGTAAAAGAAATCCCAGTAACACCGTCAAATGGC  
 GAAATCACTGTTAGCAAAC TTTGGGACAAAGGTTTCAGATTTAGAGAATGCGAATGTTGTTTATACCCTTAAA  
 GATGGTGGAACAGCTGTTGCCTCAGTTTCATTGACAAAACAACACCAAATGGCGAAATCAACTTAGGTAAT  
 GGTATTAAATTTACAGTTACTGGAGCGTTTGTGTTAAATTCAGTGGTCTGACTGATAGTAAAACATACATG  
 ATCTCAGAACGTATCGCTGGTTATGGTAATACAATCACTACTGGTGTGTTAGTGCAGCTATCACCAATACT  
 30 CCAGATTCAGACAACCCAACACCCTTAATCCAAC TGAACCAAAGTTGTGACACACGGTAAAAAATTCGTC  
 AAAACAAGTTTCGACTGAAACAGAACGCTTGCAAGGTGCACAGTTCTGTTGTTAAAGATTCAGCTGGTAAATAC  
 CTTGCATTGAAATCATCTGCGACAATATCAGCTCAAACAACAGCTTACACAAATGCTAAAAC TGTCTTTGAC  
 GCTAAAATCGCAGCTTACAACAAACTTTTCAGCAGACGATCAAAAAGGTACTAAAGGTGAAACAGCTAAAGCA  
 GAAATCAAAC TGTCAAGACGCTTACAATGCAGCCTTCATCGTAGCTCGTACAGCTTACGAGTGGGTAAC T  
 35 AATAAAGAAGATGCTAACGTTGTTAAAGTGACTTCAAACGCTGACGGTCAATTTGAAGTTAGCGGTCTTGCA  
 ACTGGTGATTATAAACTTGAAGAAACACAAGCTCCAGCTGGTTACGCTAAATTAGCAGGTGATGTTGATTTT  
 AAAGTTGGAAACAGCTCAAAGCAGACGACTCAGGTAACATTTGATTACACTGCTAGCAGCAATAAAAAAGAC  
 GCTCAACGCATAGAAAACAAAAAAGTGACTATTTCCACAACAGGTGGTATTGGTACAATTTCTTTTCACAATT  
 ATTTGGTTTAAAGCATTATGCTTGGAGCGGTAATTATCATGAAAAGACGTCAATCAGAGGAAGCTTAA  
 40 ORF amino acid sequence (SEQ ID NO: 96):  
 MKRINKYFAMFSALLLILTSLLSVAPVFAAEMGNITKTVTLHKIVQTS DN LAKPNFPINGLNGTKYMGQKL  
 TDISGYFGQSKEIAGAFFAVMNESQTKYITESGTEVESIDAAGVLKGLTTENGITFNTANLKGTYQIVELL  
 DKSNYKNGDKVLADSKAVPVKITLPLYNEEGIIVDAEVYPKNTEEAPQIDKNFAKANKLLNDSDNSAIAGGA  
 45 DYDKYQAEKAKATAEIGQEIPIYEVKTKIQKGSKYKNLAWVD TMSNGLTMGNTVNLEASSGSFVEGTDYNVER  
 DDRGFTLKFDTGLTKLQKEAETHAVEFTLTYSATVNGAAIDDKPESNDIKLQYGNKPGKVKVEIPVTPSNG  
 EITVSKTWDKGS DLENANVVYTLKDG GTAVASVSLTKTTPNGEINLNGNIKFTVTGAFAGKFSGLTDSKTYM  
 ISERIAGYGNITTTGAGSAAITNTPDSDNPTPLNPTEPKVVTHGKKFVKTSSTETERLQGAQFVVKDSAGKY  
 LALKSSATISAQTTAYTNAKTALDAKIAAYNKLSADDQKGTKGETAKAEIKTAQDAYNAAFIVARTAYEWVT  
 NKEDANVVKVT SNADGQFEV SGLATGDYKLEETQAPAGYAKLAGDVDFKVGNSKADDSGNI DYTASSNKKD  
 50 AQRIENKKVTIPQTGGIGTILFTIIGLSIMLGAVIIMKRRQSEEA

**Strain 6313**

55 ORF DNA sequence (SEQ ID NO: 139):  
 ATGAAAAAATCAACAAATGTCTTACAGTGTCTCGACACTGCTATTGATCTTAAACGTC ACTATTCTCAGTT  
 GCACCAGCGTTTTCGGACGACGTAACAACTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATTAAGGGTGCCTTTCTTTGTTTTCAAAAATGAAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGCCTGTT  
 CTTTCAGGGTTAACAAAAGACACTGGTTTTGCTTTTAACTGCTAAGTTAAAAGGAACTTACCAAATCGTT  
 60 GAATTGAAAGAAAATCAAAC TACGATAACAACGGTTCATCTTGGCTGATTCAAAGCAGTTCAGTTAAA



ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGAAGCTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATTCTTAAAGGCTCAGACTATAAG  
 5 AAAGATTTTCCCTGTTTTAACTACAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCTGCTGCAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAGTTCCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAAGTCATTAAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGTTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAAACGGATGGTACATGG  
 10 GTGAACGTTGCTTACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTCACAGGTTTGGATAATACT  
 AAACTTACCGCGTTGTGCAACGTGTTAGCGGCTACACTCCAGAATATGTATCATTTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAAACCTCAAATGATCCAACCTCCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTTGTAAAGAAA  
 GAAGGAAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 15 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 ACTGGTTTGGATAAAGGCACTTATAGCTTGGAAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGA  
 20 TCTGTAAAAAAGATGCCCAACAAGTTCAAACAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATCTTTTTCACAATTATTGGTTTAAAGCATTATGCTTGGAGCAGTAGTTGTCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 97):

MKKINKCLTVFSTLLILLTSLFSVAPAFADDVTTDTVTLHKIVMPQAAFDFNFTEGTKGKNDSDYVGKQINDL  
 25 KSYFGSTDAKEIKGAFFVFKNETGTFKFIENGEKVDLEAKDAEGGAVLSGLTKDTGFNFNTAKLKGTYQIV  
 ELKEKSNYDNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTETKPVQVDKNFADKDLDDYTDNRKDKGV  
 VSATVGDKEYIVGTKILKGSYKLVWTDMSMTKGLTFNNNVKVTLDGKDFPVLNYKLVTDQGFRLALNAT  
 GLAAVAAAADKDKDVEIKITYSATVNGSTTVEVPETNDVKLDYGNPTEESEPEQEGTPANQEIKVIKDWAVDG  
 TITDENVAVKAIFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNTKTYRVVERVSGYTPYVVSFKNGVV  
 30 TIKNNKNSNDPTPINPSEPKVVVYGRKFVKTNQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYSLLEETQAPAGYATLSGDVNFVETATSYSKGATTDIAYDKGSVKKDAQQVQNKKVTIPQTGGIGT  
 ILFTIIGLSIMLGAVVVMKKRQSEEA

### 35 Strain BAA23

ORF DNA sequence (SEQ ID NO: 140):

ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCACTATTCTCAGTT  
 GCACCAGCGTTTGCAGGACGACGCAACAACTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 40 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCCTTCTTTGTTTTCAAAAATGAAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTGTT  
 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 GAATTGAAAGAAAAATCAAACCTACGATAACAACGGTTCATCTTGGCTGATTCAAAGCAGTTCCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 45 AAACCACAAGTAGATAAGAAGCTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATTCTTAAAGGCTCAGACTATAAG  
 AAAGTGGTTTGGACTGATAGCATGACTAAAGGTTTGCAGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 GAAGATTTTCCCTGTTTTAACTACAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 50 AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAAGTCATTAAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAAACGGATGGTACATGG  
 GTGAACGTTGCTTACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTCACAGGTTTGGATAATGCT  
 AAACTTACCGCGTTGTGCAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 55 ACTATCAAGAACAACAAAACCTCAAATGATCCAACCTCCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAAGAAA  
 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 60 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT



ACTGGTTTGGATAAAGGCACTTATGGCTTGGGAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAAAACAAAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATTCTTTTTCACAATTATTGGTTTAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 5 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 98):

MKKINKCLTMFSTLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGTEGKGNDSYVVGKQINDL  
 KSYFGSTDAKEIKGAFFVFKNETGTFKITENGKEVDLEAKDAEGGAVLSGLTKDNGFVFNTAKLKGIIYQIV  
 10 ELKEKSNDNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTETKPQVDKNFADKDLDDYTDNRKDKGV  
 VSATVGDKKEYIVGTKILKGSYKLVWTDSTMKGLTFNNNVKVTLDGEDFPVLNYKLVTDQGFRLALNAT  
 GLAAVAAAADKDKDVEIKITYSATVNGSTTVEIPETNDVKLDYGNPTEESEPEQEGTPANQEIKVIKDWAVDG  
 TITDANVAVKAIFFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPHYVSFKNGVV  
 TIKNNKNSNDPTPINPSEPKVVVYGRKFVKTNQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
 15 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYGLEETQAPAGYATLSGDVNFVETATSYSKGATTDIAYDKGSVKKDAQQVQNKKVTIPQTGGIGT  
 ILFTIIGLSIMLGAVVIMKKRQSEEA

### Strain BAA611

ORF DNA sequence (SEQ ID NO: 141):

ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCACTATTCTCAGTT  
 GCACCAGCGTTTGCAGGACGACGCAACAACTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCTTTCTTTGTTTTCAAAAATGAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTCTGT  
 25 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 GAATTGAAAGAAAAATCAAACTACGATAACAACGGTTCCTATCTTGGCTGATTCAAAGCAGTTCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGAAGTTCGAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATCTTAAAGGCTCAGACTATAAG  
 30 AAAGTGGTTTGGACTGATAGCATGACTAAAGGTTTGGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 GAAGATTTTCTGTTTTAACTACAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAGTTCATTAAAGACTGGGCAGTAGATGGT  
 35 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAACGGATGGTACATGG  
 GTGAACGTTGCTTACACGAAGCAACAACCAATCACGCTTTGAACATACTTTTACAGGTTTGGATAATGCT  
 AAACTTACCGCGTTGTCGAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAAACTCAAATGATCCAACCTCCAATCAACCCATCAGAACCAAAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
 40 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 ACTGGTTTGGATAAAGGCACTTATGGCTTGGGAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 45 GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAAAACAAAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATTCTTTTTCACAATTATTGGTTTAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 99):

MKKINKCLTMFSTLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGTEGKGNDSYVVGKQINDL  
 KSYFGSTDAKEIKGAFFVFKNETGTFKITENGKEVDLEAKDAEGGAVLSGLTKDNGFVFNTAKLKGIIYQIV  
 ELKEKSNDNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTETKPQVDKNFADKDLDDYTDNRKDKGV  
 VSATVGDKKEYIVGTKILKGSYKLVWTDSTMKGLTFNNNVKVTLDGEDFPVLNYKLVTDQGFRLALNAT  
 GLAAVAAAADKDKDVEIKITYSATVNGSTTVEIPETNDVKLDYGNPTEESEPEQEGTPANQEIKVIKDWAVDG  
 55 TITDANVAVKAIFFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPHYVSFKNGVV  
 TIKNNKNSNDPTPINPSEPKVVVYGRKFVKTNQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYGLEETQAPAGYATLSGDVNFVETATSYSKGATTDIAYDKGSVKKDAQQVQNKKVTIPQTGGIGT  
 ILFTIIGLSIMLGAVVIMKKRQSEEA



**Strain C388/90**

ORF DNA sequence (SEQ ID NO: 142):

ATGAAAAAATCAACAAATATTTTGCAGTCTTCTCGGCATTGCTACTGACCGTAACATCATTGTTCTCAGTT  
 GCACCAGTGTGGCGGAAGAAGCAAAACTACTGACACAGTGACCTTGCACAAGATTGTCATGCCTCGAACT  
 5 GCATTTGACGGTTTTACTGCTGGTACAAAGGGTAAGGATAATACTGACTACGTTGGTAAACAAATCGAAGAC  
 CTTAAACTTACTTTGGCTCAGGCGAAGCGAAAGAAATCGCAGGTGCTTACTTTGCTTTCAAAAATGAAGCT  
 GGTACTAAATACATCACTGAAAATGGTGAAGAAGTTGATACTTTGGATAACAACAGATGCCAAAGGTGGTGCT  
 GTTCTTAAAGGTTTAAACAACAGACAATGGTTTCAAATTTAACACTTCTAAATTAACAGGAACCTACCAAATC  
 10 GTTGAATTGAAAGAAAAATCTACATAACAACGATGGTTCTATCTTGGCTGATTCAAAGCAGTTCCAGTT  
 AAAATCACTCTTCCATTGGTAAACGACAATGGTGTGTTAAAGACGCTCACGTTTATCCAAAGAACACTGAA  
 ACAAACCACAAGTAGATAAGAACTTCGCAGATAAAGAACTTGATTATGCGAACAACAAAAAGACAAAGGG  
 ACTGTCTCAGCATCTGTTGGTGATGTTAAAAAATATCATGTTGGAACAAAAATCCTTAAAGGTTCCAGACTAT  
 AAGAAATTAATCTGGACCGATAGCATGACCAAAGGTTTGACTTTCAACAACGATATTGCTGTAACATTGGAT  
 15 GGTGCAACTCTTGATGCTACAAATTACAACTTGTAGCAGATGACCAAGGTTTCCGCCTTGTCTTGACTGAC  
 AAAGGTCTTGAAGCAGTGGCAAAGCCGCAAAAACAAAAGATGTTGAAATCAAGATCACTTACTCAGCTACT  
 TTGAACGGTTCTGCTGTCGTTGAAGTTCTAGAAACCAATGATGTTAAATTGGACTACGGCAACAACCCAACA  
 ATTGAAAATGAACCAAAGAAGGTATTCCAGTTGATAAGAAAATCACTGTTAACAAAACATGGGCAGTAGAT  
 GGCAATGAAGTGAATAAAGCAGATGAAACAGTTGATGCTGTCTTACCTTGCAAGTTAAAGATGGTGACAAA  
 20 TGGGTGAATGTTGATTCAGCTAAAGCAACAGCTGCAACTAGCTTCAAACACACTTTTGAAAACCTGGATAAT  
 GCTAAAACCTACCGCGTTATCGAACGTGTTAGCGGCTACGCTCCAGAATACGTCTCATTGTAAATGGCGTT  
 GTAACCATCAAGAACAACAAGACTCAAATGAGCCAACTCCAATCAACCCATCAGAACCAAAGTGGTGACT  
 TATGGACGTAAATTTGTGAAAACAAATAAAGATGGAAAAGAACGCTTGGCAGGAGCTACCTTCTTGTAAAG  
 AAAGATGGCAAGTACTTGGCACGTAAATCAGGTGTTGCAACAGATGCAGAAAAAGCTGCTGTAGATTCAACT  
 AAATCAGCATTGGATGCTGCTGTTAAAGCTTACAATGATTTGACTAAAGAAAAACAAGAAGGTCAAGATGGT  
 25 AAATCAGCATTGGCTACCGTTAGTGAAAACAAAAGCTTACAATGATGCCTTTGTTAAAGCTAACTACTCA  
 TACGAATGGGTTGAAGATAAAAATGCTAAGAATGTTGTTAAATTGATTTCTAACGATAAAGGTCAATTTGAA  
 ATTACTGGCTTGACTGAAGGTCAATACTCATTGGAAGAAACACAAGCACCAACTGGTTATGCTAAATTATCA  
 GGTGATGTTTCGTTTAAATGTTAATGCTACTTCATACAGTAAAGGTTCTGCTCAAGATATTGAGTATACCCAA  
 GGTTCTAAAACATAAAGATGCACAACAAGTTATCAATAAGAAGGTTACTATTCCACAAACAGGTGGTATTGGT  
 30 ACAATTTTTTTTCACAATTATTGGATTAAGTATTATGCTTGGAGCGGTAGTTATCATGAAAAGACGTCAATCA  
 GAGGAAGTTTAA

ORF amino acid sequence (SEQ ID NO: 100):

MKKINKYFAVFSALLLTVTSLFSVAPVFEEAKTTDTVTLLHKIVMPRTAFDGF TAGTKGKDNTDYVGKQIED  
 LKTYFGSGEAKEIAGAYFAFKNEAGTKYITENGEVDLDTTDAKGGAVLKGLTTDNGFKFNTSKLTGTYQI  
 35 VELKEKSTYNNDSILADSKAVPVKITLPLVNDNGVVKDAHVYPKNTETKPQVDKNFADKELDYANNKKDKG  
 TVSASVGDVKKYHVGTKILKGSYKLIWTD SMTKGLTFNNDIAVTLDGATLDATNYKLVADDQGFRLVLD  
 KGLEAVAKAAKTKDVEIKITYSATLNGSAVVEVLETNDVKLDYGNPTIENEPKEGIPVDKKITVNKTWAVD  
 GNEVNKADETVDAVFTLQVKDGDKWNVDSAKATAATSFKHTFENLDNAKTYRVIERSGYAPEYVSFVNGV  
 VTIKNNKDSNEPTPINPSEPKVVITYGRKFVKTNKDGKERLAGATFLVKKDGKYLARKSGVATDAEKA AVDST  
 40 KSALDAVKAYNDLTKEKQEGQDGKSALATVSEKQKAYNDAFVKANYSYEWVEDKNAKNVVKLISNDKGQFE  
 ITGLTEGQYSLEETQAPTGYAKLSGDVSNVFNATSYSKGSAQDIEYTGQSKTKDAQQVINKKVTIPQTGGIG  
 TIFFTIIGLSIMLGAVVIMKRRQSEEV

**Strain IC97**

ORF DNA sequence (SEQ ID NO: 143):

ATGAAAAGAATCAACAAATATTTTGCATGTTCTCGGCATTGTTATTAATTTTAAACATCGTTGTTATCGGTA  
 GCTCCGGTATTTGCTGCTGAGATGGGAAATATCACTAAAACAGTAACCTTACACAAAATTTGTTCAAACATCC  
 GATAATTTGGCTAAGCCAAATTTCCAGGAATAAATGGATTGAATGGAACGAAGTATATGGGTCAAAAACCTT  
 ACTGACATTTTCAGGATATTTTGGGCAAGGTTCTAAAGAAATCGCCGGTGCCTTTCTTTGCGGTTATGAATGAA  
 50 AGTCAGACAAAATATATCACAGAAAGTGGTACTGAAGTAGAAAGTATCGATGCAGCAGGTGTCCTTAAAGGT  
 TTGACAACTGAAAACGGCATTACATTTAATACTGCAAACTTAAAAGGAACATACCAAATCGTTGAGTTGCTT  
 GACAAATCTAATTATAAAAATGGTGACAAAGTTCTTGCTGACTCAAAGCTGTCCAGTGAAAATCACTCTT  
 CCTTTGTATAACGAAGAAGGAATTGTCGTGGACGCTGAAGTGTATCCAAAGAATACAGAAGAAGCACCACAA  
 ATCGACAAAACCTTTGCTAAAGCAAATAAATTGTTGAATGACAGTGATAATTCAGCTATTGCAGGTGGGGCA  
 55 GACTACGACAAATATCAGGCAGAAAAGCAAAAAGCTACTGCTGAAATCGGTCAAGAAATCCCTTACGAAGTT  
 AAAACAAAATCCAAAAGGGTCTAAATACAAAACCTTGCTTGGGTGATACCATGTCAAATGGTTTGACA  
 ATGGGTAACACTGTTAACTTAGAAGCATCGTCAGGCTCTTTTGTAGAAGGTACAGATTACAATGTTGAACGT  
 GATGACCGTGGTTTCACTTTGAAATTCACAGATACAGGTTTGACTAAGCTACAAAAGAAGCGGAAACACAA  
 GCTGTTGAATTACATTGACATATAGCGCAACAGTTAACGGTGCGGCTATTGATGACAAGCCAGAAAGCAAT  
 60 GATATCAAACCTTCAATACGGTAACAAACCAGGTAAAAAGTAAAAGAAATCCAGTAACACCGTCAAATGGC



GAAATCACTGTTAGCAAACTTGGGACAAAGGTTTCAGATTTAGAGAATGCGAATGTTGTTTATACCCTTAAA  
 GATGGTGGAACAGCTGTTGCCTCAGTTTCATTGACAAAAACAACACCAAATGGCGAAATCAACTTAGGTAAT  
 GGTATTAATTTACAGTTACTGGAGCGTTTGCTGGTAAATTCAGTGGTCTGACTGATAGTAAAACATACATG  
 ATCTCAGAACGTATCGCTGGTTATGGTAATACAATCACTACTGGTGTCTGGTAGTGCAGCTATCACCAATACT  
 5 CCAGATTCAGACAACCCAACACCCTTAATCCAACCTGAACCAAAAGTTGTGACACACGGTAAAAAATTCGTC  
 AAAACAAGTTCGACTGAAACAGAACGCTTGCAAGGTGCACAGTTCGTTGTTAAAGATTCAGCTGGTAAATAC  
 CTTGCATTGAAATCATCTGCGACAATATCAGCTCAAACAACAGCTTACACAAATGCTAAAACCTGCTCTTGAC  
 GCTAAAATCGCAGCTTACAACAAACTTTCAGCAGACGATCAAAAAGGTACTAAAGGTGAAACAGCTAAAGCA  
 GAAATCAAACTGCTCAAGACGCTTACAATGCAGCCTTCATCGTAGCTCGTACAGCTTACGAGTGGGTAAC  
 10 AATAAAGAAGATGCTAACGTTGTTAAAGTGACTTCAAACGCTGACGGTCAATTTGAAGTTAGCGGTCTTGCA  
 ACTGGTGATTATAAACTTGAAGAAACACAAGCTCCAGCTGGTTACGCTAAATTAGCAGGTGATGTTGATTTT  
 AAAGTTGGAAACAGCTCAAAGCAGACGACTCAGGTAACATTGATTACACTGCTAGCAGCAATAAAAAAGAC  
 GCTCAACGCATAGAAAACAAAAAGTGACTATCCACAAACAGGTGGTATTGGTACAATTCTTTTCACAATT  
 ATTGTTTTAAGCATTATGCTTGGAGCGGTAATTATCATGAAAAGACGTCAATCAGAGGAAGCTTAA  
 15 ORF amino acid sequence (SEQ ID NO: 101):  
 MKRINKYFAMFSALLLILTSLLSVAPVFAEMGNITKTVTLHKIVQTSNLAKPFPNGINGLNGTKYMGQKL  
 TDISGYFGQGSKEIAGAFFAVMNESQTKYITESGTEVESIDAAGVLKGLTTENGI TFNTANLKGTYQIVELL  
 DKSNYKNGDKVLADSKAVPVKITLPLYNEEGIVVDAEVYPKNTEEAPQIDKNFAKANKLLNDSNSAIAGGA  
 DYDKYQAEKAKATAEIGQEI PYEVKTKIQKGSKYKNLAWVDTMSNGLTMGNTVNLEASSGSFVEGTDYNVER  
 20 DDRGFLLKFTDTGLTKLQKEAETQAVEFTLTYSATVNGAAIDDKPESNDIKLQYGNKPGKKVKEIPVTPSNG  
 EITVSKTWDKGSLENANVVYTLKDGGTAVASVSLTKTTPNGEINLNGIKFTVTGAFAGKFSGLTDSKTYM  
 ISERIAGYGNTITTGAGSAAITNTPSDSNPTPLNPTEPKVVTHGKKFVKTSSTETERLQGAQFVVKDSAGKY  
 LALKSSATISAQTTAYTNAKTALDAKIAAYNKLSADDQKGTGETAKAEIKTAQDAYNAAFIVARTAYEWT  
 NKEDANVVKVTSNADGQFEVSGLATGDYKLEETQAPAGYAKLAGDVDFKVGNSKADDSGNI DYTASSNKKD  
 25 AQRIENKQVTIPQTGGIGTILFTIIGLSIMLGAVIIMKRRQSEEA

**Strain IC98**  
ORF DNA sequence (SEQ ID NO: 144):  
 ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCACTATTCTCAGTT  
 30 GCACCAGCGTTTGCAGGACGACGCAACAACCTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCCTTCTTTGTTTTCAAAAATGAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTCTGTT  
 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 35 GAATTGAAAGAAAAATCAAACCTACGATAACAACGGTCTATCTTGGCTGATTCAAAGCAGTTCCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGAATTTGCAGATAAAGATCTTGATTACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATCTTAAAGGCTCAGACTATAAG  
 AAACCTGGTTTGGACTGATAGCATGACTAAAGGTTTGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 40 GAAGATTTTCCCTGTTTTAACTACAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAGTCATTAAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAAACGGATGGTACATGG  
 45 GTGAACGTTGCTTACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTCACAGGTTTGGATAATGCT  
 AAAACTTACCGCGTTGTGCAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAAACCTCAAATGATCCAACCTCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAACCTGCTAAA  
 50 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 ACTGGTTTGGATAAAGGCACTTATGGCTTGGAAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAACCTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 55 TCTGTAAAAAAGATGCCCAACAAGTTCAAACAAAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATTCTTTTCACAATTATTGGTTTTAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA  
ORF amino acid sequence (SEQ ID NO: 102):  
 MKKINKCLTMFSTLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGTKGKNDSDYVGKQINDL  
 60 KSYFGSTDAKEIKGAFFVFKNETGTFITENGKEVDLLEAKDAEGGAVLSGLTKDNGFVFNTAKLKGIIYQIV



ELKEKSNYDNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTETKPQVDKNFADKDLDDYTDNRKDKGV  
 VSATVGDKEYIVGTKILKGSYKLVWTDSDMTKGLTFNNNVKVTLDGEDFPVLNYKLVTDQGFRLALNAT  
 GLAAVAAAADKDKDVEIKITYSATVNGSTTVEIPETNDVKLDYGNPTEESEPEQEGTPANQEIKVIKDWAVDG  
 TITDANVAVKAIFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPHYVSFKNGVV  
 5 TIKNNKNSNDPTPINPSEPKVVITYGRKFVKTQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYGLEETQAPAGYATLSGDVNFVETATSYSKGATTDIAYDKGSVKKDAQVQNKVVTIPQTGGIGT  
 ILFTIIGLSIMLGAVVIMKRRQSEEA

10 **Strain IC105**

ORF DNA sequence (SEQ ID NO: 145):

ATGAAAAGAATCAACAAATATTTTGCATGTTCTCGGCATTGTTATTAATTTTAAACATCGTTGTTATCGGTA  
 GCTCCGGTATTTGCTGCTGAGATGGGAAATATCACTAAAACAGTAACCTTACACAAAATTGTTCAAACATCC  
 GATAATTTGGCTAAGCCAAATTTCCCAGGAATAAATGGATTGAATGGAACGAAGTATATGGGTCAAAAACCTT  
 15 ACTGACATTTTCAGGATATTTTGGGCAAGGTTCTAAAGAAATCGCCGGTGTCTTTCTTTGCGGTTATGAATGAA  
 AGTCAGACAAAATATATCACAGAAAGTGGTACTGAAGTAGAAAGTATCGATGCAGCAGGTGTCTTAAAGGT  
 TTGACAACGAAAACGGCATTACATTTAATACTGCAAACCTAAAAGGAACATAACCAAATCGTTGAGTTGCTT  
 GACAAATCTAATTATAAAAATGGTGACAAAGTTCTTGCTGACTCAAAGCTGTCCCAGTGAAAATCACTCTT  
 CCTTTGTATAACGAAGAAGGAATTGTCGTGGACGCTGAAGTGTATCCAAAGAATACAGAAGAAGCACCACAA  
 20 ATCGACAAAACCTTTGCTAAAGCAAATAAATTGTTGAATGACAGTGATAATTCAGCTATTCAGGTTGGGGCA  
 GACTACGACAAATATCAGGCAGAAAAGCAAAGCTACTGCTGAAATCGGTCAAGAAATCCCTTACGAAGTT  
 AAAACAAAATCCAAAAGGGTCTAAATACAAAACCTTGCCTGGGTGCGATACCATGTCAAATGGTTTGACA  
 ATGGGTAACTGTTAACTTAGAAGCATCGTCAGGCTCTTTTGTAGAAGGTACAGATTACAATGTTGAACGT  
 GATGACCGTGGTTTTCACTTTGAAATTCACAGATACAGGTTTGACTAAGCTACAAAAGAAGCGGAAACACAC  
 25 GCTGTTGAATTCACATTGACATATAGCGCAACAGTTAACGGTGCAGGCTATTTGATGACAAGCCAGAAAGCAAT  
 GATATCAAACCTTCAATACGGTAACAAACCAGGTAAAAAGTAAAAGAAATCCCAGTAACACCGTCAAATGGC  
 GAAATCACTGTTAGCAAACCTTGGGACAAAGGTTGAGATTTAGAGAATGCGAATGTTGTTTATACCCTTAAA  
 GATGGTGGAAACAGCTGTTGCCTCAGTTTCATTGACAAAACAACCAAATGGCGAAATCACTTAGGTAAT  
 GGTATTAATTTACAGTTACTGGAGCGTTTGCTGGTAAATTCAGTGGTCTGACTGATAGTAAAACATACATG  
 30 ATCTCAGAACGTATCGCTGGTTATGGTAATACAATCACTACTGGTGTGGTAGTGCAGCTATCACCAATACT  
 CCAGATTCAGACAACCCAACCACTTAATCCAACCTGAACCAAAGTTGTGACACACGGTAAAAAATTCGTC  
 AAAACAAGTTCGACTGAAACAGAACGCTTGCAAGGTGCACAGTTCGTTGTTAAAGATTCAGCTGGTAAATAC  
 CTTGCATTGAAATCATCTGCGACAATATCAGCTCAAACAACAGCTTACACAAATGCTAAAACCTGCTCTTGAC  
 GCTAAAATCGCAGCTTACAACAACTTTCAGCAGACGATCAAAGGTACTAAAGGTGAAACAGCTAAAGCA  
 35 GAAATCAAACCTGCTCAAGACGCTTACAATGCAGCCTTCATCGTAGCTCGTACAGCTTACGAGTGGGTAAC  
 AATAAAGAAGATGCTAACGTTGTTAAAGTGACTTCAAACGCTGACGGTCAATTTGAAGTTAGCGGTCTTGCA  
 ACTGGTGATTATAAACTTGAAGAAACACAAGCTCCAGCTGGTTACGCTAAATTAGCAGGTGATGTTGATTTT  
 AAAGTTGGAAACAGCTCAAAGCAGACGACTCAGGTAACATTTGATTACACTGCTAGCAGCAATAAAAAAGAC  
 GCTCAACGCATAGAAAACAAAAAGTGACTATTCCACAACAGGTGGTATTGGTACAATTCTTTTACAATT  
 40 ATTGGTTTAAAGCATTATGCTTGGAGCGGTAATTATCATGAAAAGACGTCAATCAGAGGAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 103):

MKRINKYFAMFSALLLILTSLLSVAPVFAEMGNITKTVTLHKIVQTSNLAKPFPNGINGLNGTKYMGQKL  
 TDISGYFGQGSKEIAGAFFAVMNESQTKYITESGTEVESIDAAGVLKGLTTENGI TFNTANLKGTYQIVELL  
 DKSNYKNGDKVLADSKAVPVKITLPLYNEEGIVVDAEVYPKNTTEAPQIDKNFAKANKLLNDSNSAIAGGA  
 45 DYDKYQAEKAKATAEIGQEIPEYVKTQKQKSKYKNLAWVDTMSNGLTMGNTVNLEASSGSFVEGTDYNVER  
 DDRGFTLKFDTGLTKLQKEAETHAVEFTLTYSATVNGAAIDDKPESNDIKLQYGNKPGKKVKEIPVTPSNG  
 EITVSKTWDKGSDELNANVVYTLKDGGTAVASVSLTKTTPNGEINLNGIKFTVTGAFAGKFSGLTDSKTYM  
 ISERIAGYGNITTTGAGSAAITNTPDSDNPTPLNPTEPKVVTHGKKFVKTSSTETERLQGAQFVVKDSAGKY  
 LALKSSATISAQTTAYTNAKTALDAKIAAYNKLSADDQKGTGETAKAEIKTAQDAYNAAFIVARTAYEWT  
 50 NKEDANVVKVTSNADGQFEVSGLATGDYKLEETQAPAGYAKLAGDVDFKVGNSKADDSGNI DYTASSNKKD  
 AQRIENKVTIPQTGGIGTILFTIIGLSIMLGAVIIMKRRQSEEA

55 **Strain IC216**

ORF DNA sequence (SEQ ID NO: 146):

ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCACCTATTCTCAGTT  
 GCACCAGCGTTTGCAGACGACGCAACAACTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGTCTTTCTTTGTTTTCAAAAATGAAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTCTGTT  
 60 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTTAACTGCTAAGTTAAAAGGAATTTACCAAATCGTT



GAATTGAAAGAAAAATCAAACCTACGATAACAACGGTTCTATCTTGGCTGATTCAAAGCAGTTCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGAACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 5 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGTAACAAAAATTTCTTAAAGGCTCAGACTATAAG  
 AAAGTTTGGACTGATAGCATGACTAAAGGTTTGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 GAAGATTTTCCCTGTTTTAACTACAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 10 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAGTCACTTAAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAAACGGATGGTACATGG  
 GTGAACGTTGCTTACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTCACAGGTTTGGATAATGCT  
 AAAACTTACCGCGTTGTGCAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAAACTCAAATGATCCAACCTCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
 15 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 20 ACTGGTTTGGATAAAGGCACTTATGGCTTGGGAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAAAACAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATTCTTTTCACAATTATTGGTTTAAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 104):

25 MKKINKCLTMFSTLLLI L T S L F S V A P A F A D D A T T D T V T L H K I V M P Q A A F D N F T E G T K G K N D S D Y V G K Q I N D L  
 K S Y F G S T D A K E I K G A F F V F K N E T G T K F I T E N G K E V D T L E A K D A E G G A V L S G L T K D N G F V F N T A K L K G I Y Q I V  
 E L K E K S N Y D N N G S I L A D S K A V P V K I T L P L V N N Q G V V K D A H I Y P K N T E T K P Q V D K N F A D K D L D Y T D N R K D K G V  
 V S A T V G D K K E Y I V V T K I L K G S D Y K K L V W T D S M T K G L T F N N N V K V T L D G E D F P V L N Y K L V T D D Q G F R L A L N A T  
 30 G L A A V A A A K D K D V E I K I T Y S A T V N G S T T V E I P E T N D V K L D Y G N N P T E E S E P Q E G T P A N Q E I K V I K D W A V D G  
 T I T D A N V A V K A I F T L Q E K Q T D G T W V N V A S H E A T K P S R F E H T F T G L D N A K T Y R V V E R V S G Y T P E Y V S F K N G V V  
 T I K N N K N S N D P T P I N P S E P K V V T Y G R K F V K T N Q A N T E R L A G A T F L V K K E G K Y L A R K A G A A T A E A K A A V K T A K  
 L A L D E A V K A Y N D L T K E K Q E G Q E G K T A L A T V D Q K Q A Y N D A F V K A N Y S Y E W V A D K K A D N V V K L I S N A G G Q F E I  
 T G L D K G T Y G L E E T Q A P A G Y A T L S G D V N F E V T A T S Y S K G A T T D I A Y D K G S V K K D A Q Q V Q N K K V T I P Q T G G I G T  
 I L F T I I G L S I M L G A V V I M K K R Q S E E A

### Strain IC245

ORF DNA sequence (SEQ ID NO: 147):

40 ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCACTATTCTCAGTT  
 GCACCAGCGTTTGCAGCAGCACAACCAACTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCCTTCTTTGTTTTCAAAAATGAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTGTT  
 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 45 GAATTGAAAGAAAAATCAAACCTACGATAACAACGGTTCTATCTTGGCTGATTCAAAGCAGTTCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGAACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATTTCTTAAAGGCTCAGACTATAAG  
 AAAGTTTGGACTGATAGCATGACTAAAGGTTTGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 50 GAAGATTTTCCCTGTTTTAACTACAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAGTCACTTAAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAAACGGATGGTACATGG  
 GTGAACGTTGCTTACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTCACAGGTTTGGATAATGCT  
 55 AAAACTTACCGCGTTGTGCAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAAACTCAAATGATCCAACCTCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 60 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT



- 88 -

GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 ACTGGTTTGGATAAAGGCACTTATGGCTTGGGAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAAACAAAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 5 ATTCTTTTACAATTATTGGTTTAAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 105):

MKKINKCLTMFSTLLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGTKGKNDSYVVGKQINDL  
 KSYFGSTDAKEIKGAFFVFKNETGTFKITENGKEVDLEAKDAEGGAVLSGLTKDNGFVFNTAKLKGIIYQIV  
 10 ELKEKSNDNNGSILADSKAVPVKITLPLVNNQGVVKAHIIYPKNETETKPQVDKNFADKDLDDYTDNRKDKGV  
 VSATVGDKKEYIVGTKILKGSYKLVWTDGMTKGLTFNNNVKVTLDGEDFPVLNYKLVTDQGFRLALNAT  
 GLAAVAAAADKDKDVEIKITYSATVNGSTTVEIPEITNDVKLDYGNPTEESEPEQEGTPANQEIKVIKDWAVDG  
 TITDANVAVKAIFFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPYVVSFKNGVV  
 TIKNNKNSNDPTPINPSEPKVVVYGRKFKVKNQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
 15 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYGLEETQAPAGYATLSGDVNFVETATSYSKGATTDIAYDKGSVKKDAQQVQNKKVTIPQTGGIGT  
 ILFTIIIGLSIMLGAVVIMKKRQSEA

### Strain IC250

ORF DNA sequence (SEQ ID NO: 148):

ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCACACTATTCTCAGTT  
 GCACCAGCGTTTGCAGGACGACGCAACAACTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCCTTTCTTTGTTTTCAAAAATGAAACTGGT  
 25 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTCTGTT  
 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 GAATTGAAAGAAAATCAAACCTACGATAACAACGGTCTATCTTGGCTGATTCAAAGCAGTTCCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGAACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 30 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATCTTAAAGGCTCAGACTATAAG  
 AAACGGTTTTGGACTGATAGCATGACTAAAGGTTTGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 GAAGATTTTCCCTGTTTTAACTACAAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 35 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAGTCATTAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAAACGGATGGTACATGG  
 GTGAACGTTGCTTACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTACAGGTTTGGATAATGCT  
 AAAACTTACCGCGTTGTGCAACGTTGTTAGCGGCTACACTCCAGAATACGTATCATTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAAACCTCAAATGATCCAACCTCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
 40 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 45 ACTGGTTTGGATAAAGGCACTTATGGCTTGGGAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAAACAAAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATTCTTTTACAATTATTGGTTTAAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 106):

MKKINKCLTMFSTLLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGTKGKNDSYVVGKQINDL  
 KSYFGSTDAKEIKGAFFVFKNETGTFKITENGKEVDLEAKDAEGGAVLSGLTKDNGFVFNTAKLKGIIYQIV  
 ELKEKSNDNNGSILADSKAVPVKITLPLVNNQGVVKAHIIYPKNETETKPQVDKNFADKDLDDYTDNRKDKGV  
 VSATVGDKKEYIVGTKILKGSYKLVWTDGMTKGLTFNNNVKVTLDGEDFPVLNYKLVTDQGFRLALNAT  
 55 GLAAVAAAADKDKDVEIKITYSATVNGSTTVEIPEITNDVKLDYGNPTEESEPEQEGTPANQEIKVIKDWAVDG  
 TITDANVAVKAIFFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPYVVSFKNGVV  
 TIKNNKNSNDPTPINPSEPKVVVYGRKFKVKNQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYGLEETQAPAGYATLSGDVNFVETATSYSKGATTDIAYDKGSVKKDAQQVQNKKVTIPQTGGIGT  
 60 ILFTIIIGLSIMLGAVVIMKKRQSEA



**Strain IC251**

ORF DNA sequence (SEQ ID NO: 149):

5 ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCCTATTCTCAGTT  
 GCACCAGCGTTTGC GGACGACGCAACAACTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCCTTCTTTGTTTTCAAAAATGAAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGCTGTT  
 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 10 GAATTGAAAGAAAATCAAACACTACGATAACAACGGTCTATCTTGGCTGATTCAAAGCAGTTCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGA ACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATCTTAAAGGCTCAGACTATAAG  
 AA ACTGGTTTGGACTGATAGCATGACTAAAGGTTTGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 15 GAAGATTTTCTGTTTTAACTACAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAGTCAATAAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAACGGATGGTACATGG  
 20 GTGAACGTTGCTTACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTCACAGGTTTGGATAATGCT  
 AAACTTACCGCGTTGTGCAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAAACCTCAAATGATCCAACCTCAATCAACCCATCAGAACCAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAACTGCTAAA  
 25 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 ACTGGTTTGGATAAAGGCACTTATGGCTTGGAAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 30 TCTGTAAAAAAGATGCCCAACAAGTTCAAAAACAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATTCTTTTCACAATTATTGGTTTAAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 107):

35 MKKINKCLTMFSTLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGKTKGNDSYVVGKQINDL  
 KSYFGSTDAKEIKGAFFVFKNETGTFKIFITENGKEVDLEAKDAEGGAVLSGLTKDNGFVFNTAKLKG IYQIV  
 ELKEKSNDNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTETKPQVDKNFADKDLDYTDNRKDKGV  
 VSATVGDKEYIVGTKILKGSYKLVWTD SMTKGLTFNNNVKVTLDGEDFPVLNYKLVTD DQGFRLALNAT  
 GLAAVAAA AKDKDVEIKITYSATVNGSTTVEIPE TNDVKLDYGNPTEESEPEQEGTPANQEIKVIKDWAVDG  
 40 TITDANVAVKAI FTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPEYVSFKNGVV  
 TIKNNKNSNDPTPINPSEPKVVVYGRKFVKT NQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTA  
 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYGLEETQAPAGYATLSGDVNF EVTATSYSKGATTDIAYDKGSVKKDAQVQNKKVTIPQTGGIGT  
 ILFTIIGLSIMLGAVVIMKKRQSEEA

**Strain IC252**

ORF DNA sequence (SEQ ID NO: 150):

45 ATGAAAAAATCAACAAATATTTTGCAGTCTTCTCGGCATTGCTACTGACCGTAACATCATTGTTCTCAGTT  
 GCACCAGTGT TTTGC GGGAAGAAGCAAAA ACTACTGACACAGTGACCTTGCACAAGATTGTCATGCCTCGAACT  
 GCATTTGACGGTTTTACTGCTGGTACAAAGGTAAGGATAATACTGACTACGTTGGTAAACAAATCGAAGAC  
 50 CTTAAACTTACTTTGGCTCAGGCGAAGCGAAAGAAATCGCAGGTGCTTACTTTGCTTTCAAAAATGAAGCT  
 GGTACTAAATACATCACTGAAAATGGTGAAGAAGTTGATACTTTGGATAACAACAGATGCCAAAGGTGGTGCT  
 GTTCTTAAAGGTTTAAACAACAGACAATGGTTTCAAATTTAACACTTCTAAATTAACAGGAACTTACCAAATC  
 GTTGAATTGAAAGAAAATCTACATAACAACGATGGTTCATCTTGGCTGATTCAAAGCAGTTCAGTT  
 AAAATCACTCTTCCATTGGTAAACGACAATGGTGTGTTAAAGACGCTCACGTTTATCCAAAGAACACTGAA  
 55 ACAAACCACAAGTAGATAAGA ACTTTCGAGATAAAGA ACTTGATTATGCGAACAACAAAAAGACAAAGGG  
 ACTGTCTCAGCATCTGTTGGTGTGTTAAAAAATATCATGTTGGAACAAAAATCCTTAAAGGTTTCAGACTAT  
 AAGAAATTAATCTGGACCGATAGCATGACCAAAGGTTTGACTTTCAACAACGATATGCTGTAACATTGGAT  
 GGTGCAACTCTTGATGCTACAAATTACAACTTGTAGCAGATGACCAAGGTTTCCGCCTTGTCTTGACTGAC  
 AAAGGTCTTGAAGCAGTGGCAAAAGCCGCAAAAACAAAAGATGTTGAAATCAAGATCACTTACTCAGCTACT  
 60 TTGAACGGTTCTGCTGTCGTTGAAGTTCTAGAAACCAATGATGTTAAATTGGACTACGGCAACAACCCAACA



ATTGAAAATGAACCAAAGAAGGTATTCCAGTTGATAAGAAAATCACTGTAAACAAAACATGGGCAGTAGAT  
 GGCAATGAAGTGAATAAAGCAGATGAAACAGTTGATGCTGTCTTCACCTTGCAAGTTAAAGATGGTGACAAA  
 TGGGTGAATGTTGATTCAGCTAAAGCAACAGCTGCAACTAGCTTCAAACACACTTTTGAAAACCTGGATAAT  
 GCTAAAACCTACCGCGTTATCGAACGTGTTAGCGGCTACGCTCCAGAATACGTCTCATTTGTAAATGGCGTT  
 5 GTAACCATCAAGAACAACAAAGACTCAAATGAGCCAACTCCAATCAACCCATCAGAACCAAAGTGGTGACT  
 TATGGACGTAAATTTGTGAAAACAAATAAAGATGGAAAAGAACGCTTGGCAGGAGCTACCTTCCTTGTTAAG  
 AAAGATGGCAAGTACTTGGCACGTAAATCAGGTGTTGCAACAGATGCAGAAAAAGCTGCTGTAGATTCAACT  
 AAATCAGCATTGGATGCTGCTGTTAAAGCTTACAATGATTTGACTAAAGAAAAACAAGAAGGTCAAGATGGT  
 AAATCAGCATTGGCTACCGTTAGTGAAAAACAAAAGCTTACAATGATGCCTTTGTTAAAGCTAACTACTCA  
 10 TACGAATGGGTTGAAGATAAAAATGCTAAGAATGTTGTTAAATTGATTTCTAACGATAAAGGTCAATTTGAA  
 ATTACTGGCTTGACTGAAGGTCAATACTCATTGGAAGAAACACAAGCACCAACTGGTTATGCTAAATTATCA  
 GGTGATGTTTCGTTTAATGTTAATGCTACTTCATACAGTAAAGGTTCTGCTCAAGATATTGAGTATACCCAA  
 GGTTCTAAAACATAAGATGCACAACAAGTTATCAATAAGAAGGTTACTATTCACAAACAGGTGGTATTGGT  
 ACAATTTTTTTTCACAATTATTGGATTAAGTATTATGCTTGGAGCGGTAGTTATCATGAAAAGACGTCAATCA  
 15 GAGGAAGTTTAA

ORF amino acid sequence (SEQ ID NO: 108):

MKKINKCLTMFSTLLLI L T S L F S V A P A F A D D A T T D T V T L H K I V M P Q A A F D N F T E G T K G K N D S D Y V G K Q I N D L  
 K S Y F G S T D A K E I K G A F F V F K N E T G T K F I T E N G K E V D T L E A K D A E G G A V L S G L T K D N G F V F N T A K L K G I Y Q I V  
 E L K E K S N Y D N N G S I L A D S K A V P V K I T L P L V N N Q G V V K D A H I Y P K N T E T K P Q V D K N F A D K D L D Y T D N R K D K G V  
 20 V S A T V G D K K E Y I V G T K I L K G S D Y K K L V W T D S M T K G L T F N N N V K V T L D G E D F P V L N Y K L V T D D Q G F R L A L N A T  
 G L A A V A A A A K D K D V E I K I T Y S A T V N G S T T V E I P E T N D V K L D Y G N N P T E E S E P Q E G T P A N Q E I K V I K D W A V D G  
 T I T D A N V A V K A I F T L Q E K Q T D G T W V N V A S H E A T K P S R F E H T F T G L D N A K T Y R V V E R V S G Y T P E Y V S F K N G V V  
 T I K N N K N S N D P T P I N P S E P K V V T Y G R K F V K T N Q A N T E R L A G A T F L V K K E G K Y L A R K A G A A T A E A K A A V K T A K  
 L A L D E A V K A Y N D L T K E K Q E G Q E G K T A L A T V D Q K Q A Y N D A F V K A N Y S Y E W V A D K K A D N V V K L I S N A G G Q F E I  
 25 T G L D K G T Y G L E E T Q A P A G Y A T L S G D V N F E V T A T S Y S K G A T T D I A Y D K G S V K K D A Q Q V Q N K K V T I P Q T G G I G T  
 I L F T I I G L S I M L G A V V I M K K R Q S E E A

### Strain IC253

ORF DNA sequence (SEQ ID NO: 151):

ATGAAAAAATCAACAAATGTCTTACAGTGTCTCGACACTGCTATTGATCTTAACGTCACTATTCTCAGTT  
 GCACCAGCGTTTGC GGACGACGTAACAACCTGATACTGTGACCTTG CACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATTAAGGGTGCTTTCTTTGTTTTCAAAAATGAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGCTGTT  
 35 CTTTCAGGGTTAACAAAAGACACTGGTTTTGCTTTTAACTGCTAAGTTAAAAGGAACTTACCAAATCGTT  
 GAATTGAAAGAAAAATCAAACTACGATAACAACGGTTCATCTTGGCTGATTCAAAGCAGTTCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGAACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATCTTAAAGGCTCAGACTATAAG  
 40 AAAGTGGTTTGGACTGATAGCATGACTAAAGGTTTGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 AAAGATTTTCTGTTTTTAACTACAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCTGCTGCAAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAGTTCCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAGTCACTTAAAGACTGGGCAGTAGATGGT  
 45 ACAATTACTGATGTTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAACGGATGGTACATGG  
 GTGAACGTTGCTTACACGAAGCAACAACCAATCACGCTTTGAACATACTTTACAGGTTTGGATAATACT  
 AAACTTACCGCGTTGTGCAACGTGTTAGCGGCTACACTCCAGAATATGTATCATTTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAAACCTCAAATGATCCAACCTCCAATCAACCCATCAGAACCAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTTGTTAAGAAA  
 50 GAAGGAAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 ACTGGTTTGGATAAAGGCACCTTATAGCTTGGAAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 55 GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACTGACATCGCATATGATAAAGGA  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAACAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATTCTTTTCACAATTATTGGTTTAAAGCATTATGCTTGGAGCAGTAGTTGTTCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 109):



MKKINKCLTVFSTLLLLILTSLFSVAPAFADDVTTDTVTLHKIVMPQAAFDNFTEGKKGKNDSDYVGKQINDL  
 KSYFGSTDAKEIKGAFFVFKNETGTKFITENGKEVDLEAKDAEGGAVLSGLTKDTGFAFNTAKLKGTYQIV  
 ELKEKSNYDNNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTETKPQVDKNFADKDLDDYTDNRKDKGV  
 VSATVGDKEYIVGTKILKGSYKLVWTDGMTKGLTFNNNVKVTLDGKDFPVLNYKLVTDQGFRLALNAT  
 5 GLAAVAAAADKDKDVEIKITYSATVNGSTTVEVPETNDVKLDYGNPNTEESEPEQEGTPANQEIKVIKDWAVDG  
 TITDENVAVKAIFFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNTKTYRVVERVSGYTPYVVSFKNGVV  
 TIKNNKNSNDPTPINPSEPKVVVYGRKFVKTQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 10 TGLDKGTYSLEETQAPAGYATLSGDVNFVETATSYSKGATTDIAYDKGSVKKDAQVQNKVVTIPQTGGIGT  
 ILFTIIGLSIMLGAVVVMKKRQSEEA

**Strain IC254**

ORF DNA sequence (SEQ ID NO: 152):

ATGAAAAGAATCAACAAATATTTTGCATGTTCTCGGCATTGTTACTGACTTTAACGTCATTGCTCTCAGTT  
 15 GCACCAGCGTTTGC GGATGAAGCAACAATAACAGTGACTTTGCACAAGATTTTGCAAACCGAATCAAAT  
 CTTAACAAAAGTAACTTCCCAGGAACTACAGGTCTTAACGGAAAAGACTACAAAGGTGGAGCTATTTCTGAC  
 CTTGCTGGTTACTTTGGCGAGGGATCTAAAGAAATCGAAGGTGCGTTCTTTGCTTTAGCTTTGAAAGAAGAT  
 AAAAGTGGTAAAGTGCAATATGTTAAGGCAAAAGAAGGTAACAAATTAACACCAGCCTTAATTAATAAAGAT  
 20 GGTACTCCTGAAATAACAGTAAATATTGATGAGGCCGTGTCTGGATTGACACCAGAGGGAGATACTGGACTT  
 GTTTTCAACACCAAAGGATTGAAAGGCGAGTTTAAAATTGTTGAAGTTAAATCAAATCTACTTACAACAAT  
 AATGGTTCCTCCTGGCTGCTTCAAAGCGGTTCCAGTTAACATCACTCTTCCATTGGTAAATGAAGATGGT  
 GTTGTGCTGATGCCCATGTTTATCCAAAGAACACTGAAGAAAACCAGAAATTGATAAAAACCTTTGCTAAA  
 ACAACGATTTGACAGCATTGACAGATGTTAATAGACTTTTGACAGCTGGCGCAAATTATGGTAATTATGCA  
 25 CGTGACAAAGCAACTGCTACTGCTGAAATCGGTAAAGTTGTTCCATTATGAAGTTAAAACAAAAATTCACAAA  
 GGTTCTAAATACGAAAACCTGGTTTGGACAGATATAATGTCAAATGGTTTGACAATGGGTTCAACTGTTAGC  
 CTTAAAGCTTCAGGAACTACAGAACTTTTGCTAAGGATACAGACTATGAACTTAGCATTGATGCCCGTGGT  
 TTCACATTAATAATTCACAGCTGATGGATTGGGCAAATTGGAAAAGCAGCTAAAACAGCTGATATTGAATTT  
 ACATTGACTTATAGTGCTACTGTTAATGGTCAAGCAATTATTGATAATCCAGAATCCAATGATATCAAATTTG  
 TCGTATGGTAACAAACCAGGTAAGACTTGACTGAACTTCCGTGTTACACCTTCAAAGGGTGAAGTAACAGTT  
 30 GCTAAAACCTGGTCTGACGGAATTGCACCTGATGGTGTAAACGTTGTTTACACATTGAAAGATAAAGATAAA  
 ACTGTTGCTTCAGTATCATTGACAAAAACATCTAAAGGTACAATCGACCTTGGAAATGGTATCAAATTTGAA  
 GTATCTGGTAACTTCTCGGGTAAATTCAGTGGTCTAGAAAACAAATCATAACATGATCTCAGAACGTGTTTCT  
 GGTTACGGAAGTGCAATAAATCTAGAAAATGGTAAAGTAACCATTACCAATACCAAAGATTCTGATAACCCA  
 ACACCATTGAACCCAACTGAACCAAAGTTGAACTCATGGTAAGAAATTTGTCAAACCTAATGAACAAGGT  
 35 GACCGTTTGGCTGGTGCACAATTCGTTGTGAAAAACTCAGCAGGTAATACTTGGCTCTTAAAGCAGATCAA  
 TCAGAAGGTCAAAAAACTTTAGCTGCTAAGAAAATAGCTTTAGATGAAGCTATCGCTGCTTATAACAAGTTG  
 TCTGCAACAGACCAAAGGTGAAAAGGAATTACTGCAAAAGAACTTATCAAACCTAAACAAGCAGATTAC  
 GATGCAGCCTTCATTGAGGCTCGTACAGCTTATGAGTGGATAACAGATAAGGCTAGAGCCATTACCTACACT  
 TCAAACGATCAAGGTCAATTTGAAGTTACAGGTCTTGCAGACGGTACTTACAACCTTGAAGAAACACTTGGT  
 40 CCAGCAGGATTTGCTAAGTTGGCAGGTAATATTAAGTTTGTAGTTAATCAAGGGTCATACATAACAGGTGGT  
 AACATTGACTACGTTGCTAACAGCAACCAAAGATGCGACACGTGTAGAAAATAAAAAGGTAACAATCCCA  
 CAAACAGGTGGTATTGGTACAATTCTTTTACAATTATTGGTTTAAAGCATTATGCTTGGAGCAGTAGTTATC  
 ATGAAAAGACGCCAATCAAAGGAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 110):

MKRINKYFAMFSALLLTLTSLLSVAPAFADDEATTNTVTLHKILQTESNLNKS NFPGTTGLNGKDYKGGAI SD  
 45 LAGYFGEKSKEIEGAFFALALKEDKSGKVQYVKAKEGNKLT PALINKDGTPEITVNI DEAVSGLTPEGDTGL  
 VFNTKGLKGEFKIVEVKSKSTYNNNGSLLAASKAVPVNITLPLV NEDGVVADAHVYPKNTEEKPEIDKNFAK  
 TNDLTALTDVNRLLTAGANYGNYARDKATATAEIGKVVPYEVKTKIHKGSKYENLVWTDIMSNGLTMGSTVS  
 LKASGTTETFADTDYELSIDARGFTLKF TADGLGKLEKAAKTADIEFTLTYSATVNGQAIIDNPESNDIKL  
 50 SYGNKPGKDLTELPVTPSKGEVTVAKTWS DGIAPDGVNVVYTLKDKDKTVASVSLTKTSKGTIDLNGIKFE  
 VSGNFSGKFTGLENKSYMISERVSGYGSAINLENGKVTITNTKDSNDPTPLNPTEPKVETHGKKFVKTNEQG  
 DRLAGAQFVVKNSAGKYLALKADQSEGQKTLA AKKIALDEAIAAYNKLSATDQKGEKGITAKELIKTKQADY  
 DAAFI EARTAYEWITDKARAITYTSNDQGF EVTGLADGTYNLEETLAPAGFAKLAGNIKFVVNQGSYITGG  
 55 NIDYVANSNQKDATRVENKVTIPQTGGIGTILFTIIGLSIMLGAVVIMKRRQSKEA

**Strain IC255**

ORF DNA sequence (SEQ ID NO: 153):

ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCACTATTCTCAGTT  
 60 GCACCAGCGTTTGC GGACGACGCAACAATACTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT



AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCTTTCTTTGTTTTCAAAAATGAAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTCTGTT  
 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 GAATTGAAAGAAAATCAAACACTACGATAACAACGGTTCCTATCTTGGCTGATTCAAAGCAGTTCCAGTTAAA  
 5 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGAACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATTTCTTAAAGGCTCAGACTATAAG  
 AAAGTGGTTTGGACTGATAGCATGACTAAAGGTTTGGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 GAAGATTTTCTGTTTTAACTACAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 10 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAGTCAATTAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAAACGGATGGTACATGG  
 GTGAACGTTGCTTCACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTTACAGGTTTGGATAATGCT  
 15 AAACTTACCGCGTTGTGCAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAAACACTCAAATGATCCAACCTCCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 20 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 ACTGGTTTGGATAAAGGCACCTTATGGCTTGGAAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACTGACATCGCATATGATAAAGGC  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAAACAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 25 ATCTTTTACAATTATTGGTTTAAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 111):

MKKINKCLTMFSTLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGTKGKNDSDYVGKQINDL  
 KSYFGSTDAKEIKGAFFVFKNETGTFKFI TENGKEVDTLKAKDAEGGAVLSGLTKDNGFVFNTAKLKIYQIV  
 30 ELKEKSNYDNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTEKPKQVDKNFADKDLDDYTDNRKDKGV  
 VSATVGDKEYIVGTKILKGSYKLVWTDMSMTKGLTFNNNVKVTLDGEDFPVLNYKLVTDQGFRLALNAT  
 GLAAVAAAADKDKVEIKITYSATVNGSTTVEI PETNDVKLDYGNPTEESEPEQEGTPANQEI KVIKDWAVDG  
 TITDANVAVKAI FTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPYVVSFKNGVV  
 TIKNNKNSNDPTPINPSEPKVVVYGRKFVKTNQANTERLAGATFLVKKEGKY LARKAGAATAEAKAAVKTAK  
 35 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYGLEETQAPAGYATLSGDVNFVETATSYSKGATTDIAYDKGSVKKDAQQVQNKKVTI PQTGIGT  
 ILFTIIGLSIMLGAVVIMKKRQSEEA

### Strain IC287

ORF DNA sequence (SEQ ID NO: 154):  
 ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCACTATTCTCAGTT  
 GCACCAGCGTTTGCAGGACGACGCAACAACTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 45 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCTTTCTTTGTTTTCAAAAATGAAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTCTGTT  
 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 GAATTGAAAGAAAATCAAACACTACGATAACAACGGTTCCTATCTTGGCTGATTCAAAGCAGTTCCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGAACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 50 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATTTCTTAAAGGCTCAGACTATAAG  
 AAAGTGGTTTGGACTGATAGCATGACTAAAGGTTTGGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 GAAGATTTTCTGTTTTAACTACAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 55 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAGTCAATTAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAAACGGATGGTACATGG  
 GTGAACGTTGCTTCACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTTACAGGTTTGGATAATGCT  
 AAACTTACCGCGTTGTGCAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAAACACTCAAATGATCCAACCTCCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
 60 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA



GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 5 ACTGGTTTGGATAAAGGCACTTATGGCTTGGGAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAAAACAAAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATTCTTTTTCACAATTATTGGTTTAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA

10 ORF amino acid sequence (SEQ ID NO: 112):

MKKINKCLTMFSTLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGTEGKGNDSYVVGKQINDL  
 KSYFGSTDAKEIKGAFFVFKNETGTFKITENGKEVDLEAKDAEGGAVLSGLTKDNGFVFNTAKLKGIVQIV  
 ELKEKSNDYDNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTETKPQVDKNFADKDLDTDNRKDKGV  
 VSATVGDKEYIVGTKILKGSYKLVWTDSTMTKGLTFNNNVKVTLDGEDFPVLNYKLVTDQGFRLALNAT  
 15 GLAAVAAAADKDKDVEIKITYSATVNGSTTVEIPETNDVKLDYGNPTEESEPEQEGTPANQEIKVIKDWAVDG  
 TITDANVAVKAIFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPHYVSFKNGVV  
 TIKNNKNSNDPTPINPSEPKVVVYGRKFVKTQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYLEETQAPAGYATLSGDVNFVETATSYSKGATTDIAYDKGSVKKDAQQVQNKKVTIPQTGGIGT  
 20 ILFTIIGLSIMLGAVVIMKKRQSEEA

### Strain IC289

ORF DNA sequence (SEQ ID NO: 155):

ATGAAAAGAATCAACAAATATTTTGCATGTTCTCGGCATTGTTATTAATTTTAAACATCGTTGTTATCGGTA  
 25 GCTCCGGTATTTGCTGCTGAGATGGGAAATATCACTAAAACAGTAACCTTACACAAAATTGTTCAAACATCC  
 GATAATTTGGCTAAGCCAAATTTCCCAGGAATAAATGGATTGAATGGAACGAAGTATATGGGTCAAAAACCTT  
 ACTGACATTTTCAGGATATTTTGGGCAAGGTTCTAAAGAAATCGCCGGTGCCTTCTTTGCGGTTATGAATGAA  
 AGTCAGACAAAATATATCACAGAAAGTGGTACTGAAGTAGAAAGTATCGATGCAGCAGGTGTCCCTTAAAGGT  
 TTGACAACTGAAAACGGCATTACATTTAATACTGCAAACTTAAAAGGAACATAACCAATCGTTGAGTTGCTT  
 30 GACAAATCTAATTATAAAAATGGTGACAAAGTTCCTTGCTGACTCAAAAAGCTGTCCCAGTGAAAATCACTCTT  
 CCTTTGTATAACGAAGAAGGAATTGTCGTGGACGCTGAAGTGTATCCAAAGAATACAGAAGAAGCACCACAA  
 ATCGACAAAACCTTTGCTAAAGCAAATAAATGTTGAATGACAGTGATAATTCAGCTATTCAGGTTGGGGCA  
 GACTACGACAAATATCAGGCAGAAAAGCAAAGCTACTGCTGAAATCGGTCAAGAAATCCCTTACGAAGTT  
 AAAACAAAATCCAAAAGGGTCTAAATACAAAACCTTGCCTGGGTTCGATACCATGTCAAATGGTTTGACA  
 35 ATGGGTAACACTGTAACTTAGAAGCATCGTCAGGCTCTTTTGTAGAAGGTACAGATTACAATGTTGAACGT  
 GATGACCGTGGTTTCACTTTGAAATTCACAGATACAGGTTTGACTAAGCTACAAAAGAAGCGGAAACACAA  
 GCTGTTGAATTCACATTGACATATAGCGCAACAGTTAACGGTGCAGGCTATTGATGACAAGCCAGAAAGCAAT  
 GATATCAAACCTTCAATACGGTAACAAACCAGGTAAAAAAGTAAAAGAAATCCCAGTAACACCGTCAAATGGC  
 GAAATCAAACTGTTAGCAAACTTGGGACAAAGGTTTCAGATTTAGAGAATGCGAATGTTGTTTATACCCTTAAA  
 40 GATGGTGGAAACAGCTGTTGCCTCAGTTTCATTGACAAAACAACACCAAATGGCGAAATCAACTTAGGTAAT  
 GGTATTAATTTACAGTTACTGGAGCGTTTGCTGGTAAATTCAGTGGTCTGACTGATAGTAAAACATACATG  
 ATCTCAGAACGTATCGCTGGTTATGGTAATACAATCACTACTGGTGCCTGGTAGTGCAGCTATCACCAATACT  
 CCAGATTCAGACAACCCAACCACTTAATCCAACCTGAACCAAAGTTGTGACACACGGTAAAAAATTCGTC  
 AAAACAAGTTCGACTGAAACAGAACGCTTGCAAGGTGCACAGTTCGTTGTTAAAGATTTCAGCTGGTAAATAC  
 45 CTTGCATTGAAATCATCTGCGACAATATCAGCTCAAACAACAGCTTACACAAATGCTAAAACCTGCTCTTGAC  
 GCTAAAATCGCAGCTTACAACAACTTTCAGCAGACGATCAAAAAGGTACTAAAGGTGAAACAGCTAAAGCA  
 GAAATCAAACTGCTCAAGACGCTTACAATGCAGCCTTCATCGTAGCTCGTACAGCTTACGAGTGGGTAAC  
 AATAAAGAAGATGCTAACGTTGTTAAAGTGACTTCAAACGCTGACGGTCAATTTGAAGTTAGCGGTCTTGCA  
 ACTGGTGAATTATAAACTTGAAGAAACACAAGCTCCAGCTGGTTACGCTAAATTAGCAGGTGATGTTGATTT  
 50 AAAGTTGGAAACAGCTCAAAGCAGACGACTCAGGTAACATTGATTACACTGCTAGCAGCAATAAAAAAGAC  
 GCTCAACGCATAGAAAACAAAAAAGTGACTATTCCACAAACAGGTGGTATTGGTACAATCTTTTTCACAATT  
 ATTGTTTAAGCATTATGCTTGGAGCGGTAATTATCATGAAAAGACGTCAATCAGAGGAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 113):

MKRINKYFAMFSALLLILTSLLSVAPVFAEMGNITKTVTLHKIVQTSNLAKPFPNGINGLNGTKYMGQKL  
 55 TDISGYFGQSKEIAGAFFAVMNESQTKYITESGTEVESIDAAGVLKGLTTENGITFNTANLKGTYQIVVELL  
 DKSNYKNGDKVLADSKAVPVKITLPLYNEEGIVVDAEVYPKNTTEAPQIDKNFAKANKLLNDSNSAIAGGA  
 DYDKYQAEKAKATAEIGQEIPEYVTKIKQKSKYKNLAWVDTMSNGLTMGNTVNLEASSGSFVEGTDYNVER  
 DDRGFTLKFDTGLTKLQKEAETQAVEFTLTYSATVNGAAIDDKPESNDIKLQYGNKPGKKVKEIPVTPSNG  
 EITVSKTWDKGSDELNANVVYTLKDGGTAVASVSLTKTTPNGEINLNGIKFTVTGAFAGKFSGLTDSKTYM  
 60 ISERIAGYGNITTTGAGSAAITNTPDSDNPTPLNPTEPKVVTHGKKFVKTSSTETERLQGAQFVVKDSAGKY



LALKSSATISAQTTAYTNAKTALDAKIAAYNKLSADDQKGTKGETAKAEIKTAQDAYNAAFIVARTAYEWVT  
 NKEDANVVKVTSNADGQFEVSGLATGDYKLEETQAPAGYAKLAGDVDFKVGNSKADDSGNIDYTASSNKKD  
 AQRIENKQVTIPQTGGIGTILFTIIGLSIMLGAVIIMKRRQSEEA

5 **Strain IC291**

ORF DNA sequence (SEQ ID NO: 156):

ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCACTATTCTCAGTT  
 GCACCAGCGTTTGC GGACGACGCAACAACCTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 10 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCCTTCTTTGTTTTCAAAAATGAAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTCTGTT  
 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 GAATTGAAAGAAAAATCAAACCTACGATAACAACGGTCTATCTTGGCTGATTCAAAGCAGTTCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 15 AAACCACAAGTAGATAAGAACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATCTTAAAGGCTCAGACTATAAG  
 AAAGTTGGTTTGGACTGATAGCATGACTAAAGGTTTGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 GAAGATTTTCCCTGTTTTAACTACAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 20 AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAGTCAATAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAACGGATGGTACATGG  
 GTGAACGTTGCTTACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTCACAGGTTTGGATAATGCT  
 AAACTTACCGCGTTGTCGAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 25 ACTATCAAGAACAACAAAACCTCAAATGATCCAACCTCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 30 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 ACTGGTTTGGATAAAGGCACTTATGGCTTGGAAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAAAACAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATTTCTTTTACAATTATTGGTTTAAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 35 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 114):

MKKINKCLTMFSTLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGKTKGNDSYVVGKQINDL  
 KSYFGSTDAKEIKGAFFVFKNETGTFKITENGKEVDLLEAKDAEGGAVLSGLTKDNGFVFNTAKLKGIIYQIV  
 ELKEKSNYDNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTETKPQVDKNFADKDLDTDNRKDKGV  
 40 VSATVGDKEYIVGTKILKGSYKLVWTD SMTKGLTFNNNVKVTLDGEDFPVLNYKLVTD DQGFRLALNAT  
 GLAAVAAA AKDKDVEIKITYSATVNGSTTVEI PETNDVKLDYGNPTEESEPEQEGTPANQEIKVIKDWAVDG  
 TITDANVAVKAI FTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTP EYVSFKNGVV  
 TIKNNKNSNDPTPINPSEPKVVTYGRKFVKTN QANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 45 TGLDKGTYGLEETQAPAGYATLSGDVNFVETATSYSKGATTDIAYDKGSVKKDAQQVQNKKVTIPQTGGIGT  
 ILFTIIGLSIMLGAVVIMKRRQSEEA

50 **Strain IC304**

ORF DNA sequence (SEQ ID NO: 157):

ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCACTATTCTCAGTT  
 GCACCAGCGTTTGC GGACGACGCAACAACCTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 50 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCCTTCTTTGTTTTCAAAAATGAAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTCTGTT  
 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 55 GAATTGAAAGAAAAATCAAACCTACGATAACAACGGTCTATCTTGGCTGATTCAAAGCAGTTCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGAACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATCTTAAAGGCTCAGACTATAAG  
 60 AAAGTTGGTTTGGACTGATAGCATGACTAAAGGTTTGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT



GAAGATTTTCCTGTTTTAAACTACAAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAATTCCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAAGTCATTAAAGACTGGGCAGTAGATGGT  
 5 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAAACGGATGGTACATGG  
 GTGAACGTTGCTTCACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTTACAGGTTTGGATAATGCT  
 AAAACTTACCGCGTTGTTCGAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAAACCTCAAATGATCCAACCTCCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
 10 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 ACTGGTTTGGATAAAGGCACTTATGGCTTGGAAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 15 GATGTAAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAAAACAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATCTTTTTCACAATTATTGGTTTAAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 115):

MKKINKCLTMFSTLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGTKGKNDSDYVGKQINDL  
 KSYFGSTDAKEIKGAFFVFKNETGTFKFI TENGKEVDLEAKDAEGGAVLSGLTKDNGFVFNTAKLKIYQIV  
 ELKEKSNYDNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNETETKPVQVDKNFADKDLDDYTDNRKDKGV  
 VSATVGDKEYIVGTKILKGSYKLVWTDMSMTKGLTFNNNVKVTLDGEDFPVLNYKLVTDQGFRLALNAT  
 GLAAVAAAAKDKDVEIKITYSATVNGSTTVEI PETNDVKLDYGNPTEESEPEQEGTPANQEI KVIKDWAVDG  
 25 TITDANVAVKAI FTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPYVVSFKNGVV  
 TIKNNKNSNDPTPINPSEPKVVVYGRKFVKTNQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYGLEETQAPAGYATLSGDVNFVETATSYSKGATTDIAYDKGSVKKDAQQVQNKKVTI PQTGGIGT  
 ILFTIIGLSIMLGAVVIMKKRQSEEA

### Strain IC305

ORF DNA sequence (SEQ ID NO: 158):

ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCACTATTCTCAGTT  
 GCACCAGCGTTTGCAGGACGACGCAACAACTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 35 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCCTTCTTTGTTTTCAAAAATGAAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTGTT  
 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 GAATTGAAAGAAAAATCAAACCTACGATAACAACGGTCTATCTTGGCTGATTCAAAGCAGTTCCAGTTAAA  
 40 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGAATTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATCTTAAAGGCTCAGACTATAAG  
 AAACCTGGTTTGGACTGATAGCATGACTAAAGGTTTGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 GAAGATTTTCCTGTTTTAAACTACAAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 45 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAATTCCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAAGTCATTAAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAAACGGATGGTACATGG  
 GTGAACGTTGCTTCACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTTACAGGTTTGGATAATGCT  
 50 AAAACTTACCGCGTTGTTCGAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAAACCTCAAATGATCCAACCTCCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 55 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 ACTGGTTTGGATAAAGGCACTTATGGCTTGGAAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAAAACAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA



ATTCTTTTCACAATTATTGGTTTAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 116):

5 MKKINKCLTMFSTLLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGKTKGNDSYVVGKQINDL  
KSYFGSTDAKEIKGAFFVFKNETGTFKITENGKEVDLEAKDAEGGAVLSGLTKDNGFVFNTAKLKGIIYQIV  
ELKEKSNDNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTETKPQVDKNFADKDLDDYTDNRKDKGV  
VSATVGDKKEYIVGTKILKGSYKLVWTDGMTKGLTFNNNVKVTLDGEDFPVLNYKLVTDQGFRLALNAT  
10 GLAAVAAAADKDKDVEIKITYSATVNGSTTVEIPETNDVKLDYGNPTEESEPEQEGTPANQEIKVIKDWAVDG  
TITDANVAVKAIFFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPYVVSFKNGVV  
TIKNNKNSNDPTPINPSEPKVVVYGRKFVKTNQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
TGLDKGTYGLEETQAPAGYATLSGDVNFVETATSYSGGATTDIAYDKGSVKKDAQQVQNKKVTIPQTGGIGT  
ILFTIIIGLSIMLGAVVIMKKRQSEEA

15 **Strain IC306**

ORF DNA sequence (SEQ ID NO: 159):

ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCACACTATTCTCAGTT  
GCACCAGCGTTTGCAGGACGACGCAACAACTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
20 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
AAATCCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCCTTCTTTGTTTTCAAAAATGAAACTGGT  
ACAAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTCTGT  
CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
GAATTGAAAGAAAAATCAAACACTACGATAACAACGGTCTATCTTGGCTGATTCAAAGCAGTTCCAGTTAAA  
25 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
AAACCACAAGTAGATAAGAACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATCTTAAAGGCTCAGACTATAAG  
AACTGGTTTTGGACTGATAGCATGACTAAAGGTTTGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
GAAGATTTTCCCTGTTTTAACTACAAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
30 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAGTCATTAAGACTGGGCAGTAGATGGT  
ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAAACGGATGGTACATGG  
GTGAACGTTGCTTACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTACAGGTTTGGATAATGCT  
35 AAACTTACCGCGTTGTCGAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
ACTATCAAGAACAACAAAACTCAAATGATCCAACCTCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
40 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
ACTGGTTTTGGATAAAGGCACTTATGGCTTGGAAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
TCTGTAAAAAAGATGCCCAACAAGTTCAAAAACAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
45 ATCTTTTTCACAATTATTGGTTTAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 117):

50 MKKINKCLTMFSTLLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGKTKGNDSYVVGKQINDL  
KSYFGSTDAKEIKGAFFVFKNETGTFKITENGKEVDLEAKDAEGGAVLSGLTKDNGFVFNTAKLKGIIYQIV  
ELKEKSNDNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTETKPQVDKNFADKDLDDYTDNRKDKGV  
VSATVGDKKEYIVGTKILKGSYKLVWTDGMTKGLTFNNNVKVTLDGEDFPVLNYKLVTDQGFRLALNAT  
GLAAVAAAADKDKDVEIKITYSATVNGSTTVEIPETNDVKLDYGNPTEESEPEQEGTPANQEIKVIKDWAVDG  
TITDANVAVKAIFFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPYVVSFKNGVV  
TIKNNKNSNDPTPINPSEPKVVVYGRKFVKTNQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
55 TGLDKGTYGLEETQAPAGYATLSGDVNFVETATSYSGGATTDIAYDKGSVKKDAQQVQNKKVTIPQTGGIGT  
ILFTIIIGLSIMLGAVVIMKKRQSEEA

**Strain IC361**

ORF DNA sequence (SEQ ID NO: 160):



ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCACTATTCTCAGTT  
 GCACCAGCGTTTGC GGACGACGCAACAACTGATACTGTGACCTTG CACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 5 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCTTTCTTTGTTTTCAAAAATGAAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTCTGTT  
 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 GAATTGAAAGAAAAATCAAACTACGATAACAACGGTTCCTATCTTGGCTGATTCAAAGCAGTTCCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 10 AAACCACAAGTAGATAAGAACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATCTTAAAGGCTCAGACTATAAG  
 AAAGTGGTTTGGACTGATAGCATGACTAAAGGTTTGGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 GAAGATTTTCTGTTTTAACTACAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 15 AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAGTCACTTAAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAAACGGATGGTACATGG  
 GTGAACGTTGCTTACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTTACAGGTTTGGATAATGCT  
 AAAACTTACCGCGTTGTCGAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAAACCTCAATGATCCAACCTCAATCAACCCATCAGAACCAAAAAGTGGTGACTTAT  
 20 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 25 ACTGGTTTGGATAAAGGCCTTATGGCTTGGAAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAAAACAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATTCTTTTCACAATTATTGGTTTAAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA  
 30 ORF amino acid sequence (SEQ ID NO: 118):  
 MKKINKCLTMFSTLLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGTKGKNDSYVVGKQINDL  
 KSYFGSTDAKEIKGAFFVFKNETGTFKITENGKEVDLEAKDAEGGAVLSGLTKDNGFVFNTAKLKIYQIV  
 ELKEKSNDNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTETKQPVDKNFADKDLDDYTDNRKDKGV  
 VSATVGDKEYIVGTKILKGSYKKLVWTDGMTKGLTFNNNVKVTLDGEDFPVLNYKLVTDQGFRLALNAT  
 35 GLAAVAAAAKDKDVEIKITYSATVNGSTTVEIPETNDVKLDYGNPTEESEPEQEGTPANQEIKVIKDWAVDG  
 TITDANVAVKAIFFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPYVVSFKNGVV  
 TIKNNKNSNDPTPINPSEPKVVVYGRKFVKTNQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTA  
 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYGLEETQAPAGYATLSGDVNFVETATSYSGATTDIAYDKGSVKKDAQVQNKKVTIPQTGGIGT  
 40 ILFTIIGLSIMLGAVVIMKKRQSEEA

**Strain IC363**  
ORF DNA sequence (SEQ ID NO: 161):  
 ATGAAAAGAATCAACAAATATTTTGAATGTTCTCGGCATTGTTACTGACTTTAACGTCATTGCTCTCAGTT  
 45 GCACCAGCGTTTGC GGATGAAGCAACAACTAATACAGTGACTTTGCACAAGATTTTGCAAACCGAATCAAAT  
 CTTAACAAAAGTAACTTCCCAGGAACTACAGGTCTTAACGGAAAAGACTACAAAGGTGGAGCTATTTCTGAC  
 CTTGCTGGTTACTTTGGCGAGGGATCTAAAGAAATCGAAGGTGCGTTCTTTGCTTTAGCTTTGAAAGAAGAT  
 AAAAGTGGTAAAGTGCAATATGTTAAGGCAAAAAGAAGGTAACAAATTAACACCAGCCTTAATTAATAAAGAT  
 50 GGTACTCCTGAAATAACAGTAAATATTGATGAGGCCGTGCTGGATTGACACCAGAGGGAGATACTGGACTT  
 GTTTTCAACACCAAAGGATTGAAAGGCGAGTTTAAAATTGTTGAAGTTAAATCAAATCTACTTACAACAAT  
 AATGGTTCCTCCTGGCTGCTTCAAAGCGGTTCCAGTTAACATCACTCTTCCATTGGTAAATGAAGATGGT  
 GTTGTGCTGATGCCCATGTTTATCCAAAGAACACTGAAGAAAACCAGAAATTGATAAAAACCTTTGCTAAA  
 ACAAACGATTTGACAGCATTGACAGATGTTAATAGACTTTTGACAGCTGGCGCAAATTTATGGTAATTATGCA  
 55 CGTGACAAAGCAACTGCTACTGCTGAAATCGGTAAAGTTGTTCCCTTATGAAGTTAAAACAAAATTCACAAA  
 GGTTCTAAATACGAAAACCTGGTTTGGACAGATATAATGTCAAATGGTTTGGACAATGGGTTCAACTGTTAGC  
 CTTAAAGCTTCAGGAACTACAGAACTTTTGCTAAGGATACAGACTATGAAGTTAGCATTGATGCCCGTGGT  
 TTCACATTAATAATTCACAGCTGATGGATTGGGCAAATTTGGAAAAGCAGCTAAAACAGCTGATATTGAATTT  
 ACATTGACTTATAGTGCTACTGTTAATGGTCAAGCAATTATTGATAATCCAGAATCCAATGATATCAAATTG  
 60 TCGTATGGTAACAAACCAGGTAAAGACTTGACTGAACTTCCGTGTTACACCTTCAAAGGGTGAAGTAACAGTT  
 GCTAAAACCTGGTCTGACGGAATTGCACCTGATGGTGTAAACGTTGTTTACACATTGAAAGATAAAGATAAA



ACTGTTGCTTCAGTATCATTGACAAAAACATCTAAAGGTACAATCGACCTTGGAAATGGTATCAAATTTGAA  
 GTATCTGGTAACTTCTCGGGTAAATTCAGTGGTCTAGAAAAACAAATCATAACATGATCTCAGAACGTGTTTCT  
 GGTTACGGAAGTGCAATAAATCTAGAAAAATGGTAAAGTAACCATTACCAATACCAAAGATTCTGATAACCCA  
 ACACCATTTGAACCCAACCTGAACCAAAAAGTTGAAACTCATGGTAAGAAATTTGTCAAAACTAATGAACAAGGT  
 5 GACCGTTTGGCTGGTGCACAATTCGTTGTGAAAACTCAGCAGGTAAATACCTTGCTCTTAAAGCAGATCAA  
 TCAGAAGGTCAAAAAACTTTAGCTGCTAAGAAAATAGCTTTAGATGAAGCTATCGCTGCTTATAACAAGTTG  
 TCTGCAACAGACCAAAAAGGTGAAAAAGGAATTACTGCAAAAGAACTTATCAAAACTAAACAAGCAGATTAC  
 GATGCAGCCTTCATTGAGGCTCGTACAGCTTATGAGTGGATAACAGATAAGGCTAGAGCCATTACCTACACT  
 TCAAACGATCAAGGTCAATTTGAAGTTACAGGTCTTGCAGACGGTACTTACAACCTTGAAGAAACACTTGCT  
 10 CCAGCAGGATTTGCTAAGTTGGCAGGTAATATTAAGTTTGTAGTTAATCAAGGGTCATACATAACAGGTGGT  
 AACATTGACTACGTTGCTAACAGCAACCAAAAAGATGCGACACGTGTAGAAAATAAAAAGGTAACAATCCCA  
 CAAACAGGTGGTATTGGTACAATCTTTTTCACAATTATTGGTTTAAAGCATTATGCTTGGAGCAGTAGTTATC  
 ATGAAAAGACGCCAATCAAAGGAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 119):

15 MKRINKYFAMFSALLLTLTSLLSVAPAFADDEATTNTVTLHKILQTESNLNKS NFPGTTGLNGKDYKGGAI SD  
 LAGYFGEKSKEIEGAFFALALKEDKSGKVQYVKAKEGNKLT PALINKDGTPEITVNI DEAVSGLTPEGDTGL  
 VFNTKGLKGEFKIVEVKSSTYNNNGSLLAASKAVPVNITLPLV NEDGVVADAHVYPKNTEEKPEIDKNFAK  
 TNDLTALTDVNRLLTAGANYGN YARDKATATAEIGKVVPYEVKTKIHKGSKYENLVWTDIMSNGLTMGSTVS  
 20 LKASGTTETFAKDDYELSIDARGFTLKF TADGLGKLEKAAKTADIEFTLTYSATVNGQAIIDNPESNDIKL  
 SYGNKPGKDLTELPVTPSKGEVTVAKTWS DGIAPDGVNVVYTLKDKDKTVASVSLTKT SKGTIDLNGIKFE  
 VSGNFSGKFTGLENKSYMISERVSGYGSAINLENGKVTITNTK DSDNPTPLNPTEPKVETHGKKFVKTNEQG  
 DRLAGAQFVVKNSAGKYLALKADQSEGQKTLAAKKIALDEAIAAYNKLSATDQKGEKGITAKELIKTKQADY  
 DAAFI EARTAYEWITDKARAITYTSNDQGFVETGLADGTYNLEETLAPAGFAKLAGNIK FVVNQGSYITGG  
 25 NIDYVANSNQKDATRVENKKVTIPQTGGIGTILFTIIGLSIMLGAVVIMKRRQSKEA

### Strain IC365

ORF DNA sequence (SEQ ID NO: 162):

ATGAAAAAATCAACAAATATTTTGCAGTCTTCTCGGCATTGCTACTGACCGTAACATCATTGTTCTCAGTT  
 GCACCAGTGTTTGC GGAAGAAGCAAAA ACTACTGACACAGTGACCTTGCACAAGATTGTCATGCCTCGAACT  
 30 GCATTTGACGGTTTTACTGCTGGTACAAAGGGTAAGGATAATACTGACTACGTTGGTAAACAAATCGAAGAC  
 CTTAAACTTACTTTGGCTCAGGCGAAGCGAAAGAAATCGCAGGTGCTTACTTTGCTTTCAAAAATGAAGCT  
 GGTACTAAATACATCACTGAAAATGGTGAAGAAGTTGATACTTTGGATAACAACAGATGCCAAAGGTTGTGCT  
 GTTCTTAAAGGTTTAAACAACAGACAATGGTTTCAAATTTAACACTTCTAAATTAACAGGAACTTACCAAATC  
 GTTGAATTGAAAGAAAAATCTACATAACAACGATGGTTCTATCTTGGCTGATTCAAAGCAGTTCAGTT  
 35 AAAATCACTCTTCCATTGGTAAACGACAATGGTGTGTTAAAGACGCTCACGTTTATCCAAAGAACACTGAA  
 ACAAACCACAAGTAGATAAGA ACTTTCGAGATAAAGA ACTTGATTATGCGAACAACAAAAAAGACAAAGGG  
 ACTGTCTCAGCATCTGTTGGT GATGTTAAAAAATACCATGTTGGAACAAAAATCCTTAAAGGTT CAGACTAT  
 AAGAAATTAATCTGGACCGATAGCATGACCAAAGGTTTGACTTTCAACAACGATATTGCTGTAACATTGGAT  
 GGTGCAACTCTTGATGCTACAAATTACAAACTTGTAGCAGATGACCAAGGTTTCCGCCTTGTCTTGACTGAC  
 40 AAAGGTCTTGAAGCAGTGGCAAAAAGCCGCAAAAACAAAAGATGTTGAAATCAAGATCACTTACTCAGCTACT  
 TTGAACGGTTCTGCTGTCGTTGAAGTTCTAGAAACCAATGATGTTAAATTGGACTACGGCAACAACCCAACA  
 ATTGAAATGAACCAAAAAGAAGGTATTCCAGTTGATAAGAAAATCACTGTTAACAAAACATGGGCAGTAGAT  
 GGCAATGAAGTGAATAAAGCAGATGAAACAGTTGATGCTGTCTTACCTTGCAAGTTAAAGATGGTGACAAA  
 TGGGTGAATGTTGATTCAGCTAAAGCAACAGCTGCAACTAGCTTCAAACACACTTTTGAAACTTGGATAAT  
 45 GCTAAACTTACCGCGTTATCGAACGTGTTAGCGGCTACGCTCCAGAATACGTCTCATTGTAAATGGCGTT  
 GTAACCATCAAGAACAACAAGACTCAAATGAGCCA ACTCCAATCAACCCATCAGAACC AAAAGTGGTGACT  
 TATGGACGTAAATTTGTGAAAACAAATAAAGATGGAAAAGAACGCTTGGCAGGAGCTACCTTCCTTGTTAAG  
 AAAGATGGCAAGTACTTGGCACGTAAATCAGGTGTTGCAACAGATGCAGAAAAAGCTGCTGTAGATTCAACT  
 AAATCAGCATTGGATGCTGCTGTTAAAGCTTACAATGATTTGACTAAAGAAAAACAAGAAGGTCAAGATGGT  
 50 AAATCAGCATTGGCTACCGTTAGTGAAAAACAAAAGCTTACAATGATGCCTTTGTAAAGCTAACTACTCA  
 TACGAATGGGTTGAAGATAAAAATGCTAAGAATGTTGTTAAATTGATTTCTAACGATAAAGGTCAATTTGAA  
 ATTACTGGCTTGACTGAAGGTCAATACTCATTGGAAGAAACACAAGCACC AACTGGTTATGCTAAATTATCA  
 GGTGATGTTTCGTTTAAATGTTAATGCTACTTCATACAGTAAAGGTTCTGCTCAAGATATTGAGTATACCCAA  
 GGTTCTAAACTAAAGATGCACAACAAGTTATCAATAAGAAGGTTACTATTCCACAACAGGTGGTATTGGT  
 55 ACAATTTTTTTTACAATTATTGGATTAAGTATTATGCTTGGAGCGGTAGTTATCATGAAAAGACGTCAATCA  
 GAGGAAGTTTAA

ORF amino acid sequence (SEQ ID NO: 120):

60 MKKINKYFAVFSALLLTVTSLFSVAPVFAEEAKTTDTVTLHKIVMPRTAFDGF TAGTKGKDNTDYVGKQIED  
 LKTYFGSGEAKEIAGAYFAFKNEAGTKYITENGEVDLDTTDAKGC AVLKGLTTDNGFKFN TSKLTGTYQI  
 VELKEKSTYNN DGSILADSKAVPVKITLPLVNDNGVVKDAHVYPKNTE TPKQVDKNFADKELDYANNKKDKG



TVSASVGDVKKYHVGTKILKGSYKLIWTD SMTKGLTFNNDIAVTL DGATLDATNYKLVADDQGFRLVLT D  
KGLEAVAKAAKTKDVEIKITYSATLNGSAVVEVLETNDVKLDYGNPTIENEPKEGIPVDKKITV NKTWAVD  
GNEVNKADETVDAVFTLQVKDGDKWNVD SAKATAATSFKHTFENLDNAKTYRVIERSGYAPEYVSFVNGV  
VTIKNNKDSNEPTPINPSEPKVVTYGRKFVKT NKDGKERLAGATFLVKKDGKYLARKSGVATDAEKA AVDST  
5 KSALDAAVKAYNDLTKEKQEGQDGKSALATVSEKQKAYNDAFVKANYSYEWVEDKNAKNVVKLISNDK GQFE  
ITGLTEGQYSLEETQAPTGYAKLSGDVSNFN NATSYSKGSAQDIEYTGQSKTKDAQQVINKKVTIPQTGGIG  
TIFFTIIGLSIMLGAVVIMKRRQSEEV

**Strain IC367**

10 ORF DNA sequence (SEQ ID NO: 163):  
ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTC ACTATTCTCAGTT  
GCACCAGCGTTTGC GGACGACGCAACA ACTGATACTGTGACCTTG CACAAGATTGTCATGCCACAAGCTGCA  
TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCTTTCTTTGTTTTCAAAAATGAACTGGT  
15 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTCTGTT  
CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
GAATTGAAAGAAAAATCAA CTACGATAACAACGGTTCATCTTGGCTGATTCAAAGCAGTTC CAGTTAAA  
ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
AAACCACAAGTAGATAAGA ACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
20 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATCTTAAAGGCTCAGACTATAAG  
AACTGGTTTTGACTGATAGCATGACTAAAGGTTTGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
GAAGATTTTCTGTTTTTAACTACAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
GGTCTTGCAGCAGTAGCAGCAGCTGCAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
25 GAAAGTGAACCACAAGAAGTACTCCAGCTAACCAAGAAATTAAGTCAATAAGACTGGGCAGTAGATGGT  
ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAACGGATGGTACATGG  
GTGAACGTTGCTTCACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTCACAGGTTTGGATAATGCT  
AAAACTTACCGCGTTGTGCAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
ACTATCAAGAACAACAAA ACTCAAATGATCCAACCTCAATCAACCCATCAGAACCAAAGTGGTGACTTAT  
30 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
35 ACTGGTTTGGATAAAGGCACTTATGGCTTGGAAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAAC TGACATCGCATATGATAAAGGC  
TCTGTAAAAAAGATGCCCAACAAGTTCAAAAACAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
ATTCTTTTACAATTATTGGTTTAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
GAAGCTTAA

40 ORF amino acid sequence (SEQ ID NO: 121):  
MKKINKCLTMFSTLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGTKGKNDS DYVGKQINDL  
KSYFGSTDAKEIKGAFFVFKNETGTKFITENGKEVD TLEAKDAEGGAVLSGLTKDNGFVFNTAKLKG IYQIV  
ELKEKSNYDNNGSILADSKAVPVKITLPLVNNQGVV KDAHIYPKNETETKPQVDKNFADKDL DYT DNRKDKGV  
VSATVGDKKEYIVGTKILKGSYKLVWTD SMTKGLTFNNDIAVTL DGEDFPVLNYKLVTD DQGFRLALNAT  
45 GLAAVAAA AKDKDVEIKITYSATVNGSTTVEI PETNDVKLDYGNPTIENEPKEGIPVDKKITV NKTWAVD  
TITDANVAVKAI FTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVERVSGYTP EYVSFKNGVV  
TIKNNKNSNDPTPINPSEPKVVTYGRKFVKT NQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVK TAK  
LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
TGLDKGTYGLEETQAPAGYATLSGDVNF EVTATSYSKGATTDIAYDKGSVKKDAQQVQNKKVTIPQTGGIGT  
50 ILFTIIGLSIMLGAVVIMKRRQSEEA

**Strain IC377**

ORF DNA sequence (SEQ ID NO: 164):  
ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTC ACTATTCTCAGTT  
55 GCACCAGCGTTTGC GGACGACGCAACA ACTGATACTGTGACCTTG CACAAGATTGTCATGCCACAAGCTGCA  
TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCTTTCTTTGTTTTCAAAAATGAACTGGT  
ACAAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTCTGTT  
CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
60 GAATTGAAAGAAAAATCAA CTACGATAACAACGGTTCATCTTGGCTGATTCAAAGCAGTTC CAGTTAAA



ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGAACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATTCTTAAAGGCTCAGACTATAAG  
 5 AAAGTGGTTTGGACTGATAGCATGACTAAAGGTTTGGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 GAAGATTTTCCCTGTTTTAACTACAAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAATTCCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAAGTCATTAAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAAACGGATGGTACATGG  
 10 GTGAACGTTGCTTACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTCACAGGTTTGGATAATGCT  
 AAACTTACCGCGTTGTGCAACGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAAACCTCAAATGATCCAACCTCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 15 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 ACTGGTTTGGATAAAGGCACTTATGGCTTGGAAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 20 TCTGTAAAAAAGATGCCCAACAAGTTCAAACAAAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATCTTTTTCACAATTATTGGTTTAAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 122):

MKKINKCLTMFSTLLILLTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGTKGKNDSDYVGKQINDL  
 25 KSYFGSTDAKEIKGAFFVFKNETGTFITENGKEVDLEAKDAEGGAVLSGLTKDNGFVFNTAKLKIYQIV  
 ELKEKSNYDNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTETKPVQVDKNFADKDLDDYTDNRKDKGV  
 VSATVGDKEYIVGTKILKGSYKLVWTDMSMTKGLTFNNNVKVTLDGEDFPVLNYKLVTDDQGFRLALNAT  
 GLAAVAAAADKDKDVEIKITYSATVNGSTTVEIPETNDVKLDYGNPTEESEPEQEGTPANQEIKVIKDWAVDG  
 TITDANVAVKAIFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPYVVSFKNGVV  
 30 TIKNNKNSNDPTPINPSEPKVVVYGRKFKTNQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYGLEETQAPAGYATLSGDVNFVETATSYSKGATTDIAYDKGSVKKDAQQVQNKKVTIPQTGGIGT  
 ILFTIIGLSIMLGAVVIMKKRQSEEA

### 35 Strain IC379

ORF DNA sequence (SEQ ID NO: 165):

ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCACTATTCTCAGTT  
 GCACCAGCGTTTGCAGGACGACGCAACAACTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 40 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCCTTCTTTGTTTTCAAAAATGAAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTGTT  
 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 GAATTGAAAGAAAAATCAAACCTACGATAACAACGGTTCATCTTGGCTGATTCAAAGCAGTTCCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 45 AAACCACAAGTAGATAAGAACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATTCTTAAAGGCTCAGACTATAAG  
 AAAGTGGTTTGGACTGATAGCATGACTAAAGGTTTGGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 GAAGATTTTCCCTGTTTTAACTACAAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 50 AACGGCTCCACTACTGTTGAAATTCCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAAGTCATTAAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAAACGGATGGTACATGG  
 GTGAACGTTGCTTACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTCACAGGTTTGGATAATGCT  
 AAACTTACCGCGTTGTGCAACGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 55 ACTATCAAGAACAACAAAACCTCAAATGATCCAACCTCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 60 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT



- 101 -

ACTGGTTTGGATAAAGGCACTTATGGCTTGGGAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAAAACAAAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATTCTTTTTCACAATTATTGGTTTAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 5 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 123):

MKKINKCLTMFSTLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGTEGKGNDSYVVGKQINDL  
 KSYFGSTDAKEIKGAFFVFKNETGTFKITENGKEVDLEAKDAEGGAVLSGLTKDNGFVFNTAKLKGIIYQIV  
 10 ELKEKSNDNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTETKPQVDKNFADKDLDTDNRKDKGV  
 VSATVGDKKEYIVGTKILKGSYKLVWTDGMTKGLTFNNNVKVTLDGEDFPVLNYKLVTDQGFRLALNAT  
 GLAAVAAAADKDKDVEIKITYSATVNGSTTVEIPETNDVKLDYGNPTEESEPEQEGTPANQEIKVIKDWAVDG  
 TITDANVAVKAIFFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPHYVSFKNGVV  
 TIKNNKNSNDPTPINPSEPKVVVYGRKFVKTNQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
 15 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYGLEETQAPAGYATLSGDVNFVETATSYSKGATTDIAYDKGSVKKDAQQVQNKVTIIPQTGGIGT  
 ILFTIIGLSIMLGAVVIMKKRQSEEA

### Strain IC432

ORF DNA sequence (SEQ ID NO: 166):

ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCCTATTCTCAGTT  
 GCACCAGCGTTTGCAGGACGACGCAACAACTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCTTTCTTTGTTTTCAAAAATGAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTCTGT  
 25 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 GAATTGAAAGAAAAATCAAACTACGATAACAACGGTTCCTATCTTGGCTGATTCAAAGCAGTTCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGAACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATCTTAAAGGCTCAGACTATAAG  
 30 AAAGTGGTTTGGACTGATAGCATGACTAAAGGTTTGGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 GAAGATTTTCTGTTTTAACTACAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAGTCACTTAAAGACTGGGCAGTAGATGGT  
 35 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAACGGATGGTACATGG  
 GTGAACGTTGCTTACACGAAGCAACAACCAATCACGCTTTGAACATACTTTTACAGGTTTGGATAATGCT  
 AAACTTACCGCGTTGTCGAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAAACTCAAATGATCCAACCTCCAATCAACCCATCAGAACCAAAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
 40 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 ACTGGTTTGGATAAAGGCACTTATGGCTTGGGAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 45 GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAAAACAAAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATTCTTTTTCACAATTATTGGTTTAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 124):

MKKINKCLTMFSTLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGTEGKGNDSYVVGKQINDL  
 KSYFGSTDAKEIKGAFFVFKNETGTFKITENGKEVDLEAKDAEGGAVLSGLTKDNGFVFNTAKLKGIIYQIV  
 ELKEKSNDNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTETKPQVDKNFADKDLDTDNRKDKGV  
 VSATVGDKKEYIVGTKILKGSYKLVWTDGMTKGLTFNNNVKVTLDGEDFPVLNYKLVTDQGFRLALNAT  
 GLAAVAAAADKDKDVEIKITYSATVNGSTTVEIPETNDVKLDYGNPTEESEPEQEGTPANQEIKVIKDWAVDG  
 55 TITDANVAVKAIFFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPHYVSFKNGVV  
 TIKNNKNSNDPTPINPSEPKVVVYGRKFVKTNQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYGLEETQAPAGYATLSGDVNFVETATSYSKGATTDIAYDKGSVKKDAQQVQNKVTIIPQTGGIGT  
 ILFTIIGLSIMLGAVVIMKKRQSEEA

60



**Strain IC455**

ORF DNA sequence (SEQ ID NO: 167):

5 ATGAAAAGAATCAACAAATATTTTGC AATGTTCTCGGCATTGTTACTGACTTTAACGTCATTGCTCTCAGTT  
 GCACCAGCGTTTGC GGATGAAGCAACA ACTAATACAGTGACTTTGCACAAGATTTTGCAAACCGAATCAAAT  
 CTTAACAAAAGTAACTTCCCAGGAACTACAGGTCTTAACGGAAAAGACTACAAAGGTGGAGCTATTTCTGAC  
 CTTGCTGGTTACTTTGGCGAGGGATCTAAAGAAATCGAAGGTGCGTTCTTTGCTTTAGCTTTGAAAGAAGAT  
 AAAAGTGGTAAAGTGCAATATGTTAAGGCAAAAAGAAGGTAACAAATTAACACCAGCCTTAATTAATAAAGAT  
 GGTACTCCTGAAATAACAGTAAATATTGATGAGGCCGTGTCTGGATTGACACCAGAGGGAGATACTGGACTT  
 10 GTTTTCAACACCAAAGGATTGAAAGGCGAGTTTAAAATTGTTGAAGTTAAATCAAAATCTACTTACAACAAT  
 AATGGTTCCTCCTGGCTGCTTCAAAGCGGTTCCAGTTAACATCACTCTTCCATTGGTAAATGAAGATGGT  
 GTTGTGCTGATGCCCATGTTTATCCAAAGAACACTGAAGAAAAACCAGAAATTGATAAAAACTTTGCTAAA  
 ACAAACGATTTGACAGCATTGACAGATGTTAATAGACTTTTGACAGCTGGCGCAAATTATGGTAATTATGCA  
 CGTGACAAAGCAACTGCTACTGCTGAAATCGGTAAAGTTGTTCCATTATGAAGTTAAAACAAAAATTCACAAA  
 15 GGTTCTAAATACGAAA ACTTGGTTTGGACAGATATAATGTCAAATGGTTT GACAATGGGTTCAACTGTTAGC  
 CTTAAAGCTTCAGGAACTACAGAACTTTTGCTAAGGATACAGACTATGAACTTAGCATTGATGCCCGTGGT  
 TTCACATTA AAATTCACAGCTGATGGATTGGGCAAATTGGAAAAAGCAGCTAAAACAGCTGATATTGAATTT  
 ACATTGACTTATAGTGCTACTGTTAATGGTCAAGCAATTATTGATAATCCAGAATCCAATGATATCAAATTG  
 TCGTATGGTAACAAACCAGGTAAAGACTTGACTGAACTTCTGTACACCTTCAAAGGGTGAAGTAACAGTT  
 20 GCTAAA ACTTGGTCTGACGGAATTGCACCTGATGGTGTAAACGTTGTTTACACATTGAAAGATAAAGATAAA  
 ACTGTTGCTTCAGTATCATTGACAAAAACATCTAAAGGTACAATCGACCTTGGAAATGGTATCAAATTTGAA  
 GTATCTGGTAACTTCTCGGGTAAATTCACTGGTCTAGAAAACAAATCATA CATGATCTCAGAACGTGTTTCT  
 GGTTACGGAAGTGCAATAAATCTAGAAAATGGTAAAGTAACCATTACCAATACCAAAGATTCTGATAACCCA  
 ACACCATTGAACCCA ACTGAACCAAAAAGTTGAAACTCATGGTAAGAAATTTGTCAA AACTAATGAACAAGGT  
 25 GACCGTTTGGCTGGTGCACAATTCGTTGTGAAAACTCAGCAGGTAAATACCTTGCTCTTAAAGCAGATCAA  
 TCAGAAGGTCAAAAA ACTTTAGCTGCTAAGAAAATAGCTTTAGATGAAGCTATCGCTGCTTATAACAAGTTG  
 TCTGCAACAGACCAAAAAGGTGAAAAGGAATTACTGCAAAAAGAACTTATCAA AACTAAACAAGCAGATTAC  
 GATGCAGCCTTCATTGAGGCTCGTACAGCTTATGAGTGGATAACAGATAAGGCTAGAGCCATTACCTACACT  
 TCAAACGATCAAGGTCAATTTGAAGTTACAGGTCTTGCAGACGGTACTTACAACCTTGAAGAAACACTTGCT  
 30 CCAGCAGGATTTGCTAAGTTGGCAGGTAATATTAAGTTTGTAGTTAATCAAGGGTCATACATAACAGGTGGT  
 AACATTGACTACGTTGCTAACAGCAACCAAAAAGATGCGACACGTGTAGAAAATAAAAAGGTAACAATCCCA  
 CAAACAGGTGGTATTGGTACAATTCTTTT CACAATTATTGGTTTAAGCATTATGCTTGGAGCAGTAGTTATC  
 ATGAAAAGACGCCAATCAAAGGAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 125):

35 MKRINKYFAMFSALLLTLTSLLSVAPAFAD EATTNTVTLHKILQTESNLNKS NFPGTTGLNGKDYKGGAI SD  
 LAGYFGEKSKEIEGAFFALALKEDKSGKVQYVKAKEGNKLT PALINKDGTPEITVNI DEAVSGLTPEGDTGL  
 VFNTKGLKGEFKIVEVKSSTYNNNGSLLAASKAVPVNITL PLVNEGDVVADAHVY PKNTEEKPEIDKNFAK  
 TNDLTALTDVNRLLTAGANYGN YARDKATATAEIGKVVPYEVKTKIHKGSKYENLVWTDIMSNGLTMGSTVS  
 LKASGTTETFADTDYELSIDARGFTLKF TADGLGKLEKAAKTADIEFTLTYSATVNGQAI IDNPESNDIKL  
 40 SYGNKPGKDLTELPVTPSKGEVTVAKTWS DGIAPDGVNVVYTLKDKDKTVASVSLTKT SKGTIDLNGIKFE  
 VSGNFSGKFTGLENKSYMISERVSGYGSAINLENGKVTITNTK DSDNPTPLNPTEPKVETHGKKFVKTNEQG  
 DRLAGAQFVVKNSAGKYLALKADQSEGQKTLA AKKIALDEAIAAYNKLSATDQKGEKGITAKELIKTKQADY  
 DAAFIEARTAYEWITDKARAIT YTSNDQGQFEVTGLADGTYNLEETLAPAGFAKLAGNIK FVVNQGSYITGG  
 NIDYVANSNQKDATRVENKKVTIPQTGGIGTILFTI IGLSIMLGAVVIMKRRQSKEA

**Strain IC457**

ORF DNA sequence (SEQ ID NO: 168):

45 ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTC ACTATTCTCAGTT  
 GCACCAGCGTTTGC GGACGACGCAACA ACTGATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 50 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGT GCTTTCTTTGTTTTCAA AATGAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTGTT  
 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTTAACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 GAATTGAAAGAAAAATCAA ACTACGATAACAACGGTTCATCTTGGCTGATTCAA AAGCAGTTCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 55 AAACCACAAGTAGATAAGA ACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAAT TCTTAAAGGCTCAGACTATAAG  
 AAAGTTGTTGGACTGATAGCATGACTAAAGGTTT GACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 GAAGATTTTCTGTTTTAACTACAACTCGTAA CAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 60 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA



GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAAGTCATTAAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAAACGGATGGTACATGG  
 GTGAACGTTGCTTCACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTCACAGGTTTGGATAATGCT  
 AAAACTTACCGCGTTGTGCAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 5 ACTATCAAGAACAACAAAACTCAAATGATCCAACCTCCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 10 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 ACTGGTTTGGATAAAGGCACTTATGGCTTGGGAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAAAACAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATTCTTTTTCACAATTATTGGTTTAAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 15 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 126):

MKKINKCLTMFSTLLLI L T S L F S V A P A F A D D A T T D T V T L H K I V M P Q A A F D N F T E G T K G K N D S D Y V G K Q I N D L  
 K S Y F G S T D A K E I K G A F F V F K N E T G T K F I T E N G K E V D T L E A K D A E G G A V L S G L T K D N G F V F N T A K L K G I Y Q I V  
 E L K E K S N Y D N N G S I L A D S K A V P V K I T L P L V N N Q G V V K D A H I Y P K N T E T K P Q V D K N F A D K D L D Y T D N R K D K G V  
 20 V S A T V G D K K E Y I V G T K I L K G S D Y K K L V W T D S M T K G L T F N N N V K V T L D G E D F P V L N Y K L V T D D Q G F R L A L N A T  
 G L A A V A A A K D K D V E I K I T Y S A T V N G S T T V E I P E T N D V K L D Y G N N P T E E S E P Q E G T P A N Q E I K V I K D W A V D G  
 T I T D A N V A V K A I F T L Q E K Q T D G T W V N V A S H E A T K P S R F E H T F T G L D N A K T Y R V V E R V S G Y T P E Y V S F K N G V V  
 T I K N N K N S N D P T P I N P S E P K V V T Y G R K F V K T N Q A N T E R L A G A T F L V K K E G K Y L A R K A G A A T A E A K A A V K T A K  
 L A L D E A V K A Y N D L T K E K Q E G Q E G K T A L A T V D Q K Q A Y N D A F V K A N Y S Y E W V A D K K A D N V V K L I S N A G G Q F E I  
 25 T G L D K G T Y G L E E T Q A P A G Y A T L S G D V N F E V T A T S Y S K G A T T D I A Y D K G S V K K D A Q Q V Q N K K V T I P Q T G G I G T  
 I L F T I I G L S I M L G A V V I M K K R Q S E E A

#### Strain IC458

ORF DNA sequence (SEQ ID NO: 169):

ATGAAAAAATCAACAAATATTTTGCAGTCTTCTCGGCCTTGCTACTGACCGTAACATCATTGCTCTCAGTT  
 GCACCAGCGTTTGCAGGACGAAGCAACAACTAATACAGTGACTTTGCACAAGATCTTGCAAACTGAATCAAAT  
 CTTAATAAAAAGTAACTTCCCAGGAACCTACAGGCCTTAACGGAGATGACTATAAAGGTGAATCTATTTCTGAC  
 CTTGCTGAATACTTTGGATCAGGTTCTAAAGAAATTGACGGTGCCTTTCTTTGCTTTGGCTTTAGAAGAGGAA  
 AAAGATGGTGTGCTACAATATGTTAAGGCAAAAGCAAATGACAAATTAACACCAGACTTAATTACTAAAGGT  
 35 ACACCTGCAACAACAACAAAAGTTGAAGAAGCTGTAGGTGGTTTGACAACCTGGTACGGGTATTGTTTTCAAT  
 ACAGCTGGTTTGAAGGTAATTTCAAATTTATTGAATTGAAAGACAAATCAACTTACAACAATAATGGTTCC  
 CTCTTAGCAGCTTCAAAGCAGTTCGGGTGAAAATCACTCTTCCATTGGTAAGCAAAGATGGTGTGTTAAA  
 GATGCACACGTTTATCCAAAGAACACTGAAACAAAACCAGAAGTAGACAAGAAGTTCGCTAAAACAAACGAT  
 TTGACAGCTCTCAAAGACGCTACTCTTCTTAAGGCTGGTGCAGACTACAAAACCTATTCAGCGACTAAAGCT  
 40 ACTGTAACAGCTGAAATCGGTAAAGTTATCCCTTACGAAGTTAAAACAAAAGTTCTTAAAGGTTCTAAATAC  
 GAAAACTGGTTTGGACCGATACCATGTCAAATGGTTTGAACAATGGGTGATGATGTTAACCTTGCAGTTTCA  
 GGGACTACAACAACCTTTCATTAAAGATATAGATTACACTCTTAGCATTGATGACCGTGGTTTTCACATTGAAA  
 TTCAAAGCTACTGGATTGGACAAATTGGAAGAAGCAGCTAAAGCATCTGATGTTGAATTTACATTGACTTAT  
 AAAGCTACTGTTAATGGCCAAGCAATTATTGACAACCCAGAAGTCAATGACATCAAATTTGGACTATGGTAAT  
 45 AAACCTGGTACAGATTTATCAGAACAACCTGTGACACCTGAAGATGGTGAAGTTAAAGTCACTAAAACATGG  
 GCAGCAGGTGCTAATAAAGCAGACGCTAAAGTTGTCTACACACTTAAAAATGCTACTAAACAAGTCGTAGCT  
 TCTGTGCGATTGACCGCAGCTGATACAAAAGGTACGATTAATCTTGGTAAAGGCATGACCTTTGAAATCACA  
 GGAGCTTTCTCAGGTACATTCAAAGGCCTTCAAATAAAGCTTACACTGTTTCTGAACGTGTTGCAGGTTAT  
 ACTAATGCTATTAATGTTACTGGTAATGCTGTTGCTATCACCAATACACCAGACAGTGACAATCCAACGCCA  
 50 CTTAACCCAACTCAACCAAAAGTTGAAACACATGGTAAGAAATTTGTCAAAGTTGGCGATGCAGATGCCCGC  
 TTAGCTGGTGCACAATTCGTTGTGAAAAATTCAGCTGGTAAATTCCTTGCTCTTAAAGAAGATGCAGCTGTA  
 TCAGGAGCTCAAACCTGAATTGGCAACTGCTAAAACAGACTTGGATAATGCCATCAAAGCTTACAACGGTTTG  
 ACAAAGCGCAGCAAGAAGGTGCTGATGGTACATCAGCAAAAAGAACTTATCAACACTAAACAGTCAGCTTAC  
 GACGCAGCCTTCATCAAAGCACGTACAGCTTATACATGGGTAGATGAAAAAATAAAGCTATTACCTTCACT  
 55 TCAAATAATCAAGGTCAATTTGAAGTTACTGGTCTTGAAGTAGGTTCTTACAAACTTGAAGAACTCTTGCA  
 CCAGCAGGTTATGCTAAATTGTCAGGCGACATTGAGTTTACAGTTGGACACGATTCTTACACAAGTGGTGAC  
 ATCAAGTACAAGACAGATGATGCTAGCAACAATGCACAAAAGTTTTCAATAAAAAAAGTAACCATCCCACAA  
 ACAGGTGGTATTGGTACAATTCTTTTTCACAATTATTGGTTTAAAGCATTATGCTTGGAGCGGTAGTTATCATG  
 AAAAGACGTCAATCAGAGGAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 127):



MKKINKYFAVFSALLLVTSLLSVAPAFADDEATTNTVTLHKILQTESNLNKS NFPGTTGLNGDDYKGESISD  
 LAEYFGSGSKEIDGAFFALALEEEKDGVVQYVKAKANDKLTPLDITKGT PATTTKVEEAVGGLTTGTGIVFN  
 TAGLKGNFKI IELKDKSTYNNNGSLLAASKAVPVKITLPLVSKDGVVKDAHVPKNTETKPEVDKNFAKTND  
 LTALKDATLLKAGADYKNYSATKATVTAEIGKVI PYEVKTKVLKGSKYEKLWTD TMSNGLTMGDDVNLA VS  
 5 GTTTTFIKDIDYTLSIDDRGFTLKF KATGLDKLEEAAKASDVEFTLTYKATVNGQAI IDNPEVNDIKLDYGN  
 KPGTDLSEQPVT PEDGEVKVTKTWAAGANKADAKVVYTLKNATKQVVASVALTAADTKGTINLKGGMTFEIT  
 GAFSGTFKGLQNKAYTVSERVAGYTNAINVTGNAVAITNTPDSDNPTPLNPTQPKVETHGKKFVKVGDADAR  
 LAGAQFVVKNSAGKFLALKEDAAVSGAQTE LATAKTDLDNAIKAYNGLTKAQQEGADGTSAKELINTKQSAY  
 10 DAAFIKARTAYTWVDEKTKAITFTSNNQGF EVTGLEVGSYKLEETLAPAGYAKLSGDIEFTVGHDSYTSGD  
 IKYKTDDASNNAQKVFNKVTI PQTGGIGTILFTI IGLSIMLGAVVIMKRRQSEA

**Strain IC459**

ORF DNA sequence (SEQ ID NO: 170):

ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTC ACTATTCTCAGTT  
 15 GCACCAGCGTTTGC GGACGACGCAACA ACTGATACTGTGACCTTG CACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCTTTCTTTGTTTTCAAAAATGAAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTGTT  
 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 20 GAATTGAAAGAAAAATCAA CTACGATAACAACGGTCTATCTTGGCTGATTCAAAGCAGTTCCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGA ACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATCTTAAAGGCTCAGACTATAAG  
 AAAGTGGTTTTGGACTGATAGCATGACTAAAGGTTTGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 25 GAAGATTTTCCCTGTTTTAACTACAACTCGTAAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAGTCAATTAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAACGGATGGTACATGG  
 30 GTGAACGTTGCTTCACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTACAGGTTTGGATAATGCT  
 AAAACTTACCGCGTTGTGCAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAA ACTCAAATGATCCAACCTCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCTCGTTAAGAAA  
 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 35 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 ACTGGTTTTGGATAAAGGCACTTATGGCTTGGAAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACTGACATCGCATATGATAAAGGC  
 40 TCTGTAAAAAAGATGCCCAACAAGTTCAAAAACAAAAGTAACCATCCACAAACAGGTGGTATTGGTACA  
 ATCTTTTTCACAATTATTGGTTTTAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 128):

MKKINKCLTMFSTLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEG TKGKNDS DYVGKQINDL  
 45 KSYFGSTDAKEIKGAFFVFKNETGTFITENGKEVD TLEAKDAEGGAVLSGLTKDNGFVFNTAKLKG IYQIV  
 ELKEKSNDNNGSILADSKAVPVKITLPLVNNQGVVKDAHIYPKNTETK PQVDKNFADKDL DYT DNRKDKGV  
 VSATVGDKKEYIVGTKILKGS DYKKLVWTD SMTKGLTFNNNVKVTLDGEDFPVLNYKLVTD DQGFRLALNAT  
 GLAAVAAA AKDKDVEIKITYSATVNGSTTVEI PETNDVKLDYGN NPTEESE PQEGTPANQEIKVIKDWAVDG  
 TITDANVAVKAI FTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTP EYVSFKNGVV  
 50 TIKNNKNSNDPTPINPSEPKVV TYGRKFVKTNQAN TERLAGATFLVKKEGKY LARKAGAATAEAKAAVKTAK  
 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYGLEETQAPAGYATLSGDVNF EVTATSYSKGATTDIAYDKGSVKKDAQQVQNKV TI PQTGGIGT  
 ILFTI IGLSIMLGAVVIMKRRQSEA

**Strain IC460**

ORF DNA sequence (SEQ ID NO: 171):

ATGAAAAGAATCAACAAATATTTTGC AATGTTCTCGGCATTGTTACTGACTTTAACGTCATTGCTCTCAGTT  
 55 GCACCAGCGTTTGC GGATGAAGCAACA ACTAATACAGTGACTTTGCACAAGATTTTGC AAACCGAATCAAAT  
 CTTAACAAAAGTAACTTCCCAGGAACTACAGGTCTTAACGGAAAAGACTACAAAGGTGGAGCTATTTCTGAC  
 60 CTTGCTGGTTACTTTGGCGAGGGATCTAAAGAAATCGAAGGTGCGTTCTTTGCTTTAGCTTTGAAAGAAGAT



AAAAGTGGTAAAGTGCAATATGTTAAGGCCAAAAGAAGGTAACAAATTAACACCAGCCTTAATTAATAAAGAT  
 GGTACTCCTGAAATAACAGTAAATATTGATGAGGCCGTGTCTGGATTGACACCAGAGGGAGATACTGGACTT  
 GTTTTCAACACCAAAGGATTGAAAGGCGAGTTTAAAATTGTTGAAGTTAAATCAAATCTACTTACAACAAT  
 AATGGTTCCTCCTGGCTGCTTCAAAGCGGTTCCAGTTAACATCACTCTTCCATTGGTAAATGAAGATGGT  
 5 GTTGTGCTGATGCCCATGTTTATCCAAAGAACACTGAAGAAAAACCAGAAATTGATAAAAACTTTGCTAAA  
 ACAAACGATTTGACAGCATTGACAGATGTTAATAGACTTTTGACAGCTGGCGCAAATTTATGGTAATTATGCA  
 CGTGACAAAGCAACTGCTACTGCTGAAATCGGTAAAGTTGTTCCCTTATGAAGTTAAAACAAAAATTCACAAA  
 GGTTCTAAATACGAAAACCTTGGTTTGGACAGATATAATGTCAAATGGTTTGGACAATGGGTTCAACTGTTAGC  
 CTTAAAGCTTCAGGAACACAGAACTTTTGCTAAGGATACAGACTATGAACTTAGCATTGATGCCCGTGGT  
 10 TTCACATTAATAATTCACAGCTGATGGATTGGGCAAATTTGGAAAAAGCAGCTAAAACAGCTGATATTGAATTT  
 ACATTGACTTATAGTGCTACTGTTAATGGTCAAGCAATTATTGATAATCCAGAATCCAATGATATCAAATTTG  
 TCGTATGGTAACAAACCAGGTAAAGACTTGACTGAACTTCCCTGTTACACCTTCAAAGGGTGAAGTAACAGTT  
 GCTAAAACCTTGGTCTGACGGAATTGCACCTGATGGTGTAAACGTTGTTTACACATTGAAAGATAAAGATAAA  
 ACTGTTGCTTCAGTATCATTGACAAAAACATCTAAAGGTACAATCGACCTTGGAAATGGTATCAAATTTGAA  
 15 GTATCTGGTAACTTCTCGGGTAAATTCAGTGGTCTAGAAAACAAATCATAACATGATCTCAGAACGTGTTTCT  
 GGTTACGGAAGTGCAATAAATCTAGAAAATGGTAAAGTAACCATTACCAATACCAAAGATTCTGATAACCCA  
 ACACCATTTGAACCAACTGAACCAAAAAGTTGAAACTCATGGTAAGAAATTTGTCAAACTAATGAACAAGGT  
 GACCGTTTGGCTGGTGCACAATTCGTTGTGAAAAACTCAGCAGGTAAATACCTTGTCTTAAAGCAGATCAA  
 TCAGAAGGTCAAAAAACTTTAGCTGCTAAGAAAATAGCTTTAGATGAAGCTATCGCTGCTTATAACAAGTTG  
 20 TCTGCAACAGACCAAAAAGGTGAAAAAGGAATTACTGCAAAAGAACTTATCAAACTAAACAAGCAGATTAC  
 GATGCAGCCTTCATTGAGGCTCGTACAGCTTATGAGTGGATAACAGATAAGGCTAGAGCCATTACCTACACT  
 TCAAACGATCAAGGTCAATTTGAAGTTACAGGTCTTGCAGACGGTACTTACAACCTTGAAGAAACACTTGCT  
 CCAGCAGGATTTGCTAAGTTGGCAGGTAATATTAAGTTTGTAGTTAATCAAGGGTCATACATAACAGGTGGT  
 AACATTGACTACGTTGCTAACAGCAACCAAAAAGATGCGACACGTGTAGAAAATAAAAAGGTAACAATCCCA  
 25 CAAACAGGTGGTATTGGTACAATTCCTTTTACAATTATTGGTTTAAAGCATTATGCTTGGAGCAGTAGTTATC  
 ATGAAAAGACGCCAATCAAAGGAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 129):

MKRINKYFAMFSALLLTLTSLLSVAPAFADDEATTNTVTLHKILQTESNLNKS NFPGTTGLNGKDYKGGAI SD  
 LAGYFGEKSKEIEGAFFALALKEDKSGKVQYVKAKEGNKLT PALINKDGTPEITVNI DEAVSGLTPEGDTGL  
 30 VFNTKGLKGEFKIVEVKSSTYNNNGSLLAASKAVPVNITLPLV NEDGVVADAHVYPKNTEEKPEIDKNFAK  
 TNDLTALTDVNRLLTAGANYGNYARDKATATAEIGKVVPYEVKTKIHKGSKYENLVWTDIMSNGLTMGSTVS  
 LKASGTTETFADTDYELSIDARGFTLKFADGLGKLEKAAKTADIEFTLTYSATVNGQAIIDNPESNDIKL  
 SYGNKPGKDLTELPVTPSKGEVTVAKTWS DGIAPDGVNVVYTLKDKDKTVASVSLTKTSKGTIDLNGIKFE  
 VSGNFSGKFTGLENKSYMISERVSGYGSAINLENGKVTITNTKSDNPTPLNPTEPKVETHGKKFVKTNEQG  
 35 DRLAGAQFVVKNSAGKYLALKADQSEGQKTLAAKKIALDEAIAAYNKLSATDQKGEKGITAKELIKTKQADY  
 DAAFIEARTAYEWITDKARAITYTSNDQGFVETGLADGTYNLEETLAPAGFAKLAGNIKFFVNQGSYITGG  
 NIDYVANSNQKDATRVENKKVTIPQTGGIGTILFTIIGLSIMLGAVVIMKRRQSKEA

#### Strain IC461

ORF DNA sequence (SEQ ID NO: 172):  
 ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCACTATTCTCAGTT  
 GCACCAGCGTTTGC GGACGACGCAACAAC T GATACTGTGACCTTGCACAAGATTGTCATGCCACAAGCTGCA  
 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 40 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCCTTCTTTGTTTTCAAAAATGAAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTGTT  
 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 GAATTGAAAGAAAAATCAAAC TACGATAACAACGGTTCATCTTGGCTGATTCAAAGCAGTTCCAGTTAAA  
 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGA ACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 50 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATCTTAAAGGCTCAGACTATAAG  
 AAAC TGGTTTGGACTGATAGCATGACTAAAGGTTT GACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 GAAGATTTTCCCTGTTTTAAACTACAAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 55 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAGTCATTAAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAGAAAAACAAACGGATGGTACATGG  
 GTGAACGTTGCTTACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTCACAGGTTTGGATAATGCT  
 AAAACTTACCGCGTTGTGCAACGTTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTTG  
 ACTATCAAGAACAACAAAAC TCAAATGATCCAAC TCCAATCAACCCATCAGAACCAAAAAGTGGTGACTTAT  
 60 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCCCTCGTTAAGAAA



GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTAAAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 ACAGCATTGGCTACTGTTGATCAAAAACAAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 5 ACTGGTTTGGATAAAGGCACTTATGGCTTGGAAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAAAACAAAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 ATTCTTTTTCACAATTATTGGTTTAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA

10 ORF amino acid sequence (SEQ ID NO: 130):

MKKINKCLTMFSTLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDFNFTEGTEGKGNDSYVVGKQINDL  
 KSYFGSTDAKEIKGAFFVFKNETGTFKITENGKEVDLEAKDAEGGAVLSGLTKDNGFVFNTAKLKGIVQIV  
 ELKEKSNYDNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTETKPQVDKNFADKDLDTDNRKDKGV  
 VSATVGDKEYIVGTKILKGSYKLVWTDGMTKGLTFNNNVKVTLDGEDFPVLNYKLVTDQGFRLALNAT  
 15 GLAAVAAAADKDKDVEIKITYSATVNGSTTVEIPETNDVKLDYGNPTEESEPEQEGTPANQEIKVIKDWAVDG  
 TITDANVAVKAIFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPHYVSFKNGVV  
 TIKNNKNSNDPTPINPSEPKVVVYGRKFVKTQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYGLEETQAPAGYATLSGDVNFVETATSYSKGATTDIAYDKGSVKKDAQQVQNKKVTIPQTGGIGT  
 20 ILFTIIGLSIMLGAVVIMKKRQSEEA

### Strain IC462

ORF DNA sequence (SEQ ID NO: 173):

ATGAAAAGAATCAACAAATATTTTGCATGTTCTCGGCATTGTTACTGACTTTAACGTCATTGCTCTCAGTT  
 25 GCACCAGCGTTTGCAGGATGAAGCAACAATAACAGTGACTTTGCACAAGATTTTGCAAACCGAATCAAAT  
 CTTAACAAAAGTAACTTCCCAGGAACCTACAGGTCTTAACGGAAAAGACTACAAAGGTGGAGCTATTTCTGAC  
 CTTGCTGGTTACTTTGGCGAGGGATCTAAAGAAATCGAAGGTGCGTTCTTTGCTTTAGCTTTGAAAGAAGAT  
 AAAAGTGGTAAAGTGCAATATGTTAAGGCAAAAAGAAGGTAACAAATTAACACCAGCCTTAATTAATAAAGAT  
 GGTACTCCTGAAATAACAGTAAATATTGATGAGGCCGTGTCTGGATTGACACCAGAGGGAGATACTGGACTT  
 30 GTTTTCAACACCAAAGGATTGAAAGGCGAGTTTAAAATTGTTGAAGTTAAATCAAAATCTACTTACAACAAT  
 AATGGTTCCCTCCTGGCTGCTTCAAAGCGGTTCCAGTTAACATCACTCTTCCATTGGTAAATGAAGATGGT  
 GTTGTGCTGATGCCCATGTTTATCCAAAGAACAACCTGAAGAAAACCAGAAATTGATAAAAACTTTGCTAAA  
 ACAACGATTTGACAGCATTGACAGATGTTAATAGACTTTTGACAGCTGGCGCAAATTATGGTAATTATGCA  
 CGTGACAAAGCAACTGCTACTGCTGAAATCGGTAAAGTTGTTCCCTTATGAAGTTAAAACAAAAATTCACAAA  
 35 GGTTCTAAATACGAAAACCTGGTTTGGACAGATATAATGTCAAATGGTTTGAACAATGGGTTCAACTGTTAGC  
 CTTAAAGCTTCAGGAACCTACAGAACTTTTGCCTAAGGATACAGACTATGAACTTAGCATTTGATGCCCGTGGT  
 TTCACATTAATAATTCACAGCTGATGGATTGGGCAAATTGGAAAAGCAGCTAAAACAGCTGATATTGAATTT  
 ACATTGACTTATAGTGCTACTGTTAATGGTCAAGCAATTATTGATAATCCAGAATCCAATGATATCAAATTG  
 TCGTATGGTAACAAACCAGGTAAAGACTTGACTGAACTTCCCTGTTACACCTTCAAAGGGTGAAGTAACAGTT  
 40 GCTAAAACCTGGTCTGACGGAATTGCACCTGATGGTGTAAACGTTGTTTACACATTGAAAGATAAAGATAAA  
 ACTGTTGCTTCAGTATCATTGACAAAAACATCTAAAGGTACAATCGACCTTGGAATGGTATCAAATTTGAA  
 GTATCTGGTAACTTCTCGGGTAAATTCACTGGTCTAGAAAACAAATCATAACATGATCTCAGAACGTGTTTCT  
 GGTTACGGAAGTGCAATAAATCTAGAAAATGGTAAAGTAACCATTACCAATACCAAAGATTCTGATAACCCA  
 ACACCATTGAACCCAACCTGAACCAAAAAGTTGAAACTCATGGTAAGAAATTTGTCAAACCTAATGAACAAGGT  
 45 GACCGTTTGGCTGGTGCACAATTCGTTGTGAAAACCTCAGCAGGTAAATACCTTGCTCTTAAAGCAGATCAA  
 TCAGAAGGTCAAAAACCTTTAGCTGCTAAGAAAATAGCTTTAGATGAAGCTATCGCTGCTTATAACAAGTTG  
 TCTGCAACAGACCAAAAAGGTGAAAAGGAATTACTGCAAAGAACTTATCAAACCTAAACAAGCAGATTAC  
 GATGCAGCCTTCATTGAGGCTCGTACAGCTTATGAGTGGATAACAGATAAGGCTAGAGCCATTACCTACACT  
 TCAAACGATCAAGGTCAATTTGAAGTTACAGGTCTTGCAGACGGTACTTACAACCTTGAAGAAACACTTGCT  
 50 CCAGCAGGATTTGCTAAGTTGGCAGGTAATATTAAGTTTGTAGTTAATCAAGGGTCATACATAACAGGTGGT  
 AACATTGACTACGTTGCTAACAGCAACCAAAAAGATGCGACACGTGTAGAAAATAAAAAGGTAACAATCCCA  
 CAAACAGGTGGTATTGGTACAATCTTTTTCACAATTATTGGTTTAAGCATTATGCTTGGAGCAGTAGTTATC  
 ATGAAAAGACGCCAATCAAAGGAAGCTTAA

55 ORF amino acid sequence (SEQ ID NO: 131):

MKRINKYFAMFSALLLTLTSLLSVAPAFADDEATTNTVTLHKILQTESNLNKS NFPGTTGLNGKDYKGGAI SD  
 LAGYFGEKSKEIEGAFFALALKEDKSGKVQYVKAKEGNKLT PALINKDGTPEITVNI DEAVSGLTPEGDTGL  
 VFNTKGLKGEFKIVEVKSSTYNNNGSLLAASKAVPVNITLPLV NEDGVVADAHVYPKNTEEKPEIDKNFAK  
 TNDLTALTDVNRLLTAGANYGN YARDKATATAEIGKVVPYEVKTKIHKGSKYENLVWTDIMSNGLTMGSTVS  
 LKASGTTETF AKD TDYELSIDARGFTLKF TADGLGKLEKAAKTADIEFTLTYSATVNGQAIIDNPESNDIKL  
 60 SYGNKPGKDLTELPVTPSKGEVTVAKTWS DGIAPDGVNVVYTLKDKDKTVASVSLTKTSKGTIDLNGIKFE



VSGNFSGKFTGLENKSYMISERVSGYGSAINLENGKVTITNTKSDNPTPLNPTEPKVETHGKKFVKTNEQG  
 DRLAGAQFVVKNSAGKYLALKADQSEGQKTLAACKIALDEAIAAYNKLSATDQKGEKGITAKELIKTKQADY  
 DAAFIEARTAYEWITDKARAITYTSNDQGQFEVTGLADGTYNLEETLAPAGFAKLAGNIKFVFNQGSYITGG  
 NIDYVANSNQKDATRVENKKVTIPQTGGIGTILFTIIGLSIMLGAVVIMKRRQSKEA

5

**Strain IC470**

ORF DNA sequence (SEQ ID NO: 174):

ATGAAAAAATCAACAAATGTCTTACAATGTTCTCGACACTGCTATTGATCTTAACGTCACTATTCTCAGTT  
 GCACCAGCGTTTGC GGACGACGCAACAACTGATACTGTGACCTTGACACAAGATTGTCATGCCACAAGCTGCA  
 10 TTTGATAACTTTACTGAAGGTACAAAAGGTAAGAATGATAGCGATTATGTTGGTAAACAAATTAATGACCTT  
 AAATCTTATTTTGGCTCAACCGATGCTAAAGAAATCAAGGGTGCTTTCTTTGTTTTCAAAAATGAAACTGGT  
 ACAAATTCATTACTGAAAATGGTAAGGAAGTCGATACTTTGGAAGCTAAAGATGCTGAAGGTGGTGTCTGTT  
 CTTTCAGGGTTAACAAAAGACAATGGTTTTGTTTTAACACTGCTAAGTTAAAAGGAATTTACCAAATCGTT  
 GAATTGAAAGAAAAATCAAACTACGATAACAACGGTTCATCTTGGCTGATTCAAAGCAGTTCAGTTAAA  
 15 ATCACTCTGCCATTGGTAAACAACCAAGGTGTTGTTAAAGATGCTCACATTTATCCAAAGAATACTGAAACA  
 AAACCACAAGTAGATAAGAACTTTGCAGATAAAGATCTTGATTATACTGACAACCGAAAAGACAAAGGTGTT  
 GTCTCAGCGACAGTTGGTGACAAAAAGAATACATAGTTGGAACAAAAATCTTAAAGGCTCAGACTATAAG  
 AAAGTGGTTTGGACTGATAGCATGACTAAAGGTTTGACGTTCAACAACAACGTTAAAGTAACATTGGATGGT  
 GAAGATTTTCTGTTTTAACTACAACTCGTAACAGATGACCAAGGTTTCCGTCTTGCCTTGAATGCAACA  
 20 GGTCTTGCAGCAGTAGCAGCAGCTGCAAAAAGACAAAGATGTTGAAATCAAGATCACTTACTCAGCTACGGTG  
 AACGGCTCCACTACTGTTGAAATTCAGAAACCAATGATGTTAAATTGGACTATGGTAATAACCCAACGGAA  
 GAAAGTGAACCACAAGAAGGTACTCCAGCTAACCAAGAAATTAAGTCAATTAAGACTGGGCAGTAGATGGT  
 ACAATTACTGATGCTAATGTTGCAGTTAAAGCTATCTTTACCTTGCAAGAAAAACAACGGATGGTACATGG  
 GTGAACGTTGCTTACACGAAGCAACAAAACCATCACGCTTTGAACATACTTTTACAGGTTTGGATAATGCT  
 25 AAACTTACCGCGTTGTCGAACGTGTTAGCGGCTACACTCCAGAATACGTATCATTTAAAAATGGTGTGTG  
 ACTATCAAGAACAACAAAACCTCAATGATCCAACCTCAATCAACCCATCAGAACCAAAAGTGGTGACTTAT  
 GGACGTAAATTTGTGAAAACAAATCAAGCTAACACTGAACGCTTGGCAGGAGCTACCTTCTCGTTAAGAAA  
 GAAGGCAAATACTTGGCACGTAAAGCAGGTGCAGCAACTGCTGAAGCAAAGGCAGCTGTA AAAACTGCTAAA  
 CTAGCATTGGATGAAGCTGTTAAAGCTTATAACGACTTGACTAAAGAAAAACAAGAAGGCCAAGAAGGTAAA  
 30 ACAGCATTGGCTACTGTTGATCAAAAACAAAAGCTTACAATGACGCTTTTGTAAAGCTAACTACTCATAT  
 GAATGGGTTGCAGATAAAAAGGCTGATAATGTTGTTAAATTGATCTCTAACGCCGGTGGTCAATTTGAAATT  
 ACTGGTTTGGATAAAGGCACCTTATGGCTTGGAAAGAACTCAAGCACCAGCAGGTTATGCGACATTGTCAGGT  
 GATGTAACTTTGAAGTAACTGCCACATCATATAGCAAAGGGGCTACAACCTGACATCGCATATGATAAAGGC  
 TCTGTAAAAAAGATGCCCAACAAGTTCAAAACAAAAGTAACCATCCCACAAACAGGTGGTATTGGTACA  
 35 ATTCTTTTACAATTATTGGTTTAAAGCATTATGCTTGGAGCAGTAGTTATCATGAAAAACGTCAATCAGAG  
 GAAGCTTAA

ORF amino acid sequence (SEQ ID NO: 132):

MKKINKCLTMFSTLLLLILTSLFSVAPAFADDATTDVTLHKIVMPQAAFDNFTEGTKGKNDSYVVGKQINDL  
 KSYFGSTDAKEIKGAFFVFKNETGTFKITENGKEVDLEAKDAEGGAVLSGLTKDNGFVFNTAKLKGIIYQIV  
 40 ELKEKSNYDNNGSILADSKAVPVKITLPLVNNQGVVKAHIYPKNTETKPKQVDKNFADKDLDDYTDNRKDKGV  
 VSATVGDKKEYIVGTKILKGSYKLVWTD SMTKGLTFNNNVKVTLDGEDFPVLNYKLVTDDQGFRLALNAT  
 GLAAVAAAAKDKDVEIKITYSATVNGSTTVEIPETNDVKLDYGNPTEESEPEGTPANQEIKVIKDWAVDG  
 TITDANVAVKAIFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLDNAKTYRVVERVSGYTPYVVSFKNGVV  
 TIKNNKNSNDPTPINPSEPKVVVYGRKFVKTNQANTERLAGATFLVKKEGKYLARKAGAATAEAKAAVKTAK  
 45 LALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWVADKKADNVVKLISNAGGQFEI  
 TGLDKGTYGLEETQAPAGYATLSGDVNFVETATSYSKGATTDIAYDKGSVKKDAQVQNKKVTIPQTGGIGT  
 ILFTIIGLSIMLGAVVIMKRRQSEEA

50 **Table 11: Amino acid and encoding DNA sequences of gbs1478 proteins derived from different strains of *S. agalactiae***

**Strain 12401**

ORF DNA sequence (SEQ ID NO: 204):

ATGAGAAAATACCAAAAATTTCTAAAATATTGACGTTAAGTCTTTTTTGTGTCGCAAATACCGCTTAAT  
 55 ACCAATGTTTTAGGGGAAAGTACCGTACCGGAAAATGGTGCTAAAGGAAAGTTAGTTGTTAAAAAGACAGAT  
 GACCAGAACAAACCACTTTCAAAGCTACCTTTGTTTTAAAACCTACTTCACACTCAGAAAGCAAAGTAGAA  
 AAAGTAACTACTGAGGTAACAGGTGAAGCTACTTTTGATAATCTCACACCTGGAGATTACACTTTATCAGAA  
 GAAACGGCACCCGAAGGATACAAAAGACTACCCAGACTTGGCAAGTTAAGGTTGAGAGTAATGGAAAACT



ACGATACAAAATAGTGATGATAAAAAATCTATAATTGAACAAAGGCAAGAGGAACTAGATAAGCAGTATCCC  
 CTTACAGGAGCTTATGAAGATACAAAAGAATCTTATAATCTTGAGCATGTTAAAAATTC AATTCCAAATGGG  
 AAATTAGAGGCAAAAGCAGTTAATCCATATTC AAGTGAAGGTGAGCACATAAGAGAAATTC AAGAGGGAACA  
 TTATCTAAACGTATTT CAGAAGTAAATGATTTGGATCATAATAAATATAAAAATTGAGTTAACTGTTAGCGGT  
 5 AAATCCATAATAAAAACTATAAATAAAGATGAACCTCTGGATGTTGTTTTTGTCTTGATAAATTC A AATCT  
 ATGAAGAATAATGGAAAAATAACAAGGCAAAAAAGGCAGGTGAAGCAGTAGAAACAATTATAAAAAGATGTT  
 TTAGGAGCAAATGTTGAAAACCGAGCAGCTTTAGTTACTTATGGTTCAGATATTTTTGATGGAAGGACAGTT  
 AAAGTTATAAAAGGTTTTAAAGAGGATCCTTATTATGGACTTGAAACTAGTTTCACAGTTCAGACAAATGAT  
 TATAGCTATAAAAAGTTCACTAATATTGCTGCTGATATTATAAAAAAGATCCCTAAAGAAGCTCCAGAAGCT  
 10 AAGTGGGGGGGACAAGTCTAGGATTA ACTCCAGAAAAAAGAGGGAATATGATTTAAGTAAAGTAGGTGAG  
 ACCTTTACAATGAAAGCTTTTTATGGAGGCAGATACCTTGTTAAGTAGTATACAGCGTAAGAGTAGAAAGATT  
 ATTGTTTCATCTAACTGACGGTGTTC AACAAGATCATATGCCATTAATAGTTTTGTAAAAGGTTCAACATAC  
 GCAATCAATTTGAGAGAATAAAAAGAAAAAGGTTATTTAGACAAAAATAATTATTTTATAACTGATGATCCA  
 GAAAAGATCAAAGGCAATGGGGAGAGTTACTTTTTGTTTTCCCTTAGATAGTTATCAAACACAGATAATTTCT  
 15 GGAACTTACAAAACTTCATTATTTAGATTTAAATCTTAATTACCTAAAGGTACAATTTATAGAAATGGA  
 CCAGTAAGAGAACATGGAACACCAACCAAACTTTATATAAATAGTTTAAAACAGAAAAATTATGACATCTTT  
 AATTTTGGTATAGATATATCTGGTTTTAGACAAGTTTTATAATGAGGATTATAAGAAAAATCAAGATGGTACT  
 TTTCAAAAATTGAAGGAGGAAGCTTTTTGAACTTTCAGATGGGGAAATAACAGAACTAATGAATTCATTCTCT  
 TCTAAACCTGAGTATTATACCCCGATAGTAACTTCAGCTGATGTATCTAATAATGAAATTTTATCTAAAATT  
 20 CAGCAACAATTTGAAAAGATTTTAAACAAAGGAAAACTCAATTGTTAATGGA ACTATAGAAGATCCTATGGGT  
 GATAAAATCAATTTACATCTTGGCAACGGACAAACATTGCAACCAAGTGATTATACTTTACAGGGAAATGAT  
 GGAAGTATAATGAAAGATAGCATTGCAACTGGAGGGCCTAATAATGATGGTGGGATACTTAAAGGGGTTAAA  
 TTAGAATACATCAAAAATAAACTCTACGTTAGAGGTTTGA ACTTAGGGGAGGGGCAAAAAGTAACACTCACA  
 TATGATGTGAACTAGATGACAGTTTTATTAGTAACAAATTCTATGACACTAATGGTAGAACAAACATTGAAT  
 25 CCTAAATCAGAGGAACCCGATACACTTAGAGATTTTCCAATCCCTAAAATTCGTGATGTGAGAGAATATCCT  
 ACAATAACGATTAAAAACGAGAAGAAGTTAGGTGAAATTTGAATTTACAAAAGTTGATAAAGATAATAAAG  
 TTGCTTCTCAAAGGAGCTACATTTGAACTTCAAGAATTTAATGAAGATTATAAACTTTATTTACCAATAAAA  
 AATAATAATTCAAAAGTAGTGACGGGAGAAAACGGCAAAAATTTCTTACAAAGATTTGAAAGATGGCAAATAT  
 CAGTTAATAGAAGCAGTTTCGCCGAAGGATTATCAAAAATTACTAATAAAACCAATTTTAACTTTTGAAGTT  
 30 GTTAAAGGATCGATACAAAATATAATAGCTGTTAATAAACAGATTTCTGAATATCATGAGGAAGGTGACAAG  
 CATTTAATTACCAACACGCATATTCACCAAAAAGGAATTATTCGATGACAGGTGGGAAAGGAATTCATCT  
 TTCATTTTAAATAGGTGGAGCTATGATGTCTATTGCAGGTGGAATTTATATTTGGAAAAGACATAAGAAATCT  
 AGTGATGCATCAATCGAGAAAGATTAA

ORF amino acid sequence (SEQ ID NO: 185):

35 MRKYQKFSKILTLSLFCLSQIPLNTNVLGESTVPENGAKGKLVVKKTDQNKPLSKATFVLKPTSHSESKVE  
 KVTTEVTGEATFDNLTPGDYTLSEETAPEGYKKTQTWQVKVESNGKTTIQNSDDKKSIEQRQEELDKQYP  
 LTGAYEDTKESYNLEHVKN SIPNGKLEAKAVNPYSSEGEHIREIQEGTLSKR ISEVNDLDHNKYKIELTVSG  
 KSI IKTINKDEPLDVVFLDNSNSMKNNNGKNNKAKKAGEAVETI IKDVLGANVENRAALVTYGSDFDGR TV  
 40 KVIKGFKEDPYGLETSTVQTDYSYKKFTNIAADI IKKIPKEAPEAKWGGTSLGLTPEKKREYDLSKVG  
 TFTMKAFMEADTLLSSIQRKSRK IIVHLTDGVPTRSYAINSFVKGSTYANQFERIKEKGYLDKNNYFITDDP  
 EKIKNGESYFLFPLDSYQTQ IISGNLQKLHYLDLNLNYPKGTIYRNGPVREHGTPTKLYINSLKQKNYDIF  
 NFGIDISGFRQVYNEDYKKNQDGT FQKLKEEFELSDGEITELMNSFSSKPEYYTPIVTSADVSNNEILSKI  
 QQQFEKILTKENSIVNGTIEDPMDKINLHLGNGQTLQPSDYTLQGN DGSIMKDSIATGGPNNDGGILKGVK  
 45 LEYIKNKLYVRGLNLGEGQKVTLT YDVKLDDSFISNKFYDTNGR TTLNPKSEEPDTLRDFPIPKIRDVREYP  
 TITIKNEKKLGEIEFTKVDKDNK LLLKGATFELQEFNEDYKLYLP IKNNSKVVTGENGKISYKDLKDGKY  
 QLIEAVSPKDYQKITNKPI LTFEVVKGSIQNI IAVNKQISEYHEEGDKHLITNTHIPP KGIIPMTGGKGI LS  
 FILIGGAMMSIAGGIYIWK RHKKSSDASIEKD

### Strain BAA23

50 ORF DNA sequence (SEQ ID NO: 205):  
 ATGAGAAAATACCAAAAATTTTCTAAAATATTGACGTTAAGTCTTTTTTGTGTCGCAAATACCGCTTAAT  
 ACCAATGTTTTAGGGGAAAGTACCGTACCGGAAAATGGTGCTAAAGGAAAGTTAGTTGTTAAAAAGACAGAT  
 GACCAGAACAAACCACTTTCAAAGCTACCTTTGTTTTAAAACTACTGCTCATCCAGAAAGTAAAATAGAA  
 AAAGTAACTGCTGAGCTAACAGGTGAAGCTACTTTTGATAATCTCATACTGGAGATTATACTTTATCAGAA  
 55 GAAACAGCGCCCGAAGGTTATAAAAAGACTAACCAGACTTGGCAAGTTAAGGTTGAGAGTAATGGAAAACT  
 ACGATACAAAATAGTGGTGATAAAAAATCCACAATTGGACAAAATCAGGAAGAACTAGATAAGCAGTATCCC  
 CCCACAGGAATTTATGAAGATACAAAGGAATCTTATAAACTTGAGCATGTTAAAGGTT CAGTTCCAAATGGA  
 AAGTCAGAGGCAAAAGCAGTTAACCCATATTC AAGTGAAGGTGAGCATATAAGAGAAATTCAGAGGGAACA  
 TTATCTAAACGTATTT CAGAAGTAGGTGATTTAGCTCATAATAAATATAAAAATTGAGTTAACTGTCAGTGA  
 60 AAAACCATAGTAAAACCAAGTGGACAAACAAAAGCCGTTAGATGTTGTCTTCGTACTCGATAATTC TAACTCA



ATGAATAACGATGGCCCAAATTTTCAAAGGCATAATAAAGCCAAGAAAGCTGCCGAAGCTCTTGGGACCGCA  
 GTAAAAGATATTTTAGGAGCAAACAGTGATAATAGGGTTGCATTAGTTACCTATGGTTCAGATATTTTGGAT  
 GGTAGGAGTGTAGATGTCGTAAAAGGATTTAAAGAAGATGATAAATATTATGGCCTTCAAACCTAAGTTCACA  
 ATTCAGACAGAGAATTATAGTCATAACAATTAACAAATAATGCTGAAGAGATTATAAAAAGGATTCCGACA  
 5 GAAGCTCCTAAAGCTAAGTGGGGATCTACTACCAATGGATTAACCTCCAGAGCAACAAAAGGAGTACTATCTT  
 AGTAAAGTAGGAGAAACATTTACTATGAAAGCCTTCATGGAGGCAGATGATATTTTGGAGTCAAGTAAATCGA  
 AATAGTCAAAAATTATTGTTTCATGTAAGTGGTGTTCCTACGAGATCATATGCTATTAATAATTTTAAA  
 CTGGGTGCATCATATGAAAGCCAATTTGAACAAATGAAAAAAATGGATATCTAAATAAAAGTAATTTTCTA  
 CTTACTGATAAGCCCGAGGATATAAAAAGGAAATGGGGAGAGTTACTTTTTGTTTCCCTTAGATAGTTATCAA  
 10 ACACAGATAATCTCTGGAACTTACAAAACCTTCATTATTTAGATTTAAATCTTAATTACCCTAAAGGTACA  
 TTTTATCGAAATGGACCAGTAAGAGAACATGGAACACCAACCAACTTTATATAAATAGTTTAAAACAGAAA  
 AATTATGACATCTTTAATTTTGGTATAGATATATCTGGTTTTAGACAAGTTTATAATGAGGATTATAAGAAA  
 AATCAAGATGGTACTTTTCAAAAATTGAAAGAGGAAGCTTTTGAACCTTCAGATGGGGAAATAACAGAATA  
 ATGAAGTCATTCTCTTCTAAACCTGAGTATTATACCCCGATAGTAACTTCATCCGATGCATCTAACAAATGAA  
 15 ATTTTATCTAAAATTCAGCAACAATTTGAAAAGATTTTAAACAAAAGAAAACCTCAATTGTTAATGGAACATA  
 GAAGATCCTATGGGTGACAAAATCAATTTACAGCTTGGCAACGGACAAACATTGCAACCAAGTGATTATACT  
 TTACAGGGAAATGATGGAAGTATAATGAAAGATAGCATTGCAACTGGTGGGCCATAAATGATGGTGGAAATA  
 CTTAAAGGGGTAAATTAGAATACATCAAAAATAAACTCTACGTTAGAGGTTTGAACCTTAGGGGAGGGACAA  
 AAAGTAACACTCACATATGATGTGAAACTAGATGACAGTTTTATAAGTAACAAATCTATGACACTAATGGT  
 20 AGAACAACTTGAATCCTAAATCAGAGGATCCTAATACACTTAGAGATTTTCCAATCCCTAAAATTCGTGAT  
 GTGAGAGAATATCCTACAATAACGATTAAAAACGAGAAGAAGTTAGGTGAAATTGAATTTACAAAAGTTGAT  
 AAAGATAATAAAGTTGCTTCTCAAAGGAGCTACGTTTGAACCTCAAGAATTTAATGAAGATTATAAACTT  
 TATTTACCAATAAAAAATAAATTCAAAAGTAGTGACGGGAGAAAACGGCAAAATTTCTTACAAAGATTTG  
 AAAGATGGCAAATATCAGTTAATAGAAGCAGTTTCGCCGAAGGATTATCAAAAATTAATAAAACCAATT  
 25 TTAACCTTTGAAGTTGTTAAAGGATCGATACAAAATATAATAGCTGTTAATAAACAGATTTCTGAATATCAT  
 GAGGAAGGTGACAAGCATTAAATTACCAACACGCATATTCACCAAAAAGGAATTATTCGATGACAGGTGGG  
 AAAGGAATCTATCTTTCAATTTAATAGGTGGATCTATGATGCTATTGCAGGTGGAATTTATATTTGGAAA  
 AGATATAAGAAATCTAGTGATATATCTAGAGAAAAAGATTAA  
 ORF amino acid sequence (SEQ ID NO: 186):  
 30 MRKYQKFSKILTLFLSFLSFIPLNTNVLGESTVPENGAKGKLVVKKTDQNKPLSKATFVLKTTAHPESKIE  
 KVTAELTGEATFDNLI PGDYTLSEETAPEGYKKTNQWQVKVESNGKTTIQNSGDKNSTIGQNQEELDKQYP  
 PTGIYEDTKESYKLEHVKGSVPNGKSEAKAVNPYSSEGEHIREIPEGTLISKRISEVGDLAHNKYKIELTVSG  
 KTIVKPVDKQKPLDVVFLDNSNSMNDGPNFQRHNKAKKAAEALGTAVKDI LGANSNDRVALVTYGSDFD  
 35 GRSVDVVKGFKEDDKYYGLQTKFTIQTENYSHKQLTNNAAEIIKRIPTAPKAKWGSTTNGLTPEQQKEYYL  
 SKVGETFTMKAFMEADDILSQVNRNSQKIVHVTGDVPTRSYAINNFKLGASYESQFEQMKNKGYLNKSNFL  
 LTDKPEDIKNGESYFLPLDSYQTQII SGNLQKLHYLDLNLNYPKGTFYRNGPVREHGTPTKLYINSLKQK  
 NYDIFNFGIDISGFRQVYNEKYKNDGTFQKLKEEAFELSDGEITELMKSFSSKPEYYTPIVTSDDASNE  
 ILSKIQQFQFEKILTKENSIVNGTIEDPMGDKINLQLGNGQTLQPSDYTLQNDGSIKDSIATGGPNDGGI  
 LKGVKLEYIKNKLYVRGLNLGEGQKVTLYDVKLLDSFI SNKFYDTNGRITLNPKSEDPNTRDFPIPKIRD  
 40 VREYPTITIKNEKKLGEIEFTKVDKDNKLLLKGATFELQEFNEDYKLYLPIKNNNSKVVTGENGKISYKDL  
 KDGYQLIEAVSPKDYQKINTKPIILFEVVKGSIQNI IAVNKQISEYHEEGDKHLITNTHIPPKGIIPMTGG  
 KGILSFILIGGSMMSIAGGIYIWKRYKSSDISREKD

**Strain IC98**  
 45 ORF DNA sequence (SEQ ID NO: 206):  
 ATGAAAAAGAGACAAAAAATATGGAGAGGGTTATCAGTTACTTTACTAATCCTGTCCCAAATTCATTTGGT  
 ATATTGGTACAAGGTGAAACCAAGATACCAATCAAGCACTTGGAAAAGTAATTGTTAAAAAACGGGAGAC  
 AATGCTACACCATTAGGCAAAGCGACTTTTGTGTTAAAAAATGACAATGATAAGTCAGAAACAAGTCACGAA  
 50 ACGGTAGAGGGTTCTGGAGAAGCAACCTTTGAAAACATAAAACCTGGGAGACTACACATTAAGAGAAGAAACA  
 GCACCAATTGGTTATAAAAAAAGTATAAAACCTGGAAAGTTAAAGTTGCAGATAACGGAGCAACAATAATC  
 GAGGGTATGGATGCAGATAAAGCAGAGAAACGAAAAGAAGTTTTGAATGCCCAATATCCAAAATCAGCTATT  
 TATGAGGATACAAAAGAAAATTACCCATTAGTTAATGTAGAGGGTTCCAAAGTTGGTGAACAATACAAAGCA  
 TTGAATCCAATAAATGGAAAAGATGGTCAAGAGAGATTGCTGAAGGTTGGTTATCAAAAAAATTACAGGG  
 55 GTCAATGATCTCGATAAGAATAAATATAAAATTGAATTAACCTGTTGAGGGTAAAACCACTGTTGAAACGAAA  
 GAACTTAATCAACCACTAGATGTCGTTGTGCTATTAGATAATTCAAATAGTATGAATAATGAAAGAGCCAAT  
 AATTCTCAAAGAGCATTAAAAGCTGGGGAAAGCAGTTGAAAAGCTGATTGATAAAATTACATCAAATAAAGAC  
 AATAGAGTAGCTCTTGTGACATATGCCTCAACCAATTTTTGATGGTACTGAAGCGACCGTATCAAAGGGAGTT  
 GCCGATCAAAATGGTAAAGCGCTGAATGATAGTGTATCATGGGATTATCATAAAACTACTTTTACAGCAACT  
 60 ACACATAATTACAGTTATTTAAATTTAAACAAATGATGCTAACGAAGTTAATATTCTAAAGTCAAGAATTCCA  
 AAGGAAGCGGAGCATATAAATGGGGATCGCACGCTCTATCAATTTGGTGCAGATTTACTCAAAAAGCTCTA



- 110 -

ATGAAAGCAAATGAAATTTTAGAGACACAAAGTTCTAATGCTAGAAAAAACTTATTTTTACGTAAGTATGAT  
 GGTGTCCCTACGATGTCCTTATGCCATAAATTTTAATCCTTATATATCAACATCTTACCAAACCAGTTTAAAT  
 TCTTTTTTAAATAAAAATACCAGATAGAAGTGGTATTCTCCAAGAGGATTTTATAATCAATGGTGATGATTAT  
 CAAATAGTAAAAGGAGATGGAGAGAGTTTTAAACTGTTTTCGGATAGAAAAGTTCCTGTTACTGGAGGAACG  
 5 ACACAAGCAGCTTATCGAGTACCGCAAATCAACTCTCTGTAATGAGTAATGAGGGATATGCAATTAATAGT  
 GGATATATTTATCTCTATTGGAGAGATTACAACCTGGGTCTATCCATTTGATCCCTAAGACAAAGAAAGTTTCT  
 GCAACGAAACAAATCAAACTCATGGTGAGCCAACAACATTATACTTTAATGGAAATATAAGACCTAAAGGT  
 TATGACATTTTTACTGTTGGGATTGGTGTAACCGGAGATCCTGGTGCAACTCCTCTTGAAGCTGAGAAATTT  
 ATGCAATCAATATCAAGTAAAACAGAAAATTATACTAATGTTGATGATACAAAATAAAATTTATGATGAGCTA  
 10 AATAAATACTTTAAAACAATTGTTGAGGAAAACATTCTATTGTTGATGGAAATGTGACTGATCCTATGGGA  
 GAGATGATTGAATTCCAATTAATAAATGGTCAAAGTTTTACACATGATGATTACGTTTTGGTTGGAAATGAT  
 GGCAGTCAATTAATAAATGGTGTGGCTCTTGGTGGACCAAACAGTATGGGGGAATTTTAAAAGATGTTACA  
 GTGACTTATGATAAGACATCTCAAACCATCAAAATCAATCATTGAACTTAGGAAGTGGACAAAAGTAGTT  
 CTTACCTATGATGTACGTTTTAAAAGATAACTATATAAGTAACAAATTTTACAATACAAATAATCGTACAACG  
 15 CTAAGTCCGAAGAGTGAAAAAGAACCATACTATTCGTGATTTCCCAATTCCAAATTCGTGATGTTCTGT  
 GAGTTTCCGGTACTAACCATCAGTAATCAGAAGAAAATGGGTGAGGTTGAATTTATTAAAGTTAATAAAGAC  
 AAACATTCAGAATCGCTTTTGGGAGCTAAGTTTCAACTTCAGATAGAAAAGATTTTTCTGGGTATAAGCAA  
 TTTGTTCCAGAGGAAGTGATGTTACAACAAAGAATGATGGTAAAATTTATTTTAAAGCACTTCAAGATGGT  
 AACTATAAATTATATGAAATTTCAAGTCCAGATGGCTATATAGAGGTTAAAACGAAACCTGTTGTGACATTT  
 20 ACAATTCAAATGGAGAAGTTACGAACCTGAAAGCAGATCCAAATGCTAATAAAAATCAAATCGGGTATCTT  
 GAAGGAAATGGTAAACATCTTATTACCAACACTCCCAAACGCCACCAGGTGTTTTCTTAAACAGGGGGA  
 ATTGGTACAATTGTCTATATATTAGTTGGTCTACTTTTATGATACTTACCATTTGTTCTTTCCGTCGTAAA  
 CAATTGTAA

ORF amino acid sequence (SEQ ID NO: 187):

MKKRQKIWRGLSVTLILLSQIPFGILVQGETQDTNQLGKVIKKTGDNATPLGKATFVLKNDNDKSETSHE  
 TVEGSGEATFENIKPGDYTLREETAPIGYKKTDKTWKVKVADNGATIIEGMDADKAEKRKEVLNAQYPKSAI  
 YEDTKENYPLVNVEGSKVGEQYKALNPINGKDRREIAEGWLSKKITGVNDLKNKYKIELTVEGKTTVETK  
 ELNQPLDVVLLDNSNSMNERANNSQRALKAGEAVEKLIDKITSNKDNRVALVTYASTIFDGTEATVSKGV  
 ADQNGKALNDSVSWDYHKTTFATTHNYSYLNLTNDANEVNILKSRIKPEAEHINGDRTLYQFGATFTQKAL  
 30 MKANEILETQSSNARKKLI FHVTDGVP TMSYAINFNPIYSTSYQNQFNSFLNKIPDRSGILQEDFIINGDDY  
 QIVKGDGESFKLFSDRKVPVTGGTTQAA YRVPQNQLSVMSEGYAINSGYIYLYWRDYNWVYPFDPKTKKVS  
 ATKQIKTHGEPTTLYFNGNIRPKGYDIFTVGI VNGDPGATPLEAEKFMQSISSKTENYTNVDDTNKIYDEL  
 NKYFKTIVEEKHSIVDGNVTDPMGEMIEFQLKNGQSFTHDDYVLVGN DGSQLKNGVALGGPNSDGGILKDV  
 VTYDKTSQTIKINHLNLGSGQKVVLTYDVRLK DNYISNKFYNTNRR TLLSPKSEKEPNTIRDFPIPKIRDVR  
 35 EFPVLTISNQKMGVEFEIKV NKDKHSESLGAKFQLQIEKDFSGYKQFVPEGS DVTTKNDGKIYFKALQDG  
 NYKLYEISSPDGYIEVKTKPVVFTFIQNGEVTNLKADPNANKNQIGYLEGNGKHLITNTPKRPPGVFPKTGG  
 IGTIVYILVGSFMILTICSFRRKQL

### Strain IC105

ORF DNA sequence (SEQ ID NO: 207):

ATGAGAAAATACCAAATTTTCTAAAATATTGACGTTAAGTCTTTTTTGTGTTGTCGCAAATACCGCTTAAT  
 ACCAATGTTTTAGGGGAAAGTACCGTACCGGAAAATGGTGCTAAAGGAAAGTTAGTTGTTAAAAGACAGAT  
 GACCAGAACAAACCACTTTCAAAGCTACCTTTGTTTTAAAACACTACTGCTCATCCAGAAAGTAAAATAGAA  
 AAAGTAACTGCTGAGGTAACAGGTGAAGCTACTTTTGATAATCTCACACCTGGAGATTACACTTTATCAGAA  
 45 GAAACGGCACCCGAAGGATACAAAAGACTACCCAGACTTGGCAAGTTAAGGTTGAGAGTAATGGAAAACCT  
 ACGATACAAAATAGTGATGATAAAAATCTATAATTGAACAAAGGCAAGAGGAACTAGATAAGCAGTATCCC  
 CTTACAGGAGCTTATGAAGATACAAAAGAATCTTATAATCTTGAGCATGTTAAAATTTCAATTCCAAATGGG  
 AAATTAGAGGCAAAGCAGTTAATCCATATTCAAGTGAAGGTGAGCACATAAGAGAAATTCAGAGGGAACA  
 TTATCTAAACGTATTTCAGAAGTAAATGATTTGGATCATAATAAATAAAAATTTAGTTAACTGTTAGCGGT  
 50 AAATCCATAATAAAAACCTATAAATAAAGATGAACCTCTGGATGTTGTTTTGTTCTTGATAATTCAAATCT  
 ATGAAGAATAATGGAAAAATAACAAGGCAAAAAGGCAGGTGAAGCAGTAGAAACAATTATAAAAGATGTT  
 TTAGGAGCAAATGTTGAAAACCGAGCAGCTTTAGTTACTTATGGTTCAGATATTTTTGATGGAAGGACAGTT  
 AAAGTTATAAAAGTTTTTAAAGAGGATCCTTATTATGGACTTGAACTAGTTTCACAGTTCAGACAAATGAT  
 TATAGCTATAAAAAGTTCACTAATATTGCTGCTGATATTATAAAAAGATCCCTAAAGAAGCTCCAGAAGCT  
 55 AAGTGGGGGGGACAAGTCTAGGATTAACCTCAGAAAAAAGAGGGAATATGATTTAAGTAAAGTAGGTGAG  
 ACCTTTACAATGAAAGCTTTTATGGAGGCAGATACCTTGTTAAGTAGTATACAGCGTAAGAGTAGAAAGATT  
 ATTGTTTCACTAACTGACGGTGTTCACAACAGATCATATGCCATTAATAGTTTTGTAACAGGTTCAACATAC  
 GCAAATCAATTTGAGAGAATAAAAAGAAAAGGTTATTTAGACAAAATAATTTATTTTATAACTGATGATCCA  
 GAAAAGATCAAAGGCAATGGGGAGAGTTACTTTTTGTTTCCCTTAGATAGTTATCAAACACAGATAATTTCT  
 60 GGAAACTTACAAAACCTTCAATTATTAGATTTAAATCTTAATTACCTAAAGGTACAATTTATAGAAATGGA



- 111 -

CCAGTAAGAGAACATGGAACACCAACCAAACTTTATATAAATAGTTTAAAACAGAAAAATTATGACATCTTT  
 AATTTTGGTATAGATATATCTGGTTTTAGACAAGTTTATAATGAGGATTATAAGAAAAATCAAGATGGTACT  
 TTTCAAAAATTGAAGGAGGAAGCTTTTGAACCTTCAGATGGGGAAATAACAGAATAATGAATTCATTCTCT  
 TCTAAACCTGAGTATTATACCCCGATAGTAACTTCAGCTGATGTATCTAATAATGAAATTTTATCTAAAATT  
 5 CAGCAACAATTTGAAAAGATTTTAAACAAAGGAAAACTCAATTGTTAATGGAACATAGAAAGATCCTATGGGT  
 GATAAAATCAATTTACATCTTGGCAACGGACAAACATTGCAACCAAGTGATTATACTTTACAGGGAAATGAT  
 GGAAGTATAATGAAAGATAGCATTGCAACTGGAGGGCCTAATAATGATGGTGGGATACTTAAAGGGGTAAA  
 TTAGAATACATCAAAAATAAACTCTACGTTAGAGGTTTGAACCTTAGGGGAGGGGCAAAAAGTAACACTCACA  
 TATGATGTGAAACTAGATGACAGTTTTATTAGTAACAAATTCTATGACACTAATGGTAGAACAAACATTGAAT  
 10 CCTAAATCAGAGGAACCCGATACACTTAGAGATTTTCCAATCCCTAAAATTCGTGATGTGAGAGAATATCCT  
 ACAATAACGATTAAAAACGAGAAGAAGTTAGGTGAAATTGAATTTACAAAAGTTGATAAAGATAATAAAG  
 TTGCTTCTCAAAGGAGCTACATTTGAACCTCAAGAATTTAATGAAGATTATAAACTTTATTTACCAATAAAA  
 AATAAATTTCAAAGTAGTGACGGGAGAAAACGGCAAAAATTTCTTACAAAAGATTTGAAAGATGGCAAATAT  
 15 CAGTTAATAGAAGCAGTTTCGCCGAAGGATTATCAAAAAATTACTAATAAACCAATTTTAACTTTTGAAGTT  
 GTTAAAGGATCGATACAAAATATAATAGCTGTTAATAAACAGATTTCTGAATATCATGAGGAAGGTGACAAG  
 CATTTAATTACCAACACGCATATTCACCAAAAAGGAATTATTCGGATGACAGGTGGGAAAGGAATTCATCT  
 TTCATTTTAAATAGGTGGAGCTATGATGTCTATTGCAGGTGGAATTTATATTTGAAAAGACATAAGAAATCT  
 AGTGATGCATCAATCGAGAAAGATTAA

ORF amino acid sequence (SEQ ID NO: 188):

20 MRKYQKFSKILTLFLSLSQIPLNTNVLGESTVPENGAAGKGLVVKKTDDQNKPLSKATFVLKTTAHPESKIE  
 KVTA EVTGEATFDNLTPGDYTLSEETAPEGYKKTQTWQVKVESNGKTTIQNSDDKKSIEQRQEELDKQYP  
 LTGAYEDTKESYNLEHVKNISIPNGKLEAKAVNPYSSEGEHIREIQEGTLSKRISVNDLDHNKYKIELTVSG  
 KSI IKTINKDEPLDVVFLDNSNSMKNNNGKNNKAKKAGEAVETI IKDVLGANVENRAALVTYGSDFDGRTV  
 KVIKGFKEDPYYGLETSFTVQTDYSYKKFTNIAADI IKKIPKEAPEAKWGGTSLGLTPEKKREYDLSKVGE  
 25 TFTMKAFMEADTLLSSIQRKSRKIVHLTDGVPTRSYAINSFVTGSTYANQFERIKEKGYLDKNNYFITDDP  
 EKIKNGESYFLFPLDSYQTQIISGNLQKLHYLDLNLNYPKGTIYRNGPVREHGTPTKLYINSLKQKNYDIF  
 NFGIDISGFRQVYNEDYKKNQDGTQKLKEEAFELSDGEITELMNSFSSKPEYYTPIVTSADVSNNEILSKI  
 QQQFEKILTKENSIVNGTIEDPMGDKINLHLGNGQTLQPSDYTLQNDGSIMKDSIATGGPNNDGGILKGVK  
 LEYIKNKLYVRGLNLGEGQVTLTYDVKLDDSFISNKFYDTNGRITLNPKSEEPDTLRDFPIPKIRDVREYP  
 30 TITIKNEKKLGEIEFTKVVDKDNKLLKGFELQEFNEDYKLYLP IKNNSKVVTGENGKISYKDLKDGKY  
 QLIEAVSPKDYQKITNKPILTFEVVKGSIQNI IAVNKQISEYHEEGDKHLITNTHIPPKGIIPMTGGKGILS  
 FILIGGAMMSIAGGIYIWKRRHKSSDASIEKD

**Strain IC108**

ORF DNA sequence (SEQ ID NO: 208):

35 ATGAAAAAGAGACAAAAAATATGGAGAGGGTTATCAGTTACTTTACTAATCCTGTCCCAAATTCATTTGGT  
 ATATTGGTACAAGGTGAAACCCAAGATACCAATCAAGCACTTGGAAAAGTAATTGTAAAAAACGGGAGAC  
 AATGCTACACCATTAGGCAAAGCGACTTTTGTGTTAAAAAATGACAATGATAAGTCAGAAACAAGTCACGAA  
 ACGGTAGAGGGTCTGGAGAAGCAACCTTTGAAAACATAAAACCTGGAGACTACACATTAAGAGAAGAAACA  
 40 GCACCAATTGGTTATAAAAAAATGATAAAACCTGGAAAGTTAAAGTTGCAGATAACGGAGCAACAATAATC  
 GAGGGTATGGATGCAGATAAAGCAGAGAAACGAAAAGAAGTTTGAATGCCCAATATCCAAAATCAGCTATT  
 TATGAGGATACAAAAGAAAATTACCCATTAGTTAATGTAGAGGGTTCCAAAGTTGGTGAACAATACAAAGCA  
 TTGAATCCAATAAATGGAAAAGATGGTCGAAGAGAGATTGCTGAAGGTTGGTTATCAAAAAAATTACAGGG  
 GTCAATGATCTCGATAAGAATAAATATAAAATTTGAATTAACCTGTTGAGGGTAAAACCACTGTTGAAACGAAA  
 45 GAACTTAATCAACCCTAGATGTCGTTGTGCTATTAGATAATTCAAATAGTATGAATAATGAAAGAGCCAAT  
 AATTCTCAAAGAGCATTAAAAGCTGGGGAAGCAGTTGAAAAGCTGATTGATAAAATTACATCAATAAAGAC  
 AATAGAGTAGCTCTTGTGACATATGCCTCAACCTTTTTGATGGTACTGAAGCGACCGTATCAAAGGGAGTT  
 GCCGATCAAAATGGTAAAGCGCTGAATGATAGTGTATCATGGGATTATCATAAAACTACTTTTACAGCAACT  
 ACACATAATTACAGTTATTTAAATTTAAACAAATGATGCTAACGAAGTTAATATCTAAAGTCAAGAATCCA  
 50 AAGGAAGCGGAGCATATAAATGGGGATCGCACGCTCTATCAATTTGGTGCACATTTACTCAAAAAGCTCTA  
 ATGAAAGCAAATGAAATTTTAGAGACACAAAGTTCTAATGCTAGAAAAAACTTATTTTTCACGTAACCTGAT  
 GGTGTCCCTACGATGTCCTTATGCCATAAATTTTAATCCTTATATATCAACATCTTACCAAACCAAGTTAAT  
 TCTTTTTTAAATAAAATACCAGATAGAAGTGGTATTCTCCAAGAGGATTTTATAATCAATGGTGATGATTAT  
 CAAATAGTAAAAGGAGATGGAGAGAGTTTTAAACTGTTTTCGGATAGAAAAGTTCCCTGTTACTGGAGGAACG  
 55 ACACAAGCAGCTTATCGAGTACCGCAAAATCAACTCTCTGTAATGAGTAATGAGGGATATGCAATTAATAGT  
 GGATATATTTATCTCTATTGGAGAGATTACAACCTGGGTCTATCCATTTGATCCCTAAGACAAAGAAAGTTTCT  
 GCAACGAAACAAATCAAACTCATGGTGAGCCAACAACATTATACTTTAATGGAAATATAAGACCTAAAGGT  
 TATGACATTTTTACTGTTGGGATTGGTGTAAACGGAGATCCTGGTGCAACTCCTCTTGAAGCTGAGAAATTT  
 ATGCAATCAATATCAAGTAAAACAGAAAATTATACTAATGTTGATGATACAAAATAAAATTTATGATGAGCTA  
 60 AATAAATACTTTAAAACAATTGTTGAGGAAAACATTCTATTGTTGATGGAAATGTGACTGATCCTATGGGA



- 112 -

GAGATGATTGAATTCCAATTAATAAATGGTCAAAGTTTTACACATGATGATTACGTTTTGGTTGGAAATGAT  
 GGCAGTCAATTAATAAATGGTGTGGCTCTTGGTGGACCAAACAGTGTGGGGGAATTTTAAAAGATGTTACA  
 GTGACTTATGATAAGACATCTCAAACCATCAAAATCAATCATTGAACTTAGGAAGTGGACAAAAAGTAGTT  
 CTTACCTATGATGTACGTTTTAAAAGATAACTATATAAGTAACAAATTTTACAATACAAATAATCGTACAACG  
 5 CTAAGTCCGAAGAGTGAAAAAGAACCAAATACTATTCGTGATTTCCCAATTCCCAAAATTCGTGATGTTTCGT  
 GAGTTTCCGGTACTAACCATCAGTAATCAGAAGAAAATGGGTGAGGTTGAATTTATTAAAGTTAATAAAGAC  
 AAACATTCAGAATCGCTTTTGGGAGCTAAGTTTCAACTTCAGATAGAAAAAGATTTTTCTGGGTATAAGCAA  
 TTTGTTCCAGAGGGAAGTGATGTTACAACAAAGAATGATGGTAAAATTTATTTTAAAGCACTTCAAGATGGT  
 AACTATAAATTATATGAAATTTCAAGTCCAGATGGCTATATAGAGGTTAAAACGAAACCTGTTGTGACATTT  
 10 ACAATTCAAAATGGAGAAGTTACGAACCTGAAAGCAGATCCAAATGCTAATAAAAATCAAATCGGGTATCTT  
 GAAGGAAATGGTAAACATCTTATTACCAACACTCCCAAACGCCACCAGGTGTTTTCTTAAACAGGGGGA  
 ATTGGTACAATTGTCTATATATTAGTTGGTTCTACTTTTATGATACTTACCATTTGTTCTTTCCGTCGTAAA  
 CAATTGTAA

ORF amino acid sequence (SEQ ID NO: 189):

15 MKKRQKIWRGLSVTLILLISQIPFGILVQGETQDTNQLGKVIKKTGDNATPLGKATFVLKNDNDKSETSH  
 TVEGSGEATFENIKPGDYTLREETAPIGYKKTDKTWKVKVADNGATIIEGMDADKAEKRKEVLNAQYPKSAI  
 YEDTKENYPLVNVESKVGQYKALNPINGKDGREIAEGWLSKKITGVNDLDDKNKYKIELTVEGKTTVETK  
 ELNQPLDVVLLDNSNSMNERANNSQRALKAGEAVEKLIKITSNKDNRVALVTYASTIFDGTEATVSKGV  
 ADQNGKALNDSVSWDYHKTTFTATTHNYSYLNLTNDANEVNI LKSRIPKEAHINGDRTLYQFGATFTQKAL  
 20 MKANEILETQSSNARKKLI FHVTDGVPTMSYAINFNPIYISTSYQNQFNSFLNKIPDRSGILQEDFIINGDDY  
 QIVKGDGESFKLFSDRKVPVTGGTTQAAAYRVPQNQLSVMSNEGYAINSGYIYLYWRDYNWVYPFDPKTKKVS  
 ATKQIKTHGEPTTLYFNGNIRPKGYDIFTVIGVNGDPGATPLEAEKFMQSISSKTENYTNVDDTNKIYDEL  
 NKYFKTIVEEKHSIVDGNVTDPMGEMIEFQLKNGQSFTHDDYVLVGNDSQLKNGVALGGPNSDGGILKDV  
 VTYDKTSQTIKINHLNLGSGQKVVLTYDVRLKDNYSNKFYNTNNRTT LSPKSEKEPNTIRDFPIPKIRDVR  
 25 EFPVLTISNQKMGVEFEIKVNDKHSESLGAKFQLQIEKDFSGYKQFVPEGSVDVTTKNDGKIYFKALQDG  
 NYKLYEISSPDGYIEVKTKPVVFTTIQNGEVTNLKADPNANKNQIGYLEGNGKHLITNTPKRPPGVFPKTTG  
 IGTIVYILVGSFMILTICSFRRKQL

### Strain IC216

ORF DNA sequence (SEQ ID NO: 209):

30 ATGAAAAAGAGACAAAAATATGGAGAGGGTTATCAGTTACTTTACTAATCCTGTCCCAAATTCATTTGGT  
 ATATTGGTACAAGGTGAAACCCAAGATACCAATCAAGCACTTGGAAAAGTAATTGTTAAAAAACGGGAGAC  
 AATGCTACACCATTAGGCAAAGCGACTTTTGTGTTAAAAATGACAATGATAAGTCAGAAACAAGTCACGAA  
 ACGGTAGAGGGTTCTGGAGAAGCAACCTTTGAAAACATAAAACCTGGGAGACTACACATTAAGAGAAGAAACA  
 35 GCACCAATTGGTTATAAAAAACTGATAAAACCTGGAAAGTTAAAGTTGCAGATAACGGAGCAACAATAATC  
 GAGGGTATGGATGCAGATAAAGCAGAGAAACGAAAAGAAGTTTTGAATGCCCAATATCCAAAATCAGCTATT  
 TATGAGGATACAAAAGAAAATTACCCATTAGTTAATGTAGAGGGTTCCAAAGTTGGTGAACAATACAAAGCA  
 TTGAATCCAATAAATGAAAAGATGGTTCGAAGAGAGATTGCTGAAGGTTGGTTATCAAAAAAATTAACAGGG  
 GTCAATGATCTCGATAAGAATAAATATAAAATTGAATTAACCTGTTGAGGGTAAAACCACTGTTGAAACGAAA  
 40 GAACTTAATCAACCACTAGATGTCGTTGTGCTATTAGATAAATCAAATAGTATGAATAATGAAAGAGCCAAT  
 AATTCTCAAAGAGCATTAAAAGCTGGGGAAGCAGTTGAAAAGCTGATTGATAAAATTACATCAAATAAAGAC  
 AATAGAGTAGCTCTTGTGACATATGCCTCAACCATTTTTGATGGTACTGAAGCGACCGTATCAAAGGGAGTT  
 GCCGATCAAATGGTAAAGCGCTGAATGATAGTGTATCATGGGATTATCATAAACTACTTTTACAGCAACT  
 ACACATAATTACAGTTATTTAAATTTAACAAATGATGCTAACGAAGTTAATATCTAAAGTCAAGAATCCA  
 45 AAGGAAGCGGAGCATATAAATGGGGATCGCACGCTCTATCAATTTGGTGCACATTTACTCAAAAAGCTCTA  
 ATGAAAGCAAATGAAATTTTAGAGACACAAAGTTCTAATGCTAGAAAAAACTTATTTTTTACGTAACCTGAT  
 GGTGTCCCTACGATGTCTTATGCCATAAATTTAATCCTTATATATCAACATCTTACCAAACCAAGTTAAT  
 TCTTTTTTAAATAAAATACCAGATAGAAGTGGTATTTCTCCAAGAGGATTTTATAATCAATGGTGTGATTAT  
 CAAATAGTAAAAGGAGATGGAGAGAGTTTTAAACTGTTTTCGGATAGAAAAGTTCTGTACTGGAGGAACG  
 50 ACACAAGCAGCTTATCGAGTACCGCAAATCAACTCTCTGTAATGAGTAATGAGGGATATGCAATTAATAGT  
 GGATATATTTATCTCTATTGGAGAGATTACAACCTGGGTCTATCCATTTGATCCTAAGACAAAGAAAGTTTCT  
 GCAACGAAACAAATCAAACCTCATGGTGGAGCAACAACATTATACTTTAATGGAAATATAAGACCTAAAGGT  
 TATGACATTTTTACTGTTGGGATTGGTGTAAACGGAGATCCTGGTGCACCTCTTGAAGCTGAGAAATTT  
 ATGCAATCAATATCAAGTAAAACAGAAAATTATACTAATGTTGATGATACAAATAAAATTTATGATGAGCTA  
 55 AATAAATACTTTAAAACAATTGTTGAGGAAAAACATTCTATTGTTGATGGAAATGTGACTGATCCTATGGGA  
 GAGATGATTGAATTCCAATTAATAAATGGTCAAAGTTTTACACATGATGATTACGTTTTGGTTGGAAATGAT  
 GGCAGTCAATTAATAAATGGTGTGGCTCTTGGTGGACCAAACAGTGTGGGGGAATTTTAAAAGATGTTACA  
 GTGACTTATGATAAGACATCTCAAACCATCAAAATCAATCATTGAACTTAGGAAGTGGACAAAAAGTAGTT  
 CTTACCTATGATGTACGTTTTAAAAGATAACTATATAAGTAACAAATTTTACAATACAAATAATCGTACAACG  
 60 CTAAGTCCGAAGAGTGAAAAAGAACCAAATACTATTCGTGATTTCCCAATTCCCAAAATTCGTGATGTTTCGT



- 113 -

GAGTTTCCGGTACTAACCATCAGTAATCAGAAGAAAATGGGTGAGGTTGAATTTATTAAAGTTAATAAAGAC  
 AAACATTCAGAATCGCTTTTGGGAGCTAAGTTTCAACTTCAGATAGAAAAAGATTTTCTGGGTATAAGCAA  
 TTTGTTCCAGAGGGAAGTGATGTTACAACAAAGAATGATGGTAAAATTTATTTTAAAGCACTTCAAGATGGT  
 AACTATAAATTATATGAAATTTCAAGTCCAGATGGCTATATAGAGGTTAAAACGAAACCTGTTGTGACATTT  
 5 ACAATTCAAAATGGAGAAGTTACGAACCTGAAAGCAGATCCAAATGCTAATAAAAAATCAAATCGGGTATCTT  
 GAAGGAAATGGTAAACATCTTATTACCAACACTCCCAAACGCCACCAGGTGTTTTTCTAAAACAGGGGGA  
 ATTGGTACAATTGTCTATATATTAGTTGGTTCTACTTTTATGATACTTACCATTTGTTCTTTCCGTCGTAAA  
 CAATTGTAA

ORF amino acid sequence (SEQ ID NO: 190):

10 MKKRQKIWRGLSVTLILLSQIPFGILVQGETQDTNQLGKVIKKTGDNATPLGKATFVLKNDNDKSETSHE  
 TVEGSGEATFENIKPGDYTLREETAPIGYKKTDKTWKVKVADNGATIIEGMDADKAEKRKEVLNAQYPKSAI  
 YEDTKENYPLVNVEGSKVGEQYKALNPINGKDGREIAEGWLSKKITGVNDLKNKYKIELTVEGKTTVETK  
 ELNQPLDVVLLDNSNSMNERANNSQRALKAGEAVEKLIKITSNKDNRVALVTYASTIFDGTEATVSKGV  
 ADQNGKALNDSVSWDYHKTTFTATTHNYSYLNLTNDANEVNI LKSRI PKEAEHINGDRTLYQFGATFTQKAL  
 15 MKANEI LETQSSNARKKLI FHVTDGVPTMSYAINFNPI YISTSYQNQFNSFLNKI PDRSGILQEDFI INGDDY  
 QIVKGDGESFKLFS DRKVPVTGGTTQAA YRVPQNQL SVMSNEGYA INSGYI YLYWRDYNWVY PFDPKTKKVS  
 ATKQIKTHGEPTTLYFNGNIRPKGYDI FTVGIGVNGDPGAT PLEAEKFMQSI SSKTENYTNVDDTNKI YDEL  
 NKYFKTIVEEKHSIVDGNVTDPMGEMIEFQLKNGQSFTHDDYVLVGNDSQLKNGVALGGPNSDGGILKDV  
 VTYDKTSQTIKINHLNLGSGQKVVLTYDVRLKDNYSNKFYNTNRTT LSPKSEKEPNTIRDFPIPKIRDVR  
 20 EFPVLTISNQKMGVEFEIKV NKDKHSESLLGAKFQLQIEKDFSGYKQFVPEGSDVTTKNDGKIYFKALQDG  
 NYKLYEISSPDGYIEVKTKPVVFTIQNGEVTNLKADPNANKNQIGYLEGNGKHLITNTPKRPPGVFPKTGG  
 IGTIVYILVGSTFMILTICSFRRKQL

### Strain IC244

ORF DNA sequence (SEQ ID NO: 210):

25 ATGAAAAAGAGACAAAAAATATGGAGAGGGTTATCAGTTACTTTACTAATCCTGTCCCAAATTCATTTGGT  
 ATATTGGTACAAGGTGAAACCCAAGATACCAATCAAGCACTTGAAAAGTAATTGTTAAAAAACGGGAGAC  
 AATGCTACACCATTAGGCAAAGCGACTTTTGTGTTAAAAAATGACAATGATAAGTCAGAAACAAGTCACGAA  
 ACGGTAGAGGGTCTGGAGAAGCAACCTTTGAAAACATAAAACCTGGGAGACTACACATTAAGAGAAGAAACA  
 30 GCACCAATTGGTTATAAAAAAAGTATAAAACCTGGAAAGTTAAAGTTGCAGATAACGGAGCAACAATAATC  
 GAGGGTATGGATGCAGATAAAGCAGAGAAACGAAAAGAAGTTTGAATGCCCAATATCCAAAATCAGCTATT  
 TATGAGGATACAAAAGAAAATTACCCATTAGTTAATGTAGAGGGTTCCAAAGTTGGTGAACAATACAAAGCA  
 TTGAATCCAATAAATGAAAAGATGGTCGAAGAGAGATTGCTGAAGGTTGGTTATCAAAAAAATTACAGGG  
 GTCAATGATCTCGATAAGAATAAATATAAAATTGAATTAAC TGTGAGGGTAAAACCACTGTTGAAACGAAA  
 35 GAACTTAATCAACCACTAGATGTCGTTGTGCTATTAGATAATTCAAATAGTATGAATAATGAAAGAGCCAAT  
 AATCTCAAAGAGCATTAAAAGCTGGGGAAGCAGTTGAAAAGCTGATTGATAAAATTACATCAAATAAAGAC  
 AATAGAGTAGCTCTTGTGACATATGCCTCAACCAATTTTTGATGGTACTGAAGCGACCGTATCAAAGGGAGTT  
 GCCGATCAAATGGTAAAGCGCTGAATGATAGTGTATCATGGGATTATCATAAAACTACTTTTACAGCAACT  
 ACACATAATTACAGTTATTTAAATTTAACAAATGATGCTAACGAAGTTAATATCTAAAGTCAAGAATTCCA  
 40 AAGGAAGCGGAGCATATAAATGGGGATCGCACGCTCTATCAATTTGGTGCAGCATTTACTCAAAAAGCTCTA  
 ATGAAAGCAAATGAAATTTTAGAGACACAAAGTTCTAATGCTAGAAAAAACTTATTTTTACGTAAGTATGAT  
 GGTGTCCCTACGATGTCCTTATGCCATAAATTTAATCCTTATATATCAACATCTTACCAAACCAAGTTAAT  
 TCTTTTTTAAATAAAAATACCAGATAGAAGTGGTATTCTCCAAGAGGATTTTATAATCAATGGTGATGATTAT  
 CAAATAGTAAAAGGAGATGGAGAGAGTTTTAAACTGTTTTCGGATAGAAAAGTTCCCTGTACTGGAGGAACG  
 45 ACACAAGCAGCTTATCGAGTACCGCAAAATCAACTCTCTGTAATGAGTAATGAGGGATATGCAATTAATAGT  
 GGATATATTTATCTCTATTGGAGAGATTACAACCTGGGTCTATCCATTTGATCCCTAAGACAAAGAAAGTTTCT  
 GCAACGAAACAAATCAAACCTCATGGTGAGCCAACAACATTATACTTTAATGGAAATATAAGACCTAAAGGT  
 TATGACATTTTTACTGTTGGGATTGGTGTAACGGAGATCCTGGTGCAACTCCTCTTGAAGCTGAGAAATTT  
 ATGCAATCAATATCAAGTAAAACAGAAAATTATACTAATGTTGATGATACAAATAAAATTTATGATGAGCTA  
 50 AATAAATACTTTAAAACAATTGTTGAGGAAAACATTCATTTGTTGATGGAAATGTGACTGATCCTATGGGA  
 GAGATGATTGAATTCCAATTAATAAATGGTCAAAGTTTTACACATGATGATTACGTTTTGGTTGGAAATGAT  
 GGCAGTCAATTAATAAATGGTGTGGCTCTTGGTGGACCAAACAGTGTGGGGGAATTTTAAAAGATGTTACA  
 GTGACTTATGATAAGACATCTCAAACCATCAAATCAATCATTTGAACTTAGGAAGTGGACAAAAGTAGTT  
 CTTACCTATGATGTACGTTTTAAAAGATAACTATATAAGTAACAAATTTTACAATACAAATAATCGTACAACG  
 55 CTAAGTCCGAAGAGTGAAAAGAACCAAATACTATTCGTGATTTCCCAATTCCCAAAATTCGTGATGTTCTGT  
 GAGTTTCCGGTACTAACCATCAGTAATCAGAAGAAAATGGGTGAGGTTGAATTTATTAAAGTTAATAAAGAC  
 AAACATTCAGAATCGCTTTTGGGAGCTAAGTTTCAACTTCAGATAGAAAAAGATTTTTCTGGGTATAAGCAA  
 TTTGTTCCAGAGGGAAGTGATGTTACAACAAAGAATGATGGTAAAATTTATTTTAAAGCACTTCAAGATGGT  
 AACTATAAATTATATGAAATTTCAAGTCCAGATGGCTATATAGAGGTTAAAACGAAACCTGTTGTGACATTT  
 60 ACAATTCAAAATGGAGAAGTTACGAACCTGAAAGCAGATCCAAATGCTAATAAAAAATCAAATCGGGTATCTT



- 114 -

GAAGGAAATGGTAAACATCTTATTACCAACACTCCCAAACGCCACCAGGTGTTTTTCTAAAACAGGGGGA  
ATTGGTACAATTGTCTATATATTAGTTGGTTCTACTTTTATGATACTTACCATTTGTTCTTTCCGTCGTAAA  
CAATTGTAA

ORF amino acid sequence (SEQ ID NO: 191):

5 MKKRQKIWRGLSVTLLILSQIPFGILVQGETQDTNQLGKVIVKKTGDNATPLGKATFVLKNDNDKSETSHE  
TVEGSGEATFENIKPGDYTLREETAPIGYKKTDKTWKVKVADNGATIIEGMDADKAEKRKEVLNAQYPKSAI  
YEDTKENYPLVNVESKVGQYKALNPINGKDGREIAEGWLSKKITGVNDLDDKNKYKIELTVEGKTTVETK  
ELNQLDVLVVLLDNSNSMNNERANNSQRALKAGEAVEKLIIDKITSNKDNRVALVTYASTIFDGTEATVSKGV  
10 ADQNGKALNDSVSWDYHKTTFTATTHNYSYLNLTNDANEVNI LKSRI PKEAEHINGDRTLYQFGATFTQKAL  
MKANEI LETQSSNARKKLI FHVTDGVP TMSYAINFNPI YISTSYQNQFNSFLNKI PDRSGILQEDFI INGDDY  
QIVKGDGESFKLFS DRKVPVTGGTTQAA YRVPQNQLS VMSNEGYAINSGYI YLYWRDYNWVY PFDPKTKKVS  
ATKQIKTHGEPTTLYFNGNIRPKGYDI FTVGIGVNGDPGATPLEAEKFMQSI SSKTENYTNVDDTNKIYDEL  
NKYFKTIVEEKHSIVDGNVTDPMGEMIEFQLKNGQS FTHDDYVLVGN DGSQ LKNGVALGGPNSDGGILKDV  
15 VTYDKTSQTIKINHLNLGSGQKVVLTYDVRLK DNYI SNKFYNTNRTT LSPKSEKEPNTIRDFPIPKIRDVR  
EFPVLTISNQKMGVEFEIKV NKDKHSESLGAKFQLQIEKDFSGYKQFVPEGS DVTTKNDGKIYFKALQDG  
NYKLYEISSPDGYIEVKTKPVVFTTIQNGEVTNLKADPNANKNQIGYLEGNGKHLITNTPKRPPGVFPKTGG  
IGTIVYILVGSTFMILTICSFRRKQL

### Strain IC245

20 ORF DNA sequence (SEQ ID NO: 211):  
ATGAAAAAGAGACAAAAAATATGGAGAGGGTTATCAGTTACTTTACTAATCCTGTCCCAAATTCATTTGGT  
ATATTGGTACAAGGTGAAACCCAAGATACCAATCAAGCACTTGGAAAAGTAATTGTTAAAAAACGGGAGAC  
AATGCTACACCATTAGGCAAAGCGACTTTTGTGTTAAAAAATGACAATGATAAGTCAGAAACAAGTCACGAA  
ACGGTAGAGGGTTCTGGAGAAGCAACCTTTGAAAACATAAAACCTGGGAGACTACACATTAAGAGAAGAAACA  
25 GCACCAATTGGTTATAAAAAA ACTGATAAAACCTGGAAAGTTAAAGTTGCAGATAACGGAGCAACAATAATC  
GAGGGTATGGATGCAGATAAAGCAGAGAAACGAAAAGAAGTTTTGAATGCCCAATATCCAAAATCAGCTATT  
TATGAGGATACAAAAGAAAATTACCCATTAGTTAATGTAGAGGGTTCCAAAGTTGGTGAACAATACAAAGCA  
TTGAATCCAATAAATGGAAAAGATGGTCGAAGAGAGATTGCTGAAGGTTGGTTATCAAAAAAATTACAGGG  
GTCAATGATCTCGATAAGAATAAATAAATAAATTTGAATTA ACTGTTGAGGGTAAAACCACTGTTGAAACGAAA  
30 GAACTTAATCAACCACTAGATGTCGTTGTGCTATTAGATAATTCAAATAGTATGAATAATGAAAGAGCCAAT  
AATTCTCAAAGAGCATTAAAAGCTGGGGAAGCAGTTGAAAAGCTGATTGATAAAATTACATCAAATAAAGAC  
AATAGAGTAGCTCTTGTGACATATGCCTCAACCATTTTTGATGGTACTGAAGCGACCGTATCAAAGGGAGTT  
GCCGATCAAATGGTAAAGCGCTGAATGATAGTGTATCATGGGATTATCATAAACTACTTTTACAGCAACT  
ACACATAATTACAGTTATTTAAATTTAACAAATGATGCTAACGAAGTTAATATTTCTAAAGTCAAGAATTTCCA  
35 AAGGAAGCGGAGCATATAAATGGGGATCGCACGCTCTATCAATTTGGTGCACATTTACTCAAAAAGCTCTA  
ATGAAAGCAAATGAAATTTTAGAGACACAAAGTTCTAATGCTAGAAAAAACTTATTTTTTACGTAAGTATGAT  
GGTGTCCCTACGATGTCTTATGCCATAAATTTTAATCCTTATATATCAACATCTTACCAAACCAGTTTAAAT  
TCTTTTTTAAATAAATAACCAGATAGAAGTGGTATTCTCCAAGAGGATTTTATAATCAATGGTGTGATTAT  
CAAATAGTAAAAGGAGATGGAGAGAGTTTTAAACTGTTTTTCGGATAGAAAAGTTCTGTACTGGAGGAACG  
40 ACACAAGCAGCTTATCGAGTACCGCAAATCAACTCTCTGTAATGAGTAATGAGGGATATGCAATTAATAGT  
GGATATATTTATCTCTATTGGAGAGATTACAACCTGGGTCTATCCATTTGATCCTAAGACAAAGAAAGTTTCT  
GCAACGAAACAAATCAAACCTCATGGTGAGCCAACAACATTATACTTTAATGGAAATATAAGACCTAAAGGT  
TATGACATTTTTACTGTTGGGATTGGTGTAACGGGAGATCCTGGTGCAACTCCTCTTGAAGCTGAGAAATTT  
ATGCAATCAATATCAAGTAAAACAGAAAATTTACTAATGTTGATGATACAAATAAAATTTATGATGAGCTA  
45 AATAAATACTTTAAAACAATTGTTGAGGAAAACATTCTATTGTTGATGGAAATGTGACTGATCCTATGGGA  
GAGATGATTGAATTTCAATTA AAAAATGGTCAAAGTTTTACACATGATGATTACGTTTTGGTTGGAAATGAT  
GGCAGTCAATTA AAAAATGGTGTGGCTCTTGGTGGACCAAACAGTGTGGGGAAATTTAAAAGATGTTACA  
GTGACTTATGATAAGACATCTCAAACCATCAAATCAATCAATTTGAACTTAGGAAGTGGACAAAAGTAGTT  
CTTACCTATGATGTACGTTTTAAAAGATAACTATATAAGTAACAAATTTTACAATACAAATAATCGTACAACG  
50 CTAAGTCCGAAGAGTGAAAAGAACCAAATACTATTCGTGATTTCCCAATTCCAAATTCGTGATGTTCTGT  
GAGTTTCCGGTACTAACCATCAGTAATCAGAAGAAAATGGGTGAGGTTGAATTTATTAAGTTAATAAAGAC  
AAACATTCAGAATCGCTTTTGGGAGCTAAGTTTCAACTTCAGATAGAAAAGATTTTTCTGGGTATAAGCAA  
TTTGTTCAGAGGGAAGTGATGTTACAACAAAGAATGATGGTAAAATTTATTTTAAAGCACTTCAAGATGGT  
AACTATAAATTATATGAAATTTCAAGTCCAGATGGCTATATAGAGGTTAAAACGAAACCTGTTGTGACATTT  
55 ACAATTCAAAATGGAGAAGTTACGAACCTGAAAGCAGATCCAAATGCTAATAAAAATCAAATCGGGTATCTT  
GAAGGAAATGGTAAACATCTTATTACCAACACTCCCAAACGCCACCAGGTGTTTTTCTAAAACAGGGGGA  
ATTGGTACAATTGTCTATATATTAGTTGGTTCTACTTTTATGATACTTACCATTTGTTCTTTCCGTCGTAAA  
CAATTGTAA

ORF amino acid sequence (SEQ ID NO: 192):



MKKRQKIWRGLSVTLLILSQIPFGILVQGETQDTNQALGKVIKKTGDNATPLGKATFVLKNDNDKSET SHE  
 TVEGSGEATFENIKPGDYTLREETAPIGYKKTDKTWKVKVADNGATIIEGMDADKAEKRKEVLNAQYPKSAI  
 YEDTKENYPLVNVEGSKVGEQYKALNPINGKDGREIAEGWLSKKITGVNDLNDKNKYKIELTVEGKTTVETK  
 5 ELNQLDQVLLDNSNSMNERANNSQRALKAGEAVEKLIKITSNKDNRVALVYASTIFDGTEATVSKGV  
 ADQNGKALNDSVSWDYHKTTFTATTHNYSYLNLTNDANEVNIILKSRI PKEAEHINGDRTLYQFGATFTQKAL  
 MKANEI LETQSSNARKKLI FHVTDGVPTMSYAINFNPIYISTSYQNQFNSFLNKIPDRSGILQEDFIINGDDY  
 QIVKGDGESFKLFSDRKVPVTGGTTQAAYRVPQNQLSVMSNEGYAINSGYIYLYWRDYNWVYPFDPKTKKVS  
 ATKQIKTHGEPTTLYFNGNIRPKGYDIFTVIGVNGDPGATPLEAEKFMQSISSKTENYTNVDDTNKIYDEL  
 10 NKYFKTIVEEKHSIVDGNVTDPMGEMIEFQLKNGQSFTHDDYVLVGNDSQLKNGVALGGPNSDGGILKDV  
 VTYDKTSQTIKINHLNLGSGQKVVLTYDVRDKNYISNKFYNTNNRTTLPKSEKEPNTIRDFPIPKIRDVR  
 EFPVLTISNQKMGVEVEFIKVNKDKHSESLGAKFQLQIEKDFSGYKQFVPEGSVDTTKNDGKIYFKALQDG  
 NYKLYEISSPDGYIEVKTTPVVTFTIQNGEVTNLKADPNANKNQIGYLEGNGKHLITNTPKRPPGVFPKTGG  
 IGTIVYILVGSFTMILTICSFRRKQL

15

**Strain IC246**

ORF DNA sequence (SEQ ID NO: 212):

ATGAAAAAGAGACAAAAAATATGGAGAGGGTTATCAGTTACTTTACTAATCCTGTCCCAAATTCATTGGT  
 ATATTGGTACAAGGTGAAACCCAAGATACCAATCAAGCACTTGGAAAAGTAATTGTTAAAAAACGGGAGAC  
 AATGCTACACCATTAGGCAAAGCGACTTTTGTGTTAAAAAATGACAATGATAAGTCAGAAACAAGTCACGAA  
 20 ACGGTAGAGGGTCTGGAGAAGCAACCTTTGAAAACATAAAACCTGGAGACTACACATTAAGAGAAGAAACA  
 GCACCAATTGGTTATAAAAAAAGTATAAAACCTGGAAAGTTAAAGTTGCAGATAACGGAGCAACAATAATC  
 GAGGGTATGGATGCAGATAAAGCAGAGAAACGAAAAGAAGTTTTGAATGCCAATATCCAAAATCAGCTATT  
 TATGAGGATACAAAAGAAAATTACCCATTAGTTAATGTAGAGGGTTCCAAAGTTGGTGAACAATACAAAGCA  
 TTGAATCCAATAAATGGAAAAGATGGTCAAGAGAGATTGCTGAAGGTTGGTTATCAAAAAAATAACAGGG  
 25 GTCAATGATCTCGATAAGAATAAATAAATAAATTTGAATTAAGTGTGAGGGTAAAACCACTGTTGAAACGAAA  
 GAACTTAATCAACCACTAGATGTCGTTGTGCTATTAGATAATTCAAATAGTATGAATAATGAAAGAGCCAAT  
 AATTCTCAAAGAGCATTAAAAGCTGGGGAAGCAGTTGAAAAGCTGATTGATAAAATTACATCAAATAAAGAC  
 AATAGAGTAGCTCTTGTGACATATGCCTCAACATTTTTGATGGTACTGAAGCGACCGTATCAAAGGGAGTT  
 GCCGATCAAAATGGTAAAGCGCTGAATGATAGTGTATCATGGGATTATCATAAAACTACTTTTACAGCAACT  
 30 ACACATAATTACAGTTATTTAAATTTAACAAATGATGCTAACGAAGTTAATATTCTAAAGTCAAGAATTCCA  
 AAGGAAGCGGAGCATATAAATGGGGATCGCACGCTCTATCAATTTGGTGCAGACATTTACTCAAAAAGCTCTA  
 ATGAAAGCAAATGAAATTTTAGAGACACAAAGTTCTAATGCTAGAAAAAACTTATTTTTTACGTAAGTATGAT  
 GGTGTCCCTACGATGTCTTATGCCATAAATTTTAATCCTTATATATCAACATCTTACCAAACCAAGTTAAT  
 TCTTTTTTAAATAAATAACCAGATAGAAGTGGTATTCTCCAAGAGGATTTTATAATCAATGGTGTGATTAT  
 35 CAAATAGTAAAAGGAGATGGAGAGAGTTTTAAACTGTTTTTCGGATAGAAAAGTTCTGTACTGGAGGAACG  
 ACACAAGCAGCTTATCGAGTACCGCAAATCAACTCTCTGTAATGAGTAATGAGGGATATGCAATTAATAGT  
 GGATATATTTATCTCTATTGGAGAGATTACAACCTGGGTCTATCCATTTGATCCTAAGACAAAGAAAGTTTCT  
 GCAACGAAACAAATCAAACTCATGGTGAGCCAACAACATTATACTTTAATGGAAATATAAGACCTAAAGGT  
 TATGACATTTTTACTGTTGGGATTGGTGTAAACGGAGATCCTGGTGCACCTCTTGAAGCTGAGAAATTT  
 40 ATGCAATCAATATCAAGTAAAACAGAAAATTATACTAATGTTGATGATACAAATAAATTTATGATGAGCTA  
 AATAAATACTTTAAAACAATTGTTGAGGAAAACATTCTATTGTTGATGGAAATGTGACTGATCCTATGGGA  
 GAGATGATTGAATTCCAATTAATAAATGGTCAAAGTTTTACACATGATGATTACGTTTTGGTTGGAAATGAT  
 GGCAGTCAATTAATAAATGGTGTGGCTCTTGGTGGACCAAACAGTGTGGGGGAATTTTAAAAGATGTTACA  
 GTGACTTATGATAAGACATCTCAAACCATCAAATCAATCATTGAACTTAGGAAGTGGACAAAAGTAGTT  
 45 CTTACCTATGATGTACGTTTTAAAAGATAACTATATAAGTAACAAATTTTACAATACAAATAATCGTACAACG  
 CTAAGTCCGAAGAGTGAAAAGAACCAAATACTATTCGTGATTTCCCAATTCCAAATTCGTGATGTTTCGT  
 GAGTTTCCGGTACTAACCATCAGTAATCAGAAGAAAATGGGTGAGGTTGAATTTATTAAGTTAATAAAGAC  
 AAACATTCAGAATCGCTTTTGGGAGCTAAGTTTCAACTTCAGATAGAAAAGATTTTTCTGGGTATAAGCAA  
 TTTGTTCCAGAGGGAAGTGTGTTACAACAAAGAATGATGGTAAAATTTATTTTAAAGCACTTCAAGATGGT  
 50 AACTATAAATTATATGAAATTTCAAGTCCAGATGGCTATATAGAGGTTAAAACGAAACCTGTTGTGACATTT  
 ACAATTCAAAATGGAGAAGTTACGAACCTGAAAGCAGATCCAAATGCTAATAAATAAATAAATCGGGTATCTT  
 GAAGGAAATGGTAAACATCTTATTACCAACACTCCCAAACGCCACCAGGTGTTTTTCTAAAACAGGGGGA  
 ATTGGTACAATTGTCTATATATTAGTTGGTTCTACTTTTATGATACTTACCATTTGTTCTTTCCGTCGTAAA  
 CAATTGTAA

55

ORF amino acid sequence (SEQ ID NO: 193):

MKKRQKIWRGLSVTLLILSQIPFGILVQGETQDTNQALGKVIKKTGDNATPLGKATFVLKNDNDKSET SHE  
 TVEGSGEATFENIKPGDYTLREETAPIGYKKTDKTWKVKVADNGATIIEGMDADKAEKRKEVLNAQYPKSAI  
 YEDTKENYPLVNVEGSKVGEQYKALNPINGKDGREIAEGWLSKKNTGVNDLNDKNKYKIELTVEGKTTVETK  
 60 ELNQLDQVLLDNSNSMNERANNSQRALKAGEAVEKLIKITSNKDNRVALVYASTIFDGTEATVSKGV  
 ADQNGKALNDSVSWDYHKTTFTATTHNYSYLNLTNDANEVNIILKSRI PKEAEHINGDRTLYQFGATFTQKAL



MKANEIILETQSSNARKKLI FHVTDGVPTMSYAINFNPIYISTSYQNQFN SFLNKIPDRSGILQEDFI INGDDY  
 QIVKGDGESFKLFS DRKVPVTGGTTQAAYRVPQNQLSVMSNEGYAINSGYIYLYWRDYNWVYPFDPKTKKVS  
 ATKQIKTHGEPTTLYFNGNIRPKGYDIFTVGIGVNGDPGATPLEAEKFMQSISSKTENYTNVDDTNKIYDEL  
 NKYFKTIVEEKHSIVDGNVTDPMGEMIEFQLKNGQSFTHDDYVLVGN DGSQ LKNGVALGGPNSDGGILKDVT  
 5 VTYDKTSQTIKINHLNLGSGQKVVLTYDVRLKDNYSNKFYNTNNRTT LSPKSEKEPNTIRDFPIPKIRDVR  
 EFPVLTISNQKMGVEFEFIKVNKDKHSESLLGAKFQLQIEKDFSGYKQFVPEGS DVTTKNDGKIYFKALQDG  
 NYKLYEISSPDGYIEVKT KPVVFTFIQNGEVTNLKADPNANKNQIGYLEGNGKHLITNTPKRPPGVFPKTGG  
 IGTIVYILVGSTFMILTICSFRRKQL

10 **Strain IC247**

ORF DNA sequence (SEQ ID NO: 213):

ATGAAAAGAGACAAAAATATGGAGAGGGTTATCAGTTACTTTACTAATCCTGTCCCAAATTC CATT TGGT  
 ATATTGGTACAAGGTGAAACCCAAGATACCAATCAAGCACTTGGAAAAGTAATTGTTAAAAAACGGGAGAC  
 AATGCTACACCATTAGGCAAAGCGACTTTTGTGTTAAAAATGACAATGATAAGTCAGAAACAAGTCACGAA  
 15 ACGGTAGAGGGTTCTGGAGAAGCAACCTTTGAAAACATAAAACCTGGAGACTACACATTAAGAGAAGAAACA  
 GCACCAATTGGTTATAAAAAA ACTGATAAAACCTGGAAAAGTTAAAGTTGCAGATAACGGAGCAACAATAATC  
 GAGGGTATGGATGCAGATAAAGCAGAGAAACGAAAAGAAGTTTTGAATGCCCAATATCCAAAATCAGCTATT  
 TATGAGGATACAAAAGAAAATTACCCATTAGTTAATGTAGAGGGTTCCAAAGTTGGTGAACAATACAAAGCA  
 TTGAATCCAATAAATGGAAAAGATGGTCGAAGAGAGATTGCTGAAGGTTGGTTATCAAAAAAATTACAGGG  
 20 GTCAATGATCTCGATAAGAATAAATATAAAATTTGAATTAAC TGTGAGGGTAAAACCACTGTTGAAACGAAA  
 GAACCTAATCAACCCTAGATGTCGTTGTGCTATTAGATAATTCAAATAGTATGAATAATGAAAGAGCCAAT  
 AATTCTCAAAGAGCATTAAAAGCTGGGGAAGCAGTTGAAAAGCTGATTGATAAAATTACATCAAATAAAGAC  
 AATAGAGTAGCTCTTGTGACATATGCCTCAACCTTTTTGATGGTACTGAAGCGACCGTATCAAAGGGAGTT  
 GCCGATCAAATGGTAAAGCGCTGAATGATAGTGTATCATGGGATTATCATAAAACTACTTTTACAGCAACT  
 25 ACACATAATTACAGTTATTTAAATTTAACAAATGATGCTAACGAAGTTAATATTTCTAAAGTCAAGAATCCA  
 AAGGAAGCGGAGCATATAAATGGGGATCGCACGCTCTATCAATTTGGTGCACATTTACTCAAAAAGCTCTA  
 ATGAAAGCAAATGAAATTTTAGAGACACAAAGTTCTAATGCTAGAAAAAACTTATTTTTCACGTAAC TGAT  
 GGTGTCCCTACGATGTCCTTATGCCATAAATTTTAATCCCTTATATATCAACATCTTACCAAACCAGTTTAAAT  
 TCTTTTTTAAATAAAATACCAGATAGAAGTGGTATTTCTCCAAGAGGATTTTATAATCAATGGTGATGATTAT  
 30 CAAATAGTAAAAGGAGATGGAGAGAGTTTTAAACTGTTTTTCGGATAGAAAAGTTCCCTGTTACTGGAGGAACG  
 ACACAAGCAGCTTATCGAGTACCGCAAATCAACTCTCTGTAATGAGTAATGAGGGATATGCAATTAATAGT  
 GGATATATTTATCTCTATTGGAGAGATTACAACCTGGGTCTATCCATTTGATCCCTAAGACAAAGAAAGTTTCT  
 GCAACGAAACAAATCAAACCTCATGGTGAGCCAACAACATTATACTTTAATGGAAATATAAGACCTAAAGGT  
 TATGACATTTTACTGTTGGGATTGGTGTAAACGGAGATCCTGGTGCACCTCCTCTTGAAGCTGAGAAATTT  
 35 ATGCAATCAATATCAAGTAAAACAGAAAATTATACTAATGTTGATGATACAAATAAAATTTATGATGAGCTA  
 AATAAATACTTTAAAACAATTGTTGAGGAAAACATTTCTATTGTTGATGGAAATGTGACTGATCCCTATGGGA  
 GAGATGATTGAATTCCAATTA AAAAATGGTCAAAGTTTTACACATGATGATTACGTTTTGGTTGGAAATGAT  
 GGCAGTCAATTA AAAAATGGTGTGGCTCTTGGTGGACCAAACAGTGTGGGGGAATTTTAAAAGATGTTACA  
 GTGACTTATGATAAGACATCTCAAACCATCAAATCAATCATTGAACTTAGGAAGTGGACAAAAGTAGTT  
 40 CTTACCTATGATGTACGTTTTAAAAGATAACTATATAAGTAACAAATTTTACAATACAAATAATCGTACAACG  
 CTAAGTCCGAAGAGTGAAAAGAACCAAATACTATTCGTGATTTCCCAATTC CCAAATTCGTGATGTTCTGT  
 GAGTTTCCGGTACTAACCATCAGTAATCAGAAGAAAATGGGTGAGGTTGAATTTATTAAAGTTAATAAAGAC  
 AAACATTCAGAATCGCTTTTGGGAGCTAAGTTTCAACTTCAGATAGAAAAGATTTTTCTGGGTATAAGCAA  
 TTTGTTCCAGAGGAAGTGTGTTACAACAAAGAATGATGGTAAAATTTATTTTAAAGCACTTCAAGATGGT  
 45 AACTATAAATTATATGAAATTTCAAGTCCAGATGGCTATATAGAGGTTAAAACGAAACCTGTTGTGACATTT  
 ACAATTCAAAATGGAGAAGTTACGAACCTGAAAGCAGATCCAAATGCTAATAAAAATCAAATCGGGTATCTT  
 GAAGGAAATGGTAAACATCTTATTACCAACACTCCCAAACGCCACCAGGTGTTTTCTTAAAACAGGGGGA  
 ATTTGGTACAATTGTCTATATATTAGTTGGTTCTACTTTTATGATACTTACCATTTGTTCTTTCCGTCGTAA  
 CAATTGTAA

50 **ORF amino acid sequence (SEQ ID NO: 194):**

MKKRQKIWRGLSVTLLILSQIPFGILVQGETQDTNQALGKVIKKTGDNATPLGKATFVLKNDNDKSET SHE  
 TVEGSGEATFENIKPGDYTLREETAPIGYKKTDKT WKVKVADNGATIIEGMDADKA EKRKEVLNAQYPKSAI  
 YEDTKENYPLVNVEGSKVGEQYKALNPINGKDGREIAEGWLSKKITGVNDL DKNKYKIELTVEGKTTVETK  
 ELNQPLDVVLLDNSNSMNNERANNSQRALKAGEAVEKLI DKITSNKDNRVALVTYASTIFDGTEATVSKGV  
 55 ADQNGKALNDSVSWDYHKTTFTATTHNYSYLNLTNDANEVNI LKSRI PKEAEHINGDRTLYQFGATFTQKAL  
 MKANEIILETQSSNARKKLI FHVTDGVPTMSYAINFNPIYISTSYQNQFN SFLNKIPDRSGILQEDFI INGDDY  
 QIVKGDGESFKLFS DRKVPVTGGTTQAAYRVPQNQLSVMSNEGYAINSGYIYLYWRDYNWVYPFDPKTKKVS  
 ATKQIKTHGEPTTLYFNGNIRPKGYDIFTVGIGVNGDPGATPLEAEKFMQSISSKTENYTNVDDTNKIYDEL  
 NKYFKTIVEEKHSIVDGNVTDPMGEMIEFQLKNGQSFTHDDYVLVGN DGSQ LKNGVALGGPNSDGGILKDVT  
 60 VTYDKTSQTIKINHLNLGSGQKVVLTYDVRLKDNYSNKFYNTNNRTT LSPKSEKEPNTIRDFPIPKIRDVR



EFPVLTISNQKKMGEVEFIKVNKDKHSESLLGAKFQLQIEKDFSGYKQFVPEGSDVTTKNDGKIYFKALQDG  
 NYKLYEISSPDGYIEVKTKPVVFTTIQNGEVTNLKADPNANKNQIGYLEGNGKHLITNTPKRPPGVFPKTGG  
 IGTIVYILVGSTFMILTICSFRRKQL

5 **Strain IC250**

ORF DNA sequence (SEQ ID NO: 214):

ATGAGAAAATACCAAAAATTTTCTAAAATATTGACGTTAAGTCTTTTTTGTTCGCAAATACCGCTTAAT  
 ACCAATGTTTTAGGGGAAAGTACCGTACCGGAAAATGGTGCTAAAGGAAAGTTAGTTGTTAAAAAGACAGAT  
 GACCAGAACAAACCACCTTTCAAAGCTACCTTTGTTTTAAAACTACTGCTCAACCAGAAAGTAAAATAGAA  
 10 AAAGTAACTGCTGAGCTAACAGGTGAAGCTACTTTTGATAATCTCATACCTGGAGATTATACTTTATCAGAA  
 GAAACAGCGCCCGAAGGTTATAAAAAGACTAACCAGACTTGGCAAGTTAAGGTTGAGAGTAATGGAAAACT  
 ACGATACAAAATAGTGGTGATAAAAATTCACAATTTGGACAAAATCAGGAAGAACTAGATAAGCAGTATCCC  
 CCCACAGGAATTTATGAAGATACAAAGGAATCTTATAAACTTGAGCATGTTAAAGGTTTCAGTTCCAAATGGA  
 AAGTCAGAGGCAAAGCAGTTAACCATATTCAAGTGAAGGTGAGCATATAAGAGAAATTCAGAGGGAACA  
 15 TTATCTAAACGTATTTTCAAGTAGGTGATTTAGCTCATAATAAATATAAAATTTGAGTTAACTGTCAGTGGA  
 AAAACCATAGTAAAACCAGTGGACAAAACAAAAGCCGTTAGATGTTGTCTTCGTACTCGATAATTTCTAACTCA  
 ATGAATAACGATGGCCCAAATTTTCAAAGGCATAATAAAGCCAAGAAAGCTGCCGAAGCTCTTGGGACCGCA  
 GTAAAAGATATTTTAGGAGCAAACAGTGATAATAGGGTTGCATTAGTTACCTATGGTTCAGATATTTTTGAT  
 GGTAGGAGTGTAGATGTCGTAAGGATTTAAAGAAGATGATAAATATTATGGCCTTCAAACCTAAGTTCACA  
 20 ATTCAGACAGAGAATTATAGTCATAACAATTAACAAATAATGCTGAAGAGATTATAAAAAGGATTCGACACA  
 GAAGCTCCTAAAGCTAAGTGGGGATCTACTACCAATGGATTAACCTCCAGAGCAACAAAAGGAGTACTATCTT  
 AGTAAAGTAGGAGAAACATTTACTATGAAAGCCTTCATGGAGGCAGATGATATTTTGAGTCAAGTAAATCGA  
 AATAGTCAAAAATTATTGTTTCATGTAAGTGGTGTTCCTACGAGATCATATGCTATTAATAATTTTAAA  
 CTGGGTGCATCATATGAAAGCCAATTTGAACAAATGAAAAAAATGGATATCTAAATAAAAGTAATTTTCTA  
 25 CTTACTGATAAGCCCGAGGATATAAAAGGAAATGGGGAGAGTTACTTTTTGTTTCCCTTAGATAGTTATCAA  
 ACACAGATAATCTCTGGAACTTACAAAACCTTCATTATTTAGATTTAAATCTTAATTACCCCTAAAGGTACA  
 TTTTATCGAAATGGACCAGTAAGAGAACATGGAACACCAACCAACTTTATATAAATAGTTTAAAACAGAAA  
 AATTATGACATCTTTAATTTTGGTATAGATATATCTGGTTTTAGACAAGTTTATAATGAGGATTATAAGAAA  
 AATCAAGATGGTACTTTTCAAATTTGAAAGAGGAAGCTTTTGAACCTTTCAGATGGGGAAATAACAGAACTA  
 30 ATGAAGTCATTCTCTTCTAAACCTGAGTATTATACCCCGATAGTAACCTTCATCCGATGCATCTAACAAATGAA  
 ATTTTATCTAAAATTCAGCAACAATTTGAAAAGATTTTAAACAAAAGAAAACCTCAATTGTTAATGGAACATA  
 GAAGATCCTATGGGTGACAAAATCAATTTACAGCTTGGCAACGGACAAAACATTGCAACCAAGTGATTATACT  
 TTACAGGGAAATGATGGAAGTATAATGAAAGATAGCATTGCAACTGGTGGGCCATAAATGATGGTGGAAATA  
 CTTAAAGGGGTTAAATTAGAATACATCAAAAATAAACTCTACGTTAGAGGTTTGAACCTTAGGGGAGGGACAA  
 35 AAAGTAACACTCACATATGATGTGAAACTAGATGACAGTTTTATAAGTAACAAATTTCTATGACACTAATGGT  
 AGAACAACTTGAATCCTAAATCAGAGGATCCTAATACACTTAGAGATTTTCCAATCCCTAAAATTCGTGAT  
 GTGAGAGAATATCCTACAATAACGATTAAAAACGAGAAGAAGTTAGGTGAAATTTGAATTTACAAAAGTTGAT  
 AAAGATAATAATAAGTTGCTTCTCAAAGGAGCTACGTTTGAACCTTCAAGAATTTAATGAAGATTATAAACTT  
 TATTTACCAATAAAAAATAAATTTCAAAGTAGTGACGGGAGAAAACGGCAAAAATTTCTTACAAAGATTTG  
 40 AAAGATGGCAAATATCAGTTAATAGAAGCAGTTTCGCCGAAGGATTATCAAAAATTTACTAATAAACCAATT  
 TTAACCTTTTGAAGTTGTTAAAGGATCGATACAAAATATAATAGCTGTTAATAAACAGATTTCTGAATATCAT  
 GAGGAAGGTGACAAGCATTTAATTACCAACACGCATATTTCCACCAAAAAGGAATTATTCGGATGACAGGTGGG  
 AAAGGAATTCTATCTTTTCAATTTTAAATAGGTGGATCTATGATGCTATTGCAGGTGGAATTTATATTTGGAAA  
 AGATATAAGAAATCTAGTGATATATCTAGAGAAAAAGATTAA

45 ORF amino acid sequence (SEQ ID NO: 195):

MRKYQKFSKILTLFLSLSQIPLNTNVLGESTVPENGAAGKLVVKKTDQNKPLSKATFVLKTTAQPEKIE  
 KVTAELTGEATFDNLI PGDYTLSEETAPEGYKKTNTWQVKVESNGKTTIQNSGDKNSTIGQNQEELDKQYP  
 PTGIYEDTKESYKLEHVKGSVPNGKSEAKAVNPYSSEGEHIREIPEGTLSKRISEVGDLAHNKYKIELTVSG  
 KTIIVKPVDKQKPLDVVFLDNSNSMNDGPNFQRHNKAKKAAEALGTAVKDI LGANSNDRVALVTYGSDFD  
 50 GRSVDVVKGFKEDDKYYGLQTKFTIQTENYSHKQLTNNAAEIIKRIPTAPKAKWGSTTNGLTPEQQKEYYL  
 SKVGETFTMKAFMEADDILSQVNRNSQKIIHVTDGVPTRSYAINNFKLGASYESQFEQMKNKYLNKSNFL  
 LDKPEDIKNGESYFLFPLDSYQTQIISGNLQKLHYLDLNLNYPKGTfYRNGPVREHGTPTKLYINSLKQK  
 NYDIFNFGIDISGFRQVYNEDYKKNQDGTfQKLKEEAFELSDGEITELMKSFSSKPEYYTPIVTSDDASNE  
 ILSKIQQQFEKILTKENSIVNGTIEDPMGDKINLQLGNGQTLQPSDYTLQNDGSIMKDSIATGGPNDGGI  
 55 LKGVKLEYIKNKLYVRGLNLGEGQKVTLTYDVKLDDSFISNKFYDTNGRTTLNPKSEDPNTLRDFPIPKIRD  
 VREYPTITIKNEKKLGEIEFTKVDKDNKLLLKGFELQEFNEDYKLYLPIKNNNSKVVTGENGKISYKDL  
 KDGYQLIEAVSPKDYQKITNKPIILTFEVVKGSIQNI IAVNKQISEYHEEGDKHLITNTHIPKGIIPMTGG  
 KGILSFILIGGSMMSIAGGIYIWKRYKSSDISREKD

60 **Strain IC251**



## ORF DNA sequence (SEQ ID NO: 215):

ATGAAAAAGAGACAAAAAATATGGAGAGGGTTATCAGTTACTTTACTAATCCTGTCCCAAATTCATTGGT  
 ATATTGGTACAAGGTGAAACCCAAGATACCAATCAAGCACTTGGAAAAGTAATTGTTAAAAAACGGGAGAC  
 AATGCTACACCATTAGGCAAAGCGACTTTTGTGTTAAAAAATGACAATGATAAGTCAGAAACAAGTCACGAA  
 5 ACGGTAGAGGGTTCTGGAGAAGCAACCTTTGAAAACATAAAACCTGGAGACTACACATTAAGAGAAGAAACA  
 GCACCAATTGGTTATAAAAAAAGTATAAAACCTGGAAAGTTAAAGTTGCAGATAACGGAGCAACAATAATC  
 GAGGGTATGGATGCAGATAAAGCAGAGAAACGAAAAGAAGTTTTGAATGCCCAATATCCAAAATCAGCTATT  
 TATGAGGATACAAAAGAAAATTACCCATTAGTTAATGTAGAGGGTTCCAAAGTTGGTGAACAATACAAAGCA  
 TTGAATCCAATAAATGGAAAAGATGGTCGAAGAGAGATTGCTGAAGGTTGGTTATCAAAAAAATACAGGG  
 10 GTCAATGATCTCGATAAGAATAAATATAAAATTGAATTAAGTGTGAGGGTAAAACCACTGTTGAAACGAAA  
 GAACTTAATCAACCACTAGATGTCGTTGTGCTATTAGATAATTCAAATAGTATGAATAATGAAAGAGCCAAT  
 AATTCTCAAAGAGCATTAAAAGCTGGGGAAGCAGTTGAAAAGCTGATTGATAAAATTACATCAAATAAAGAC  
 AATAGAGTAGCTCTTGTGACATATGCCTCAACCATTTTTGATGGTACTGAAGCGACCGTATCAAAGGGAGTT  
 GCCGATCAAATGGTAAAGCGCTGAATGATAGTGTATCATGGGATTATCATAAACTACTTTTACAGCAACT  
 15 ACACATAATTACAGTTATTTAAATTTAACAAATGATGCTAACGAAGTTAATATTCTAAAGTCAAGAATTCCA  
 AAGGAAGCGGAGCATATAAATGGGGATCGCACGCTCTATCAATTTGGTGCACATTTACTCAAAAAGCTCTA  
 ATGAAAGCAAATGAAATTTTAGAGACACAAAGTTCTAATGCTAGAAAAAACTTATTTTTTCACGTAAGTAT  
 GGTGTCCCTACGATGTCTTATGCCATAAATTTTAATCCTTATATATCAACATCTTACCAAACAGTTTAAAT  
 TCTTTTTTAAATAAAATACCAGATAGAAGTGGTATTCTCCAAGAGGATTTTATAATCAATGGTGTGATTAT  
 20 CAAATAGTAAAAGGAGATGGAGAGAGTTTTAACTGTTTTCGGATAGAAAAGTTCTGTTACTGGAGGAACG  
 ACACAAGCAGCTTATCGAGTACCGCAAATCAACTCTCTGTAATGAGTAATGAGGGATATGCAATTAATAGT  
 GGATATATTTATCTCTATTGGAGAGATTACAACCTGGGTCTATCCATTTGATCCTAAGACAAAGAAAGTTTCT  
 GCAACGAAACAAATCAAACCTCATGGTGGGCAACAACATTATACTTTAATGGAAATATAAGACCTAAAGGT  
 TATGACATTTTTACTGTTGGGATTGGTGTAAACGGAGATCCTGGTGCACCTCTTGAAGCTGAGAAATTT  
 25 ATGCAATCAATATCAAGTAAAACAGAAAATTATACTAATGTTGATGATACAAATAAAATTTATGATGAGCTA  
 AATAAATACTTTAAAACAATTGTTGAGGAAAACATTCTATTGTTGATGGAAATGTGACTGATCCTATGGGA  
 GAGATGATTGAATTCCAATTAATAAATGGTCAAAGTTTTACACATGATGATTACGTTTTGGTTGGAAATGAT  
 GGCAGTCAATTAATAAATGGTGTGGCTCTTGGTGGACCAAACAGTATGGGGGAATTTTAAAAGATGTTACA  
 GTGACTTATGATAAGACATCTCAAACCATCAAATCAATCAATTTGAACTTAGGAAGTGGACAAAAGTAGTT  
 30 CTTACCTATGATGTACGTTTTAAAAGATAACTATATAAGTAACAAATTTTACAATACAAATAATCGTACAACG  
 CTAAGTCCGAAGAGTGAAAAGAACCAAATACTATTCGTGATTTCCCAATTCCAAATTCGTGATGTTCTGT  
 GAGTTTCCGGTACTAACCATCAGTAATCAGAAGAAAATGGGTGAGGTTGAATTTATTAAGTTAATAAAGAC  
 AACATTCAGAATCGCTTTTTGGGAGCTAAGTTTTCAACTTCAGATAGAAAAGATTTTTCTGGGTATAAGCAA  
 TTTGTTCCAGAGGGAAGTGTGTTACAACAAAGAATGATGGTAAAATTTATTTTAAAGCACTTCAAGATGGT  
 35 AACTATAAATTATATGAAATTTCAAGTCCAGATGGCTATATAGAGGTTAAAACGAAACCTGTTGTGACATTT  
 ACAATTCAAAATGGAGAAGTTACGAACCTGAAAGCAGATCCAAATGCTAATAAAATCAAATCGGGTATCTT  
 GAAGGAAATGGTAAACATCTTATTACCAACACTCCCAAACGCCACCAGGTGTTTTTCTAAAACAGGGGGA  
 ATTGGTACAATTGTCTATATATTAGTTGGTTCTACTTTTATGATACTTACCATTTGTTCTTTCCGTCGTAAA  
 CAATTGTAA

## ORF amino acid sequence (SEQ ID NO: 196):

MKKRQKIWRGLSVTLILLSQIPFGILVQGETQDTNQLGKVIKKTGDNATPLGKATFVLKNDNDKSETSH  
 TVEGSGEATFENIKPGDYTLREETAPIGYKKTDKTWKVKVADNGATIIEGMDADKAEKREVLNAQYPKSAI  
 YEDTKENYPLVNVEGSKVGEQYKALNPINGKDRREIAEGWLSKKITGVNDLTKNKYKIELTVEGKTTVETK  
 ELNQPLDVVLLDNSNSMNERANNSQRALKAGEAVEKLIKITSNKDNRVALVTYASTIFDGTEATVSKGV  
 45 ADQNGKALNDSVSWDYHKTTFTATTHNYSYLNLTNDANEVNIILKSRIKPEAEHINGDRTLYQFGATFTQKAL  
 MKANEILETQSSNARKKLI FHVTDGVPTMSYAINFNPIYSTSYQNQFNSFLNKIPDRSGILQEDFIINGDDY  
 QIVKGDGESFKLFSRDKVPVTGGTTQAAAYRVPQNQLSVMSNEGYAINSGYIYLYWRDYNWVYPFDPKTKKVS  
 ATKQIKTHGEPTTLYFNGNIRPKGYDIFTVIGVNGDPGATPLEAEKFMQSISSKTENYTNVDDTNKIYDEL  
 NKYFKTIVEEKHSIVDGNVTDPMGEMIEFQLKNGQSFTHDDYVLVGNDSQLKNGVALGGPNSDGGILKDV  
 50 VTYDKTSQTIKINHLNLGSGQKVVLTYDVRLKDNYSNKFYNTNRTTLLSPKSEKEPNTIRDFPIPKIRDVR  
 EFPVLTISNQKMGVEFEIKVNDKHSESLGAKFQLQIEKDFSGYKQFVPEGSVTTKNDGKIYFKALQDG  
 NYKLYEISSPDGYIEVKTKPVVFTFIQNGEVTNLKADPNANKNQIGYLEGNGKHLITNTPKRPPGVFPKTGG  
 IGTIVYILVGSFTMILTICSFRRKQL

55 **Strain IC253**

## ORF DNA sequence (SEQ ID NO: 216):

ATGAGAAAATACCAAAAATTTTCTAAAATATTGACGTTAAGTCTTTTTTGGTTTGTGCGCAAATACCGCTTAAT  
 ACCAATGTTTTAGGGGAAAGTACCGTACCGGAAAATGGTGTAAAGGAAAGTTAGTTGTTAAAAAGACAGAT  
 60 GACCAGAACAACCACTTTCAAAGCTACCTTTGTTTTAAAACACTGCTCATCCAGAAAGTAAAATAGAA  
 AAAGTAACTGCTGAGCTAACAGGTGAAGCTACTTTTGATAATCTCATACTGGAGATTATACTTTATCAGAA



GAAACAGCGCCCGAAGGTTATAAAAAGACTAACCAGACTTGGCAAGTTAAGGTTGAGAGTAATGGAAAACT  
 ACGATACAAAATAGTGGTGATAAAAATTCACAATTGGACAAAATCACGAAGAACTAGATAAGCAGTATCCC  
 CCCACAGGAATTTATGAAGATACAAAGGAATCTTATAAACTTGAGCATGTTAAAGGTTTCAGTTCCAAATGGA  
 AAGTCAGAGGCAAAGCAGTTAACCCATATTCAAGTGAAGGTGAGCATATAAGAGAAATTCAGAGGGGAACA  
 5 TTATCTAAACGTATTTTCAGAAGTAGGTGATTTAGCTCATAATAAATATAAAAATTGAGTTAACTGTCAGTGGA  
 AAAACCATAGTAAAACCAGTGGACAAAACAAAAGCCGTTAGATGTTGTCTTCGTAATTTCTAACTCA  
 ATGAATAACGATGGCCCAAATTTTCAAAGGCATAATAAAGCCAAGAAAGCTGCCGAAGCTCTTGGGACCGCA  
 GTAAAAGATATTTTAGGAGCAAACAGTGATAATAGGGTTGCATTAGTTACCTATGGTTCAGATATTTTTGAT  
 10 GGTTAGGAGTGTAGATGTCGTAAAAGGATTTAAAGAAGATGATAAATATTATGGCCTTCAAACCTAAGTTCACA  
 ATTCAGACAGAGAATTATAGTCATAAACAATTAACAAATAATGCTGAAGAGATTATAAAAAGGATTCCTACA  
 GAAGCTCCTAGAGCTAAATGGGGATCAACTACAAACGGACTTACTCCAGAGCAACAAAAGCAGTACTATCTT  
 AGTAAAGTAGGGGAAACATTTACTATGAAAGCCTTCATGGAGGCAGATGATATTTTGAGTCAAGTAGATCGA  
 AATAGTCAAAAAATTATTGTTTCATATAACTGATGGTGTTCACAACAGATCATATGCTATTAATAATTTTAAA  
 15 TTGGGTGCATCATATGAAAGCCAATTTGAACAAATGAAAAAAATGGATATCTAAATAAAAGTAATTTTCTA  
 CTTACTGATAAGCCCGAGGATATAAAAAGGAAATGGGGAGAGTTACTTTTTGTTTCCCTTAGATAGTTATCAA  
 ACACAGATAATCTCTGGAACTTACAAAACCTTCATTATTTAGATTTAAATCTTAATTACCTAAAGGTACA  
 ATTTATCGAAATGGACCAGTAAGAGAACATGGAACACCAACCAAACTTTATATAAATAGTTTAAAACAGAAA  
 AATTATGACATCTTTAATTTTGGTATAGATATATCTGCTTTTAGACAAGTTTATAATGAGGATTATAAGAAA  
 AATCAAGATGGTACTTTTCAAATAATTGAAAGAGGAAGCTTTTGAACCTTTCAGATGGGGAAATAACAGAACTA  
 20 ATGAAGTCATTCTCTTCTAAACCTGAGTATTATACCCCGATAGTAACCTTCATCCGATGCATCTAACAATGAA  
 ATTTTATCTAAAATTCAGCAACAATTTGAAAAGGTTTAAACAAAAGAAAACCTCAATTGTTAATGGAACATA  
 GAAGATCCTATGGGTGACAAAATCAATTTACAGCTTGGCAACGGACAAACATTGCAACCAAGTGATTATACT  
 TTACAGGGAAATGATGGAAGTATAATGAAAGATAGCATTGCAACTGGTGGGCCATAATAATGATGGTGAATA  
 CTTAAAGGGGTTAAATTAGAATACATCAAAAATAAACTCTACGTTAGAGGTTTGAACCTTAGGGGAGGGACAA  
 25 AAAGTAACACTCACATATGATGTGAAACTAGATGACAGTTTTATAAGTAACAAATTCATGACACTAATGGT  
 AGAACAACTTGAATCCTAAATCAGAGGATCCTAATACACTTAGAGATTTTCCAATCCCTAAAATTCGTGAT  
 GTGAGAGAATATCCTACAATAACGATTAAAAACGAGAAGAAGTTAGGTGAAATTGAATTTACAAAAGTTGAT  
 AAAGATAATAATAAGTTGCTTCTCAAAGGAGCTACGTTTGAACCTCAAGAATTTAATGAAGATTATAAACTT  
 TATTTACCAATAAAAAATAAATTCAAAAGTAGTGACGGGAGAAAACGGCAAAATTTCTTACAAAGATTTG  
 30 AAAGATGGCAAATATCAGTTAATAGAAGCAGTTTCGCCGAAGGATTATCAAAAATTAATAAAACCAATT  
 TTAACCTTTTGAAGTTGTTAAAGGATCGATACAAAATATAATAGCTGTTAATAAACAGATTTCTGAATATCAT  
 GAGGAAGGTGACAAGCATTTAATTACCAACACGCATATTCACCAAAAAGGAATTATTCCGATGACAGGTGGG  
 AAAGGAATTCTATCTTTTCAATTTAATAGGTGGATCTATGATGTCTATTGCAGGTGGAATTTATATTTGAAA  
 AGATATAAGAAATCTAGTGATATATCTAGAGAAAAGATTAA  
 35 ORF amino acid sequence (SEQ ID NO: 197):  
 MRKYQKFSKILTLFLSLSQIPLNTNVLGESTVPENGAAGKGLVVKKTDDQNKPLSKATFVLKTTAHPESKIE  
 KVTAELTGEATFDNLI PGDYTLSEETAPEGYKKTNTQWQVKVESNGKTTIQNSGDKNSTIGQNHEELDKQYP  
 PTGIYEDTKESYKLEHVKGSVPNGKSEAKAVNPYSSEGEHIREIPEGTLSKRISEVGDLAHNKYKIELTVSG  
 KTIVKPVDKQKPLDVFVLDNSNSMNDGPNFQRHNKAKKAAEALGTAVKDI LGANSNDRVALVTYGSDFD  
 40 GRSVDVVKGFKEDDKYYGLQTKFTIQTENYSHKQLTNNAEEI IKRI PTEAPRAKWGSTTNGLTPEQQKQYYL  
 SKVGETFTMKAFMEADDILSQVDRNSQKIIVHITDGVPTRSYAINNFKLGASYESQFEQMKNKYLNKSNFL  
 LDKPEDIKNGESYFLFPLDSYQTQII SGNLQKLHYLDLNLNYPKGTIYRNGPVREHGTPTKLYINSLKQK  
 NYDIFNFGIDISAFRQVYNEDYKKNQDGTFOKLKEEAFELSDGEITELMKSFSKPEYYTPIVTSSDASNE  
 ILSKIQQQFEKVLTKENSIVNGTIEDPMGDKINLQLGNGQTLQPSDYTLQNDGSIMKDSIATGGPNNDGGI  
 45 LKGVKLEYIKNKLYVRGLNLGEGQKVTLTYDVKLDDSFISNKFYDTNGRITLNPKSEDPNTRDFPIPKIRD  
 VREYPTITIKNEKKLGEIEFTKVVDKDNKLLLKGATFELQEFNEDYKLYLPIKNNNSKVVTGENGKISYKDL  
 KDGYQLIEAVSPKDYQKITNKPIILTFEVVKGSIQNI IAVNKQISEYHEEGDKHLITNTHIPKGIIPMTGG  
 KGILSFILIGSMMSIAGGIYIWKRYKSSDISREKD

50 **Strain IC289**  
ORF DNA sequence (SEQ ID NO: 217):  
 ATGAGAAAATACCAAAAATTTTCTAAAATATTGACGTTAAGTCTTTTTTGTGTTGTCGCAAATACCGCTTAAT  
 ACCAATGTTTTAGGGGAAAGTACCGTACCGGAAAATGGTGCTAAAGGAAAGTTAGTTGTTAAAAAGACAGAT  
 GACCAGAACAAACCACTTTCAAAGCTACCTTTGTTTTAAAACCTACTTCACACTCAGAAAGCAAAGTAGAA  
 55 AAAGTAACTACTGAGGTAACAGGTGAAGCTACTTTTGATAATCTCACACCTGGAGATTACACTTTATCAGAA  
 GAAACGGCACCCGAAGGATACAAAAGACTACCCAGACTTGGCAAGTTAAGGTTGAGAGTAATGGAAAACT  
 ACGATACAAAATAGTGATGATAAAAATCTATAATTGAACAAAGGCAAGAGGAAGTACTAGATAAGCAGTATCCC  
 CTTACAGGAGCTTATGAAGATACAAAAGAATCTTATAATCTTGAGCATGTTAAAAATTCAATTCCAAATGGG  
 AAATTAGAGGCAAAGCAGTTAATCCATATTCAAGTGAAGGTGAGCACATAAGAGAAATTCAGAGGGGAACA  
 60 TTATCTAAACGTATTTTCAGAAGTAAATGATTTGGATCATAATAAATATAAAAATTGAGTTAACTGTTAGCGGT



AAATCCATAATAAAAACTATAAATAAAGATGAACCTCTGGATGTTGTTTTGTTCTTGATAATTCAAATTCT  
 ATGAAGAATAATGGAAAAATAACAAGGCAAAAAAGGCAGGTGAAGCAGTAGAAACAATTATAAAAGATGTT  
 TTAGGAGCAAATGTTGAAAACCGAGCAGCTTTAGTTACTTATGGTTCAGATATTTTTGATGGAAGGACAGTT  
 AAAGTTATAAAAGGTTTTAAAGAGGATCCTTATTATGGACTTGAACTAGTTTCACAGTTCAGACAAATGAT  
 5 TATAGCTATAAAAAGTTCACATAATATTGCTGCTGATATTATAAAAAAGATCCCTAAAGAAGCTCCAGAAGCT  
 AAGTGGGGGGGACAAGTCTAGGATTAACCTCCAGAAAAAAGAGGGAATATGATTTAAGTAAAGTAGGTGAG  
 ACCTTTACAATGAAAGCTTTTTATGGAGGCAGATACCTTGTTAAGTAGTATACAGCGTAAGAGTAGAAAGATT  
 ATTGTTTCACTAACTGACGGTGTTCACAACAAGATCATATGCCATTAATAGTTTTGTAAAAGGTTCAACATAC  
 10 GCAAATCAATTTGAGAGAATAAAAGAAAAAGGTTATTTAGACAAAAATAATTATTTTATAACTGATGATCCA  
 GAAAAGATCAAAGGCAATGGGGAGAGTTACTTTTTGTTTCCCTTAGATAGTTATCAAACACAGATAATTTCT  
 GGAACTTACAAAACTTCATTATTTAGATTTAAATCTTAATTACCTAAAGGTACAATTTATAGAAATGGA  
 CCAGTAAGAGAACATGGAACACCAACCAAACTTTATATAAATAGTTTAAAACAGAAAAATTATGACATCTTT  
 AATTTTGGTATAGATATATCTGGTTTTAGACAAGTTTTATAATGAGGATTATAAGAAAAATCAAGATGGTACT  
 15 TTTCAAAAATTGAAGGAGGAAGCTTTTGAACCTTTCAGATGGGGAAATAACAGAACTAATGAATTCATTCTCT  
 TCTAAACCTGAGTATTATACCCCGATAGTAACTTCAGCTGATGTATCTAATAATGAAATTTTATCTAAAATT  
 CAGCAACAATTTGAAAAGATTTTAAACAAAGGAAAACTCAATTGTTAATGGAACATAGAAGATCCTATGGGT  
 GATAAAATCAATTTACATCTTGGCAACGGACAAACATTGCAACCAAGTGATTATACTTTACAGGGAAATGAT  
 GGAAGTATAATGAAAGATAGCATTGCAACTGGAGGGCCTAATAATGATGGTGGGATACTTAAAGGGGTTAAA  
 20 TTAGAATACATCAAAAATAAACTCTACGTTAGAGGTTTGAACCTTAGGGGAGGGGCAAAAAGTAACACTCACA  
 TATGATGTGAACTAGATGACAGTTTTATTAGTAACAAATTCTATGACACTAATGGTAGAACAAACATTGAAT  
 CCTAAATCAGAGGAACCCGATACACTTAGAGATTTTCCAATCCCTAAAATTCGTGATGTGAGAGAATATCCT  
 ACAATAACGATTAAAAACGAGAAGAAGTTAGGTGAAATTTGAATTTACAAAAGTTGATAAAGATAATAAAG  
 TTGCTTCTCAAAGGAGCTACATTTGAACCTTCAAGAATTTAATGAAGATTATAAACTTTATTTACCAATAAAA  
 25 AATAATAATTCAAAAGTAGTGACGGGAGAAAACGGCAAAATTTCTTACAAAGATTTGAAAGATGGCAAATAT  
 CAGTTAATAGAAGCAGTTTCGCCGAAGGATTATCAAAAATTAATAAAACCAATTTTAACTTTTGAAGTT  
 GTTAAAGGATCGATACAAAATATAATAGCTGTTAATAAACAGATTTCTGAATATCATGAGGAAGGTGACAAG  
 CATTTAATTACCAACACGCATATTCACCAAAAGGAATTATTCGATGACAGGTGGGAAAGGAATTCATCT  
 TTCATTTTAAATAGGTGGAGCTATGATGTCTATTGCAGGTGGAATTTATATTTGGAAAAGACATAAGAAATCT  
 30 AGTGATGCATCAATCGAGAAAGATTAA  
 ORF amino acid sequence (SEQ ID NO: 198):  
 MRKYQKFSKILTLFLSFLSFIPLNTNVLGESTVPENGAKGKLVVKKTDQNKPLSKATFVLKPTSHSESKVE  
 KVTTEVTGEATFDNLTPGDYTLSEETAPEGYKKTQTWQVKVESNGKTTIQNSDDKKSIEQRQEELDKQYP  
 LTGAYEDTKESYNLEHVKNISIPNGKLEAKAVNPYSSEGEHIREIQEGTLSKRISEVNDLDHNKYKIELTVSG  
 35 KSI IKTINKDEPLDVVFLDNSNSMKNNNGKNNKAKKAGEAVETI IKDVLGANVENRAALVTYGSDFIDGRTV  
 KVIKGFKEPYYGLETSFTVQTNDSYKFTNIAADI IKKIPKEAPEAKWGGTSLGLTPEKKREYDLSKVG  
 TFTMKAFMEADTLLSSIQRKSRKIVHLTDGVPTRSYAINSFVKGSTYANQFERIKEKGYLDKNNYFITDDP  
 EKIKNGESYFLFPLDSYQTQIIISGNLQKLHYLDLNLNYPKGTIYRNGPVREHGTPTKLYINSLKQKNYDIF  
 NFGIDISGFRQVYNEDYKKNQDGTFOKLKEEFELSDGEITELMNSFSSKPEYYTPIVTSADVSNNEILSKI  
 40 QQQFEKILTKENSIVNGTIEDPMGDKINLHLGNGQTLQPSDYTLQNGDGSIMKDSIATGGPNNDGGILKGVK  
 LEYIKNKLYVRGLNLGEGQKVTLTVDVKLDDSFISNKFYDTNGRITLNPKSEEPDTRLDFPIPKIRDVREYP  
 TITIKNEKKLGEIEFTKVDKDNKLLKLGATFELQEFNEDYKLYLPKNNNSKVVTGENGKISYKDLKDGKY  
 QLIEAVSPKDYQKITNKPIILTFEVVKGSIQNI IAVNKQISEYHEEGDKHLITNTHIPPKGIIPMTGGKGI  
 LS  
 FILIGGAMMSIAGGIYIWKRRHKKSSDASIEKD

45 **Strain IC291**  
 ORF DNA sequence (SEQ ID NO: 218):  
 ATGAGAAAATACCAAAAATTTTCTAAAATATTGACGTTAAGTCTTTTTTGTTCGCAAATACCGCTTAAT  
 ACCAATGTTTTAGGGGAAAGTACCGTACCGGAAAATGGTGCATAAGGAAAGTTAGTTGTTAAAAGACAGAT  
 50 GACCAGAACAAACCCTTTCAAAGCTACCTTTGTTTTAAAACACTGCTCATCCAGAAAGTAAAATAGAA  
 AAAGTAACTGCTGAGCTAACAGGTGAAGCTACTTTTGATAATCTCATACTGGAGATTATACTTTATCAGAA  
 GAAACAGCGCCCGAAGGTTATAAAAAGACTAACCAGACTTGGCAAGTTAAGGTTGAGAGTAATGGAAAACT  
 ACGATACAAAATAGTGGTGATAAAAATTCACAATTTGGACAAAATCAGGAAGAAGTAAAGCAGTATCCC  
 CCCACAGGAATTTATGAAGATACAAAGGAATCTTATAAACTTGAGCATGTTAAAGGTTTCCAAATGGA  
 55 AAGTCAGAGGCAAAGCAGTTAACCCTATTTCAAGTGAAGGTGAGCATATAAGAGAAATTCAGAGGGAACA  
 TTATCTAAACGTATTTTCAAGTAGGTGATTTAGCTCATAATAAATAAAAATTTGAGTTAACTGTCAGTGG  
 AAAACCATAGTAAAACAGTGGACAAAACAAAAGCCGTTAGATGTTGTCTTCGTAATCTTAACTCA  
 ATGAATAACGATGGCCCAAATTTTCAAAGGCATAATAAAGCCAAGAAAGCTGCCGAAGCTCTTGGGACCGCA  
 GTAAAAGATATTTTAGGAGCAAACAGTGATAATAGGGTTGCATTAGTTACCTATGGTTCAGATATTTTTGAT  
 60 GGTAGGAGTGTAGATGTCGTAAAAGGATTTAAAGAAGATGATAAATATTATGGCCTTCAAACCTAAGTTCACA  
 ATTCAGACAGAGAATTATAGTCATAACAATTAACAATAATGCTGAAGAGATTATAAAAAGGATTCGACA



- 121 -

GAAGCTCCTAAAGCTAAGTGGGGATCTACTACCAATGGATTAACCTCCAGAGCAACAAAAGGAGTACTATCTT  
 AGTAAAGTAGGAGAAACATTTACTATGAAAGCCTTCATGGAGGCAGATGATATTTTGAGTCAAGTAAATCGA  
 AATAGTCAAAAAATTATTGTTTCATGTAACCTGATGGTGTTCCTACGAGATCATATGCTATTAATAATTTTAAA  
 CTGGGTGCATCATATGAAAGCCAATTTGAACAAATGAAAAAAATGGATATCTAAATAAAAGTAATTTTCTA  
 5 CTTACTGATAAGCCCGAGGATATAAAAAGGAAATGGGGAGAGTTACTTTTTGTTTCCCTTAGATAGTTATCAA  
 ACACAGATAATCTCTGGAAACTTACAAAACTTCATTATTTAGATTTAAATCTTAATTACCTAAAGGTACA  
 TTTTATCGAAATGGACCAGTAAGAGAACATGGAACACCAACCAAACTTTATATAAATAGTTTAAAACAGAAA  
 AATTATGACATCTTTAATTTTGGTATAGATATATCTGGTTTTAGACAAGTTTATAATGAGGATTATAAGAAA  
 AATCAAGATGGTACTTTTCAAAAATTGAAAGAGGAAGCTTTTGAACCTTCAGATGGGGAAATAACAGAACTA  
 10 ATGAAGTCATTCTCTTCTAAACCTGAGTATTATACCCCGATAGTAACCTTCATCCGATGCATCTAACAATGAA  
 ATTTTATCTAAAATTCAGCAACAATTTGAAAAGATTTTAAACAAAAGAAAACCTCAATTGTTAATGGAACATA  
 GAAGATCCTATGGGTGACAAAATCAATTTACAGCTTGGCAACGGACAAACATTGCAACCAAGTGATTATACT  
 TTACAGGGAAATGATGGAAGTATAATGAAAGATAGCATTGCAACTGGTGGGCCATAAATGATGGTGGAAATA  
 CTTAAAGGGGTAAATTAGAATACATCAAAAATAAACTCTACGTTAGAGGTTTGAACCTTAGGGGAGGGACAA  
 15 AAAGTAACACTCACATATGATGTGAACTAGATGACAGTTTTATAAGTAACAAATCTATGACACTAATGGT  
 AGAACAACTTGAATCCTAAATCAGAGGATCCTAATACACTTAGAGATTTTCCAATCCCTAAAATTCGTGAT  
 GTGAGAGAATATCCTACAATAACGATTAAAAACGAGAAGAAGTTAGGTGAAATTGAATTTACAAAAGTTGAT  
 AAAGATAATAATAAGTTGCTTCTCAAAGGAGCTACGTTTGAACCTCAAGAATTTAATGAAGATTATAAACTT  
 TATTTACCAATAAAAAATAATAATTCAAAAGTAGTGACGGGAGAAAACGGCAAAAATTTCTTACAAAGATTTG  
 20 AAAGATGGCAAATATCAGTTAATAGAAGCAGTTTCGCCGAAGGATTATCAAAAAATTACTAATAAACCAATT  
 TTAACCTTTTGAAGTTGTTAAAGGATCGATACAAAATATAATAGCTGTTAATAAACAGATTTCTGAATATCAT  
 GAGGAAGGTGACAAGCATTTAATTACCAACACGCATATTCACCAAAAAGGAATTATTCGATGACAGGTGGG  
 AAAGGAATTCTATCTTTCATTTTAATAGGTGGATCTATGATGTCTATTGCAGGTGGAATTTATATTTGGAAA  
 AGATATAAGAAATCTAGTGATATATCTAGAGAAAAAGATTAA  
 25 ORF amino acid sequence (SEQ ID NO: 199):  
 MRKYQKFSKILTLFLSFLSFIPLNTNVLGESTVPENKAKGLVVKKTDQNKPLSKATFVLKTTAHPESKIE  
 KVTAELTGEATFDNLI PGDYTLSEETAPEGYKKTNQWQVKVESNGKTTIQNSGDKNSTIGQNQEELDKQYP  
 PTGIYEDTKESYKLEHVKGSVPNGKSEAKAVNPYSSEGEHIREIPEGTLISKRISEVGDLAHNKYKIELTVSG  
 30 KTIVKPVDKQKPLDVVFLDNSNSMNDGPNFQRHNKAKKAAEALGTAVKDI LGANSNDRVALVTYGSDFD  
 GRSVDVVKGFKEDDKYYGLQTKFTIQTENYSHKQLTNNAEEI IKRI PTEAPKAKWGSTTNGLTPEQQKEYYL  
 SKVGETFTMKAFMEADDILSQVNRNSQKIIVHVTGDVPTRSYAINNFKLGASYESQFEQMKNKGYLNKSNFL  
 LTDKPEDIKNGESYFLFPLDSYQTQII SGNLQKLHYLDLNLNYPKGTFYRNGPVREHGTPTKLYINSLKQK  
 NYDIFNFGIDISGFRQVYNEYDKKNQDGTFOKLKEEAFELSDGEITELMKSFSKPEYYTPIVTSDDASNE  
 35 ILSKIQQQFEKILTKENSIVNGTIEDPMGDKINLQLNGQTLQPSDYTLQNDGSIMKDSIATGGPNDGGI  
 LKGVKLEYIKNKLYVRGLNLGEGQKVTLYDVKLLDSFI SNKFYDTNGRTTLNPKSEDPNTRDFPIPKIRD  
 VREYPTITIKNEKKLGEIEFTKVDKDNKLLLKGATFELQEFNEDYKLYLPIKNNNSKVVTGENGKISYKDL  
 KDGYQLIEAVSPKDYQKITNKPILTFEVVKGSIQNI IAVNKQISEYHEEGDKHLITNTHIPPKGIIPMTGG  
 KGILSFILIGGSMMSIAGGIYIWKRYKSSDISREKD

40 **Strain IC304**  
ORF DNA sequence (SEQ ID NO: 219):  
 ATGAGAAAATACCAAAAATTTTCTAAAATATTGACGTTAAGTCTTTTTTGTGTCGCAAATACCGCTTAAT  
 ACCAATGTTTTAGGGGAAAGTACCGTACCGGAAAATGGTGCTAAAGGAAAGTTAGTTGTAAAAAGACAGAT  
 45 GACCAGAACAAACCACTTTCAAAGCTACCTTTGTTTTAAAACTACTGCTCAACCAGAAAGTAAAATAGAA  
 AAAGTAACTGCTGAGCTAACAGGTGAAGCTACTTTTGATAATCTCATACTGGAGATTATACTTTATCAGAA  
 GAAACAGCGCCCGAAGGTTATAAAAAGACTAACCAGACTTGGCAAGTTAAGGTTGAGAGTAATGGAAAACT  
 ACGATACAAAATAGTGGTGATAAAAATTCACAATTGGACAAAATCAGGAAGAACTAGATAAGCAGTATCCC  
 CCCACAGGAATTTATGAAGATACAAAGGAATCTTATAAACTTGAGCATGTTAAAGGTTCCAGTTCCAAATGGA  
 50 AAGTCAGAGGCAAAGCAGTTAACCATATTCAAGTGAAGGTGAGCATATAAGAGAAATTCAGAGGGAACA  
 TTATCTAAACGTATTTCAGAAGTAGGTGATTTAGCTCATAATAAATAAAAATTTGAGTTAACTGTCAGTGGA  
 AAAACCATAGTAAAACCAGTGGACAAAACAAAAGCCGTTAGATGTTGTCTTCGTACTCGATAATTTCTAACTCA  
 ATGAATAACGATGGCCCAAATTTTCAAAGGCATAATAAAGCCAAGAAAGCTGCCGAAGCTCTTGGGACCGCA  
 GTAAAAGATATTTTAGGAGCAAACAGTGATAATAGGGTTGCATTAGTTACCTATGGTTCAGATATTTTGTGAT  
 55 GGTAGGAGTGTAGATGTCGTAAGGATTTAAAGAAGATGATAAATATTATGGCCTTCAAACCTAAGTTCACA  
 ATTCAGACAGAGAATTATAGTCATAAACAATTAACAAATAATGCTGAAGAGATTATAAAAAGGATTCGACA  
 GAAGCTCCTAAAGCTAAGTGGGGATCTACTACCAATGGATTAACCTCCAGAGCAACAAAAGGAGTACTATCTT  
 AGTAAAGTAGGAGAAACATTTACTATGAAAGCCTTCATGGAGGCAGATGATATTTTGAGTCAAGTAAATCGA  
 AATAGTCAAAAAATTATTGTTTCATGTAACCTGATGGTGTTCCTACGAGATCATATGCTATTAATAATTTTAAA  
 60 CTGGGTGCATCATATGAAAGCCAATTTGAACAAATGAAAAAAATGGATATCTAAATAAAAGTAATTTTCTA  
 CTTACTGATAAGCCCGAGGATATAAAAAGGAAATGGGGAGAGTTACTTTTTGTTTCCCTTAGATAGTTATCAA



ACACAGATAATCTCTGGAACTTACAAAACTTCATTATTTAGATTTAAATCTTAATTACCTAAAGGTACA  
 TTTTATCGAAATGGACCAGTAAGAGAACATGGAACACCAACCAAACCTTTATATAAATAGTTTAAAACAGAAA  
 AATTATGACATCTTTAATTTTGGTATAGATATATCTGGTTTTAGACAAGTTTATAATGAGGATTATAAGAAA  
 AATCAAGATGGTACTTTTCAAAAATTGAAAGAGGAAGCTTTTGAACCTTTCAGATGGGGAAATAACAGAACTA  
 5 ATGAAGTCATTCTCTTCTAAACCTGAGTATTATACCCCGATAGTAACTTCATCCGATGCATCTAACAATGAA  
 ATTTTATCTAAAATTCAGCAACAATTTGAAAAGATTTTAAACAAAAGAAAACCTCAATTGTTAATGGAACATA  
 GAAGATCCTATGGGTGACAAAATCAATTTACAGCTTGGCAACGGACAAACATTGCAACCAAGTGATTATACT  
 TTACAGGGAAATGATGGAAGTATAATGAAAGATAGCATTGCAACTGGTGGGCCTAATAATGATGGTGAATA  
 CTTAAAGGGGTTAAATTAGAATACATCAAAAATAAACTCTACGTTAGAGGTTTGAACCTTAGGGGAGGGACAA  
 10 AAAGTAACACTCACATATGATGTGAAACTAGATGACAGTTTTATAAGTAACAAATTCTATGACACTAATGGT  
 AGAACAACTTGAATCCTAAATCAGAGGATCCTAATACACTTAGAGATTTTCCAATCCCTAAAATTCGTGAT  
 GTGAGAGAATATCCTACAATAACGATTAAAAACGAGAAGAAGTTAGGTGAAATTGAATTTACAAAAGTTGAT  
 AAAGATAATAATAAGTTGCTTCTCAAAGGAGCTACGTTTGAACCTCAAGAATTTAATGAAGATTATAAACTT  
 TATTTACCAATAAAAAATAATAATTCAAAAGTAGTGACGGGAGAAAACGGCAAAAATTTCTTACAAAGATTTG  
 15 AAAGATGGCAAATATCAGTTAATAGAAGCAGTTTCGCCGAAGGATTATCAAAAATTACTAATAAACCAATT  
 TTAACCTTTTGAAGTTGTTAAAGGATCGATACAAAATATAATAGCTGTTAATAAACAGATTTCTGAATATCAT  
 GAGGAAGGTGACAAGCATTTAATTACCAACACGCATATCCACCAAAGGAATTATCCGATGACAGGTGGG  
 AAAGGAATTCTATCTTTTCAATTTAATAGGTGGATCTATGATGTCTATTGCAGGTGGAATTTATATTTGGAAA  
 AGATATAAGAAATCTAGTGATATATCTAGAGAAAAAGATTAA

20 ORF amino acid sequence (SEQ ID NO: 200):

MRKYQKFSKILTLFLSLSQIPLNTNVLGESTVPENGAQKGLVVKKTDQNKPLSKATFVLKTTAQPEKIE  
 KVTAEELTGEATFDNLI PGDYTLSEETAPEGYKKTNTQWQVKVESNGKTTIQNSGDKNSTIGQNOEELDKQYP  
 PTGIYEDTKESYKLEHVKGSVPNGKSEAKAVNPYSSEGEHIREIPEGTLSKRISEVGDLAHNKYKIELTVSG  
 KTIVKPVDKQKPLDVVFLDNSNSMNNNDGPNFQRHNKAKKAAEALGTAVKDI LGANSNDRVALVITYGSDIFD  
 25 GRSVDVVKGFKEDDKYYGLQTKFTIQTENYSHKQLTNNAEEIKRIPTAPKAKWGSTTNGLTPEQQKEYYL  
 SKVGETFTMKAFMEADDILSQVNRNSQKIVHVTGVPTRSYAINNFKLGASYESQFEQMKNNGYLNKSNFL  
 LDKPEDIKNGESYFLFPLDSYQTQIIISGNLQKLHYLDLNLNPKGTFYRNGPVREHGTPTKLYINSLKQK  
 NYDIFNFGIDISGFRQVYNEDYKKNQDGTFOKLKEEAFELSDGEITELMKSFSKPEYYTPIVTSSDASNE  
 ILSKIQQQFEKILTKENSIVNGTIEDPMGDKINLQLGNGQTLQPSDYTLQONDGSIMKDSIATGGPNDGGI  
 30 LKGVKLEYIKNKLYVRGLNLGEGQKVTLTYDVKLDDSFISNKFYDTNGRITLNPKSEDPNTRDFPIPKIRD  
 VREYPTITIKNEKKLGEIEFTKVDKDNKLLLKGATFELQEFNEDYKLYLPIKNNNSKVVTGENGKISYKDL  
 KDGKYQLIEAVSPKDYQKITNKPIILTFEVVKGSIQNI IAVNKQISEYHEEGDKHLITNTHIPKGIIPMTGG  
 KGILSFILIGGSMMSIAGGIYIWKRYKKSSDISREKD

35 **Strain IC305**

ORF DNA sequence (SEQ ID NO: 220):

ATGAGAAAATACCAAAAATTTTCTAAAATATTGACGTTAAGTCTTTTTTGTGTCGCAAATACCGCTTAAT  
 ACCAATGTTTTAGGGGAAAGTACCGTACCGGAAAATGGTGCTAAAGGAAAGTTAGTTGTTAAAAGACAGAT  
 40 GACCAGAACAAACCCTTTCAAAGCTACCTTTGTTTTAAAACACTGCTCATCCAGAAAGTAAAATAGAA  
 AAAGTAACTGCTGAGCTAACAGGTGAAGCTACTTTTGATAATCTCATACCTGGAGATTATACTTTATCAGAA  
 GAAACAGCGCCCGAAGGTTATAAAAAGACTAACCAGACTTGGCAAGTTAAGGTTGAGAGTAATGGA AAAACT  
 ACGATACAAAATAGTGGTGATAAAAATTCACAATTTGGACAAAATCAGGAAGAAGTACTAGATAAGCAGTATCCC  
 CCCACAGGAATTTATGAAGATACAAAGGAATCTTATAAACTTGAGCATGTTAAAGGTTTCAGTTCCAAATGGA  
 45 AAGTCAGAGGCAAAGCAGTTAACCCATATTCAAGTGAAGGTGAGCATATAAGAGAAATTCAGAGGGGAACA  
 TTATCTAAACGTATTTTCAAGTAGGTGATTTAGCTCATAATAAATAAAAATTGAGTTAACTGTCAGTGGAA  
 AAAACCATAGTAAAACAGTGGACAAAACAAAAGCCGTTAGATGTTGTCTTCGTAATCTTAACCTCA  
 ATGAATAACGATGGCCCAAATTTTCAAAGGCATAATAAAGCCAAGAAAGCTGCCGAAGCTCTTGGGACCGCA  
 GTAAAAGATATTTTAGGAGCAAACAGTGATAATAGGGTTGCATTAGTTACCTATGGTTTCAGATATTTTTGAT  
 50 GGTAGGAGTGTAGATGTCGTAAGGATTTAAAGAAGATGATAAATATTATGGCCTTCAAACCTAAGTTCACA  
 ATTCAGACAGAGAATTATAGTCATAAACAATTAACAAATAATGCTGAAGAGATTATAAAAAGGATTCGACA  
 GAAGCTCCTAAAGCTAAGTGGGGATCTACTACCAATGGATTAACCTCCAGAGCAACAAAAGGAGTACTATCTT  
 AGTAAAGTAGGAGAAACATTTACTATGAAAGCCTTCATGGAGGCAGATGATATTTTGGAGTCAAGTAAATCGA  
 AATAGTCAAAAATTTATTGTTTCATGTAACCTGATGGTGTTCCTACGAGATCATATGCTATTAATAATTTTAAA  
 55 CTGGGTGCATCATATGAAAGCCAATTTGAACAAATGAAAAAAAATGGATATCTAAATAAAAGTAATTTTCTA  
 CTTACTGATAAGCCCGAGGATATAAAAAGGAAATGGGGAGAGTTACTTTTTGTTTCCCTTAGATAGTTATCAA  
 ACACAGATAATCTCTGGAACTTACAAAACTTCATTATTTAGATTTAAATCTTAATTACCTAAAGGTACA  
 TTTTATCGAAATGGACCAGTAAGAGAACATGGAACACCAACCAAACCTTTATATAAATAGTTTAAAACAGAAA  
 AATTATGACATCTTTAATTTTGGTATAGATATATCTGGTTTTAGACAAGTTTATAATGAGGATTATAAGAAA  
 AATCAAGATGGTACTTTTCAAAAATTGAAAGAGGAAGCTTTTGAACCTTTCAGATGGGGAAATAACAGAACTA  
 60 ATGAAGTCATTCTCTTCTAAACCTGAGTATTATACCCCGATAGTAACTTCATCCGATGCATCTAACAATGAA



- 123 -

ATTTTATCTAAAATTCAGCAACAATTTGAAAAGATTTTAAACAAAAGAAAACCTCAATTGTTAATGGAACATA  
 GAAGATCCTATGGGTGACAAAATCAATTTACAGCTTGGCAACGGACAAACATTGCAACCAAGTGATTATACT  
 TTACAGGGAAATGATGGAAGTATAATGAAAGATAGCATTGCAACTGGTGGGCCAATAATGATGGTGAATA  
 CTTAAAGGGGTTAAATTAGAATACATCAAAAATAAACTCTACGTTAGAGGTTTGAACCTAGGGGAGGGACAA  
 5 AAAGTAACACTCACATATGATGTGAAACTAGATGACAGTTTTATAAGTAACAAATTCTATGACACTAATGGT  
 AGAACAACTTGAATCCTAAATCAGAGGATCCTAATACACTTAGAGATTTTCCAATCCCTAAAATTCGTGAT  
 GTGAGAGAATATCCTACAATAACGATTAACAAACGAGAAGAAGTTAGGTGAAATTGAATTTACAAAAGTTGAT  
 AAAGATAATAATAAGTTGCTTCTCAAAGGAGCTACGTTTGAACCTCAAGAATTTAATGAAGATTATAAACTT  
 TATTTACCAATAAAAAATAATAATTCAAAAGTAGTGACGGGAGAAAACGGCAAAATTTCTTACAAAAGATTTG  
 10 AAAGATGGCAAATATCAGTTAATAGAAGCAGTTTCGCCGAAGGATTATCAAAAAATTACTAATAAACCAATT  
 TTAACTTTTGAAGTTGTTAAAGGATCGATACAAAATATAATAGCTGTTAATAAACAGATTTCTGAATATCAT  
 GAGGAAGGTGACAAGCATTTAATTACCAACACGCATATTCACCAAAGGAATTATTCGATGACAGGTGGG  
 AAAGGAATTCATCTTTTCAATTTAATAGGTGGATCTATGATGTCTATTGCAGGTGGAATTTATATTTGAAA  
 AGATATAAGAAATCTAGTGATATATCTAGAGAAAAGATTAA

15 ORF amino acid sequence (SEQ ID NO: 201):

MRKYQKFSKILTLFLSLSQIPLNTNVLGESTVPENGAAGKGLVVKKTDDQNKPLSKATFVLKTTAHPESKIE  
 KVTAELTGEATFDNLI PGDYTLSEETAPEGYKKTNTWQVKVESNGKTTIQNSGDKNSTIGQNQEELDKQYP  
 PTGIYEDTKESYKLEHVKGSVPNGKSEAKAVNPYSSEGEHIREIPEGTLSKRISEVGDLAHNKYKIELTVSG  
 KTIVKPVDKQKPLDVVFLDNSNSMNDGPNFQRHNKAKKAAEALGTAVKDI LGANSNDRVALVTYGSDFD  
 20 GRSVDVVKGFKEDDKYYGLQTKFTIQTENYSHKQLTNNAEEI IKRI PTEAPKAKWGSTTNGLTPEQQKEYYL  
 SKVGETFTMKAFMEADDILSQVNRNSQKIIVHVTGDVPTRSYAINNFKLGASYESQFEQMKNKYLNKSNFL  
 LDKPEDIKNGESYFLFPLDSYQTQIISGNLQKLHYLDLNLNYPKGT FYRNGPVREHGTPTKLYINSLKQK  
 NYDIFNFGIDISGFRQVYNEKYKNDGTFQKLKEEFELSDGEITELMKSFSKPEYYTPIVTSSDASNE  
 ILSKIQQQFEKILTKENSIVNGTIEDPMGDKINLQLGNGQTLQPSDYTLQNDGSIMKDSIATGGPNDGGI  
 25 LKGVKLEYIKNKLYVRGLNLGEGQKVTLYDVKLLDSFI SNKFYDTNGRDTLNPKSEDPNTRDFPIPKIRD  
 VREYPTITIKNEKKLGEIEFTKVDKDNKLLLKGATFELQEFNEDYKLYLPIKNNNSKVVTGENGKISYKDL  
 KDGYQLIEAVSPKDYQKITNKPIILTFEVVKGSIQNI IAVNKQISEYHEEGDKHLITNTHIPKGIIPMTGG  
 KGILSFILIGGSMMSIAGGIYIWKRYKSSDISREKD

30 **Strain IC306**

ORF DNA sequence (SEQ ID NO: 221):

ATGAGAAAATACCAAAAATTTTCTAAAATATTGACGTTAAGTCTTTTTTGTGTCGCAAATACCGCTTAAT  
 ACCAATGTTTTAGGGGAAAGTACCGTACCGGAAAATGGTGCTAAAGGAAAGTTAGTTGTTAAAAGACAGAT  
 35 GACCAGAACAAACCCTTTCAAAGCTACCTTTGTTTTAAAACACTGCTCATCCAGAAAGTAAAATAGAA  
 AAAGTAACTGCTGAGCTAACAGGTGAAGCTACTTTTGATAATCTCATACTGGAGATTATACTTTATCAGAA  
 GAAACAGCGCCCGAAGGTTATAAAAAGACTAACCAGACTTGGCAAGTTAAGGTTGAGAGTAATGGAAAACCT  
 ACGATACAAAATAGTGGTGATAAAAATTCACAATTGGACAAAATCAGGAAGAACTAGATAAGCAGTATCCC  
 CCCACAGGAATTTATGAAGATACAAAGGAATCTTATAAACTTGAGCATGTTAAAGGTTTCAGTTCCAAATGGA  
 40 AAGTCAGAGGCAAAGCAGTTAACCATATTCAAGTGAAGGTGAGCATATAAGAGAAATTCAGAGGGAACA  
 TTATCTAAACGTATTTTCAAGAGTAGGTGATTTAGCTCATAATAAATAAAAATTTGAGTTAACTGTCAGTGA  
 AAAACCATAGTAAAACCAGTGGACAAACAAAAGCCGTTAGATGTTGTCTTCGTAATCGATAATTTCTAACTCA  
 ATGAATAACGATGGCCCAAATTTTCAAAGGCATAATAAAGCCAAGAAAGCTGCCGAAGCTCTTGGGACCGCA  
 GTAAAAGATATTTTAGGAGCAAACAGTGATAATAGGGTTGCATTAGTTACCTATGGTTCAGATATTTTGTGAT  
 45 GGTAGGAGTGTAGATGTCGTAAGGATTTAAAGAAGATGATAAATATTATGGCCTTCAAACCTAAGTTCACA  
 ATTCAGACAGAGAATTATAGTCATAAACAATTAACAAATAATGCTGAAGAGATTATAAAAAGGATTCGACA  
 GAAGCTCCTAAAGCTAAGTGGGATCTACTACCAATGGATTAACCTCCAGAGCAACAAAAGGAGTACTATCTT  
 AGTAAAGTAGGAGAAACATTTACTATGAAAGCCTTCATGGAGGCAGATGATATTTTGTGAGTCAAGTAAATCGA  
 AATAGTCAAAAATTTATGTTTCATGTAACCTGATGGTGTTCCTACGAGATCATATGCTATTAATAATTTTAAA  
 50 CTGGGTGCATCATATGAAAGCCAATTTGAACAAATGAAAAAAATGGATATCTAAATAAAAGTAATTTTCTA  
 CTTACTGATAAGCCCGAGGATATAAAAAGGAAATGGGGAGAGTTACTTTTTGTTTCCCTTAGATAGTTATCAA  
 ACACAGATAATCTCTGGAACTTACAAAACCTTCATTATTTAGATTTAAATCTTAATTACCCTAAAGGTACA  
 TTTTATCGAAATGGACCAGTAAGAGAACATGGAACACCAACCAAACTTTATATAAATAGTTTAAAACAGAAA  
 AATTATGACATCTTTAATTTTGGTATAGATATATCTGGTTTTAGACAAGTTTATAATGAGGATTATAAGAAA  
 AATCAAGATGGTACTTTTCAAATTTGAAAGAGGAAGCTTTTGAACCTTTCAGATGGGGAAATAACAGAATA  
 55 ATGAAGTCATTCTCTTCTAAACCTGAGTATTATACCCCGATAGTAACTTCATCCGATGCATCTAACAATGAA  
 ATTTTATCTAAAATTCAGCAACAATTTGAAAAGATTTTAAACAAAAGAAAACCTCAATTGTTAATGGAACATA  
 GAAGATCCTATGGGTGACAAAATCAATTTACAGCTTGGCAACGGACAAACATTGCAACCAAGTGATTATACT  
 TTACAGGGAAATGATGGAAGTATAATGAAAGATAGCATTGCAACTGGTGGGCCAATAATGATGGTGAATA  
 60 CTTAAAGGGGTTAAATTAGAATACATCAAAAATAAACTCTACGTTAGAGGTTTGAACCTAGGGGAGGGACAA  
 AAAGTAACACTCACATATGATGTGAAACTAGATGACAGTTTTATAAGTAACAAATTCTATGACACTAATGGT



- 124 -

AGAACAAACATTGAATCCTAAATCAGAGGATCCTAATACACTTAGAGATTTTCCAATCCCTAAAATTCGTGAT  
 GTGAGAGAATATCCTACAATAACGATTAAAAACGAGAAGAAGTTAGGTGAAATTGAATTTACAAAAGTTGAT  
 AAAGATAATAATAAGTTGCTTCTCAAAGGAGCTACGTTTGAACCTCAAGAATTTAATGAAGATTATAAACTT  
 TATTTACCAATAAAAAATAATAATTCAAAAGTAGTGACGGGAGAAAACGGCAAAATTTCTTACAAAGATTTG  
 5 AAAGATGGCAAATATCAGTTAATAGAAGCAGTTTCGCCGAAGGATTATCAAAAAATTACTAATAAACCAATT  
 TTAACTTTTGAAGTTGTTAAAGGATCGATACAAAATATAATAGCTGTTAATAAACAGATTTCTGAATATCAT  
 GAGGAAGGTGACAAGCATTTAATTACCAACACGCATATTCACCACAAAAGGAATTATTCGATGACAGGTGGG  
 AAAGGAATTCTATCTTTTCAATTTAATAGGTGGATCTATGATGTCCTATTGCAGGTGGAATTTATATTTGGAAA  
 AGATATAAGAAATCTAGTGATATATCTAGAGAAAAAGATTAA  
 10 ORF amino acid sequence (SEQ ID NO: 202):  
 MRKYQKFSKILTLFLSFLSFIPLNTNVLGESTVPENGAKGKLVVKKTDQNKPLSKATFVLKTTAHPESKIE  
 KVTAEELTGEATFDNLI PGDYTLSEETAPEGYKKTNQTWQVKVESNGKTTIQNSGDKNSTIGQNQEELDKQYP  
 PTGIYEDTKESYKLEHVKGSVPNGKSEAKAVNPYSSEGEHIREIPEGTLISKRISEVGDLAHNKYKIELTVSG  
 KTIVKPVDKQKPLDVFVLDNSNSMNNNDGPNFQRHNKAKKAAEALGTAVKDI LGANSNDRVALVTYGSDFD  
 15 GRSVDVVKGFKEDDKYYGLQTKFTIQTENYSHKQLTNNAEEI IKRIPTAPKAKWGSTTNGLTPEQQKEYYL  
 SKVGETFTMKAFMEADDILSQVNRNSQKIVHVTDGVPTRSYAINNFKLGASYESQFEQMKNKNGYLNKSNFL  
 LTKPEDIKNGESYFLFPLDSYQTQII SGNLQKLHYLDLNLNYPKGTFYRNGPVREHGTPTKLYINSLKQK  
 NYDIFNFGIDISGFRQVYNEDYKKNQDGTFFQKLKEEAFELSDGEITELMKSFSKPEYYTPIVTSDDASNE  
 ILSKIQQQFEKILTKENSIVNGTIEDPMGDKINLQLNGQTLQPSDYTLQNDGSI MKDSIATGGPNNDDGGI  
 20 LKGVKLEYIKNKLYVRGLNLGEGQKVTLLTYDVKLDDSFISNKFYDTNGRTTLNPKSEDPNTRLRDFPIPKIRD  
 VREYPTITIKNEKKLGEIEFTKVDKDNKLLLKGATFELQEFNEDYKLYLPIKNNNSKVVTGENGKISYKDL  
 KDGYQLIEAVSPKDYQKITNKPIILTFEVVKGSIQNI IAVNKQISEYHEEGDKHLITNTHIPKGIIPMTGG  
 KGILSFILIGGSMMSIAGGIYIWKRYKSSDISREKD

25 **Strain IC458**

ORF DNA sequence (SEQ ID NO: 222):

ATGAGAAAATACCAAAAATTTTCTAAAATATTGACGTTAAGTCTTTTTTGTGTCGCAAATACCGCTTAAT  
 ACCAATGTTTTAGGGGAAAGTACCGTACCGGAAAATGGTGCTAAAGGAAAGTTAGTTGTTAAAAAGACAGAT  
 30 GACCAGAACAAACCACCTTTCAAAGCTACCTTTGTTTTAAAACTACTGCTCATCCAGAAAGTAAAATAGAA  
 AAAGTAACTGCTGAGGTAACAGGTGAAGCTACTTTTGATAATCTCACACCTGGAGATTACACTTTATCAGAA  
 GAAACGGCACCCGAAGGATACAAAAGACTACCCAGACTTGGCAAGTTAAGGTTGAGAGTAATGGAAAACT  
 ACGATACAAAATAGTGATGATAAAAATCTATAATTGAACAAAGGCAAGAGGAAGTAGATAAGCAGTATCCC  
 CTTACAGGAGCTTATGAAGATACAAAAGAATCTTATAATCTTGAGCATGTTAAAAATTCAATTCCAAATGGG  
 AAATTAGAGGCAAAGCAGTTAATCCATATTCAGTGAAGGTGAGCACATAAGAGAAATTCAGAGGGAACA  
 35 TTATCTAAACGTATTTTCAAGAGTAAATGATTTGGATCATAATAAATATAAAATTGAGTTAACTGTTAGCGGT  
 AAATCCATAATAAAAACCTATAAATAAAGATGAACCTCTGGATGTTGTTTTTGTCTTGTGATAATTCAAATCT  
 ATGAAGAATAATGGAAAAATAACAAGGCAAAAAGGCAAGGTGAAGCAGTAGAAACAATTATAAAGATGTT  
 TTAGGAGCAAATGTTGAAAACCGAGCAGCTTTAGTTACTTATGGTTCAGATATTTTTGATGGAAGGACAGTT  
 AAAGTTATAAAAGGTTTTAAAGAGGATCCTTATCATGGACTTGAAACTAGTTTCACAGTTCAGACAAATGAT  
 40 TATAGCTATAAAAAGTTCACATAATATTGCTGCTGATATTATAAAAAGATCCCTAAAGAAGCTCCAGAAGCT  
 AAGTGGGGGGGACAAGTCTAGGATTAACCTCAGAAAAAAGAGGGAATATGATTTAAGTAAAGTAGGTGAG  
 ACCTTTACAATGAAAGCTTTTATGGAGGCAGATACCTTGTTAAGTAGTATACAGCGTAAGAGTAGAAAGATT  
 ATTGTTTCACTAACTGACGGTGTTCACAACAGATCATATGCTATTAATAGTTTTGTAAACAGGTTCAACATAC  
 GCAAATCAATTTGAGAGAATAAAAAGAAAAGGTTATTTAGACAAAATAATTATTTTATAACTGATGATCCA  
 45 GAAAAGATCAAAGGCAATGGGGAGAGTTACTTTTTGTTTCCCTTAGATAGTTATCAAACACAGATAATTTCT  
 GGAACTTACAAAACCTTCATTATTTAGATTTAAATCTTAATTACCTAAAGGTACAATTTATAGAAATGGA  
 CCAGTAAGAGAACATGGAACACCAACCAAACTTTATATAAATAGTTTAAAACAGAAAAATTATGACATCTTT  
 AATTTTGGTATAGATATATCTGGTTTTAGACAAGTTTATAATGAGGATTATAAGAAAAATCAAGATGGTACT  
 TTTCAAATAATTGAAAGAGGAAGCTTTTGAACCTTTCAGGTGGGGAAATAACAGAACTAATGAAGTCATTCTCT  
 50 TCTAAACCTGAGTATTATACCCCGATAGTAACTTCAGCTGATGTATCTAATAATGAAATTTTATCTAAAATT  
 CAGCAACAATTTGAAAAGATTTTAAACAAAGGAAAACCTCAATTGTTAATGGAACATAGAAAGATCCTATGGGT  
 GATAAAATCAATTTACAGCTTGGCAACGGACAAACATTGCAACCAAGTGATTATACTTTACAGGGAAATGAT  
 GGAAGTATAATGAAAGATAGCATTGCAACTGGTGGGCCATAAATGATGGCGGGATACTTAAAGGGGTAAA  
 TTAGAATACATCAAAAATAAACTCTACGTTAGAGGTTTGAACCTAGGGGAGGGGCAAAAAGTAACACTCACA  
 55 TATGATGTGAAACTAGATGACAGTTTTATTAGTAACAAATCTATGACACTAATGGTAGAACAACTTGAAT  
 CCTAAATCAGAGGAACCTGATACACTTAGAGATTTTCCAATCCCTAAAATTCGTGATGTGAGAGAATATCCT  
 ACAATAACGATTAAAAACGAGAAGAAGTTAGGTGAAATTGAATTTACAAAAGTTGATAAAGATAATAAAG  
 TTGCTTCTCAAAGGAGCTACATTTGAACCTCAAGAATTTAATGAAGATTATAAACTTTATTTACCAATAAAA  
 AATAAATAATTCAAAGTAGTGACGGGAGAAAACGGCAAAATTTCTTACAAAAGATTTGAAAGATGGCAAATAT  
 60 CAGTTAATAGAAGCAGTTTCGCCGAAGGATTATCAAAAAATTACTAATAAACCAATTTTAACTTTTGAAGTT



GTTAAAGGATCGATACAAAATATAATAGCTGTTAATAAACAGATTTCTGAATATCATGAGGAAGGTGACAAG  
 CATTTAATTACCAACACGCATATTCACCAAAGGAATTATCCGATGACAGGTGGGAAAGGAATTCATCT  
 TTCATTTTAAATAGGTGGAGCTATGATGTCTATTGCAGGTGGAATTTATATTTGGAAAAACATAAGAAATCT  
 AGTGATGCATCAATCGAGAAAGATTAA

5 ORF amino acid sequence (SEQ ID NO: 203):

MRKYQKFSKILTLFLSLSQIPLNTNVLGESTVPENGAAGKLVVKKTDQNKPLSKATFVLKTTAHPESKIE  
 KVTAEVTGEATFDNLTPGDYTLSEETAPEGYKKTQTWQVKVESNGKTTIQNSDDKKSIEQRQEELDKQYP  
 LTGAYEDTKESYNLEHVKNISIPNGKLEAKAVNPYSSEGEHIREIQEGLSKRISEVNDLDHNKYKIELTVSG  
 KSI IKTINKDEPLDVVFLDNSNSMKNNKAKKAGEAVETI IKDVLGANVENRAALVITYGSDIFDGRTV  
 10 KVIKGFKEDPYHGLETSFTVQTDYSYKKFTNIAADI IKKIPKEAPEAKWGGTSLGLTPEKKREYDLSKVG  
 TFTMKAFMEADTLLSSIQRKSRKIIVHLTDGVPTRSYAINSFVTGSTYANQFERIKEKGYLDKNNYFITDDP  
 EKIKNGESYFLFPLDSYQTQII SGNLQKLHYLDLNLNYPKGTIYRNGPVREHGTPTKLYINSLKQKNYDIF  
 NFGIDISGFRQVYNEDYKKNQDGTFOQLKEEAFELSGGEITELMKSFSKPEYYTPIVTSADVSNNEILSKI  
 15 QQQFEKILTKENSIVNGTIEDPMGDKINLQLGNGQTLQPSDYTLQGNDSIMKDSIATGGPNNDGGILKGVK  
 LEYIKNKLYVRGLNLGEGQKVTLYDVKLDDSFISNKFYDTNGRITLNPKSEEPDTLRDFPIPKIRDVREYP  
 TITIKNEKKLGEIEFTKVVDKDNKLLLKGATFELQEFNEDYKLYLP IKNNSKVVTGENGKISYKDLKDGKY  
 QLIEAVSPKDYQKITNKPIILTFEVVKGSIQNI IAVNKQISEYHEEGDKHLITNTHIPPKGIIPMTGGKGILS  
 FILIGGAMMSIAGGIYIWKKHKKSSDASIEKD

20

**Table 12: Amino acid and encoding DNA sequences of gbs2018 proteins derived from different strains of *S. agalactiae***

**Strain 0176H4A**

ORF DNA sequence (SEQ ID NO: 180):

25 ATGAATAATAACGAAAAAAGTAAAATACTTTTTAAGAAAAACAGCTTATGGTTTGGCCTCAATGTCAGCA  
 GCGTTTGCTGTATGTAGTGGTATTGTACACGCGGATACTAGTTCAGGAATATCGGATTCAATTCCTCATAAG  
 AAACAAGTTAATTTAGGGGCGGTACTCTGAAGAATTTGATTTCTAAATATCGTGGTAATGACAAAGCTATT  
 GCTATACTTCTAAGTAGAGTAGATGATTTTAAATAGAGCATCACAGGATACACTTCCACAATTAATTAAGT  
 ACTGAAGCAGAAATTAACAATACTTTACCTCAGGGACGAATTATTAACAGAGTATAACCAGTCGTAAGATTA  
 30 AAAGTTGAGAGATTGGGAAGTGGTGAATTAAGGCTGAGTCGATTAATAATATTAAGCTGAATCAATTAAT  
 AAAATTCAGGGTAAATCAACTAATAACAATTAAGGCTGAGTCCATTAATAAAAATTAAGTAGAGTCTATTAAT  
 ACAATCAAAGCCGAATCAATTAATAAAAATTCAGCTAAGCCAATTAACACAATCAAAGCCGAGTCTATTAAT  
 ACAATTAAGGCTGAATCAATTCATAAAAATTAACCTCAATCAATAAAAAGTACTAGTGCTACACATGTTAAA  
 GTTAGTGATCAAGAAGTAGCTAAGCAGTCAAGACGTTCTCAAGATATCATTAATCATTAGGTTTCCTTTCA  
 35 TCAGACCAAAAAGATATTTTAGTTAAATCTATTAGCTCTTCAAAAAGATTCGCAACTTATTCTTAAATTTGTA  
 ACACAAGCCACGCAACTGAATAATGCTGAATCAACAAAAGCTAAGCACATGGCTCAAAAATGACGTGGCTTCA  
 ATAAAAAATATAAGCCTCGAAGTCTTAGAAGAATATAAAGAAAAAATTCAAAGAGCTAGCACTAAGAGTCAA  
 GTTGATGAGCTTGTAGCAGAAGCTAAAAAAGTTGTTAATTCCAATAAAGAAACATTGGTAAATCAGGCCAAT  
 GGTAAAAAGCAAGAAATTGCTAAGTTAGAAAATTTATCTAACGATGAAATGTTGAGATATAATACTGCAATT  
 40 GATAATGTAGTGAAACAGTATAATGAAGGTAAGCTCAATATTACTGATGCAATGAATGCTTTAAATAGTATT  
 AAGCAAGCAGCACAGGAAGTTGCCAGAAAACTTACAAAAGCAGTATGCTAAAAAATTGAAAGAATAAGT  
 TAAAAGGATTAGCGTTATCCAAAAGGCTAAAGAAATTTATGAAAAGCATAAAAAGTATTTTGCCCTACACCT  
 GGATATTATGCAGACTCTGTGGGAACCTATTTGAATAGGTTTAGAGATAAACGAACTTTCGGAAATAGAAGT  
 GTTTGGACTGGTCAAAGTGGACTTGATGAAGCAAAAAAATGCTTGATGAAGTCAAAAAGCTTTTAAAAGAA  
 45 CTTCAAGACCTTACCAGAGGTACTAAAGAAGATAAAAAACCAGACGTTAAGCCAGAAGCCAAACCAGAGGCC  
 AAACCAATATTCAAGTACCTAAACAAGCACCTACAGAAGCTGCAAAAACCAGCTTTGTCACCAGAAGCCTTG  
 ACAAGATTGACTACATGGTATAATCAAGCTAAAGATCTGCTTAAAGATGATCAAGTAAAGGACAAATATGTA  
 GATATACTTGCAGTTCAAAAAGCTGTTGACCAAGCTTATGATCATGTGGAAGAGGGAAAATTTATTACCACT  
 GATCAAGCAAATCAATTAGCTAACAAGCTACGTGATGCTTTACAAAGTTTAGAATTAAGAATAAAAAGTA  
 50 GCCAAACCAGAAGCTAAGCCAGAAGTTAAACCAGAAGCCAAACCAGATGTTAAGCCAGACGTTAAGCCAGAA  
 GCTAAGCCAGAAGCCAAACCAGAGGCCAAACCAGAAGCCAAACCAGAGGCCAAACCAGAAGCCAAACCAGAG  
 GCTAAGCCAGAAGTTAAACCAGACGTTAAGCCAGAGGCCAAACCAGACGTTAAGCCAGAAGCCAAACCAGAC  
 GTTAAGCCAGAGGCTAAGCCAGAAGTTAAACCAGACGTTAAGCCAGAAGTTAAACCAGAGGCTAAACCAGAA  
 ATTAACCAGACGTTAAGCCAGAGGCCAGACCAGAGGCTAAGCCAGAAGTTAAACCAGACGTTAAGCCAGAG  
 55 GCCAAACCAGAAGTTAAACCAGACGTTAAGCCAGAGGCCAAACCAGAGGCTAAGCCAGAAGTTAAACCAGAC  
 GTTAAGCCAGAGGCTAAGCCAGAAGCCAAACCAGCAACCAAAAATTCGGTTAATACTAGCGGAAACTTGGCG  
 GCTAAAAAAGCTATTGAAAACAAAAGTATAGTAAAAAATTACCATCAACGGGTGAAGCCGCAAGTCCACTC



- 126 -

TTAGCAATTGTATCACTAATTGTTATGTTAAGTGCAGGTCTTATTACGATAGTTTTAAAGCATAAAAAAAT  
TAA

ORF amino acid sequence (SEQ ID NO: 175):

5 MNNNEKKVKYFLRKTAYGLASMSAAFAVCSGIVHADTSSGISDSIPHKKQVNLGAVTLKNLISKYRGNDKAI  
AILLSRVDDFNRSQDTLPQLINSTEAEINNTLPQGRRIKQSI PVVRLKVERLGSGAIKAESINNIKAESIN  
KIQKSTNTIKAESINKIKVESINTIKAESINKIQAKPINTIKAESINTIKAESIHKIKPQSIKSTSATHVK  
VSDQELAKQSRRSQDIKSLGFLSSDQKDILVKSISSSKDSQLILKFVTQATQLNNAESTKAKHMAQNDVAS  
10 IKNISLEVLEEYKEKIQRASTKSQVDELVAEAKKVNSNKETLVNQANGKKQEI AKLENLSNDEMLRYNTAI  
DNVVKQYNEGKLNITDAMNALNSIKQAAQEVAQKNLQKQYAKKIERISLKGLALS KKAKEIYEKHKSI LPTP  
GYADSVGTYLNRFRDKRTFGNRSVWTGQSGLDEAKKMLDEVKLLKELQDLTRGTKEDKKPDVKPEAKPEA  
KPNIQVPKQAPTEAAKPALSPEALTRLTTWYNQAKDLLKDDQVKDKYVDILAVQKAVDQAYDHVEEGKFITT  
DQANQLANKLRDALQSLELKDKKVAKPEAKPEVKPEAKPDVKPDVKPEAKPEAKPEAKPEAKPEAKPE  
15 AKPEVKPDVKPEAKPEAKPEVKPDVKPEAKPEAKPATKKSVENTSGNLAAKKAIENKKYSKLPSTGEAASPL  
LAIVSLIVMLSAGLITIVLKHKKN

### Strain 12401

ORF DNA sequence (SEQ ID NO: 181):

20 ATGAATAATAACGAAAAAAGTAAAATACTTTTTAAGAAAAACAGCTTATGGTTTGGCCTCAATGTCAGCA  
GCGTTTGTCTGTATGTAGTGGTATTGTACACGCGGATACTAGTTCAGGAATATCGGATTCAATTCCTCATAAG  
AAACAAGTTAATTTAGGGGCGGTACTCTGAAGAATTTGATTTCTAAATATCGTGGTAATGACAAAGCTATT  
GCTATACTTTTAAGTAGAGTAAATGATTTTAATAGAGCATCACAGGATACACTTCCACAATTAATTAATAGT  
ACTGAAGCAGAAATTAACAATACTTTACCTCAGGACGAATTATTAACAGAGTATACCAGTCGTAAGATTA  
AAAGTTGAGAGATTGGGAAGTGGTGAATTAAGGCTGAGTCGATTAATAATATAAAGCTGAATCAATTAAT  
25 AAAATTCAGGGTAAATCAACTAATAACAATTAAGGCTGAGTCCATTAATAAAAATTAAGTAGAGTCTATTAAT  
ACAATCAAAGCCGAATCAATTAATAAAAATTCAGCTAAGCCAATTAACACAATCAAAGCCGAGTCTATTAAT  
ACAATTAAGGCTGAATCAATTCATAAAAATTAACCTCAATCAATAAAAAGTACTAGTGCTACACATGTTAAA  
GTTAGTGATCAAGAAGTAAAGCAGTCAAGACGTTCTCAAGATATCATTAAATCATTAGGTTTTCCTTTCA  
30 TCAGACCAAAAAGATATTTTAGTTAAATCTATTAGCTCTTCAAAGATTTCGCAACTTATTCTTAAATTTGTA  
ACACAAGCCACGCAACTGAATAATGCTGAATCAACAAAAGCTAAGCACATGGCTCAAAATGACGTGGCTTCA  
ATAAAAATATAAGCCTCGAAGTCTTAGAAGAATATAAAGAAAAAATTCAAAGAGCTAGCACTAAGAGTCAA  
GTTGATGAGCTTGTAGCAGAAGCTAAAAAAGTTGTTAATTCATAAAGAAACATTGGTAAATCAGGCCAAT  
GGTAAAAAGCAAGAAATTGCTAAGTTAGAAAATTTATCTAACGATGAAATGTTGAGATATAATACTGCAATT  
GATAATGTAGTGAAACAGTATAATGAAGGTAAGCTCAATATTACTGATGCAATGAATGCTTTAAATAGTATT  
35 AAGCAAGCAGCACAGGAAGTTGCCAGAAAACTTACAAAAGCAGTATGCTAAAAAATTTGAAAGAATAAGT  
TTAAAAGGATTAGCGTTATCCAAAAGGCTAAAAGAAATTTATGAAAAGCATAAAAAGTATTTTGCCCTACACCT  
GGATATTATGCAGACTCTGTGGGAAGTTATTTGAATAGGTTTAGAGATAAACGAACTTTCGGAAATAGAAGT  
GTTTGGACTGGTCAAAGTGGACTTGTGATGAAGCAAAAAAATGCTTGTGATGAAGTCAAAGCTTTTAAAAGAA  
CTTCAAGACCTTACCAGAGGTACTAAAGAAGATAAAAAACCAGACGTTAAGCCAGAAGCCAAACCAGAGGCC  
40 AAACCAAATATTCAAGTACCTAAACAAGCACCTACAGAAGCTGCAAAAACCAGCTTTGTCACCAGAAGCCTTG  
ACAAGATTGACTACATGGTATAATCAAGCTAAAGATCTGCTTAAAGATGATCAAGTAAAGGACAAATATGTA  
GATATACTTGCAGTTCAAAGCTGTTGACCAAGCTTATGATCATGTGGAAGAGGGAAAATTTATTACCACT  
GATCAAGCAAATCAATTAGCTAACAAGCTACGTGATGCTTTACAAAGTTTAGAATTAAGATAAAAAAGTA  
GCCAAACCAGAAGCTAAGCCAGAAGTTAAACCAGAAGCCAAACCAGAGGCCAAACCAGAAGCCAAACCAGAG  
45 GCTAAGCCAGAAGTTAAACCAGACGTTAAGCCAGAGGCCAAACCAGACGTTAAGCCAGAAGCCAAACCAGAC  
GTTAAGCCAGAAGTTAAACCAGACGTTAAGCCAGAGGCCAAACCAGACGTTAAGCCAGAAGCTAAGCCAGAC  
GTTAAGCCAGAAGTTAAACCAGAGGCTAAGCCAGAAGCCAAACCAGAGGCTAAGCCAGAATTAACCAGAC  
GTTAAGCCAGAGGCCAGACCAGAGGCTAAGCCAGAAGTTAAACCAGACGTTAAGCCAGAGGCCAAACCAGAG  
GTTAAGCCAGAAGTTAAACCAGACGTTAAGCCAGAGGCTAAGCCAGAAGCCAAACCAGCAACCAAAAAATCG  
50 GTTAATACTAGCGGAACTTGGCGGTTAAAAAAGCTATTGAAAACAAAAGTATAGTAAAAAATTACCATCA  
ACGGGTGAAGCCGCAAGTCCACTCTTAGCAATTGTATCACTAATTGTTATGTTAAGTGCAGGTCTTATTACG  
ATAGTTTTAAAGCATAAAAAAATTA

ORF amino acid sequence (SEQ ID NO: 176):

55 MNNNEKKVKYFLRKTAYGLASMSAAFAVCSGIVHADTSSGISDSIPHKKQVNLGAVTLKNLISKYRGNDKAI  
AILLSRVNDFNRASQDTLPQLINSTEAEINNTLPQGRRIKQSI PVVRLKVERLGSGAIKAESINNIKAESIN  
KIQKSTNTIKAESINKIKVESINTIKAESINKIQAKPINTIKAESINTIKAESIHKIKPQSIKSTSATHVK  
VSDQELGKQSRRSQDIKSLGFLSSDQKDILVKSISSSKDSQLILKFVTQATQLNNAESTKAKHMAQNDVAS  
IKNISLEVLEEYKEKIQRASTKSQVDELVAEAKKVNSNKETLVNQANGKKQEI AKLENLSNDEMLRYNTAI  
DNVVKQYNEGKLNITDAMNALNSIKQAAQEVAQKNLQKQYAKKIERISLKGLALS KKAKEIYEKHKSI LPTP  
60 GYADSVGTYLNRFRDKRTFGNRSVWTGQSGLDEAKKMLDEVKLLKELQDLTRGTKEDKKPDVKPEAKPEA







- 128 -

ACTGAAGCAGAAATTAACAATACTTTACCTCAGGGACGAATTATTAACAGAGTATAACCAGTCGTAAGATTA  
 AAAGTTGAGAGATTGGGAAGTGGTGAATTAAGGCTGAGTCGATTAATAATATTAAGCTGAATCAATTAAT  
 AAAATTCAGGGTAAATCAACTAATACAATTAAGGCTGAGTCCATTAATAAAAATTAAGTAGAGTCTATTAAT  
 5 ACAATCAAAGCCGAATCAATTAATAAAAATTCAGCTAAGCCAATTAACACAATCAAAGCCGAGTCTATTAAT  
 ACAATTAAGGCTGAATCAATTCATAAAAATTAACCTCAATCAATAAAAAGTACTAGTGCTACACATGTTAAA  
 GTTAGTGATCAAGAAGTAAAGCAGTCAAGACGTTCTCAAGATATCATTAAATCATTAGGTTTTCCTTTCA  
 TCAGACCAAAAAGATATTTTAGTTAAATCTATTAGCTCTTCAAAGATTTCGCAACTTATTCTTAAATTTGTA  
 ACACAAGCCACGCAACTGAATAATGCTGAATCAACAAAAGCTAAGCACATGGCTCAAATGACGTGGCTTCA  
 10 ATAAAAATATAAGCCTCGAAGTCTTAGAAGAATATAAAGAAAAAATTCAAAGAGCTAGCACTAAGAGTCAA  
 GTTGATGAGCTTGTAGCAGAAGCTAAAAAGTTGTTAATTCCAATAAAGAAACATTGGTAAATCAGGCCAAT  
 GGTA AAAAGCAAGAAATTGCTAAGTTAGAAAATTTATCTAACGATGAAATGTTGAGATATAATACTGCAATT  
 GATAATGTAGTGAAACAGTATAATGAAGGTAAGCTCAATATTACTGATGCAATGAATGCTTTAAATAGTATT  
 AAGCAAGCAGCACAGGAAGTTGCCAGAAAACTTACAAAAGCAGTATGCTAAAAAAATTGAAAGAATAAGT  
 15 TAAAAGGATTAGCGTTATCCAAAAGGCTAAAGAAATTTATGAAAAGCATAAAAGTATTTTGCCTACACCT  
 GGATATTATGCAGACTCTGTGGGAAGTTATTTGAATAGGTTTAGAGATAAACGAACTTTCGGAAATAGAAGT  
 GTTTGGACTGGTCAAAGTGGACTTGTGAAGCAAAAAAATGCTTGTGAAGTCAAAGCTTTTAAAAGAA  
 CTTCAAGACCTTACCAGAGGTAAGAAAGATAAAAAACCAGACGTTAAGCCAGAAGCCAAACCAGAGGCC  
 AAACCAATATTCAAGTACCTAAACAAGCACCTACAGAAGCTGCAAACCAGCTTTGTCACCAGAAGCCTTG  
 20 ACAAGATTGACTACATGGTATAATCAAGCTAAAGATCTGCTTAAAGATGATCAAGTAAAGGACAAATATGTA  
 GATATACTTGCAGTTCAAAGCTGTTGACCAAGCTTATGATCATGTGGAAGAGGGAAAATTTATTACCACT  
 GATCAAGCAAATCAATTAGCTAAACAAGCTACGTGATGCTTTACAAAGTTTAGAATTAAAAGATAAAAAAGTA  
 GCCAAACCAGAAGCTAAGCCAGAAGTTAAACCAGAAGCCAAACCAGATGTTAAGCCAGACGTTAAGCCAGAA  
 GCTAAGCCAGAAGCCAAACCAGAGGCCAAACCAGAAGCCAAACCAGAGGCCAAACCAGAAGCCAAACCAGAG  
 25 GCTAAGCCAGAAGTTAAACCAGACGTTAAGCCAGAGGCCAAACCAGACGTTAAGCCAGAAGCCAAACCAGAC  
 GTTAAGCCAGAGGCTAAGCCAGAAGTTAAACCAGACGTTAAGCCAGAGGCCAAACCAGACGTTAAGCCAGAA  
 GCTAAGCCAGACGTTAAGCCAGAAGTTAAACCAGAGGCTAAGCCAGAAGTTAAACCAGACGTTAAGCCAGAG  
 GCCAGACCAGAGGCTAAGCCAGAAGTTAAACCAGACGTTAAGCCAGAGGCCAAACCAGAGGCTAAGCCAGAA  
 GTTAACCAGACGTTAAGCCAGAGGCTAACCAGAAGCTAACCAGCAACCAAAAAATCGGTAAATACTAGC  
 30 GGAACTTGGCGGTTAAAAAGCTATTGAAAACAAAAGTATAGTAAAAAATTACCATCAACGGGTGAAGCC  
 GCAAGTCCACTCTTAGCAATTGTATCACTAATTGTTATGTTAAGTGCAGGTCTTATTACGATAGTTTTAAAG  
 CATAAAAAAATTA

ORF amino acid sequence (SEQ ID NO: 178):

MNNNEKKVKYFLRKTAYGLASMSAAFVCSGIVHADTSSGISDSIPHKKQVNLGAVTLKNLISKYRGNDKAI  
 AILLSRVDDFNRSQDTLPQLINSTEAEINNTLPQGRRIKQSI PVVRLKVERLGS GAIKAESINNIKAESIN  
 35 KIQGKSTNTIKAESINKIKVESINTIKAESINKIQAKPINTIKAESINTIKAESIHKIKPQSIKSTSATHVK  
 VSDQELGKQSRRSQDIKSLGFLSSDQKDILVKSISSSKDSQLILKFVTQATQLNNAESTKAKHMAQNDVAS  
 IKNISLEVLEEYKEKIQRASTKSQVDELVAEAKKVNSNKETLVNQANGKKQEI AKLENLSNDEM LRYNTAI  
 DNVVKQYNEGKLNITDAMNALNSIKQAAQEVAQKNLQKQYAKKIERISLKLALS KKAKEIYEKHK SILPTP  
 40 GYYADSVGTYLNRFRDKRTFGNRSVWTGQSGLDEAKKMLDEVKLLKELQDLTRGTKEDKKPDVKPEAKPEA  
 KPNIQVPKQAPTEAAKPALSPEALTRLTTWYNQAKDLLKDDQVKDKYVDILAVQKAVDQAYDHVEEGKFITT  
 DQANQLANKLRDALQSLELKD KKVAKPEAKPEVKPEAKPDVKPDVKPEAKPEAKPEAKPEAKPEAKPE  
 AKPEVKPDVKPEAKPDVKPEAKPDVKPEAKPEVKPDVKPEAKPDVKPEAKPDVKPEVKPEAKPEVKPDVKPE  
 45 ARPEAKPEVKPDVKPEAKPEAKPEVKPDVKPEAKPEAKPATKKS VNTSGNLAVKKAIENKKYSKKLPSTGEA  
 ASPLLAIVSLIVMLSAGLITIVLKHKKN

### Strain IC458

ORF DNA sequence (SEQ ID NO: 184):

ATGAATAATAACGAAAAAAGTAAAATACTTTTTAAGAAAAACAGCTTATGGTTTGGCCTCAATGTCAGCA  
 GCGTTTGTCTGTATGTAGTGGTATTGTACACGCGGATACTAGTTCAGGAATATCGGCTTCAATTCCTCATAAG  
 50 AAACAAGTTAATTTAGGGGCGGTTACTCTGAAGAATTTGATTTCTAAATATCGTGGTAATGACAAAGCTATT  
 GCTATACTTTTAAGTAGAGTAAATGATTTTAATAGAGCATCACAGGATACACTTCCACAATTAATTAATAGT  
 ACTGAAGCAGAAATTAGAAATATTTTATATCAAGGACAAATTTGGTAAGCAAAAATAACCAAGTGTA ACTACA  
 CATGCTAAAGTTAGTGATCAAGAAGTAAAGCAGTCAAGACGTTCTCAAGATATCATTAAAGTCATTAGGT  
 55 TTCCTTTCATCAGACCAAAAAGATATTTTAGTTAAATATATTAGCTCTTCAAAGATTTCGCAACTTATTCTT  
 AAATTTGTA ACTCAAGCCACGCAACTGAATAATGCTGAATCAACAAAAGCTAAGCAAATGGCTCAAATGAC  
 GTGGCCTTAATAAAAAATATAAGCCCCGAAGTCTTAGAAGAATATAAAGAAAAAATTCAAAGAGCTAGCACT  
 AAGAGTCAAGTTGATGAGTTTGTAGCAGAAGCTAAAAAGTTGTTAATTCCAATAAAGAAACGTTGGTAAAT  
 CAGGCCAATGGTAAAAAGCAAGAAATTGATAAGTTAGAAAATTTATCTAACGATGAAATGTTGAGATATAAT  
 60 ACTGCAATTGATAATGTAGTGAAACAGTATAATGAAGGTAAGCTCAATATTACTGCTGCAATGAATGCTTTA  
 AATAGTATTAAGCAAGCAGCACAGGAAGTTGCCAGAAAACTTACAAAAGCAGTATGCTAAAAAATTGAA



AGAATAAGTTCAAAGGATTAGCGTTATCTAAAAAGGCTAAAGAAATTTATGAAAAGCATAAAAGTATTTTG  
 CCTACACCTGGATATTATGCAGACTCTGTGGGAACCTATTTGAATAGGTTTAGAGATAAACAACTTTTCGGA  
 AATAGGAGTGTGGACTGGTCAAAGTGGACTTGATGAAGCAAAAAAATGCTTGATGAAGTCAAAGCTT  
 5 TAAAAGAACTTCAAGACCTTACCAGAGGTACTAAAGAAGATAAAAAACCAGACGTTAAGCCAGAAGCCAAA  
 CCAGAGGCCAAACCAATATTCAAGTACCTAAACAAGCACCTACAGAAGCTGCAAACCAGCTTTGTCACCA  
 GAAGCCTTGACAAGATTGACTACATGGTATAATCAAGCTAAAGATCTGCTTAAAGATGATCAAGTAAAGGAC  
 AAATACGTAGATATACTTGCAGTTCAAAGCTGTTGACCAAGCTTATGATCATGTGGAAGAGGGAAAATTT  
 ATTACCACTGATCAAGCAAATCAATTAGCTAACAAGCTACGTGATGCTTTACAAAGTTTAGAATTAAGAT  
 10 AAAAAAGTAGCCAAACCAGAAGCCAAACCAGAGGCCAAACCAGAAGCTAAGCCAGAAGCTAAGCCAGAAGCT  
 AAGCCAGAAGCTAAGCCAGAGGCCAAACCAGAAGCTAAGCCAGACGTTAAGCCAGAAGCTAAACCAGACGTT  
 AAACCAGAGGCTAAGCCAGAAGCTAAACCAGAGGCTAAGTCAGAAGCTAAACCAGAGGCTAAGCTAGAAGCT  
 AAACCAGAGGCCAAACCAGCAACCAAAAAATCGGTTAATACTAGCGGAACTTGGCGGCTAAAAAGCTATT  
 GAAAACAAAAAGTATAGTAAAAAATTACCATCAACGGGTGAAGCCGCAAGTCCACTCTTAGCAATTGTATCA  
 15 CTAATTGTTATGTTAAGTGCAGGTCTTATTACGATAGTTTTAAAGCATAAAAAAATTA  
 ORF amino acid sequence (SEQ ID NO: 179):  
 MNNNEKKVKYFLRKTAYGLASMSAAFVCSGIVHADTSSGISASIPHKKQVNLGAVTLKNIISKYRGNDKAI  
 AILLRVNDFNRASQDTLPQLINSTEAEIRNILYQQQIGKQNKPSVTTHAKVSDQELGKQSRRSQDIKSLG  
 20 FLSSDQKDILVKYISSSKDSQLILKFVTQATQLNNAESTKAKQMAQNDVALIKNISPEVLEEYKEKIQRAST  
 KSQVDEFVAEAKKVVNSNKETLVNQANGKKQEKIDKLENLSNDEMRLRYNTAIDNVVKQYNEGKLNITAAMNAL  
 NSIKQAAQEVAQKNLQKQYAKKIERISSKGLALSCKAKEIYEKHKSLPTPGYYADSVGTYLNRFRDKQTFG  
 NRSVWTGQSGLDKAKMLDEVKLLKELQDLTRGTEDKPKDVKPEAKPEAKPNIQVPKQAPTEAAKPALSP  
 25 EALTRLTTWYNQAKDLLKDDQVKDKYVDILAVQKAVDQAYDHVEEGKFITTDQANQLANKLRDALQSLELKD  
 KKVAKPEAKPEAKPEAKPEAKPEAKPEAKPEAKPEAKPDVKPEAKPDVKPEAKPEAKPEAKSEAKPEAKLEA  
 KPEAKPATKKSVENTSGNLAAKKAIENKKYSKKLPSTGEAASPLLAIVSLIVMLSAGLITIVLKHKKN

**Table 13 (A and B): Overview over the two runs of sequencing of the six antigens from various GBS strains; SEQ ID NOs of the corresponding proteins are listed.**

**A**

Strain name	Serotype	gbs0233	gbs1087	gbs1309
IC97 (III)	III	SEQ ID NO: 235	n.d.	n.d.
IC98 (II)	II	SEQ ID NO: 236	SEQ ID NO: 287	SEQ ID NO: 317
IC105 (IV)	IV	SEQ ID NO: 59	SEQ ID NO: 71	SEQ ID NO: 83
IC108 (III)	III	SEQ ID NO: 237	SEQ ID NO: 288	n.d.
IC216 (Ib)	Ib	SEQ ID NO: 238	SEQ ID NO: 289	SEQ ID NO: 318
IC244 (III)	III	SEQ ID NO: 239	n.d.	SEQ ID NO: 319
IC245 (Ib)	Ib	SEQ ID NO: 240	SEQ ID NO: 290	SEQ ID NO: 320
IC246 (III)	III	SEQ ID NO: 241	SEQ ID NO: 291	n.d.
IC247 (III)	III	SEQ ID NO: 242	n.d.	SEQ ID NO: 321
IC250 (Ib)	Ib	SEQ ID NO: 243	SEQ ID NO: 292	SEQ ID NO: 322
IC251 (V)	V	SEQ ID NO: 244	n.d.	SEQ ID NO: 323
IC252 (III)	III	SEQ ID NO: 245	SEQ ID NO: 293	SEQ ID NO: 324
IC252/2 (III)	III	n.d.	n.d.	n.d.
IC253 (III)	III	SEQ ID NO: 246	SEQ ID NO: 294	SEQ ID NO: 325
IC254 (II)	II	SEQ ID NO: 247	n.d.	n.d.
IC255 (V)	V	SEQ ID NO: 248	SEQ ID NO: 295	SEQ ID NO: 326
IC287 (V)	V	SEQ ID NO: 249	n.d.	SEQ ID NO: 327
IC288 (Ia)	Ia	SEQ ID NO: 250	n.d.	SEQ ID NO: 328
IC289 (Ib)	Ib	SEQ ID NO: 251	SEQ ID NO: 296	SEQ ID NO: 329
IC290 (III)	III	SEQ ID NO: 252	SEQ ID NO: 297	SEQ ID NO: 330
IC291 (V)	V	SEQ ID NO: 253	SEQ ID NO: 298	SEQ ID NO: 331
IC304 (V)	V	SEQ ID NO: 254	SEQ ID NO: 299	SEQ ID NO: 332



- 130 -

IC305 (II)	II	SEQ ID NO: 255	SEQ ID NO: 300	SEQ ID NO: 333
IC306 (Ib)	Ib	SEQ ID NO: 256	SEQ ID NO: 301	SEQ ID NO: 334
IC361 (Ib)	Ib	SEQ ID NO: 257	SEQ ID NO: 302	SEQ ID NO: 335
IC363 (III)	III	SEQ ID NO: 258	n.d.	SEQ ID NO: 336
IC364 (III)	III	SEQ ID NO: 259	SEQ ID NO: 303	SEQ ID NO: 337
IC365 (Ia)	Ia	SEQ ID NO: 260	SEQ ID NO: 304	SEQ ID NO: 338
IC366 (n.t.)	non typeable	SEQ ID NO: 261	n.d.	SEQ ID NO: 339
IC367 (II)	II	SEQ ID NO: 262	n.d.	n.d.
IC368 (Ia)	Ia	SEQ ID NO: 263	SEQ ID NO: 305	SEQ ID NO: 340
IC377 (V)	V	SEQ ID NO: 264	SEQ ID NO: 306	SEQ ID NO: 341
IC379 (Ib)	Ib	SEQ ID NO: 265	SEQ ID NO: 307	SEQ ID NO: 342
IC432 (Ib)	Ib	SEQ ID NO: 266	n.d.	SEQ ID NO: 343
IC434 (III)	III	SEQ ID NO: 267	SEQ ID NO: 308	n.d.
IC455 (III)	III	SEQ ID NO: 268	n.d.	SEQ ID NO: 344
IC457 (II)	II	SEQ ID NO: 269	SEQ ID NO: 309	SEQ ID NO: 345
IC458 (Ia)	Ia	SEQ ID NO: 60	SEQ ID NO: 72	SEQ ID NO: 84
IC459 (Ib)	Ib	SEQ ID NO: 270	n.d.	SEQ ID NO: 346
IC460 (II)	II	SEQ ID NO: 271	n.d.	SEQ ID NO: 347
IC461 (Ib)	Ib	SEQ ID NO: 272	SEQ ID NO: 310	SEQ ID NO: 348
IC462 (II)	II	SEQ ID NO: 273	n.d.	n.d.
IC463 (Ib)	Ib	SEQ ID NO: 274	n.d.	SEQ ID NO: 349
IC469 (V)	V	SEQ ID NO: 275	SEQ ID NO: 311	SEQ ID NO: 350
IC470 (V)	V	SEQ ID NO: 276	n.d.	n.d.
126H4A (Ia)	Ia	SEQ ID NO: 277	n.d.	SEQ ID NO: 351
5095S2 (Ib)	Ib	SEQ ID NO: 278	n.d.	n.d.
6313 (III)	III	SEQ ID NO: 279	SEQ ID NO: 230	SEQ ID NO: 352
12351 (IV)	IV	SEQ ID NO: 280	n.d.	SEQ ID NO: 353
NEM316, 12403 (III)	III	SEQ ID NO: 229	n.d.	SEQ ID NO: 231
12403/2 (III)	III	SEQ ID NO: 281	n.d.	SEQ ID NO: 354
12401 (Ib)	Ib	SEQ ID NO: 56	SEQ ID NO: 68	SEQ ID NO: 80
COH1 (III)	III	SEQ ID NO: 58	SEQ ID NO: 70	SEQ ID NO: 82
BAA23 (V)	V	SEQ ID NO: 57	SEQ ID NO: 69	SEQ ID NO: 81
0176H4A (II)	II	SEQ ID NO: 55	SEQ ID NO: 67	SEQ ID NO: 79
A909 (Ia/c)	Ia/c	SEQ ID NO: 282	SEQ ID NO: 312	SEQ ID NO: 355
C388/90 (Ia/c)	Ia/c	SEQ ID NO: 283	SEQ ID NO: 313	SEQ ID NO: 356
BAA22 (III)	III	SEQ ID NO: 284	SEQ ID NO: 314	SEQ ID NO: 357
2603V/R (V)	V	SEQ ID NO: 285	SEQ ID NO: 315	SEQ ID NO: 358
49447 (V)	V	SEQ ID NO: 286	SEQ ID NO: 316	SEQ ID NO: 359
BAA611 (V)	V	n.d.	n.d.	n.d.

**B**

Strain name	Serotype	gbs1477	gbs1478	gbs2018
IC97 (III)	III	SEQ ID NO: 101	n.d.	SEQ ID NO: 379
IC98 (II)	II	SEQ ID NO: 102	SEQ ID NO: 187	SEQ ID NO: 380
IC105 (IV)	IV	SEQ ID NO: 103	SEQ ID NO: 188	SEQ ID NO: 178
IC108 (III)	III	n.d.	SEQ ID NO: 189	n.d.



IC216 (Ib)	Ib	SEQ ID NO: 104	SEQ ID NO: 190	SEQ ID NO: 381
IC244 (III)	III	n.d.	SEQ ID NO: 191	SEQ ID NO: 382
IC245 (Ib)	Ib	SEQ ID NO: 105	SEQ ID NO: 192	SEQ ID NO: 383
IC246 (III)	III	n.d.	SEQ ID NO: 193	n.d.
IC247 (III)	III	n.d.	SEQ ID NO: 194	SEQ ID NO: 384
IC250 (Ib)	Ib	SEQ ID NO: 106	SEQ ID NO: 195	SEQ ID NO: 385
IC251 (V)	V	SEQ ID NO: 107	SEQ ID NO: 196	SEQ ID NO: 386
IC252 (III)	III	SEQ ID NO: 108	n.d.	SEQ ID NO: 387
IC252/2 (III)	III	SEQ ID NO: 360	n.d.	n.d.
IC253 (III)	III	SEQ ID NO: 109	SEQ ID NO: 197	SEQ ID NO: 388
IC254 (II)	II	SEQ ID NO: 110	n.d.	SEQ ID NO: 389
IC255 (V)	V	SEQ ID NO: 111	n.d.	SEQ ID NO: 390
IC287 (V)	V	SEQ ID NO: 112	n.d.	SEQ ID NO: 391
IC288 (Ia)	Ia	n.d.	n.d.	n.d.
IC289 (Ib)	Ib	SEQ ID NO: 113	SEQ ID NO: 198	SEQ ID NO: 392
IC290 (III)	III	n.d.	n.d.	SEQ ID NO: 393
IC291 (V)	V	SEQ ID NO: 114	SEQ ID NO: 199	SEQ ID NO: 394
IC304 (V)	V	SEQ ID NO: 115	SEQ ID NO: 200	SEQ ID NO: 395
IC305 (II)	II	SEQ ID NO: 116	SEQ ID NO: 201	SEQ ID NO: 396
IC306 (Ib)	Ib	SEQ ID NO: 117	SEQ ID NO: 202	SEQ ID NO: 397
IC361 (Ib)	Ib	SEQ ID NO: 118	n.d.	SEQ ID NO: 398
IC363 (III)	III	SEQ ID NO: 119	n.d.	SEQ ID NO: 399
IC364 (III)	III	n.d.	n.d.	n.d.
IC365 (Ia)	Ia	SEQ ID NO: 120	SEQ ID NO: 363	SEQ ID NO: 400
IC366 (n.t.)	non typeable	n.d.	SEQ ID NO: 364	SEQ ID NO: 401
IC367 (II)	II	SEQ ID NO: 121	SEQ ID NO: 365	SEQ ID NO: 402
IC368 (Ia)	Ia	n.d.	n.d.	SEQ ID NO: 403
IC377 (V)	V	SEQ ID NO: 122	SEQ ID NO: 366	SEQ ID NO: 404
IC379 (Ib)	Ib	SEQ ID NO: 123	SEQ ID NO: 367	SEQ ID NO: 405
IC432 (Ib)	Ib	SEQ ID NO: 124	SEQ ID NO: 368	SEQ ID NO: 406
IC434 (III)	III	n.d.	n.d.	n.d.
IC455 (III)	III	SEQ ID NO: 125	SEQ ID NO: 369	SEQ ID NO: 407
IC457 (II)	II	SEQ ID NO: 126	SEQ ID NO: 370	SEQ ID NO: 408
IC458 (Ia)	Ia	SEQ ID NO: 127	SEQ ID NO: 203	SEQ ID NO: 179
IC459 (Ib)	Ib	SEQ ID NO: 128	SEQ ID NO: 371	SEQ ID NO: 409
IC460 (II)	II	SEQ ID NO: 129	SEQ ID NO: 372	SEQ ID NO: 410
IC461 (Ib)	Ib	SEQ ID NO: 130	SEQ ID NO: 373	SEQ ID NO: 411
IC462 (II)	II	SEQ ID NO: 131	SEQ ID NO: 374	SEQ ID NO: 412
IC463 (Ib)	Ib	n.d.	n.d.	SEQ ID NO: 413
IC469 (V)	V	n.d.	SEQ ID NO: 375	SEQ ID NO: 414
IC470 (V)	V	SEQ ID NO: 132	SEQ ID NO: 376	SEQ ID NO: 415
126H4A (Ia)	Ia	SEQ ID NO: 94	n.d.	SEQ ID NO: 416
5095S2 (Ib)	Ib	SEQ ID NO: 96	n.d.	SEQ ID NO: 417
6313 (III)	III	SEQ ID NO: 97, SEQ ID NO: 232	SEQ ID NO: 233	SEQ ID NO: 418
12351 (IV)	IV	SEQ ID NO: 92	n.d.	SEQ ID NO: 419
NEM316, 12403 (III)	III	n.d.	n.d.	SEQ ID NO: 234



12403/2 (III)	III	SEQ ID NO: 361	n.d.	SEQ ID NO: 420
12401 (Ib)	Ib	SEQ ID NO: 93	SEQ ID NO: 185	SEQ ID NO: 176
COH1 (III)	III	n.d.	n.d.	SEQ ID NO: 421
BAA23 (V)	V	SEQ ID NO: 98	SEQ ID NO: 186	SEQ ID NO: 177
0176H4A (II)	II	SEQ ID NO: 91	SEQ ID NO: 377	SEQ ID NO: 175
A909 (Ia/c)	Ia/c	n.d.	n.d.	n.d.
C388/90 (Ia/c)	Ia/c	SEQ ID NO: 100	n.d.	SEQ ID NO: 422
BAA22 (III)	III	n.d.	n.d.	SEQ ID NO: 423
2603V/R (V)	V	SEQ ID NO: 362	SEQ ID NO: 378	SEQ ID NO: 424
49447 (V)	V	SEQ ID NO: 95	n.d.	SEQ ID NO: 425
BAA611 (V)	V	SEQ ID NO: 99	n.d.	n.d.



**Claims**

- 5
1. A composition comprising at least two proteins selected from the group consisting of
- i) a protective protein comprising or consisting of the protective peptide of SEQ ID NO: 1 (gbs0233p) or functionally active variant thereof;
  - 10 ii) a protective protein comprising or consisting of the protective peptide of SEQ ID NO: 2 (gbs1087p) or functionally active variant thereof;
  - iii) a protective protein comprising or consisting of the protective peptide of SEQ ID NO: 3 (gbs1309p) or functionally active variant thereof;
  - iv) a protective protein comprising or consisting of the protective peptide of SEQ ID NO: 4 (gbs1477p) or functionally active variant thereof;
  - 15 v) a protective protein comprising or consisting of the protective peptide of SEQ ID NO: 5 (gbs1478p) or functionally active variant thereof; and
  - vi) a protective protein comprising or consisting of the protective peptide of SEQ ID NO: 6 (gbs2018p) or functionally active variant thereof.
- 20 2. The composition of claim 1, wherein the composition comprises at least three proteins selected from the group consisting of subgroup i) to vi).
3. The composition of claim 1 or 2, wherein the composition comprises at least four proteins selected from the group consisting of subgroup i) to vi).
- 25 4. The composition of any of claims 1 to 3, wherein the at least two, three or four proteins are selected from different subgroups i) to vi).
5. The composition of any of claims 1 to 4, wherein at least two of the proteins are
- 30 selected from one of the subgroups i) to vi).
6. The composition of any of claims 1 to 5 comprising
- at least one protein of subgroup iv);
  - at least one protein of subgroup vi);
  - 35 - at least one protein of subgroup iv) and at least one protein of subgroup vi);



- 134 -

- at least one protein of subgroup iv), at least one protein of subgroup vi) and at least one protein of subgroup ii); or
- at least one protein of subgroup iv), at least one protein of subgroup vi), at least one protein of subgroup ii) and at least one protein of subgroup v).

5

7. The composition of any of claims 1 to 6, wherein one of the at least two proteins comprises or consists of the protective peptide of SEQ ID NO: 4 (gbs1477p) or functionally active variant thereof, preferably protective peptide of SEQ ID NO: 4 (gbs1477p) or a naturally occurring variant thereof.

10

8. The composition of any of claims 1 to 7, wherein one of the at least two proteins comprises or consists of the protective peptide of SEQ ID NO: 6 (gbs2018p) or functionally active variant thereof, preferably protective peptide of SEQ ID NO: 6 (gbs2018p) or a naturally occurring functionally active variant thereof.

15

9. The composition of any of claims 1 to 8, wherein one of the at least two proteins comprises or consists of the protective peptide of SEQ ID NO: 2 (gbs1087p) or functionally active variant thereof, preferably protective peptide of SEQ ID NO: 2 (gbs1087p) or a naturally occurring functionally active variant thereof.

20

10. The composition of any of claims 1 to 9, wherein one of the at least two proteins comprises or consists of the protective peptide of SEQ ID NO: 5 (gbs1478p) or functionally active variant thereof, preferably protective peptide of SEQ ID NO: 5 (gbs1478p) or a naturally occurring functionally active variant thereof.

25

11. The composition of any of claims 1 to 10, wherein one of the at least two proteins comprises or consists of the protective peptide of SEQ ID NO: 1 (gbs0233p) or functionally active variant thereof, preferably protective peptide of SEQ ID NO: 1 (gbs0233p) or a naturally occurring functionally active variant thereof.

30

12. The composition of any of claims 1 to 11, wherein one of the at least two proteins comprises or consists of the protective peptide of SEQ ID NO: 3 (gbs1309p) or functionally active variant thereof, preferably protective peptide of SEQ ID NO: 3 (gbs1309p) or a naturally occurring functionally active variant thereof.



13. The composition of any of claims 1 to 12, wherein the at least two proteins encompass:
- 5 - the protective peptide of SEQ ID NO: 4 (gbs1477p) and the protective peptide of SEQ ID NO: 6 (gbs2018p);
  - the protective peptide of SEQ ID NO: 4 (gbs1477p) and the protective peptide of SEQ ID NO: 6 (gbs2018p) and the protective peptide of SEQ ID NO: 2 (gbs1087p); or
  - 10 - the protective peptide of SEQ ID NO: 4 (gbs1477p) and the protective peptide of SEQ ID NO: 6 (gbs2018p) and the protective peptide of SEQ ID NO: 2 (gbs1087p) and the protective peptide of SEQ ID NO: 5 (gbs1478p).
14. The composition of any of claims 1 to 13, wherein two or more proteins of the at least two proteins are combined into at least one fusion protein.
- 15
15. The composition of any of claims 1 to 14, wherein the functionally active variant
- 20 a) is a functionally active fragment of the protective peptide, the functionally active fragment comprising at least 50% of the sequence of the protective peptide, preferably at least 70%, more preferably at least 80%, still more preferably at least 90%, even more preferably at least 95% and most preferably at least 97%, 98% or 99%;
  - b) is derived from the protective peptide by at least one amino acid substitution, addition and/or deletion, wherein the functionally active variant has a sequence identity to the protective peptide or to the functionally active fragment as defined in a) of at least 40%, preferably at least 60%, more preferably at least 75%, still more preferably at least 90%, even more preferably at least 95% and most preferably at least 97%, 98% or 99%; and/or
  - 25 c) consists of the protective peptide or a functionally active variant thereof and additionally at least one amino acid heterologous to the protective peptide,
  - 30 preferably wherein the functionally active variant is derived from or identical to any of the naturally occurring variants of any of the sequences of SEQ ID NO: 55 to 60, 67 to 72, 79 to 84, 91 to 132, 175 to 179, 185 to 203, 223 to 234, and 235 to 425.



- 136 -

16. A protective peptide consisting of the amino acid sequence selected from the group consisting of SEQ ID NO: 1, SEQ ID NO: 2, SEQ ID NO: 3, SEQ ID NO: 4, SEQ ID NO: 5, SEQ ID NO: 6, SEQ ID NO: 55 to 57, 59, 60, 68, 69, 71, 72, 79 to 84, 91 to 132, 175 to 179, 185 to 203, 223 to 234, and 235 to 425.
- 5
17. One or more nucleic acid(s) encoding the at least two proteins comprised in of the composition according to any of claims 1 to 15 and/or any of the protective peptides according to claim 16.
- 10
18. The one or more nucleic acid(s) of claim 17, comprising or consisting of at least one nucleic acid sequence selected from the group consisting of SEQ ID NO: 7, SEQ ID NO: 8, SEQ ID NO: 9, SEQ ID NO: 10, SEQ ID NO: 11, SEQ ID NO: 12, SEQ ID NO: 61 to 66, SEQ ID NO: 73 to 78, SEQ ID NO: 85 to 90, SEQ ID NO: 133 to 174, SEQ ID NO: 180 to 184 and SEQ ID NO: 204 to 222.
- 15
19. The one or more nucleic acid(s) of claim 17 to 18, wherein the nucleic acid(s) is/are located in a vector or a cell other than *S. agalactiae*.
20. A method of producing the composition of any of claims 1 to 15 or the protective peptide of claim 16, comprising
- 20
- (a) introducing the one or more nucleic acids into a host cell;
- (b) expressing the protein(s) and/or peptide(s) encoded by the nucleic acid by culturing the host cell under conditions conducive to the expression of the protein(s) and/or peptide(s); and
- 25
- (c) collecting and/or isolating the expressed protein(s) and/or peptide(s) of step (b).
21. A pharmaceutical composition, especially a vaccine, comprising
- (i) the composition according to any of claims 1 to 15 and/or at least one
- 30
- protective peptide according to claim 16; and
- (ii) optionally a pharmaceutically acceptable carrier or excipient.
22. A pharmaceutical composition comprising



- 137 -

- (i) the one or more nucleic acid(s) according to any of claims 17 to 19 or one or more nucleic acid(s) complementary thereto, and
  - (ii) optionally a pharmaceutically acceptable carrier or excipient.
- 5 23. A method for producing antibodies, characterized by the following steps:
- (a) administering an effective amount of the composition according to any of claims 1 to 15 and/or at least one protective peptide according to claim 16 to an animal; and
  - (b) isolating the antibodies produced by the animal in response to the  
10 administration of step (a) from the animal.
24. A method for producing antibodies, characterized by the following steps:
- (a) contacting a B cell with an effective amount of the composition according to any of claims 1 to 15 and/or at least one protective peptide according to claim  
15 16;
  - (b) fusing the B cell of step (a) with a myeloma cell to obtain a hybridoma cell; and
  - (c) isolating the antibodies produced by the cultivated hybridoma cell.
- 20 25. The method of claim 23 or 24, wherein the isolated antibodies are additionally purified.
26. A mixture of antibodies against the at least two proteins of the composition of any of claims 1 to 15 and/or against the at least one protective peptide according to claim  
25 16.
27. Use of the composition according to any of claims 1 to 15 and/or at least one protective peptide according to claim 16 and/or one or more the nucleic acid(s) according to any of claims 17 to 19 for the manufacture of a medicament for the  
30 immunization or treatment of a subject, preferably against *S. agalactiae*, more preferably against pneumonia, septicemia, meningitis, fever, vomiting, poor feeding, irritability, urinary tract infection and/or vaginal infection caused by *S. agalactiae*.
28. A method of diagnosing a *S. agalactiae* infection comprising the steps of:



- 138 -

- (a) contacting a sample obtained from a subject with the composition according to any of claims 1 to 15 and/or at least one protective peptide according to claim 16; and
- (b) detecting the presence of an antibody against the protective peptide, the functionally active variant and/or the composition in the sample,
- 5 wherein the presence of the antibody is indicative for the *S. agalactiae* infection.
29. A method of diagnosing a *S. agalactiae* infection comprising the steps of:
- (a) contacting a sample obtained from a subject with the mixture of antibodies
- 10 according to claim 26; and
- (b) detecting the presence of the at least two proteins of the composition of any of claims 1 to 15 and/or of the at least one protective peptide according to claim 16 in the sample,
- wherein the presence of the at least two proteins is indicative for the *S. agalactiae*
- 15 infection.
30. A method for diagnosing an infection with *S. agalactiae* comprising the steps of:
- (a) contacting a sample obtained from a subject with a primer and/or a probe specific for the one or more nucleic acid(s) of any of claims 17 to 19; and
- 20 (b) detecting the presence of one or more nucleic acid(s) of any of claims 17 to 19 in the sample,
- wherein the presence of the one or more nucleic acid(s) is indicative for the *S. agalactiae* infection.
- 25 31. A method for identifying a ligand capable of binding the composition according to any of claims 1 to 15 and/or at least one protective peptide according to claim 16 comprising:
- (a) providing a test system comprising the peptide and/or composition,
- (b) contacting the test system with a test compound, and
- 30 (c) detecting a signal generated in response to the binding of the test compound to the peptide and/or composition.



- 139 -

32. Use of the composition according to any of claims 1 to 15 and/or at least one protective peptide according to claim 16 for the isolation and/or purification and/or identification of an interaction partner of the composition and/or the peptide.



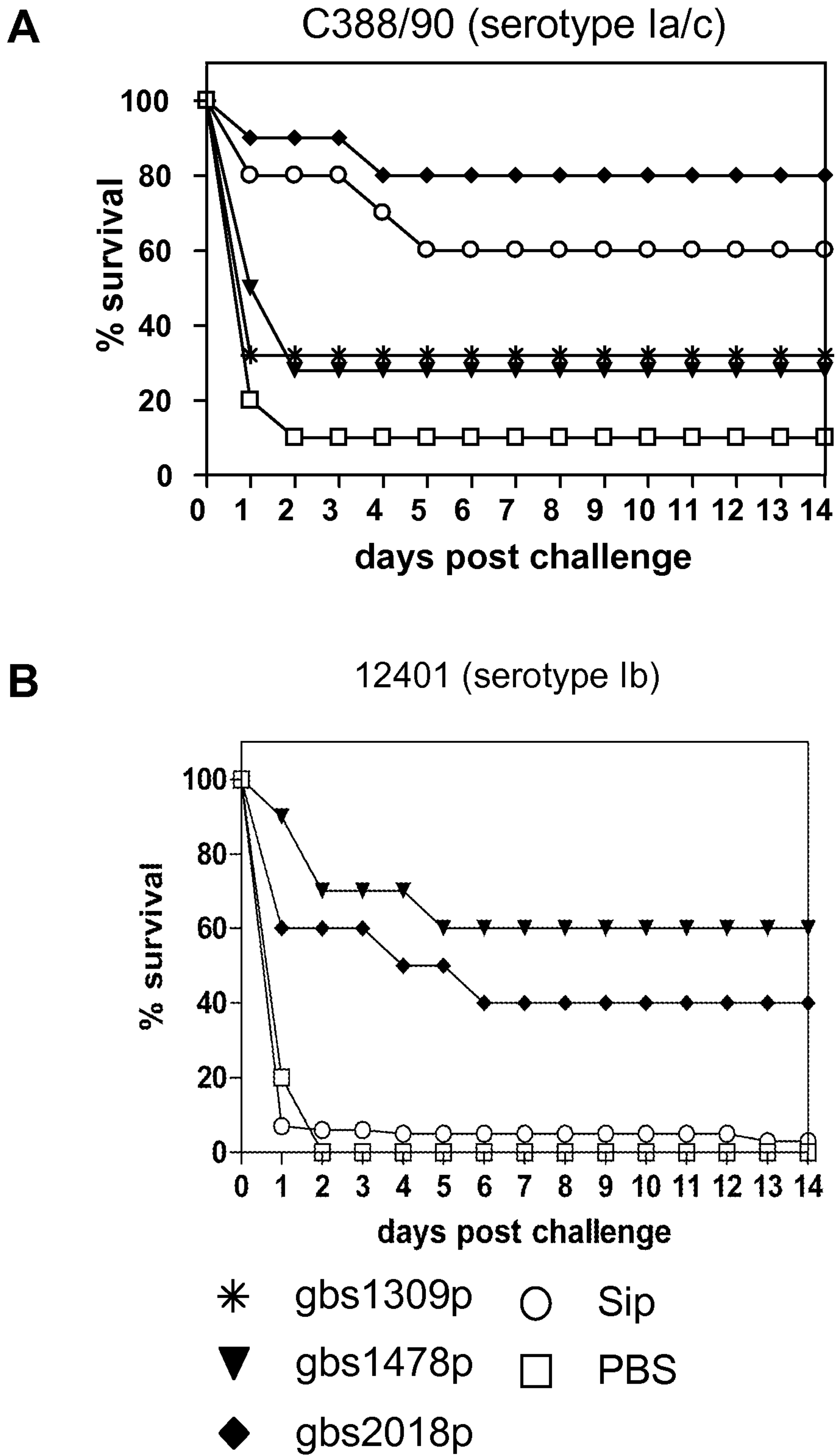
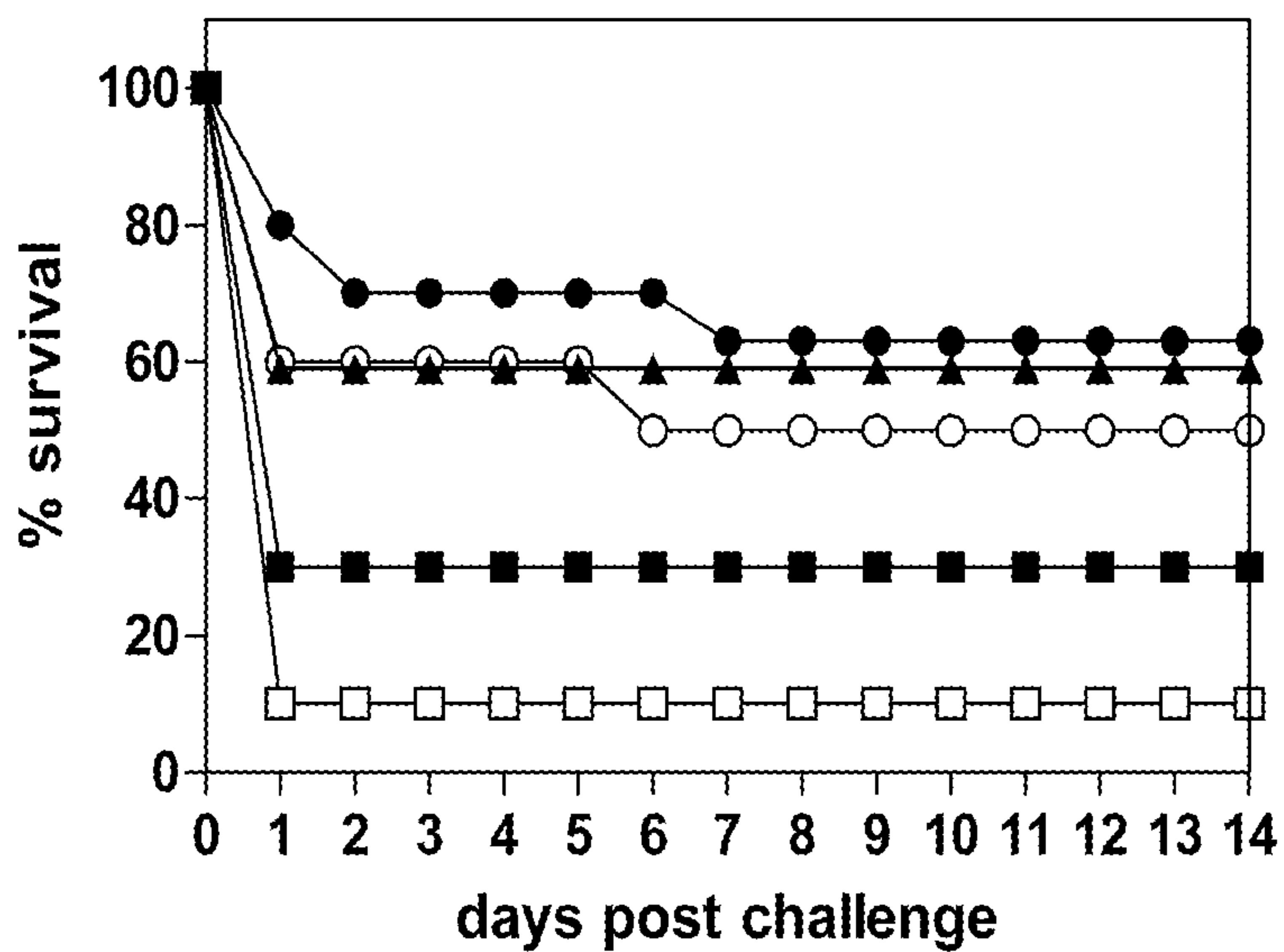


Figure 1



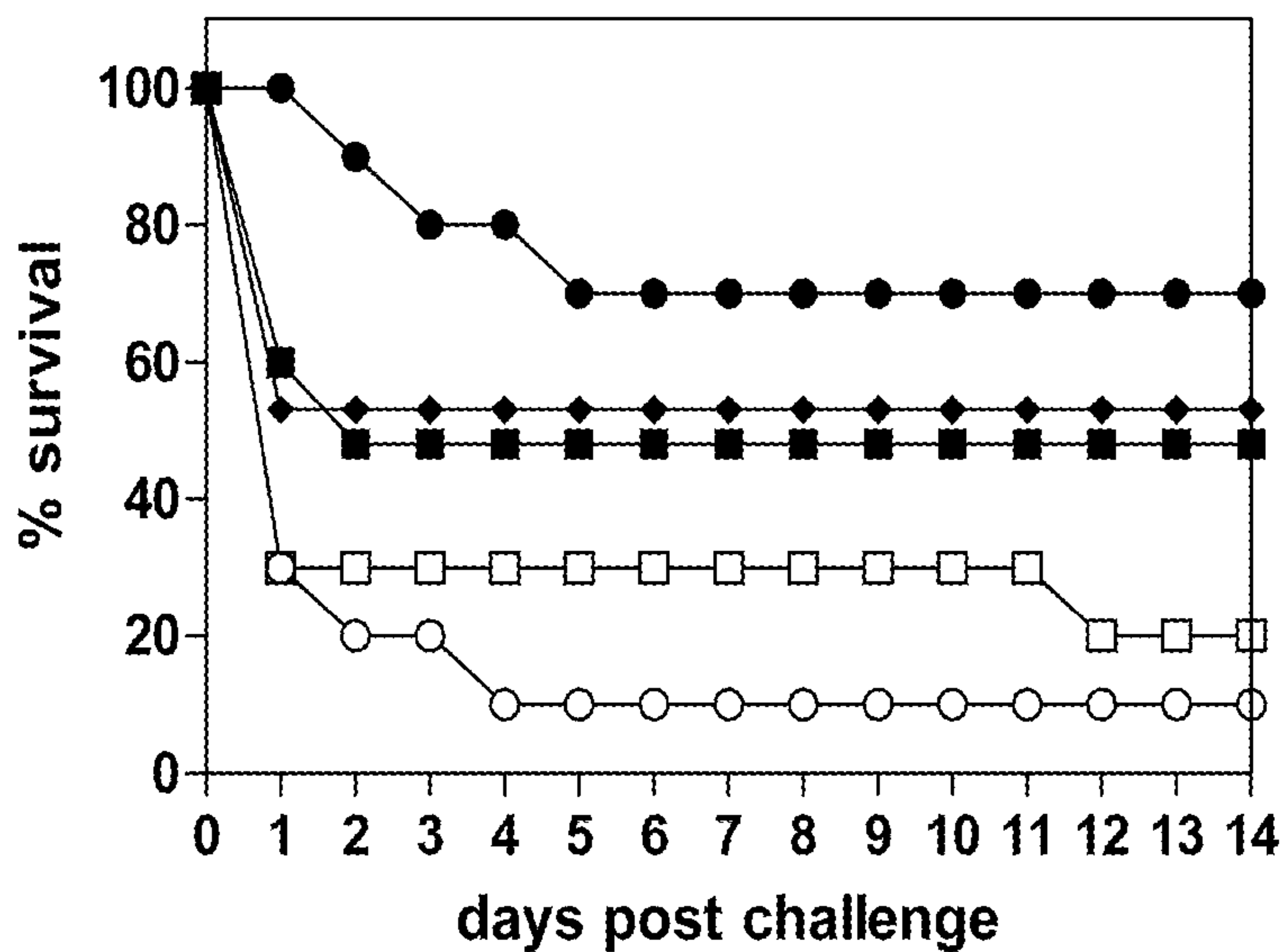
**C**

12403 (serotype III)



**D**

49447 (serotype V)

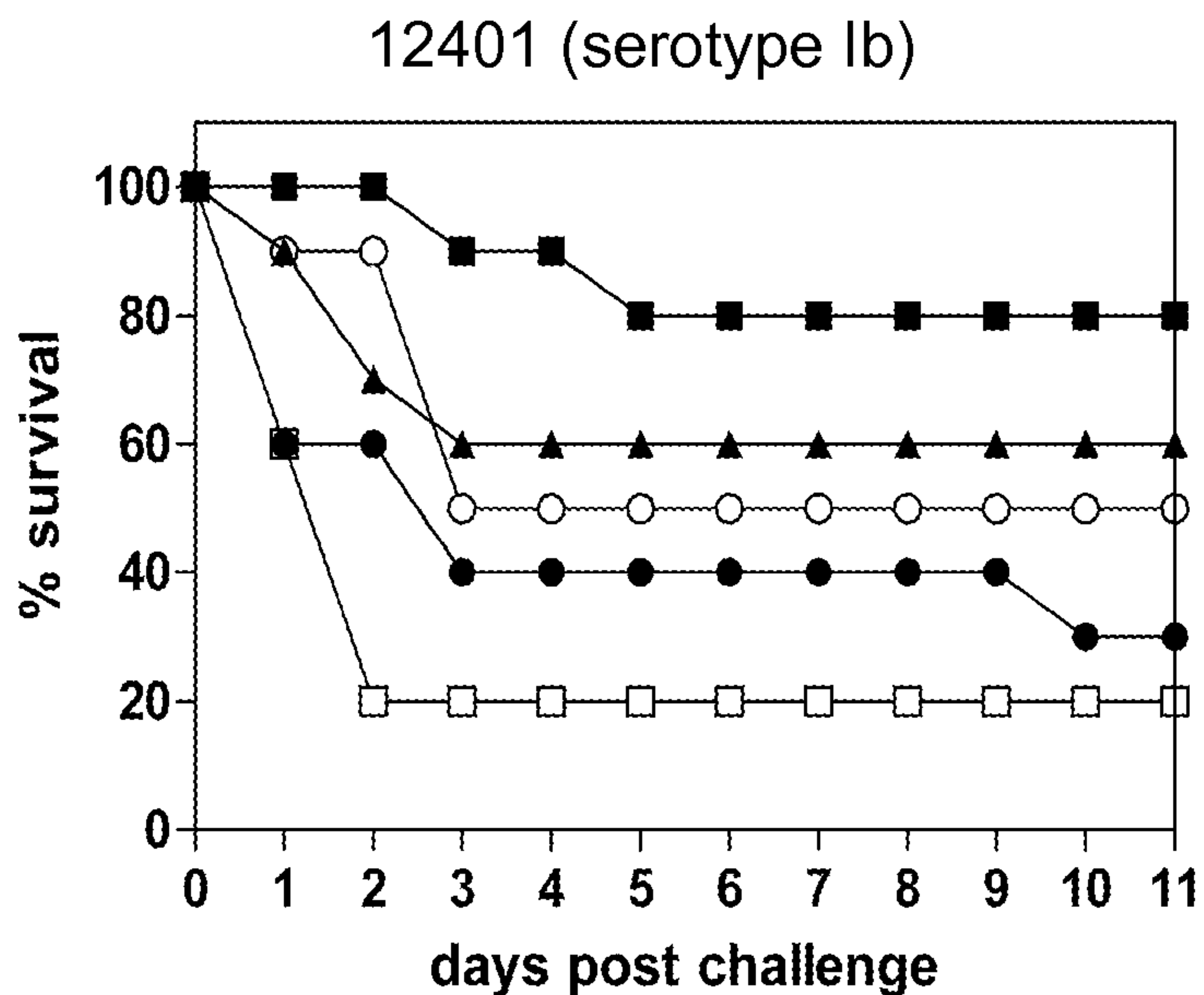


- gbs0233p      ▲ gbs1477p      ○ Sip
- gbs1087p    ◆ gbs2018p    □ PBS

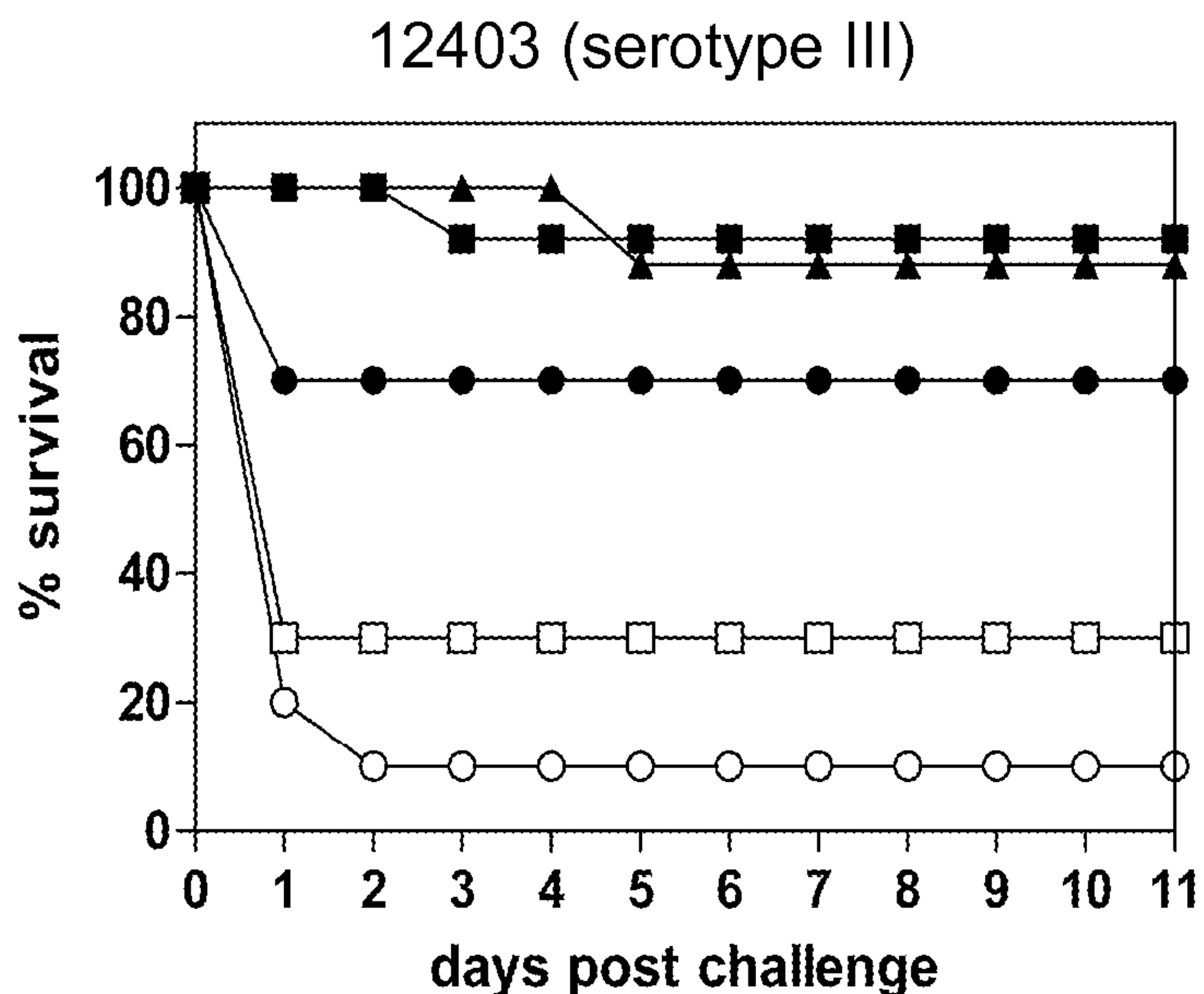
**Figure 1**



**A**



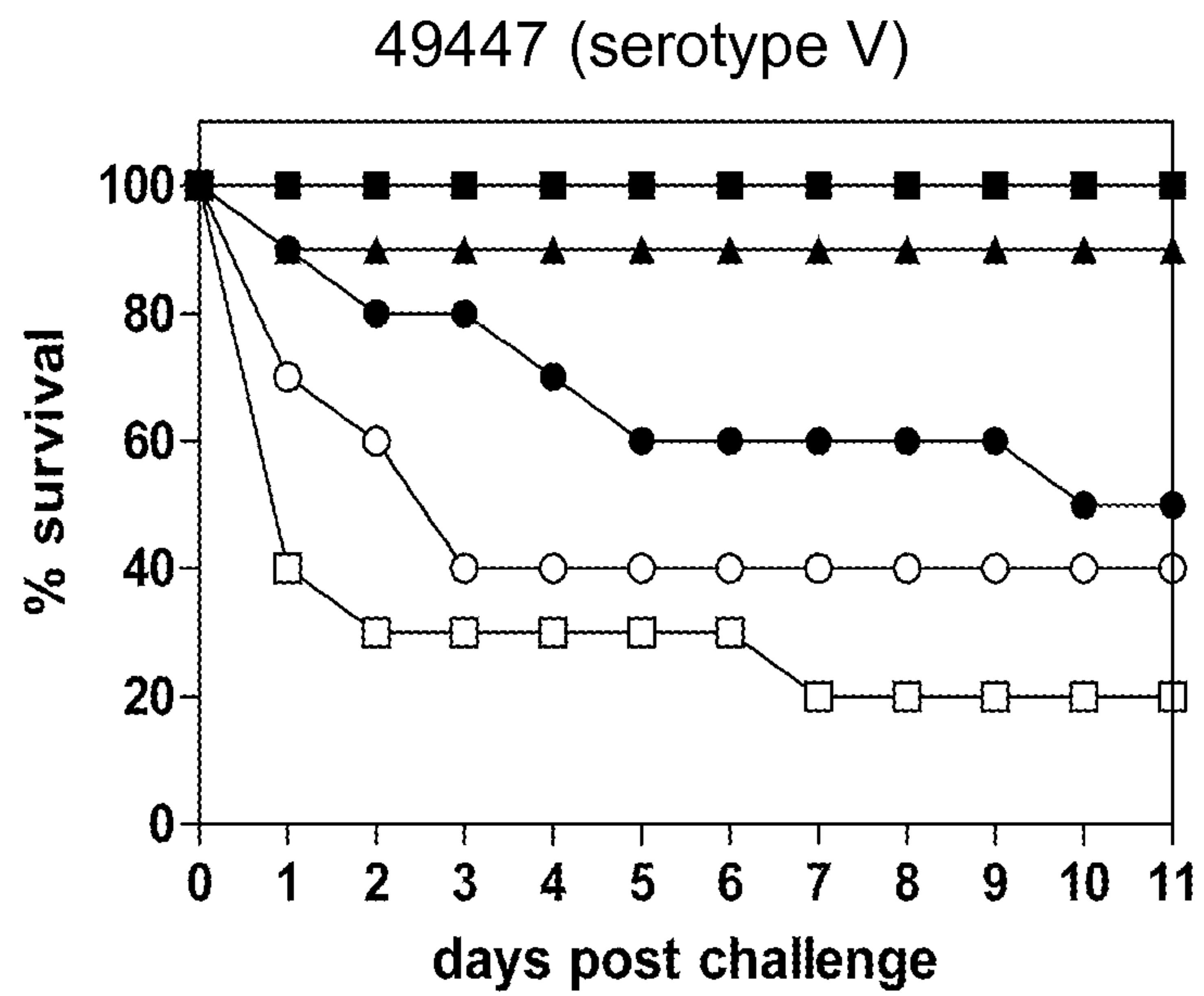
**B**



- gbs1477p + gbs2018p
- ▲ gbs1087p + gbs1477p + gbs2018p
- gbs1087p + gbs1477p + gbs1478p + gbs2018p
- Sip
- PBS

**Figure 2**



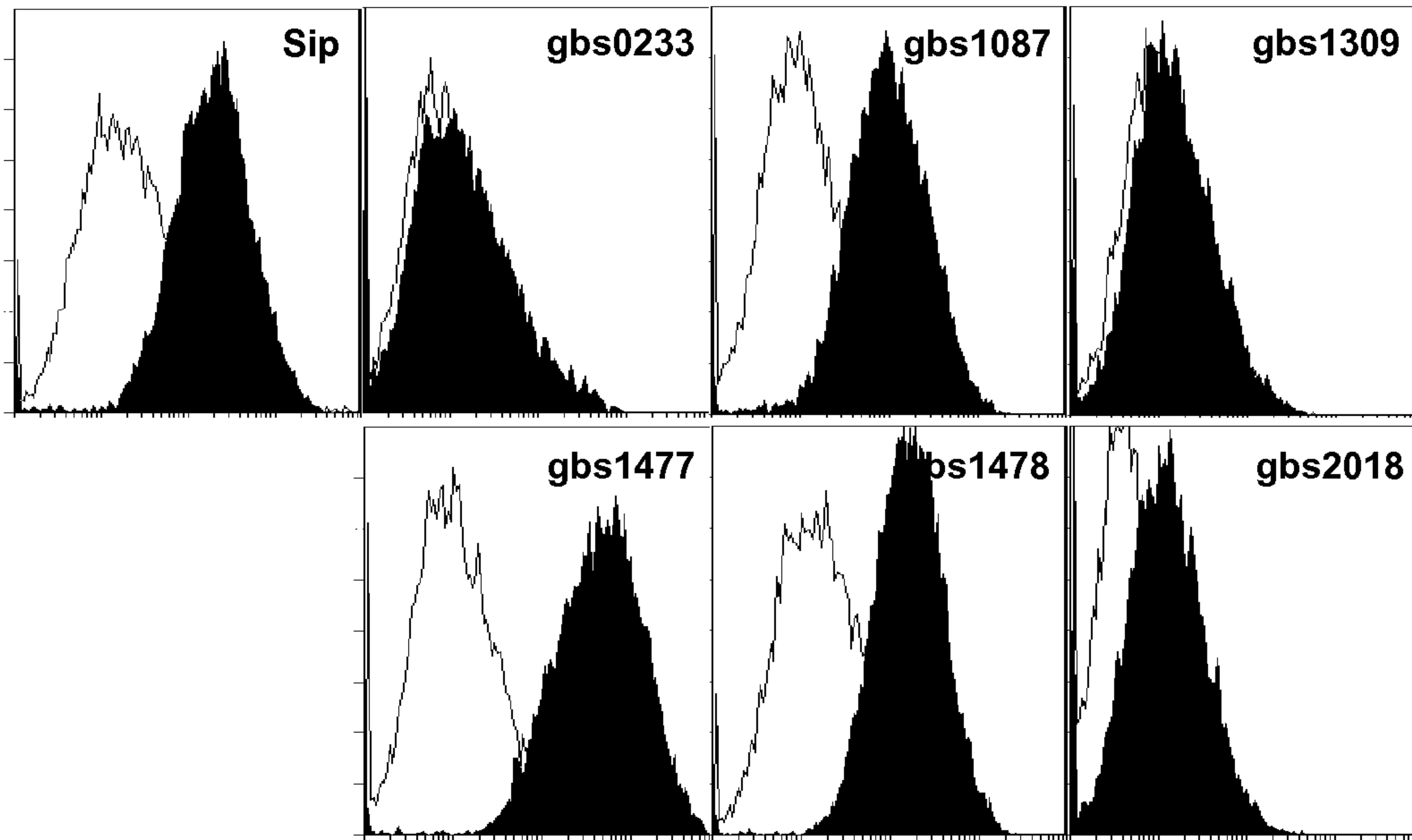
**C**

- gbs1477p + gbs2018p
- ▲ gbs1087p + gbs1477p + gbs2018p
- gbs1087p + gbs1477p + gbs1478p + gbs2018p
- Sip
- PBS

**Figure 2**



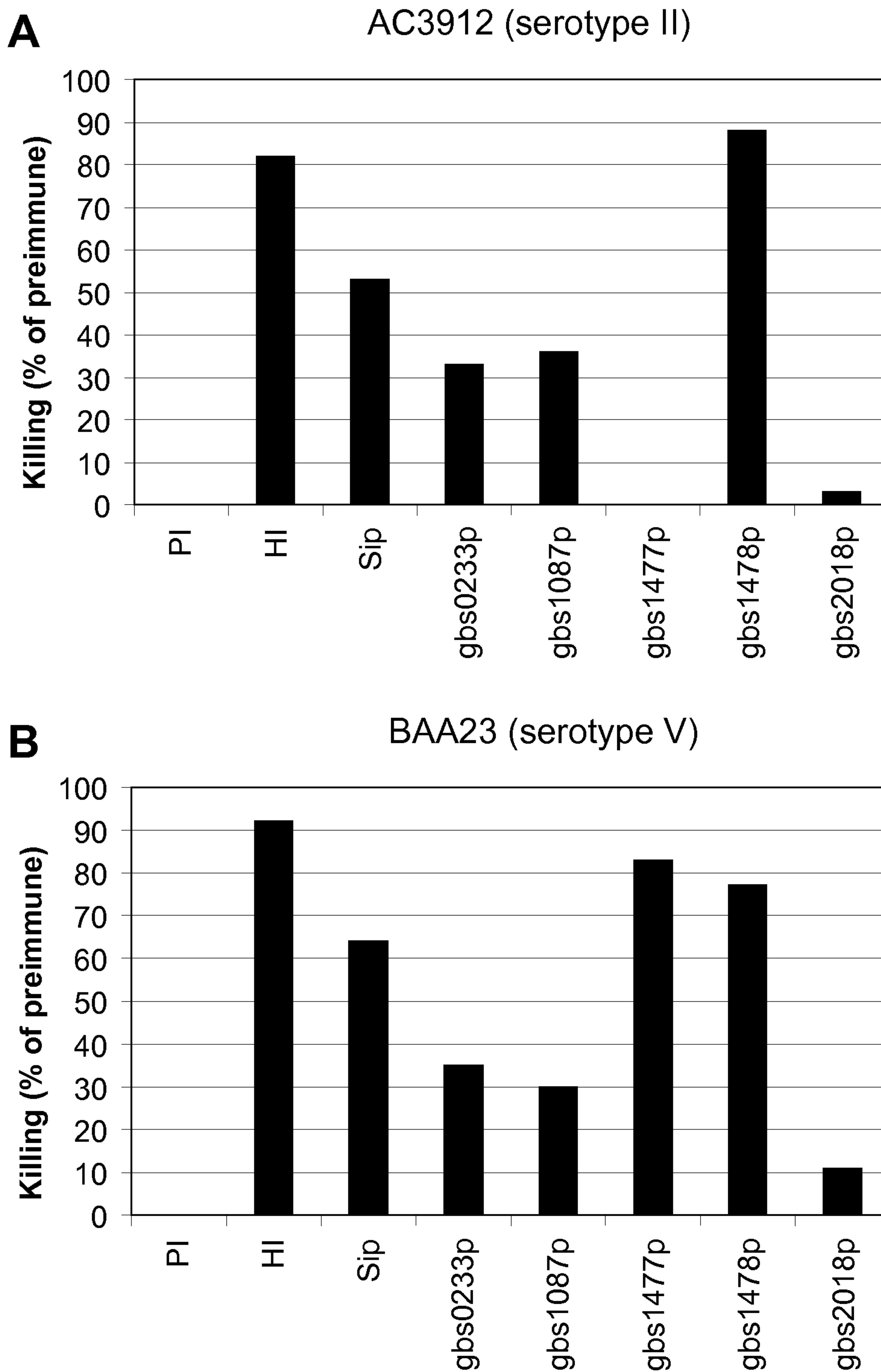
12403 (serotype III)



**Fluorescence intensity**

**Figure 3**





**Figure 4**



		1		50
NEM316	(1)	<b>MKKINKCLTVFSTLLLILTSLSVAPAFADDV--TTDTVTLHKIVMPQAA</b>		
CJB111	(1)	<b>MKKINKCLTMFSTLLLILTSLSVAPAFADDA--TTDTVTLHKIVMPQAA</b>		
BAA23	(1)	<b>MKKINKCLTMFSTLLLILTSLSVAPAFADDA--TTDTVTLHKIVMPQAA</b>		
515	(1)	<b>MKKINKYFAVFSALLLTVTSLSVAPVFAEEAK-TTDTVTLHKIVMPRTA</b>		
0176H4A	(1)	<b>MKKINKFFVAFSALLLILTSLLSVAPAFAEKEK-TTETVTLHKILQTDN</b>		
12401	(1)	<b>MKRINKYFAMFSALLLILTSLLSVAPVFAAEMGNITKTVTLHKIVQTSN</b>		
H36B	(1)	<b>MKRINKYFAMFSALLLILTSLLSVAPVFAAEMGNITKTVTLHKIVQTSN</b>		
IC105	(1)	<b>MKRINKYFAMFSALLLILTSLLSVAPVFAAEMGNITKTVTLHKIVQTSN</b>		
2603V/R	(1)	<b>MKRINKYFAMFSALLLTLTSLLSVAPAFADDA--TTNTVTLHKILQTESN</b>		
18RS21	(1)	<b>MKRINKYFAMFSALLLTLTSLLSVAPAFADDA--TTNTVTLHKILQTESN</b>		
IC458	(1)	<b>MKKINKYFAVFSALLLTVTSLLSVAPAFADDA--TTNTVTLHKILQTESN</b>		
Consensus	(1)	<b>MKKINKYFAMFSALLLILTSLLSVAPAFADDA TT TVTLHKIVQT AN</b>		
		51		100
NEM316	(49)	<b>FDN-FTEGKKGKNDSDYVGKQINDLKS YFGSTDAKEIKGAFFVFKNETGT</b>		
CJB111	(49)	<b>FDN-FTEGKKGKNDSDYVGKQINDLKS YFGSTDAKEIKGAFFVFKNETGT</b>		
BAA23	(49)	<b>FDN-FTEGKKGKNDSDYVGKQINDLKS YFGSTDAKEIKGAFFVFKNETGT</b>		
515	(50)	<b>FDG-FTAGTKGKDNTDYVGKQIEDLKTYFGSGEAKIAGAYFAFKNEAGT</b>		
0176H4A	(50)	<b>LKNSAFPGTKGLDGYDGAIDKLDYFGND-SKDIGGAYFILANSKGE</b>		
12401	(51)	<b>LAKPNFPGINGLNGTKYMGQKLTDISGYFGQG-SKEIAGAFFAVMNESQT</b>		
H36B	(51)	<b>LAKPNFPGINGLNGTKYMGQKLTDISGYFGQG-SKEIAGAFFAVMNESQT</b>		
IC105	(51)	<b>LAKPNFPGINGLNGTKYMGQKLTDISGYFGQG-SKEIAGAFFAVMNESQT</b>		
2603V/R	(49)	<b>LNKSNFPGTTGLNGKDYKGAISDLAGYFGEG-SKEIEGAFALALKEDK</b>		
18RS21	(49)	<b>LNKSNFPGTTGLNGKDYKGAISDLAGYFGEG-SKEIEGAFALALKEDK</b>		
IC458	(49)	<b>LNKSNFPGTTGLNGDDYKGESISDLAEYFGSG-SKEIDGAFFALALEEEK</b>		
Consensus	(51)	<b>L K NFPGT GLNGTDYVG ISDLA YFG G SKEI GAFFAL NES T</b>		
		101		150
NEM316	(98)	<b>KFITENGKEVDTLAKDA-----EGGAVLSGLTK--DTGFAFN</b>		
CJB111	(98)	<b>KFITENGKEVDTLAKDA-----EGGAVLSGLTK--DNGFVFN</b>		
BAA23	(98)	<b>KFITENGKEVDTLAKDA-----EGGAVLSGLTK--DNGFVFN</b>		
515	(99)	<b>KYITENGEEVDLDTTDA-----KGC AVLKGLTT--DNGFKFN</b>		
0176H4A	(99)	<b>YIKANDKNKLKPEFSGNT-----PKTTLNISEAVGGLTEE-NAGIKFE</b>		
12401	(100)	<b>KYITESGTEVESIDAA-----GVLKGLTT--ENGITFN</b>		
H36B	(100)	<b>KYITESGTEVESIDA-----AG--VLKGLTT--ENGITFN</b>		
IC105	(100)	<b>KYITESGTEVESIDAA-----GVLKGLTT--ENGITFN</b>		
2603V/R	(98)	<b>SGKVQYVKAKEGNKLT PALINKDGTPEITVNIDEAVSGLTPEGDTGLVFN</b>		
18RS21	(98)	<b>SGKVQYVKAKEGNKLT PALINKDGTPEITVNIDEAVSGLTPEGDTGLVFN</b>		
IC458	(98)	<b>DGVVQYVKAKANDKLT PDLITK-GTPATTTKVEEAVGGLTT--GTGIVFN</b>		
Consensus	(101)	<b>KYITE GKEVETLDA A AVL GLT DNGI FN</b>		
		151		200
NEM316	(134)	<b>TAKLKGTYQIVELKEKS NYDNNGS ILADSKAVPVKITLPLVNNQGVVKDA</b>		
CJB111	(134)	<b>TAKLKGTYQIVELKEKS NYDNNGS ILADSKAVPVKITLPLVNNQGVVKDA</b>		
BAA23	(134)	<b>TAKLKGTYQIVELKEKS NYDNNGS ILADSKAVPVKITLPLVNNQGVVKDA</b>		
515	(135)	<b>TSKLTGTYQIVELKEKSTYNNDGS ILADSKAVPVKITLPLVNDNGVVKDA</b>		
0176H4A	(141)	<b>TTGLRGDFQIIELKDKSTYNNGGAILADSKAVPVKITLPLINKDGVVKDA</b>		
12401	(131)	<b>TANLKGTYQIVELLDKS NYKNGDKVLADSKAVPVKITLPLYNEEGIVVDA</b>		
H36B	(131)	<b>TANLKGTYQIVELLDKS NYKNGDKVLADSKAVPVKITLPLYNEEGIVVDA</b>		
IC105	(131)	<b>TANLKGTYQIVELLDKS NYKNGDKVLADSKAVPVKITLPLYNEEGIVVDA</b>		
2603V/R	(148)	<b>TKGLKGEFKIVEVKSSTYNNGSLLAASKAVPVNITLPLVNEEDGVVADA</b>		
18RS21	(148)	<b>TKGLKGEFKIVEVKSSTYNNGSLLAASKAVPVNITLPLVNEEDGVVADA</b>		
IC458	(145)	<b>TAGLKGNFKIIELKDKSTYNNGSLLAASKAVPVKITLPLVSKDGVVKDA</b>		
Consensus	(151)	<b>TA LKG YQIVELKDKS NY NNGS ILADSKAVPVKITLPLVNEEDGVVKDA</b>		

Figure 5A



		201		250
NEM316	(184)	HIYPKNTETKPQVDKNFADK-----	DLDYTDNRKDKGVVS	
CJB111	(184)	HIYPKNTETKPQVDKNFADK-----	DLDYTDNRKDKGVVS	
BAA23	(184)	HIYPKNTETKPQVDKNFADK-----	DLDYTDNRKDKGVVS	
515	(185)	HVYPKNTETKPQVDKNFADK-----	ELDYANNKKDKGTVS	
0176H4A	(191)	HVYPKNTETKPQIDKNFADK-----	NLDYINNQKDKGTIS	
12401	(181)	EVYPKNTTEEAPQIDKNFAKANKLLNDS-	NSAIAGGADYDKYQAEKAKAT	
H36B	(181)	EVYPKNTTEEAPQIDKNFAKANKLLNDS-	NSAIAGGADYDKYQAEKAKAT	
IC105	(181)	EVYPKNTTEEAPQIDKNFAKANKLLNDS-	NSAIAGGADYDKYQAEKAKAT	
2603V/R	(198)	HVYPKNTTEEKPEIDKNFAKTNDLTALT-	DVNRLLTAGANYGNYARDKATAT	
18RS21	(198)	HVYPKNTTEEKPEIDKNFAKTNDLTALT-	DVNRLLTAGANYGNYARDKATAT	
IC458	(195)	HVYPKNTTETKPEVDKNFAKTNDLTALK-	D-ATLLKAGADYKNYSATKATVT	
Consensus	(201)	HVYPKNTETKPQIDKNFAK N L D I	AGADY Y KDKA VT	
		251		300
NEM316	(219)	ATVGDKKEYIVGTKILKGS DYKKLWTD-	SMTKGLTFN NN VKVTLD --- GK	
CJB111	(219)	ATVGDKKEYIVGTKILKGS DYKKLWTD-	SMTKGLTFN NN VKVTLD --- GE	
BAA23	(219)	ATVGDKKEYIVGTKILKGS DYKKLWTD-	SMTKGLTFN NN VKVTLD --- GE	
515	(220)	ASVGDVKKYHVGTKILKGS DYKKLWTD-	SMTKGLTFN ND IAVTLD --- GA	
0176H4A	(226)	ATVGDVKKYTVGTKILKGS DYKKLWTD-	SMTKGLTFN ND VTVTLD --- GA	
12401	(230)	AEIGQEIPYEVKTKIQKGS KYKNLAWVD-	TMSNGLTMGNTVNLEASS --- GS	
H36B	(230)	AEIGQEIPYEVKTKIQKGS KYKNLAWVD-	TMSNGLTMGNTVNLEASS --- GS	
IC105	(230)	AEIGQEIPYEVKTKIQKGS KYKNLAWVD-	TMSNGLTMGNTVNLEASS --- GS	
2603V/R	(248)	AEIGKVVPYEVKTKIHKGS KYENLVWTD-	IMSNGMTMGSTVSLKASGTTET	
18RS21	(248)	AEIGKVVPYEVKTKIHKGS KYENLVWTD-	IMSNGMTMGSTVSLKASGTTET	
IC458	(244)	AEIGKVIIPYEVKTKVLKGS KYEKLWTD-	TMSNGLTMGDDVNLA VSGTTTT	
Consensus	(251)	AEIG IPYEVKTKILKGS KYKKLWTD-	SMSNGLTMGN V L LS GS	
		301		350
NEM316	(266)	DFPVLNYKLVTD DQGFRLALNATGLAA-	VAAA AAKDKDVEIKITYSATVNGS	
CJB111	(266)	DFPVLNYKLVTD DQGFRLALNATGLAA-	VAAA AAKDKDVEIKITYSATVNGS	
BAA23	(266)	DFPVLNYKLVTD DQGFRLALNATGLAA-	VAAA AAKDKDVEIKITYSATVNGS	
515	(267)	TLDATNYKLVADDQGFRLVLT DKGLE-	AVAKAAKT KDVEIKITYSATLNGS	
0176H4A	(273)	NFEQSNYTLVADDQGFRLV LNATGLSK-	VAAEAAKT KDVEIKINYSATVNGS	
12401	(278)	FVEGTDYNVERDDRGF TLKFTDTGLTK-	LQKEAETQAVEFTLTYSATVNGA	
H36B	(278)	FVEGTDYNVERDDRGF TLKFTDTGLTK-	LQKEAETQAVEFTLTYSATVNGA	
IC105	(278)	FVEGTDYNVERDDRGF TLKFTDTGLTK-	LQKEAETHAVEFTLTYSATVNGA	
2603V/R	(298)	FAKDTDYELSIDARGF TLKFTADGLGK-	LEKAAKTADIEFTLTYSATVNGQ	
18RS21	(298)	FAKDTDYELSIDARGF TLKFTADGLGK-	LEKAAKTADIEFTLTYSATVNGQ	
IC458	(294)	FIKDIDYTLSIDDRGF TLKFKATGLDK-	LEEAASDVEFTLTYSATVNGQ	
Consensus	(301)	F TDY L DDRGF TLKFTATGL KL	KA AKT DVEFTLTYSATVNGS	
		351		400
NEM316	(316)	TTVEVPETNDVKLDYGN NPTEESE-	PQEGTPANQEIKVIKDWA VDG ----	
CJB111	(316)	TTVEIPETNDVKLDYGN NPTEESE-	PQEGTPANQEIKVIKDWA VDG ----	
BAA23	(316)	TTVEIPETNDVKLDYGN NPTEESE-	PQEGTPANQEIKVIKDWA VDG ----	
515	(317)	AVVEVLETNDVKLDYGN NPTEIENE-	PKEGIPVDKKITVNKTWA VDG ----	
0176H4A	(323)	TVVEKSENNDVKLDYGN NPTEIENE-	PQTGNPNKEITVRKTWA VDG ----	
12401	(328)	AIDDKPESNDIKLQYGNKPGK KVK-	EIPVTPSNGEITVSKTW DKGSDLENA	
H36B	(328)	AIDDKPESNDIKLQYGNKPGK KVK-	EIPVTPSNGEITVSKTW DKGSDLENA	
IC105	(328)	AIDDKPESNDIKLQYGNKPGK KVK-	EIPVTPSNGEITVSKTW DKGSDLENA	
2603V/R	(348)	AIIDNPESNDIKLSYGNKPGKDLTEL-	PVTPSKGEVTVAKTWS DGIAPDGV	
18RS21	(348)	AIIDNPESNDIKLSYGNKPGKDLTEL-	PVTPSKGEVTVAKTWS DGIAPDGV	
IC458	(344)	AIIDNPEVNDIKLDYGNKPGTDLSE-	QPVTPEDGEVKVTKTWAAGANKADA	
Consensus	(351)	AIVD PESNDIKLDYGNKPG EL E	PVTPSNGEITV KTW A GG	

Figure 5B



		401		450
NEM316	(361)	----TITDVNVAVKAIFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLD		
CJB111	(361)	---TITDANVA-VKAIFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLD		
BAA23	(361)	---TITDANVA-VKAIFTLQEKQTDGTWVNVASHEATKPSRFEHTFTGLD		
515	(363)	---EVNKADET-VDAVFTLQVKDGD-KWVNVDSAKATAATSFKHTFENLD		
0176H4A	(369)	----EVNKGDEKVDVAVFTLQVKDSD-KWVNVDSATATAATDFKYTFKNLD		
12401	(378)	NVVYTLKDGGT-AVASVSLTKTTPNGEINLGNGIKFTVTGAFAGKFSGLT		
H36B	(378)	NVVYTLKDGGT-AVASVSLTKTTPNGEINLGNGIKFTVTGAFAGKFSGLT		
IC105	(378)	NVVYTLKDGGT-AVASVSLTKTTPNGEINLGNGIKFTVTGAFAGKFSGLT		
2603V/R	(398)	NVVYTLKDKDK-TVASVSLTKTSKG-TIDLNGNGIKFEVSGNFSGKFTGLE		
18RS21	(398)	NVVYTLKDKDK-TVASVSLTKTSKG-TIDLNGNGIKFEVSGNFSGKFTGLE		
IC458	(394)	KVVYTLKNATKQVVASVALTAADTKGTINLGKGMTFEITGAFSGTFKGLQ		
Consensus	(401)	VVYTLKD VVASVSLT GTI LG GIKFTVTG FAGTFTGLD		
		451		500
NEM316	(407)	NTKTYRVVERVSGYTPEYVSFKNGVVTIKNNKNSNDPTPINPSEPKVVTY		
CJB111	(407)	NAKTYRVVERVSGYTPEYVSFKNGVVTIKNNKNSNDPTPINPSEPKVVTY		
BAA23	(407)	NAKTYRVVERVSGYTPEYVSFKNGVVTIKNNKNSNDPTPINPSEPKVVTY		
515	(408)	NAKTYRVIERVSGYAPEYVSFVNGVVTIKNNKDSNEPTPINPSEPKVVTY		
0176H4A	(414)	NAKTYRVVERVSGYAPAYVSFVGGVVTIKNNKNSNDPTPINPSEPKVVTY		
12401	(427)	DSKTYMISERIAGYG-NTITTGAGSAAITNTPDSDNPTPLNPTEPKVVTH		
H36B	(427)	DSKTYMISERIAGYG-NTITTGAGSAAITNTPDSDNPTPLNPTEPKVVTH		
IC105	(427)	DSKTYMISERIAGYG-NTITTGAGSAAITNTPDSDNPTPLNPTEPKVVTH		
2603V/R	(446)	N-KSYMISERVSGYG-SAINLENGKVTITNTKDSDNPTPLNPTEPKVETH		
18RS21	(446)	N-KSYMISERVSGYG-SAINLENGKVTITNTKDSDNPTPLNPTEPKVETH		
IC458	(444)	N-KAYTVSERVAGYT-NAINVTGNAVAITNTPDSDNPTPLNPTEPKVETH		
Consensus	(451)	NAKTY VSERVSGYG IS NG VTIITNTKDSDNPTPLNPTEPKVVTH		
		501		550
NEM316	(457)	GRKFVKTNQANTERLAGATFLVKKE-GKYLARKAGAATAEAKAAVKTAKL		
CJB111	(457)	GRKFVKTNQANTERLAGATFLVKKE-GKYLARKAGAATAEAKAAVKTAKL		
BAA23	(457)	GRKFVKTNQANTERLAGATFLVKKE-GKYLARKAGAATAEAKAAVKTAKL		
515	(458)	GRKFVKTNKDGKERLAGATFLVKKD-GKYLARKSGVATDAEAKAAVDSTKS		
0176H4A	(464)	GRKFVKTNQDGSERLAGATFLVKNSQSQYLARKSGVATNEAHKAVTDAKV		
12401	(476)	GKKFVKTSSTETERLQGAQFVVKDSAGKYLALKSSATISAQTTAYTNAKT		
H36B	(476)	GKKFVKTSSTETERLQGAQFVVKDSAGKYLALKSSATISAQTTAYTNAKT		
IC105	(476)	GKKFVKTSSTETERLQGAQFVVKDSAGKYLALKSSATISAQTTAYTNAKT		
2603V/R	(494)	GKKFVKTNEQG-DRLAGAQFVVKNSAGKYLALKADQSEGQKTAAK--KI		
18RS21	(494)	GKKFVKTNEQG-DRLAGAQFVVKNSAGKYLALKADQSEGQKTAAK--KI		
IC458	(492)	GKKFVKVGAD-ARLAGAQFVVKNSAGKFLALKEDAAVSGAQTELATAKT		
Consensus	(501)	GKKFVKTN TERLAGAQFVVK SAGKYLALKA AA S AV AKL		
		551		600
NEM316	(506)	ALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWV		
CJB111	(506)	ALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWV		
BAA23	(506)	ALDEAVKAYNDLTKEKQEGQEGKTALATVDQKQKAYNDAFVKANYSYEWV		
515	(507)	ALDAAVKAYNDLTKEKQEGQDGKSALATVSEKQKAYNDAFVKANYSYEWV		
0176H4A	(514)	QLDEAVKAYNKLTKEQQESQDGKAALNLIDEKQTAYNEAFKANYSYEWV		
12401	(526)	ALDAKIAAYNKLSADDQKGTGETAKAEIKTAQDAYNAAFIVARTAYEWV		
H36B	(526)	ALDAKIAAYNKLSADDQKGTGETAKAEIKTAQDAYNAAFIVARTAYEWV		
IC105	(526)	ALDAKIAAYNKLSADDQKGTGETAKAEIKTAQDAYNAAFIVARTAYEWV		
2603V/R	(541)	ALDEAIAAYNKLSATDQKGEKGITAKELIKTKQADYDAAFIEARTAYEWI		
18RS21	(541)	ALDEAIAAYNKLSATDQKGEKGITAKELIKTKQADYDAAFIEARTAYEWI		
IC458	(541)	DLDNAIKAYNGLTKAQQEGADGTSAKELINTKQSAFYDAAFIEARTAYTWV		
Consensus	(551)	ALDEAIAAYNKLTKE QEG DG TAKA I TKQ AYNAAFIEARTAYEWV		

Figure 5C

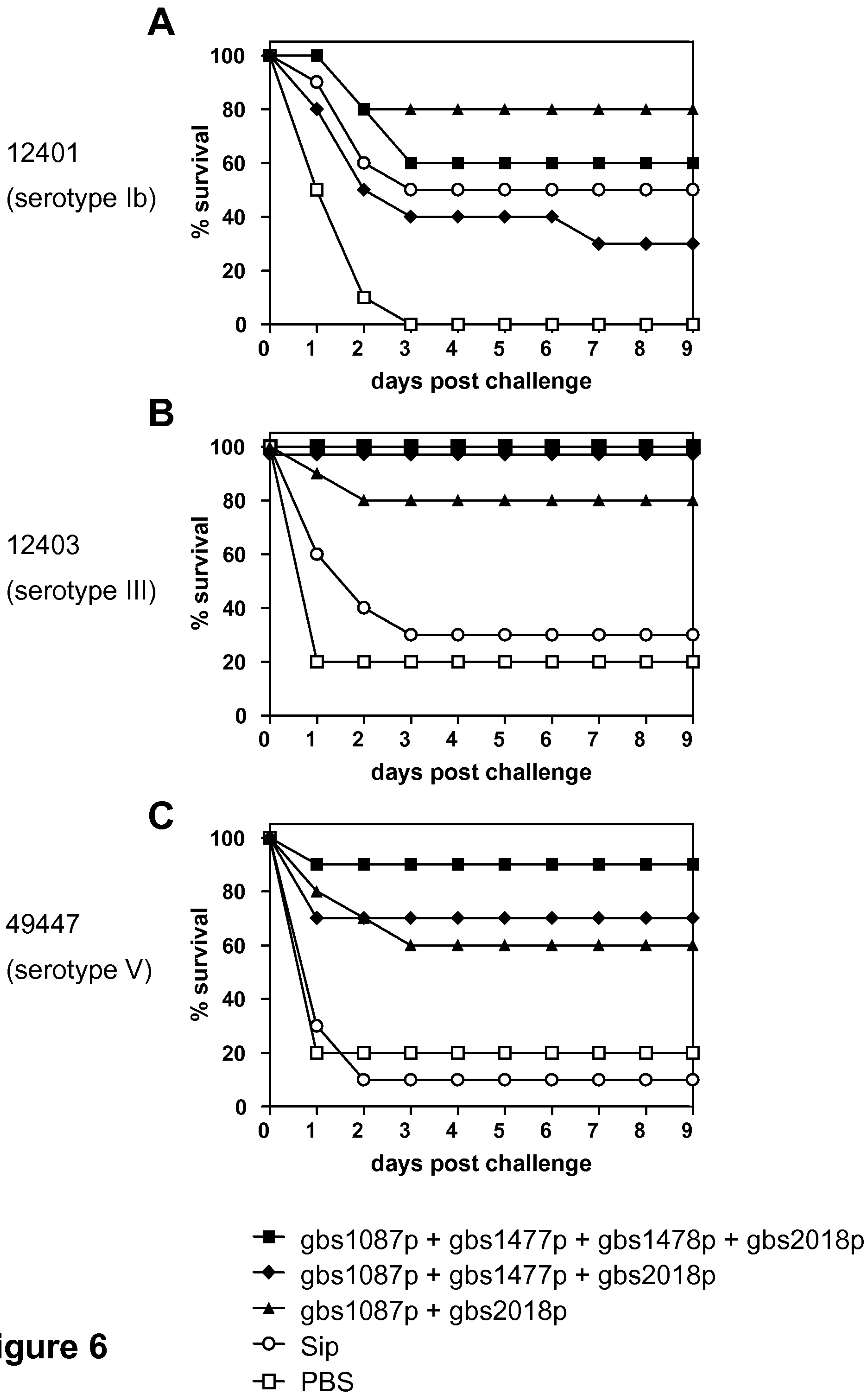


		601		650
NEM316	(556)	ADKKADNVVCLI	SNAGGQFEITGLDKGTYSLEETQAPAGYATLSGDVNFE	
CJB111	(556)	ADKKADNVVCLI	SNAGGQFEITGLDKGTYSLEETQAPAGYATLSGDVNFE	
BAA23	(556)	ADKKADNVVCLI	SNAGGQFEITGLDKGTYSLEETQAPAGYATLSGDVNFE	
515	(557)	EDKNAKNVVCLI	SNDKGQFEITGLTEGQYSLEETQAPTGYAKLSGDVSN	
0176H4A	(564)	VDKNAANVVCLI	SNTAGKFEITGLNAGEYSLEETQAPTGYAKLSGDVSN	
12401	(576)	TNKEDANVVKVTS	SNADGQFEVSGLATGDYKLEETQAPAGYAKLAGDVDFK	
H36B	(576)	TNKEDANVVKVTS	SNADGQFEVSGLATGDYKLEETQAPAGYAKLAGDVDFK	
IC105	(576)	TNKEDANVVKVTS	SNADGQFEVSGLATGDYKLEETQAPAGYAKLAGDVDFK	
2603V/R	(591)	TDK - -	ARAIYTSNDQGQFEVTGLADGTYNLEETLAPAGFAKLAGNIKFV	
18RS21	(591)	TDK - -	ARAIYTSNDQGQFEVTGLADGTYNLEETLAPAGFAKLAGNIKFV	
IC458	(591)	DEK - -	TKAITFTSNNQGQFEVTGLEVGSYKLEETLAPAGYAKLSGDIEFT	
Consensus	(601)	DK ANVVKLTSNA	GQFEVTGL GTY LEETQAPAGYAKLSGDV F	
		651		700
NEM316	(606)	VTATSYSKGATTDIAYDKGSVKKDAQQVQNKKVTIPQTGGIGTILFTIIG		
CJB111	(606)	VTATSYSKGATTDIAYDKGSVKKDAQQVQNKKVTIPQTGGIGTILFTIIG		
BAA23	(606)	VTATSYSKGATTDIAYDKGSVKKDAQQVQNKKVTIPQTGGIGTILFTIIG		
515	(607)	VNATSYSKGSAQDIEYTOGSKTKDAQQVINKKVTIPQTGGIGTIFFTIIG		
0176H4A	(614)	VNDTSYSEGASNDIAYDKDSGKTDAQKVVNKKVTIPQTGGIGTILFTIIG		
12401	(626)	VGNSSKAD-DSGNIDYTASSNKKDAQRIENKKVTIPQTGGIGTILFTIIG		
H36B	(626)	VGNSSKAD-DSGNIDYTASSNKKDAQRIENKKVTIPQTGGIGTILFTIIG		
IC105	(626)	VGNSSKAD-DSGNIDYTASSNKKDAQRIENKKVTIPQTGGIGTILFTIIG		
2603V/R	(639)	VNQGSYIT--GGNIDYVANSNQKDATRVENKKVTIPQTGGIGTILFTIIG		
18RS21	(639)	VNQGSYIT--GGNIDYVANSNQKDATRVENKKVTIPQTGGIGTILFTIIG		
IC458	(639)	VGHDSYTS---GDIKYKTDDASNNAQKVFNKKVTIPQTGGIGTILFTIIG		
Consensus	(651)	V TSYS SGDIDY S	KKDAQRV NKKVTIPQTGGIGTILFTIIG	
		701		720
NEM316	(656)	LSIMLGAVVIMKRRQSEEA*	(SEQ ID NO: 223)	
CJB111	(656)	LSIMLGAVVIMKRRQSEEA*	(SEQ ID NO: 224)	
BAA23	(656)	LSIMLGAVVIMKRRQSEEA*	(SEQ ID NO: 98)	
515	(657)	LSIMLGAVVIMKRRQSEEV*	(SEQ ID NO: 225)	
0176H4A	(664)	LSIMLGAVVIMKRRQSEEA*	(SEQ ID NO: 91)	
12401	(675)	LSIMLGAVIIMKRRQSEEA*	(SEQ ID NO: 93)	
H36B	(675)	LSIMLGAVIIMKRRQSEEA*	(SEQ ID NO: 226)	
IC105	(675)	LSIMLGAVIIMKRRQSEEA*	(SEQ ID NO: 103)	
2603V/R	(687)	LSIMLGAVVIMKRRQSKEA*	(SEQ ID NO: 227)	
18RS21	(687)	LSIMLGAVVIMKRRQSKEA*	(SEQ ID NO: 228)	
IC458	(686)	LSIMLGAVVIMKRRQSEEA*	(SEQ ID NO: 127)	
Consensus	(701)	LSIMLGAVVIMKRRQSEEA		

Figure 5D

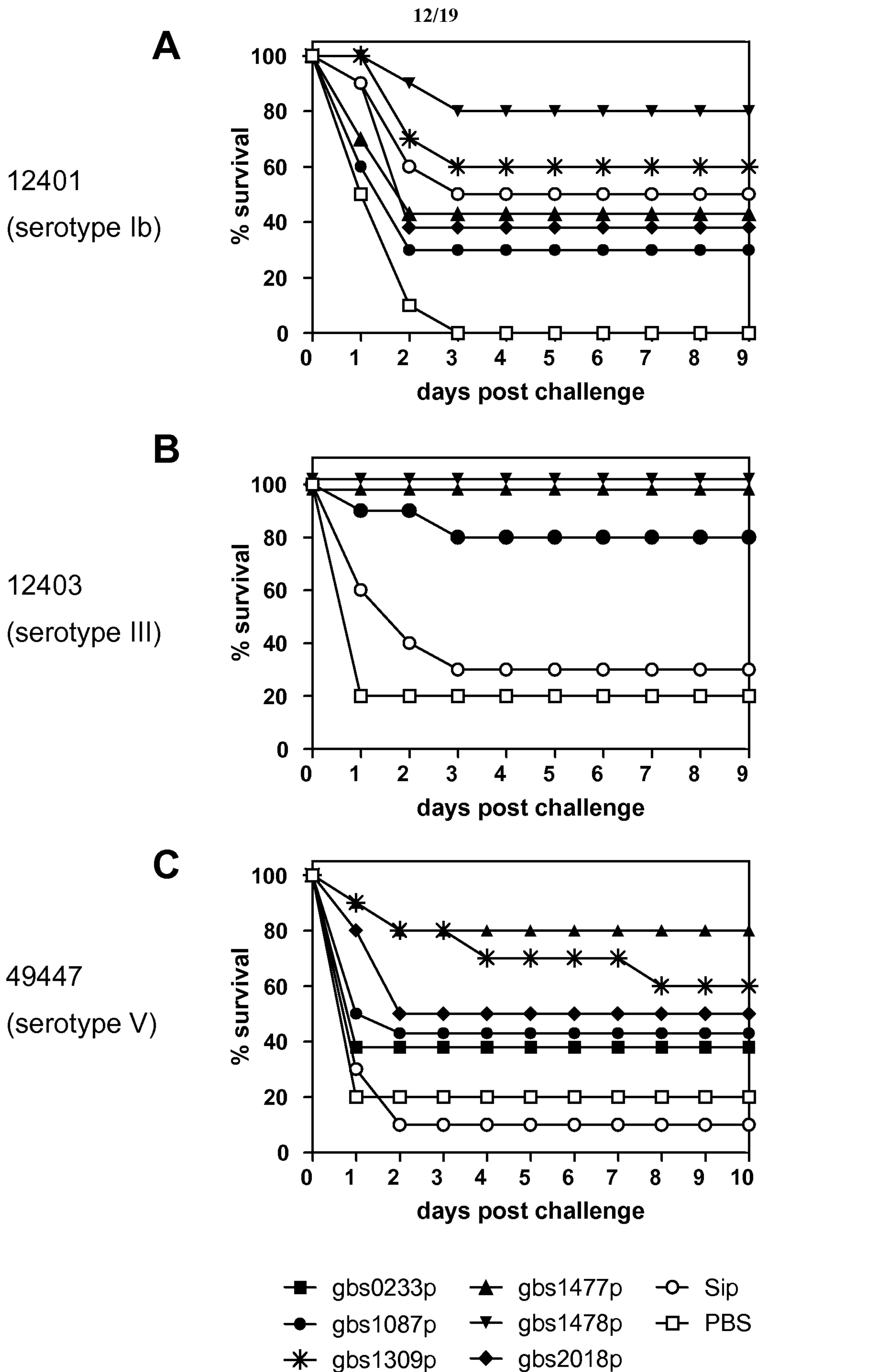


11/19



**Figure 6**





**Figure 7**



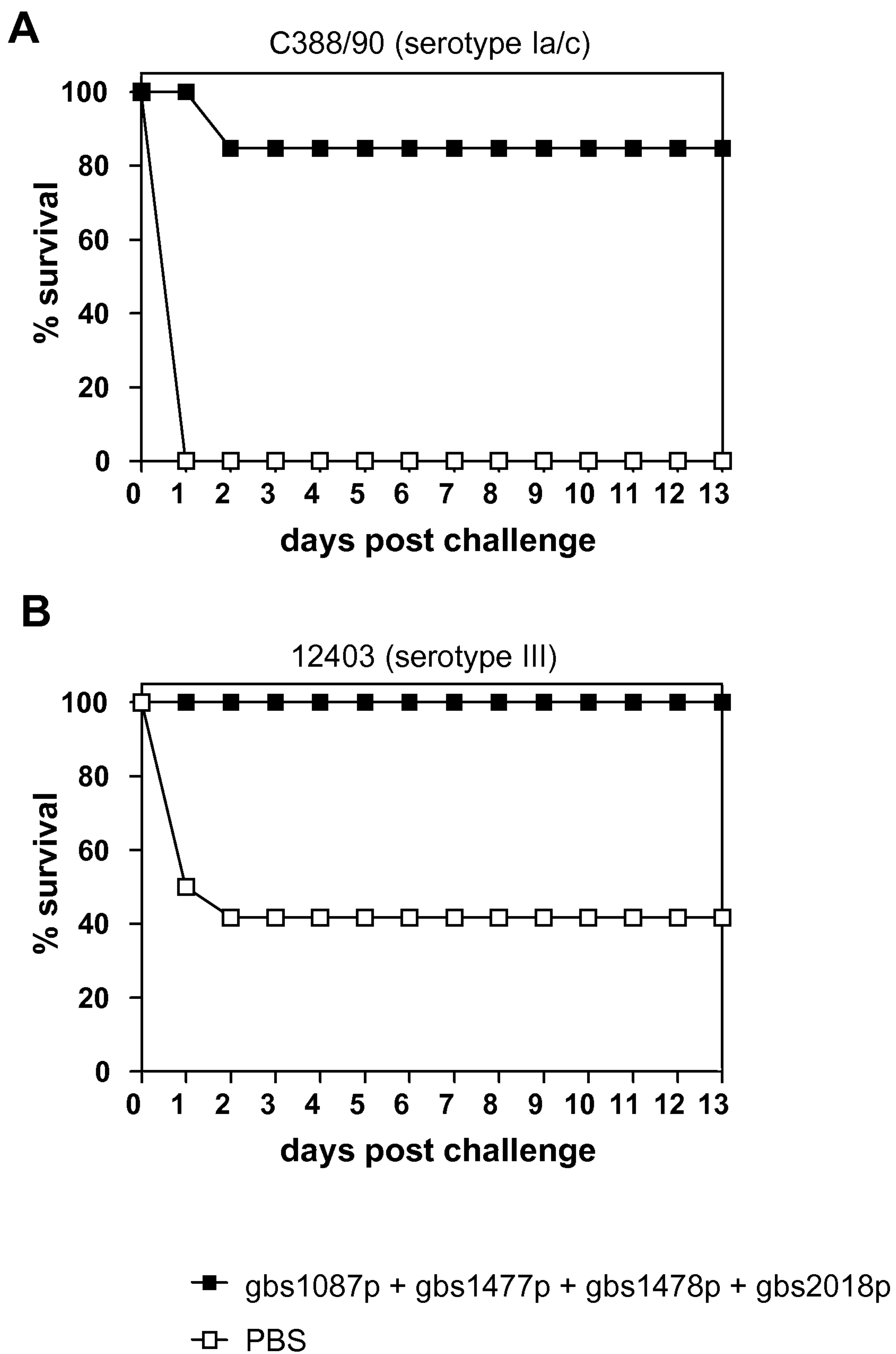
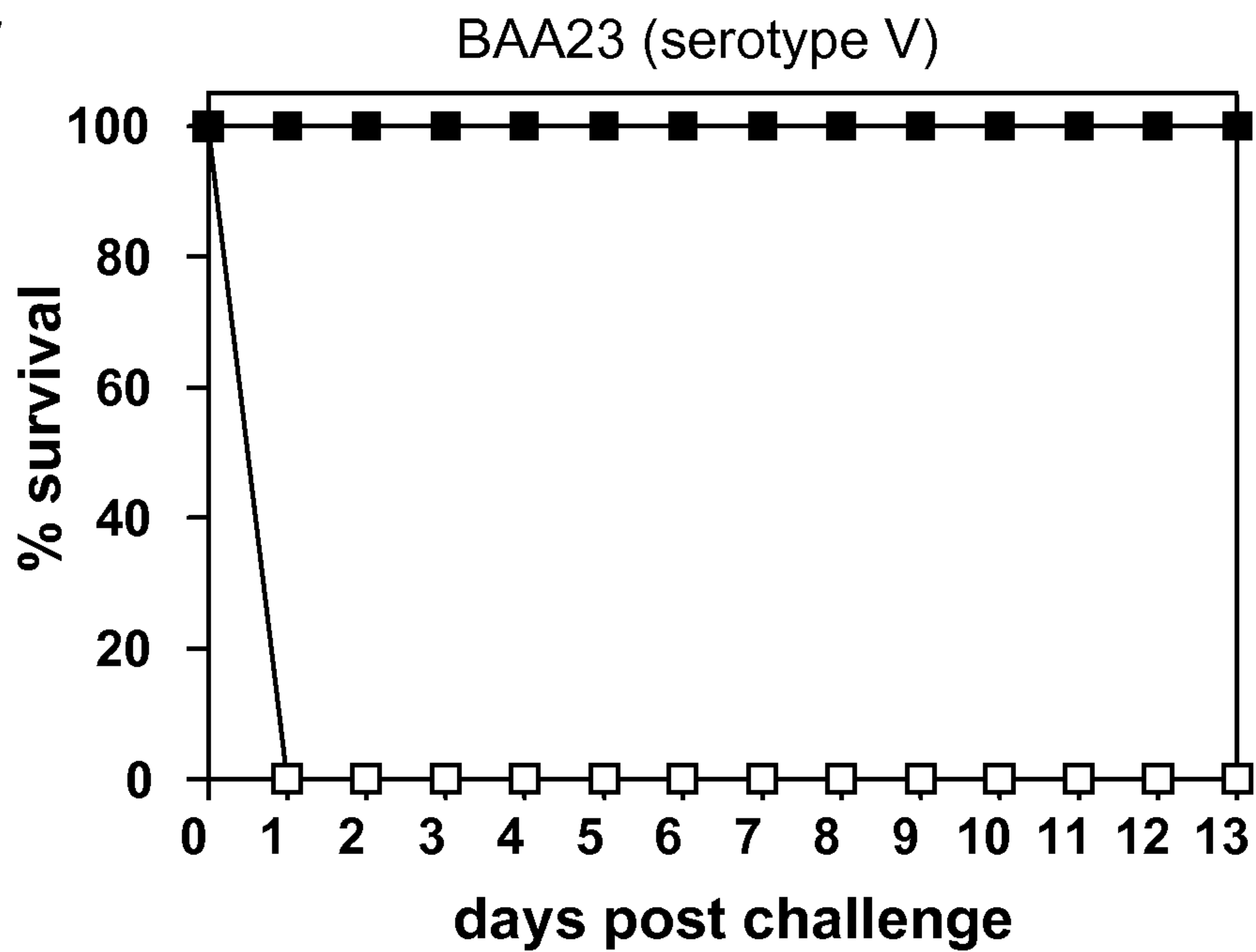


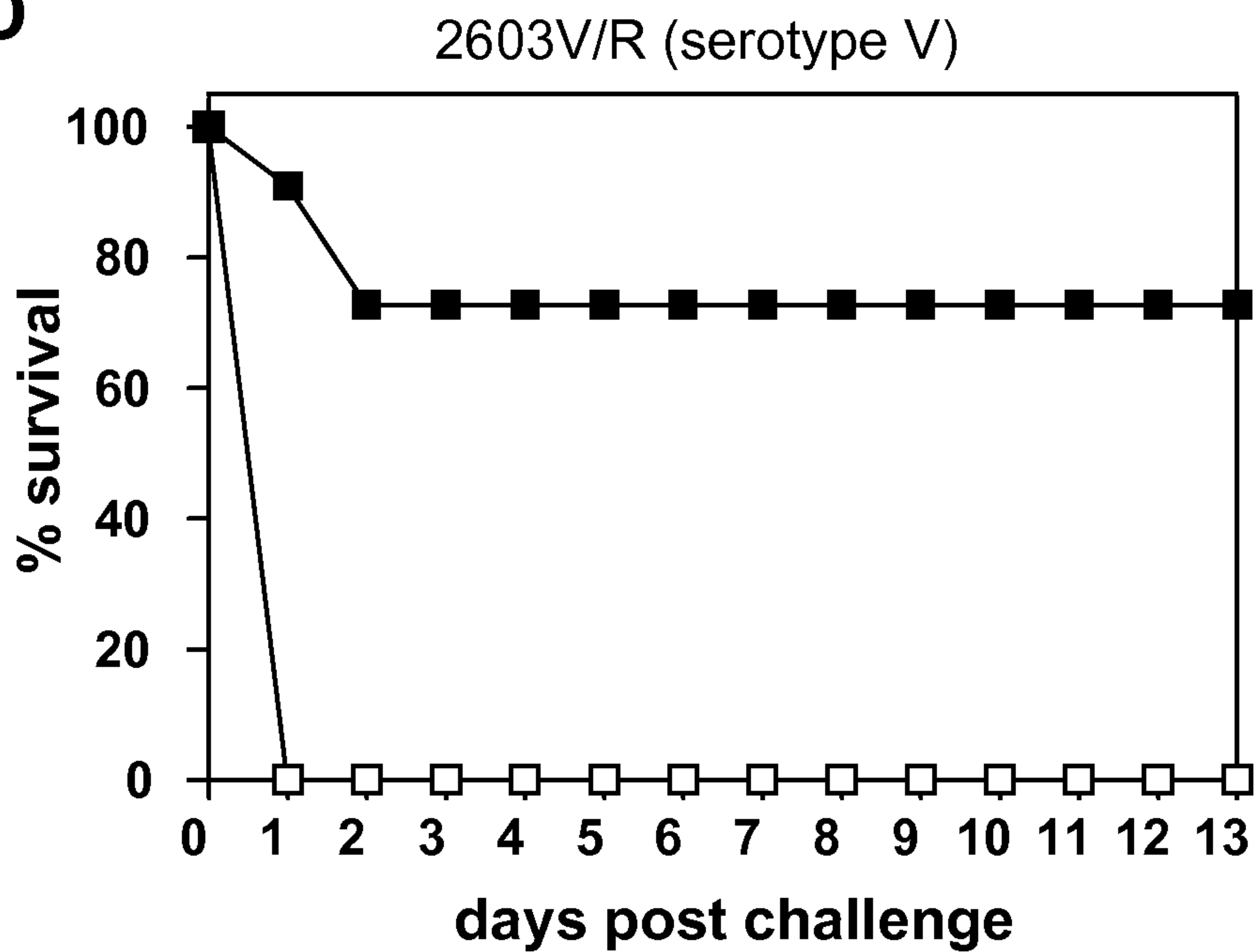
Figure 8



**C**



**D**



-■- gbs1087p + gbs1477p + gbs1478p + gbs2018p  
 -□- PBS

**Figure 8**



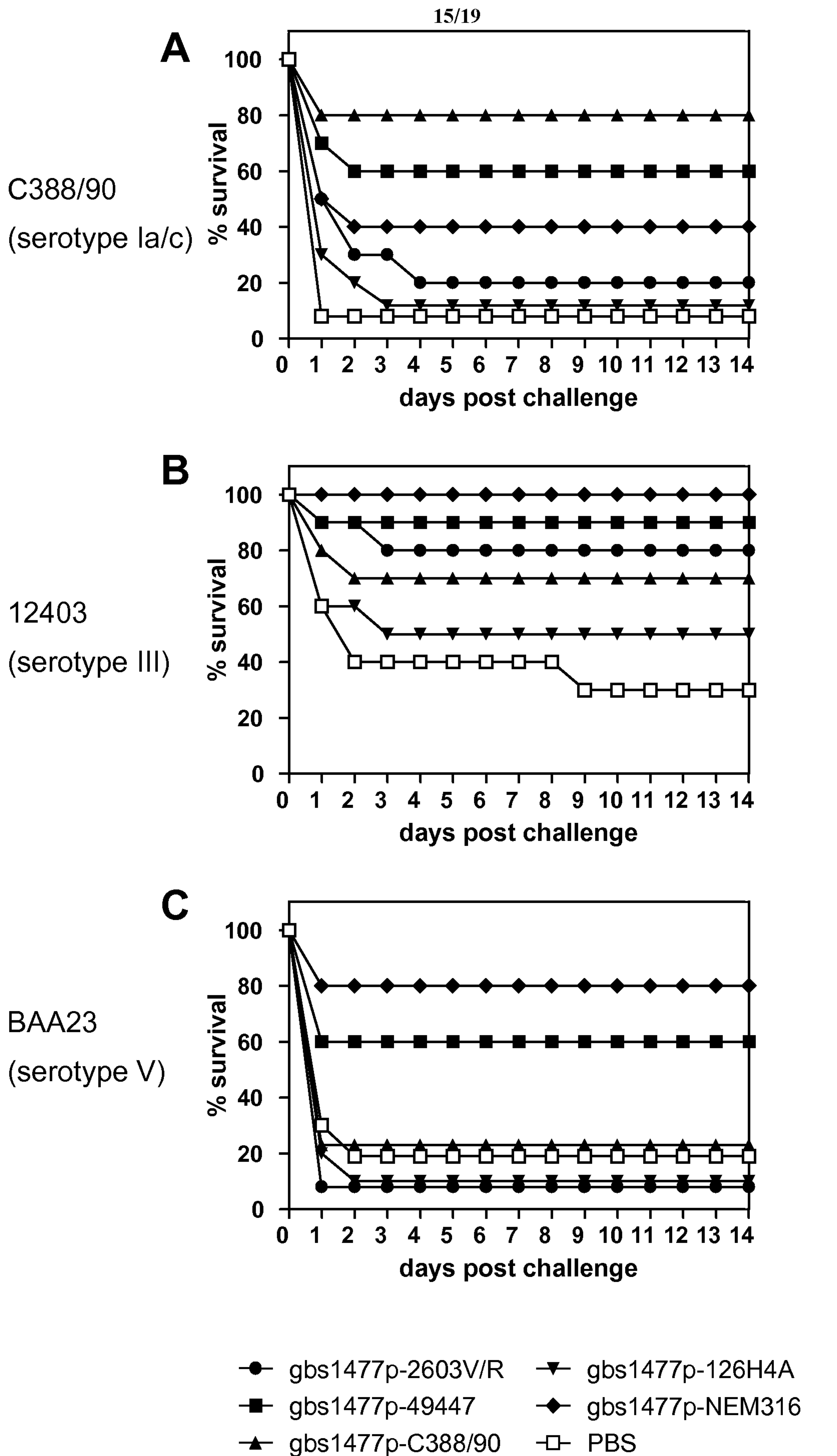


Figure 9



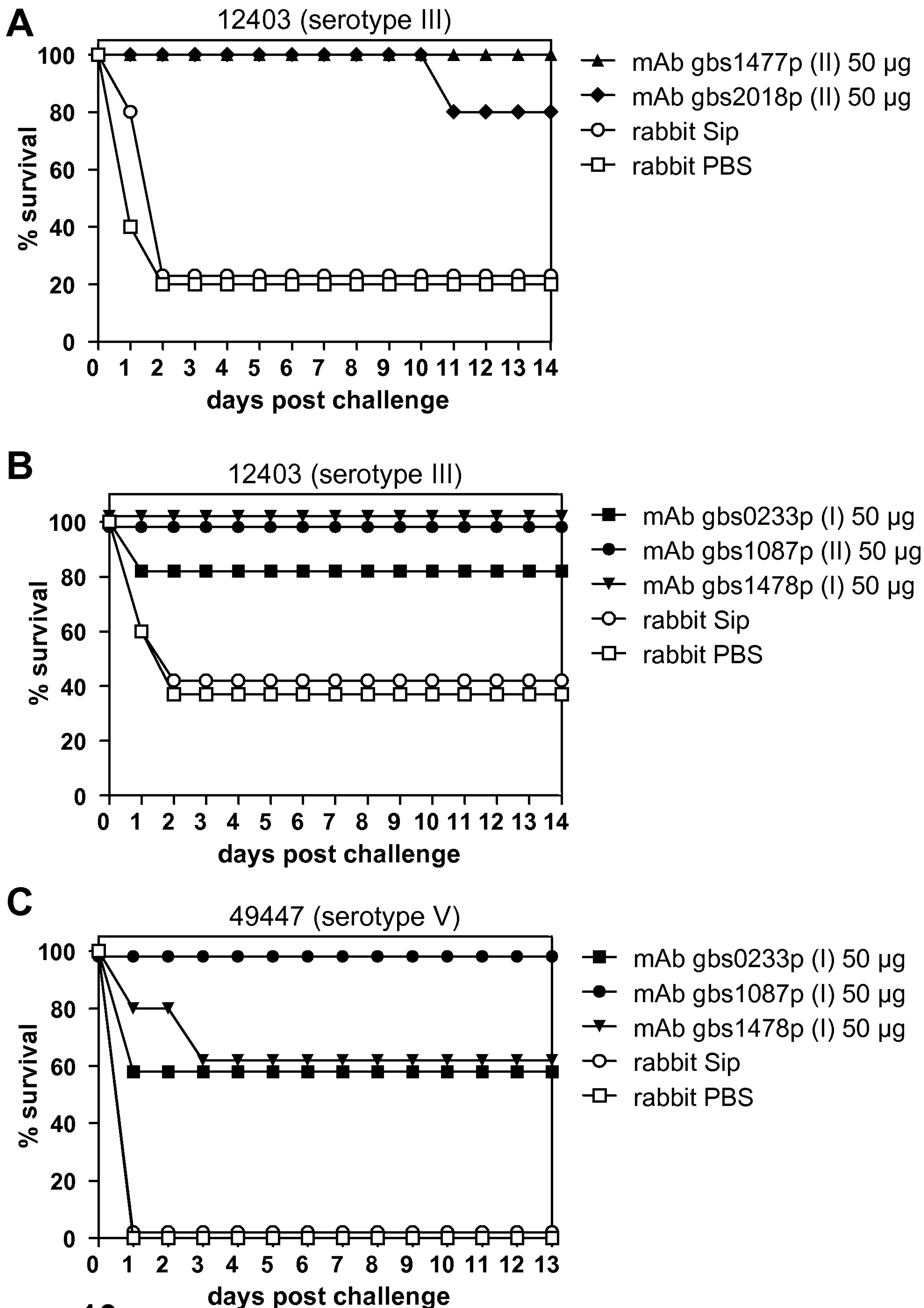
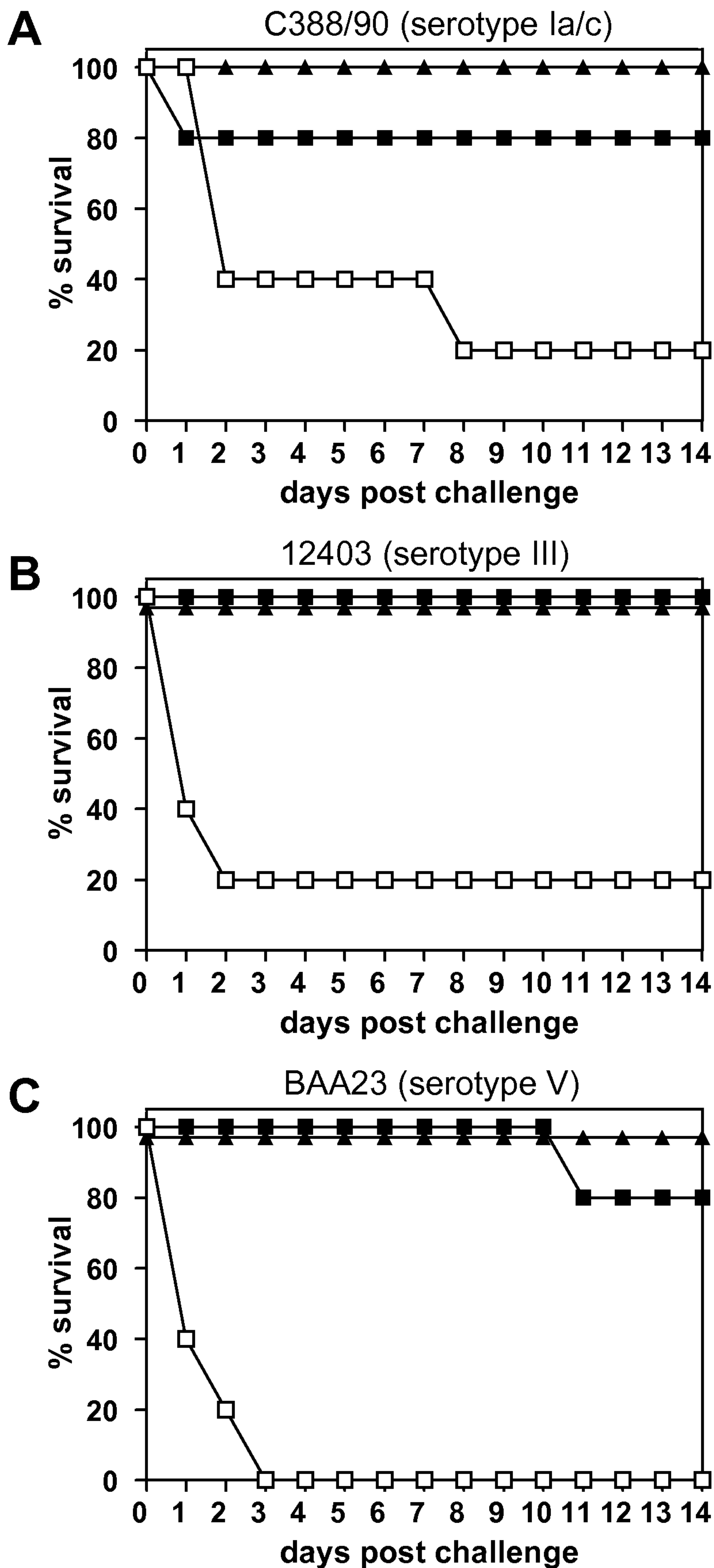


Figure 10



- mAbs (gbs0233p + gbs1087p + gbs1477p + gbs1478p + gbs2018p)
- ▲ rabbit sera (gbs0233p + gbs1087p + gbs1477p + gbs1478p + gbs2018p)
- rabbit PBS sera

**Figure 11**



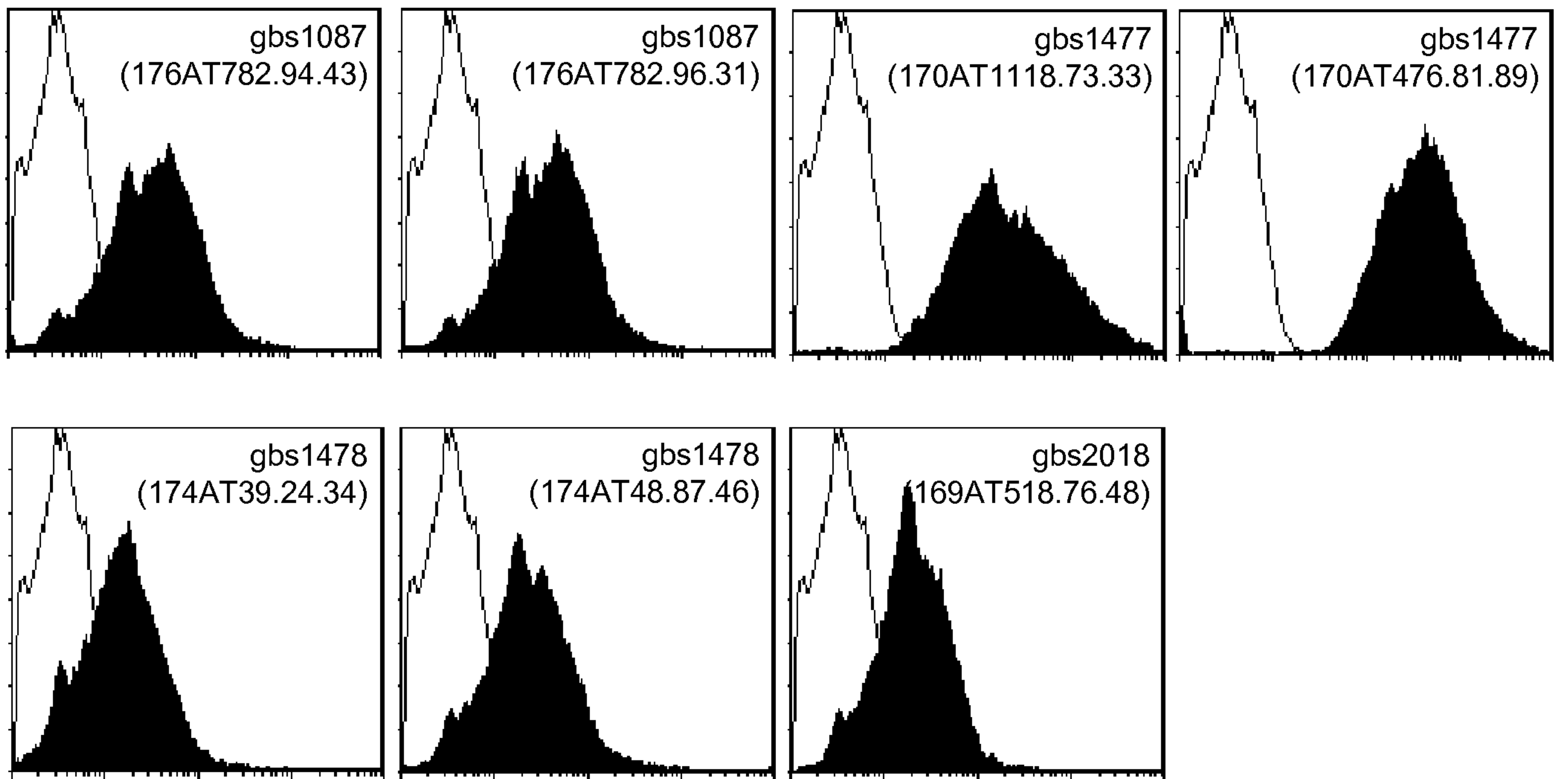


Figure 12

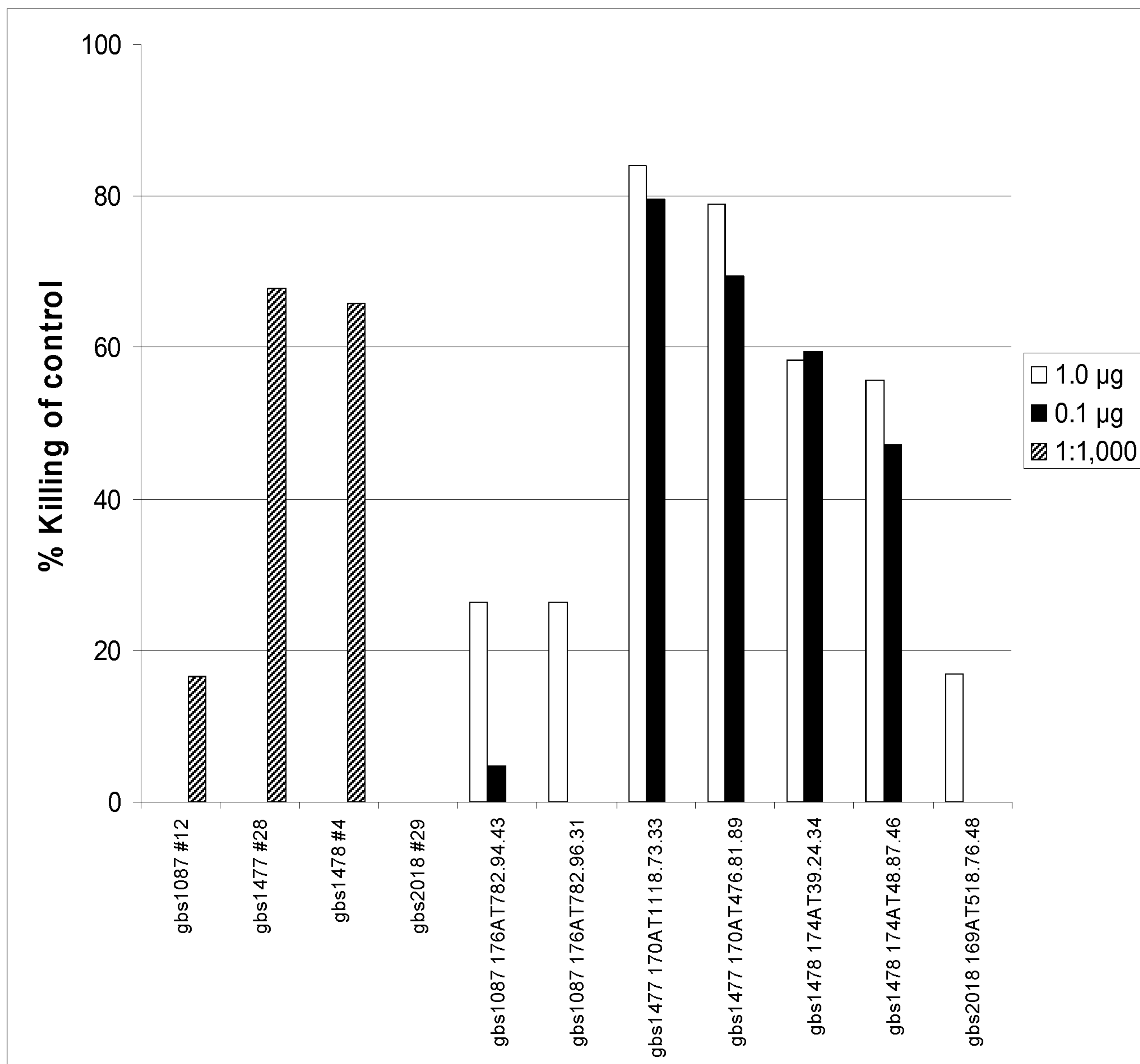


Figure 13