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Vu et al.

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(45) **Date of Patent: Aug. 11, 2020**

(54) **NON-NATURALLY OCCURRING PORCINE REPRODUCTIVE AND RESPIRATORY SYNDROME VIRUS (PRRSV) AND METHODS OF USING**

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C07K 14/005 (2006.01)

A61K 39/12 (2006.01)

C12N 7/00 (2006.01)

A61K 39/00 (2006.01)

(52) U.S. Cl.

CPC **C07K 14/005** (2013.01); **A61K 39/12** (2013.01); **C12N 7/00** (2013.01); **A61K 2039/552** (2013.01); **C12N 2770/10021** (2013.01); **C12N 2770/10022** (2013.01); **C12N 2770/10034** (2013.01)

(58) Field of Classification Search

None

See application file for complete search history.

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(57)

ABSTRACT

A non-naturally occurring porcine reproductive and respiratory syndrome virus (PRRSV) is provided herein, and methods of making and using the non-naturally occurring PRRSV also are provided.

11 Claims, 5 Drawing Sheets

Specification includes a Sequence Listing.

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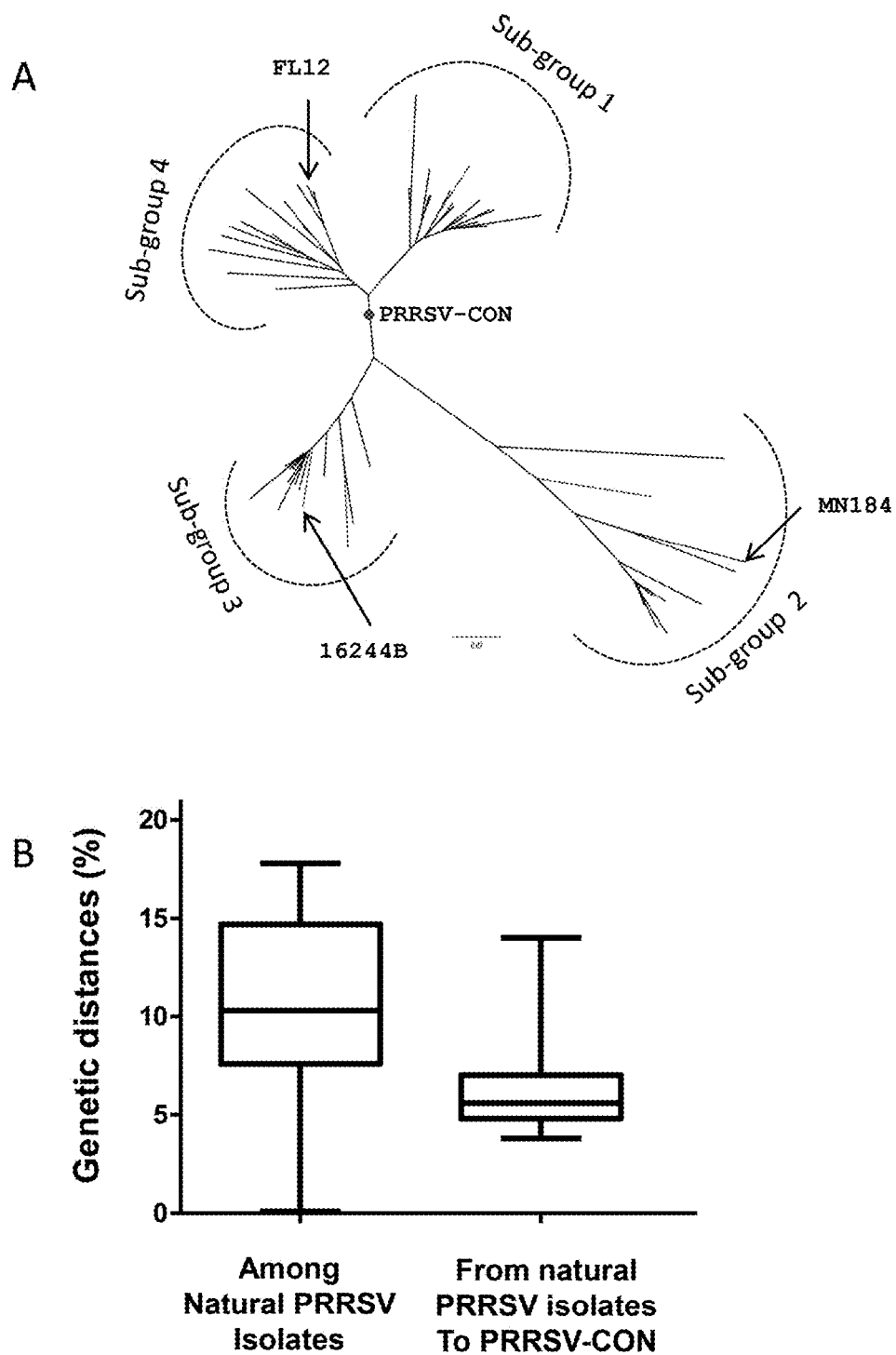
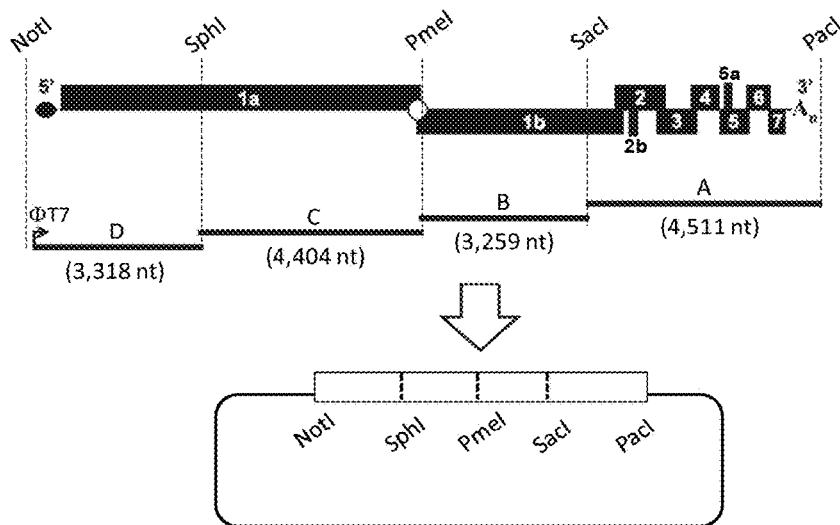
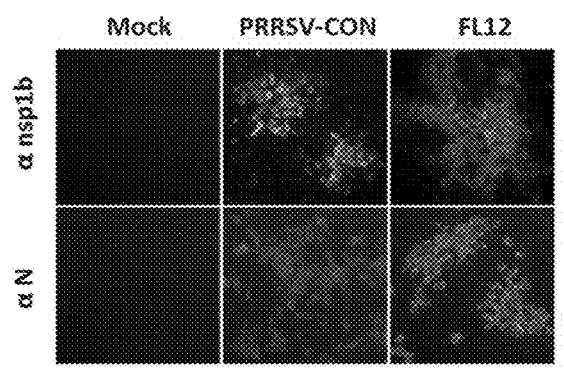


Figure 1

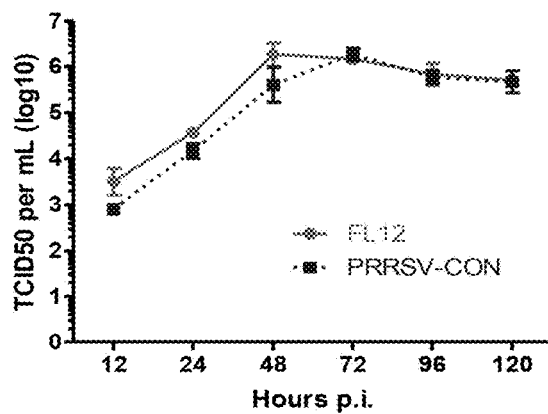
A



B



C



D

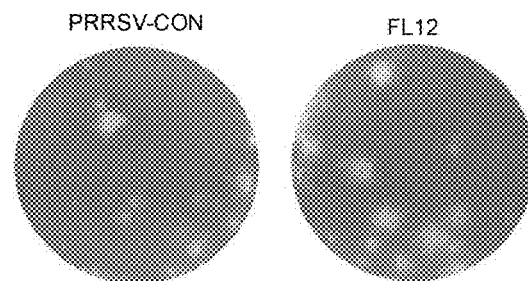


Figure 2

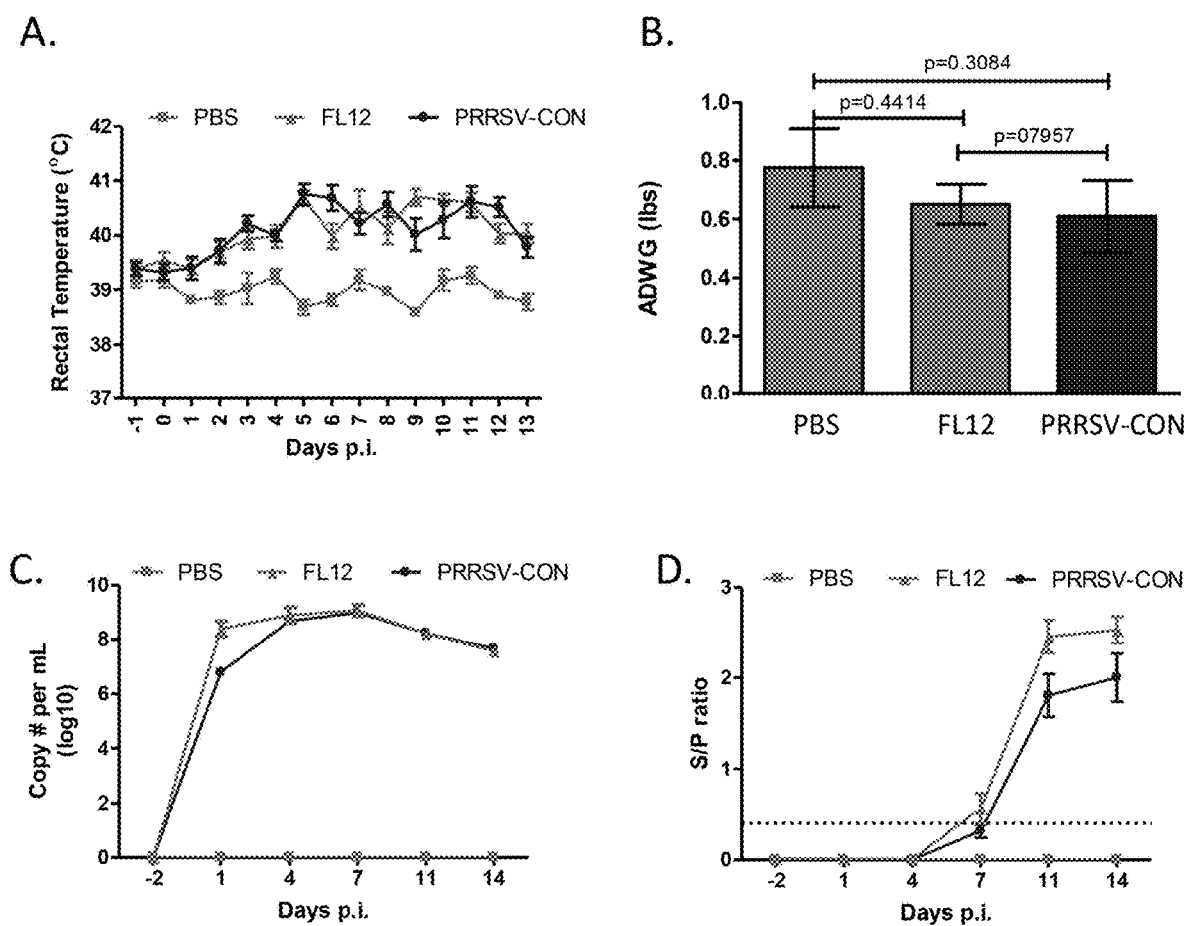


Figure 3

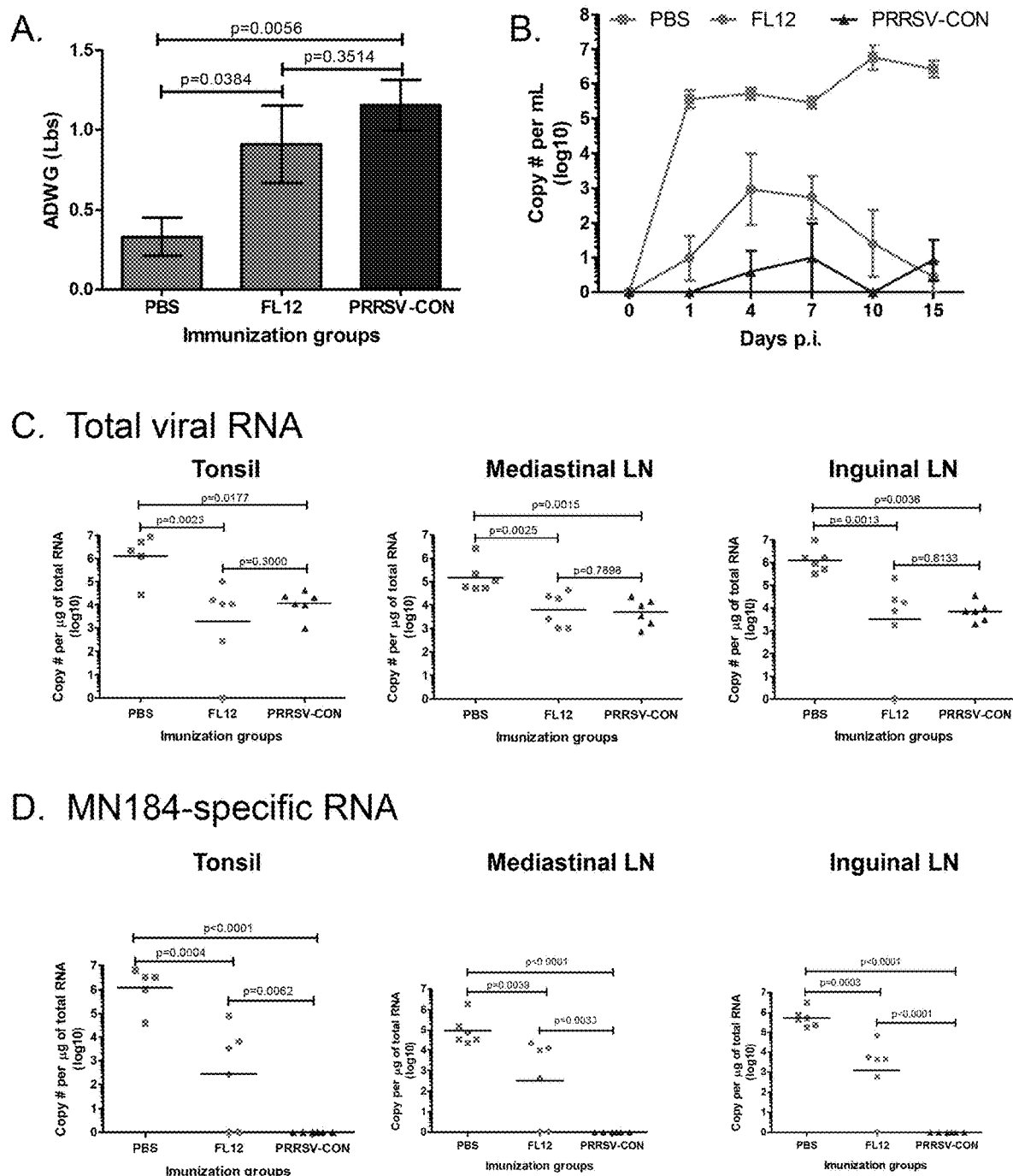
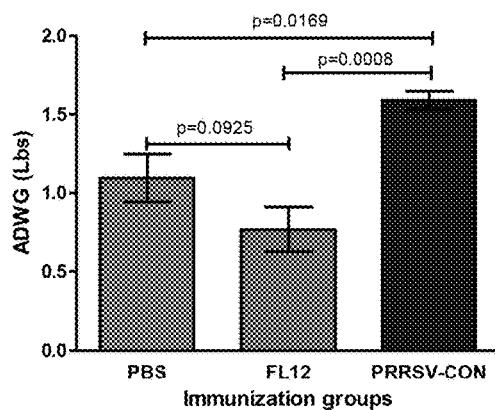
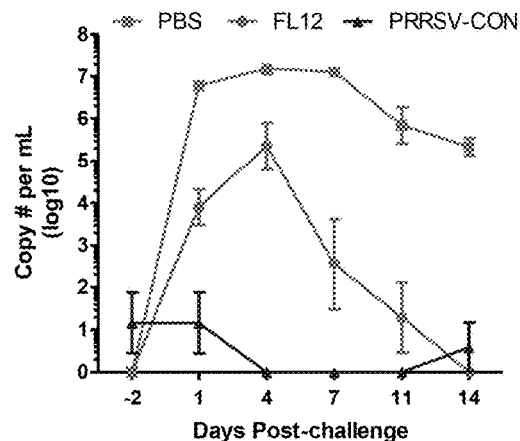


Figure 4

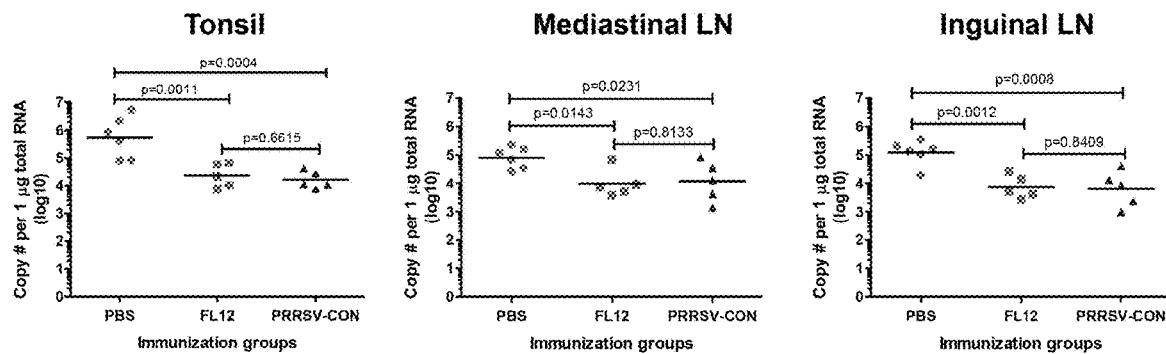
A.



B.



C. Total vRNA



D. 16244B-specific RNA

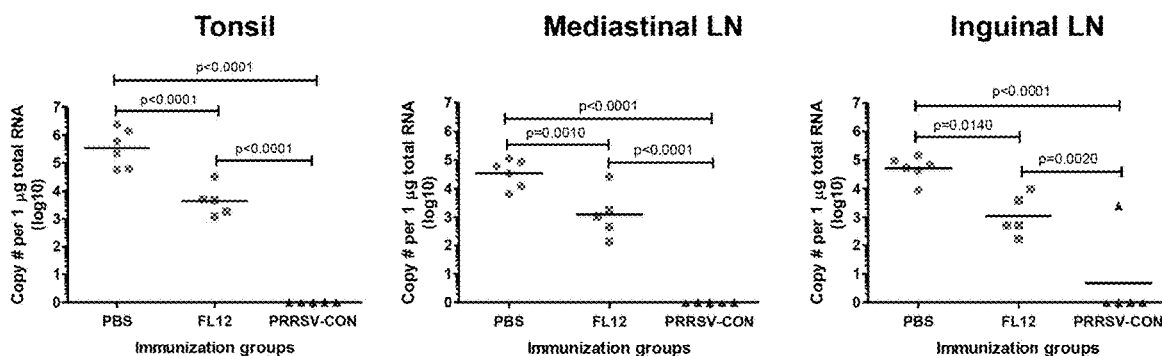


Figure 5

1

NON-NATURALLY OCCURRING PORCINE REPRODUCTIVE AND RESPIRATORY SYNDROME VIRUS (PPRSV) AND METHODS OF USING

CROSS REFERENCE TO RELATED APPLICATIONS

This application is a continuation application of U.S. patent application Ser. No. 15/127,931 filed on Sep. 21, 2016, which is a U.S. National Application from PCT Application No. PCT/IB2015/052214 filed on Mar. 25, 2015, which claims the benefit of priority under 35 U.S.C. § 119(e) to U.S. Application No. 61/968,465, filed Mar. 21, 2014.

FEDERALLY SPONSORED RESEARCH OR DEVELOPMENT

This invention was made with government support under 2013-31100-06031, 2012-31100-06031, and 2008-5562-19132 awarded by United States Department of Agriculture, National Institute of Food and Agriculture. The government has certain rights in the invention.

TECHNICAL FIELD

This disclosure generally relates to a non-naturally occurring porcine reproductive and respiratory syndrome virus (PPRSV) and methods of using.

BACKGROUND

Current porcine reproductive and respiratory syndrome virus (PPRSV) vaccines are not adequately effective for control and eradication of porcine reproductive and respiratory syndrome (PPRS). The main limitation of the current PPRSV vaccines is their sub-optimal coverage against divergent PPRSV strains. Thus far, all commercial PPRSV vaccines are formulated using natural PPRSV strains, but the substantial genetic variation among the PPRSV strains is the biggest obstacle for the development of a broadly protective PPRSV vaccine.

SUMMARY

This disclosure provides a non-naturally occurring porcine reproductive and respiratory syndrome virus (PPRSV) and methods of making and using the non-naturally occurring PPRSV.

A PPRSV-CON nucleic acid is provided, where the nucleic acid has at least 50% sequence identity (e.g., at least 75%, at least 95%, or at least 99% sequence identity) to SEQ ID NO:1. In some embodiment, the nucleic acid has the sequence shown in SEQ ID NO:1. A virus particle comprising the PPRSV-CON nucleic acid described herein. A composition comprising the PPRSV-CON nucleic acid described herein and a pharmaceutically acceptable carrier. A composition comprising the virus particle described herein and a pharmaceutically acceptable carrier. The compositions described herein, further comprising an adjuvant.

A PPRSV-CON nucleic acid also is provided, where the nucleic acid has at least 95% (e.g., at least 99%) sequence identity to a sequence selected from the group consisting of SEQ ID NO:2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, and 42. In some embodiments, the nucleic acid has a sequence selected from the group con-

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sisting of SEQ ID NO:2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, and 42. In some embodiments, the nucleic acid encodes, respectively, a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41 and 43. A virus particle comprising the PPRSV-CON nucleic acid described herein. A composition comprising the nucleic acid described herein and a pharmaceutically acceptable carrier. A composition comprising the virus particle described herein and a pharmaceutically acceptable carrier. The composition described herein, further comprising an adjuvant.

A PPRSV-CON polypeptide is provided, where the polypeptide has at least 95% (e.g., at least 99%) sequence identity to a sequence selected from the group consisting of SEQ ID NO:3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41 and 43. In some embodiments, the polypeptide has a sequence selected from the group consisting of SEQ ID NO:3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41 and 43. In some embodiments, the polypeptide is encoded by a nucleic acid, respectively, having a sequence selected from the group consisting of SEQ ID NOs:2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, or 42. A virus particle comprising the PPRSV-CON polypeptide described herein. A composition comprising the polypeptide described herein and a pharmaceutically acceptable carrier. A composition comprising the virus particle described herein and a pharmaceutically acceptable carrier. The composition described herein, further comprising an adjuvant.

A method for eliciting an immune response to PPRSV in a porcine is provided. Such a method typically includes administering, to a porcine: (i) an effective amount of any of the nucleic acids described herein; (ii) an effective amount of any of the polypeptides described herein; (iii) an effective amount of any of the virus particles described herein; or (iv) an effective amount of any of the compositions described herein. Representative routes of administration include, without limitation, intramuscularly, intraperitoneally, and orally.

A method for treating or preventing PPRS in a porcine is provided. Such a method typically includes administering, to a porcine: (i) an effective amount of any of the nucleic acids described herein; (ii) an effective amount of any of the polypeptides described herein; (iii) an effective amount of any of the virus particles described herein; or (iv) an effective amount of any of the compositions described herein. Representative routes of administration include, without limitation, intramuscularly, intraperitoneally, and orally.

Unless otherwise defined, all technical and scientific terms used herein have the same meaning as commonly understood by one of ordinary skill in the art to which the methods and compositions of matter belong. Although methods and materials similar or equivalent to those described herein can be used in the practice or testing of the methods and compositions of matter, suitable methods and materials are described below. In addition, the materials, methods, and examples are illustrative only and not intended to be limiting. All publications, patent applications, patents, and other references mentioned herein are incorporated by reference in their entirety.

DESCRIPTION OF DRAWINGS

FIG. 1, Panel (A) is a phylogenetic tree constructed from a set of 60 PPRSV full-genome sequences. These 60

PRRSV genomes are classified into 4 sub-groups. The locations of the viruses involved in the cross-protection experiments are indicated by the arrows. FIG. 1, Panel (B) is a graph showing the genetic distances among natural PRRSV strains and the genetic distance from the PRRSV-CON described herein to the natural PRRSV strains. The lower and upper boundaries of the box indicate the 25th and 75th percentile respectively. The solid line within the box represents the median. Whiskers above and below the box indicate the minimum and maximum of the data.

FIG. 2 shows the generation and characterization of the PRRSV-CON virus. Panel (A) is a schematic showing the strategy to construct the PRRSV-CON full-genome cDNA clone. The upper half of Panel (A) depicts the schematic representation of the viral genome, together with the unique restriction enzyme sites used for cloning purposes. The horizontal black lines, with the letters A-D on top, represent the DNA fragments that were synthesized. The numbers inside the parenthesis below the lines indicate the length (in nucleotides) of each corresponding fragments. Φ T7 represents the T7 RNA polymerase promoter. Individual DNA fragments of the genome were sequentially inserted into the shuttle vector (shown in the lower half of Panel (A)) in the order of fragment A to fragment D. Panel (B) are photographs showing the reactivity of the indicated viruses with different PRRSV-specific monoclonal antibodies. MARC-145 cells were mock infected or infected with PRRSV-CON or PRRSV wild type strain, FL12. At 48 hours post-infection, the cells were stained with antibodies specific to the viral nucleocapsid protein (N protein; bottom row of photographs) or to the viral nonstructural protein 1 beta (nsp1b; top row of photographs). Panel (C) shows the plaque morphology of the viruses in MARC-145 cells. Panel (D) shows a multiple step growth curve. MARC-145 cells were infected with the indicated viruses at a multiplicity of infection (MOI) of 0.01. At different timepoints post-infection (p.i.), culture supernatant was collected and viral titer was determined by titration on MARC-145 cells.

FIG. 3 is data demonstrating replication of the PRRSV-CON in pigs. Panel (A) shows the rectal temperature measured daily from 1 day before infection to 13 days post-infection (days p.i.). Panel (B) shows the average daily weight gain (ADWG) within 14 days after inoculation. Panel (C) shows the viremia levels, determined by a commercial, universal RT-qPCR (Tetracore Inc., Rockville, Md.). Panel (D) shows the levels of antibody response after inoculation, determined by IDEXX ELISA; the horizontal dotted line indicates the cut-off of the assay.

FIG. 4 is data demonstrating cross-protection provided by the PRRSV-CON described herein against the PRRSV-strain, MN-184. Panel (A) shows the average daily weight gain (ADWG) within 15 days after challenge-infection. Panel (B) shows the viremia levels after challenge determined by a commercial, universal RT-qPCR (Tetracore Inc., Rockville, Md.). Panel (C) shows total viral RNA levels in different tissues collected at 15 days post-challenge as determined by a commercial, universal RT-qPCR (Tetracore Inc., Rockville, Md.). Panel (D) shows the MN-184-specific RNA levels as determined by a differential RT-qPCR developed in-house.

FIG. 5 is data demonstrating cross-protection against PRRSV strain, 16244B. Panel (A) shows the average daily weight gain (ADWG) within 15 days after challenge-infection. Panel (B) shows the viremia levels after challenge infection determined by a commercial, universal RT-qPCR (Tetracore Inc., Rockville, Md.). Panel (C) shows total viral RNA levels in different tissues collected at 15 days post-

challenge as determined by a commercial, universal RT-qPCR (Tetracore Inc., Rockville, Md.). Panel (D) shows the 16244B-specific RNA levels as determined by a differential RT-qPCR developed in-house.

DETAILED DESCRIPTION

A non-naturally occurring porcine reproductive and respiratory syndrome virus (PRRSV) genome was designed using a large set of genomic sequences of PRRSV isolates, which represents the widest genetic diversity of PRRSV strains circulating in U.S. swine herds. The non-naturally occurring PRRSV genome was designed so that it has a high degree of genetic similarity to the PRRSV field-isolates studied when compared to any single, naturally occurring PRRSV strain.

Porcine reproductive and respiratory syndrome (PRRS) is one of the most economically important diseases in swine. Clinical signs of the disease include reproductive failure in pregnant sows and respiratory disorder in young pigs. The disease is more severe when animals are co-infected with other pathogens. The annual loss to the US swine industry was estimated to be about \$560 million in 2005 and about \$640 million in 2011.

The causative agent of PRRS is an RNA virus named PRRS virus (PRRSV). PRRSV is classified into two major genotypes: European (Type 1) and North American (Type 2). There is limited cross-protection between these two genotypes. Considerable genetic variation exists among PRRSV isolates within each of these genotypes. Importantly, genetic divergence has been shown to occur when a PRRSV strain is serially passed from pig to pig. This leads to co-circulation of multiple PRRSV variants within one herd or even within one animal that is persistently infected with PRRSV.

PRRSV vaccines have been in use since 1994. There are two types of PRRSV vaccines currently available in the market; modified-live and inactivated vaccines. In addition, several subunit vaccines against PRRSV are being tested in different laboratories worldwide, but none have been licensed for clinical application. Currently, PRRSV vaccines are prepared using naturally occurring PRRSV strains as the vaccine immunogens. The current PRRSV vaccines are not adequately effective for control and eradication of PRRS; they provide acceptable levels of homologous protection but they fail to provide consistent heterologous cross-protection. Extensive genetic diversity among PRRSV isolates is the main reason behind the sub-optimal heterologous protection of the current PRRSV vaccines.

The non-naturally occurring PRRSV-CON described herein confers superior cross-protective against different heterologous PRRSV strains, as compared to the PRRSV wild type strain FL12. Thus, the PRRSV-CON described herein can be used to formulate a universal PRRSV vaccine. In addition, the PRRSV-CON described herein provides an important tool to study the mechanism of heterologous protection against divergent PRRSV strains.

Nucleic Acids and Polypeptides

The PRRSV genome encodes at least 22 proteins; 14 non-structural proteins and 8 structural proteins. A nucleic acid is provided herein that encodes for a non-naturally occurring PRRSV. See SEQ ID NO:1 for the genomic sequence of PRRSV-CON. The non-naturally occurring PRRSV described herein possesses the highest degree of genetic identity with the naturally occurring PRRSV isolates. The PRRSV-CON genomic nucleic acid provided herein (i.e., SEQ ID NO:1) encodes for a number of different polypeptides. For example, the nucleic acid sequence shown in SEQ ID NO:2 encodes for the polypeptide sequence

having the amino acid sequence shown in SEQ ID NO:3; the nucleic acid sequence shown in SEQ ID NO:4 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:5; the nucleic acid sequence shown in SEQ ID NO:6 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:7; the nucleic acid sequence shown in SEQ ID NO:8 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:9; the nucleic acid sequence shown in SEQ ID NO:10 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:11; the nucleic acid sequence shown in SEQ ID NO:12 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:13; the nucleic acid sequence shown in SEQ ID NO:14 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:15; the nucleic acid sequence shown in SEQ ID NO:16 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:17; the nucleic acid sequence shown in SEQ ID NO:18 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:19; the nucleic acid sequence shown in SEQ ID NO:20 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:21; the nucleic acid sequence shown in SEQ ID NO:22 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:23; the nucleic acid sequence shown in SEQ ID NO:24 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:25; the nucleic acid sequence shown in SEQ ID NO:26 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:27; the nucleic acid sequence shown in SEQ ID NO:28 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:29; the nucleic acid sequence shown in SEQ ID NO:30 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:31; the nucleic acid sequence shown in SEQ ID NO:32 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:33; the nucleic acid sequence shown in SEQ ID NO:34 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:35; the nucleic acid sequence shown in SEQ ID NO:36 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:37; the nucleic acid sequence shown in SEQ ID NO:38 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:39; the nucleic acid sequence shown in SEQ ID NO:40 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:41; and the nucleic acid sequence shown in SEQ ID NO:42 encodes for the polypeptide sequence having the amino acid sequence shown in SEQ ID NO:43.

As used herein, nucleic acids can include DNA and RNA, and includes nucleic acids that contain one or more nucleotide analogs or backbone modifications. A nucleic acid can be single stranded or double stranded, which usually depends upon its intended use. Nucleic acids and polypeptides that differ from SEQ ID NOs:1-43 also are provided. Nucleic acids that differ in sequence from SEQ ID NO:1 or any of SEQ ID NOs:2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, or 42 can have at least 80% sequence identity (e.g., at least 81%, 82%, 83%, 84%, 85%, 86%, 87%, 88%, 89%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, or 99% sequence identity) to SEQ ID NO:1 or any of SEQ ID NOs:2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, or 42. Polypeptides that

differ in sequence from any of SEQ ID NOs:3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41 or 43, can have at least 80% sequence identity (e.g., at least 81%, 82%, 83%, 84%, 85%, 86%, 87%, 88%, 89%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, or 99% sequence identity) to any of SEQ ID NOs:3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41 or 43.

In calculating percent sequence identity, two sequences are aligned and the number of identical matches of nucleotides or amino acid residues between the two sequences is determined. The number of identical matches is divided by the length of the aligned region (i.e., the number of aligned nucleotides or amino acid residues) and multiplied by 100 to arrive at a percent sequence identity value. It will be appreciated that the length of the aligned region can be a portion of one or both sequences up to the full-length size of the shortest sequence. It also will be appreciated that a single sequence can align with more than one other sequence and hence, can have different percent sequence identity values over each aligned region.

The alignment of two or more sequences to determine percent sequence identity can be performed using the computer program ClustalW and default parameters, which allows alignments of nucleic acid or polypeptide sequences to be carried out across their entire length (global alignment). Chenna et al., 2003, *Nucleic Acids Res.*, 31(13): 3497-500. ClustalW calculates the best match between a query and one or more subject sequences, and aligns them so that identities, similarities and differences can be determined. Gaps of one or more residues can be inserted into a query sequence, a subject sequence, or both, to maximize sequence alignments. For fast pairwise alignment of nucleic acid sequences, the default parameters can be used (i.e., word size: 2; window size: 4; scoring method: percentage; number of top diagonals: 4; and gap penalty: 5); for an alignment of multiple nucleic acid sequences, the following parameters can be used: gap opening penalty: 10.0; gap extension penalty: 5.0; and weight transitions: yes. For fast pairwise alignment of polypeptide sequences, the following parameters can be used: word size: 1; window size: 5; scoring method: percentage; number of top diagonals: 5; and gap penalty: 3. For multiple alignment of polypeptide sequences, the following parameters can be used: weight matrix: blosum; gap opening penalty: 10.0; gap extension penalty: 0.05; hydrophilic gaps: on; hydrophilic residues: Gly, Pro, Ser, Asn, Asp, Gln, Glu, Arg, and Lys; and residue-specific gap penalties: on. ClustalW can be run, for example, at the Baylor College of Medicine Search Launcher website or at the European Bioinformatics Institute website on the World Wide Web.

Changes can be introduced into a nucleic acid molecule (e.g., SEQ ID NO:1 or any of SEQ ID NOs:2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, or 42), thereby leading to changes in the amino acid sequence of the encoded polypeptide (e.g., SEQ ID NOs:3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41 or 43). For example, changes can be introduced into nucleic acid coding sequences using mutagenesis (e.g., site-directed mutagenesis, PCR-mediated mutagenesis) or by chemically synthesizing a nucleic acid molecule having such changes. Such nucleic acid changes can lead to conservative and/or non-conservative amino acid substitutions at one or more amino acid residues. A "conservative amino acid substitution" is one in which one amino acid residue is replaced with a different amino acid residue having a similar side chain (see, for example, Dayhoff et al. (1978, in *Atlas of Protein Sequence and Structure*, 5(Suppl. 3):345-352), which pro-

vides frequency tables for amino acid substitutions), and a non-conservative substitution is one in which an amino acid residue is replaced with an amino acid residue that does not have a similar side chain.

As used herein, an "isolated" nucleic acid molecule is a nucleic acid molecule that is free of sequences that naturally flank one or both ends of the nucleic acid in the genome of the organism from which the isolated nucleic acid molecule is derived (e.g., a cDNA or genomic DNA fragment produced by PCR or restriction endonuclease digestion). Such an isolated nucleic acid molecule is generally introduced into a vector (e.g., a cloning vector, or an expression vector) for convenience of manipulation or to generate a fusion nucleic acid molecule, discussed in more detail below. In addition, an isolated nucleic acid molecule can include an engineered nucleic acid molecule such as a recombinant or a synthetic nucleic acid molecule.

As used herein, a "purified" polypeptide is a polypeptide that has been separated or purified from cellular components that naturally accompany it. Typically, the polypeptide is considered "purified" when it is at least 70% (e.g., at least 75%, 80%, 85%, 90%, 95%, or 99%) by dry weight, free from the polypeptides and naturally occurring molecules with which it is naturally associated. Since a polypeptide that is chemically synthesized is, by nature, separated from the components that naturally accompany it, a synthetic polypeptide is "purified."

Nucleic acids can be isolated using techniques routine in the art. For example, nucleic acids can be isolated using any method including, without limitation, recombinant nucleic acid technology, and/or the polymerase chain reaction (PCR). General PCR techniques are described, for example in PCR Primer: A Laboratory Manual, Dieffenbach & Dveksler, Eds., Cold Spring Harbor Laboratory Press, 1995. Recombinant nucleic acid techniques include, for example, restriction enzyme digestion and ligation, which can be used to isolate a nucleic acid. Isolated nucleic acids also can be chemically synthesized, either as a single nucleic acid molecule or as a series of oligonucleotides.

Polypeptides can be purified from natural sources (e.g., a biological sample) by known methods such as DEAE ion exchange, gel filtration, and hydroxyapatite chromatography. A polypeptide also can be purified, for example, by expressing a nucleic acid in an expression vector. In addition, a purified polypeptide can be obtained by chemical synthesis. The extent of purity of a polypeptide can be measured using any appropriate method, e.g., column chromatography, polyacrylamide gel electrophoresis, or HPLC analysis.

A vector containing a nucleic acid (e.g., a nucleic acid that encodes a polypeptide) also is provided. Vectors, including expression vectors, are commercially available or can be produced by recombinant DNA techniques routine in the art. A vector containing a nucleic acid can have expression elements operably linked to such a nucleic acid, and further can include sequences such as those encoding a selectable marker (e.g., an antibiotic resistance gene). A vector containing a nucleic acid can encode a chimeric or fusion polypeptide (i.e., a polypeptide operatively linked to a heterologous polypeptide, which can be at either the N-terminus or C-terminus of the polypeptide). Representative heterologous polypeptides are those that can be used in purification of the encoded polypeptide (e.g., 6xHis tag, glutathione S-transferase (GST)). Expression elements include nucleic acid sequences that direct and regulate expression of nucleic acid coding sequences. One example of an expression element is a promoter sequence. Expression

elements also can include introns, enhancer sequences, response elements, or inducible elements that modulate expression of a nucleic acid. Expression elements can be of bacterial, yeast, insect, mammalian, or viral origin, and vectors can contain a combination of elements from different origins. As used herein, operably linked means that a promoter or other expression element(s) are positioned in a vector relative to a nucleic acid in such a way as to direct or regulate expression of the nucleic acid (e.g., in-frame). Many methods for introducing nucleic acids into host cells, both in vivo and in vitro, are well known to those skilled in the art and include, without limitation, electroporation, calcium phosphate precipitation, polyethylene glycol (PEG) transformation, heat shock, lipofection, microinjection, and viral-mediated nucleic acid transfer.

Vectors as described herein can be introduced into a host cell. As used herein, "host cell" refers to the particular cell into which the nucleic acid is introduced and also includes the progeny of such a cell that carry the vector. A host cell can be any prokaryotic or eukaryotic cell. For example, nucleic acids can be expressed in bacterial cells such as *E. coli*, or in insect cells, yeast or mammalian cells (such as Chinese hamster ovary cells (CHO) or COS cells). Other suitable host cells are known to those skilled in the art.

Nucleic acids can be detected using any number of amplification techniques (see, e.g., PCR Primer: A Laboratory Manual, 1995, Dieffenbach & Dveksler, Eds., Cold Spring Harbor Laboratory Press, Cold Spring Harbor, N.Y.; and U.S. Pat. Nos. 4,683,195; 4,683,202; 4,800,159; and 4,965,188) with an appropriate pair of oligonucleotides (e.g., primers). A number of modifications to the original PCR have been developed and can be used to detect a nucleic acid.

Nucleic acids also can be detected using hybridization. Hybridization between nucleic acids is discussed in detail in Sambrook et al. (1989, Molecular Cloning: A Laboratory Manual, 2nd Ed., Cold Spring Harbor Laboratory Press, Cold Spring Harbor, N.Y.; Sections 7.37-7.57, 9.47-9.57, 11.7-11.8, and 11.45-11.57). Sambrook et al. discloses suitable Southern blot conditions for oligonucleotide probes less than about 100 nucleotides (Sections 11.45-11.46). The T_m between a sequence that is less than 100 nucleotides in length and a second sequence can be calculated using the formula provided in Section 11.46. Sambrook et al. additionally discloses Southern blot conditions for oligonucleotide probes greater than about 100 nucleotides (see Sections 9.47-9.54). The T_m between a sequence greater than 100 nucleotides in length and a second sequence can be calculated using the formula provided in Sections 9.50-9.51 of Sambrook et al.

The conditions under which membranes containing nucleic acids are prehybridized and hybridized, as well as the conditions under which membranes containing nucleic acids are washed to remove excess and non-specifically bound probe, can play a significant role in the stringency of the hybridization. Such hybridizations and washes can be performed, where appropriate, under moderate or high stringency conditions. For example, washing conditions can be made more stringent by decreasing the salt concentration in the wash solutions and/or by increasing the temperature at which the washes are performed. Simply by way of example, high stringency conditions typically include a wash of the membranes in 0.2xSSC at 65° C.

In addition, interpreting the amount of hybridization can be affected, for example, by the specific activity of the labeled oligonucleotide probe, by the number of probe-binding sites on the template nucleic acid to which the probe

has hybridized, and by the amount of exposure of an autoradiograph or other detection medium. It will be readily appreciated by those of ordinary skill in the art that although any number of hybridization and washing conditions can be used to examine hybridization of a probe nucleic acid molecule to immobilized target nucleic acids, it is more important to examine hybridization of a probe to target nucleic acids under identical hybridization, washing, and exposure conditions. Preferably, the target nucleic acids are on the same membrane.

A nucleic acid molecule is deemed to hybridize to a nucleic acid but not to another nucleic acid if hybridization to a nucleic acid is at least 5-fold (e.g., at least 6-fold, 7-fold, 8-fold, 9-fold, 10-fold, 20-fold, 50-fold, or 100-fold) greater than hybridization to another nucleic acid. The amount of hybridization can be quantitated directly on a membrane or from an autoradiograph using, for example, a PhosphorImager or a Densitometer (Molecular Dynamics, Sunnyvale, Calif.).

Polypeptides can be detected using antibodies. Techniques for detecting polypeptides using antibodies include enzyme linked immunosorbent assays (ELISAs), Western blots, immunoprecipitations and immunofluorescence. An antibody can be polyclonal or monoclonal. An antibody having specific binding affinity for a polypeptide can be generated using methods well known in the art. The antibody can be attached to a solid support such as a microtiter plate using methods known in the art. In the presence of a polypeptide, an antibody-polypeptide complex is formed.

Detection (e.g., of an amplification product, a hybridization complex, or a polypeptide) is usually accomplished using detectable labels. The term "label" is intended to encompass the use of direct labels as well as indirect labels. Detectable labels include enzymes, prosthetic groups, fluorescent materials, luminescent materials, bioluminescent materials, and radioactive materials.

Methods of Making and Using a PRRSV-CON Virus Particle

Methods of constructing a virus particle from a PRRSV-CON nucleic acid are known in the art and are described herein. As demonstrated herein, the PRRSV-CON described herein self-assembles into particles when appropriately expressed. The PRRSV-CON can be expressed *in vitro* or *in vivo*, for example, in a host cell. In some embodiments, a host cell can be transfected with the PRRSV-CON nucleic acid, or a host cell can be infected with a PRRSV-CON virus particle. Host cells can be, without limitation, porcine cells (e.g., porcine alveolar macrophage) or African green monkey kidney-derived cells (e.g., MARC-145). Virus particles can be isolated, for example, by ultracentrifugation.

The PRRSV-CON nucleic acids, polypeptides or virus particles described herein can be used to generate, enhance or modulate the immune response of a porcine. Such methods typically include administering a PRRSV-CON nucleic acid, polypeptide or virus particle described herein to a porcine in an amount sufficient to generate an immune response. As used herein, an "immune response" refers to the reaction elicited in an individual following administration of a PRRSV-CON nucleic acid, polypeptide or virus particle as described herein. Immune responses can include, for example, an antibody response or a cellular response (e.g., a cytotoxic T-cell response). A PRRSV-CON nucleic acid, polypeptide or virus particle can be used to prevent PRRS in porcine, e.g., as a prophylactic vaccine, or to establish or enhance immunity to PRRS in a healthy indi-

vidual prior to exposure or contraction of PRRS, thus preventing the disease or reducing the severity of disease symptoms.

Methods for administering a PRRSV-CON nucleic acid, polypeptide or virus particle to a porcine include, without limitation, intramuscular (i.m.), subcutaneous (s.c.), or intrapulmonary routes. Methods for administering a PRRSV-CON nucleic acid, polypeptide or virus particle to a porcine also include, without limitation, intratracheal, transdermal, intraocular, intranasal, inhalation, intracavity, and intravenous (i.v.) administration.

Determining an effective amount of a PRRSV-CON nucleic acid, polypeptide or virus particle depends upon a number of factors including, for example, whether the antigen is being expressed or administered directly, the age and weight of the subject, the precise condition requiring treatment and its severity, and the route of administration. Based on the above factors, determining the amount and the dosing (e.g., the number of doses and the timing of doses) are within the level of skill of an ordinary artisan.

A composition can include a PRRSV-CON nucleic acid, polypeptide or virus particle as described herein and a pharmaceutically acceptable carrier. Pharmaceutically acceptable carriers are known in the art and include, for example, buffers (e.g., phosphate buffered saline (PBS), normal saline, Tris buffer, and sodium phosphate) or diluents. The compositions described herein can be formulated as an aqueous solution, or as an emulsion, gel, solution, suspension, or powder. See, for example, Remington's Pharmaceutical Sciences, 16th Ed., Osol, ed., Mack Publishing Co., Easton, Pa. (1980), and Remington's Pharmaceutical Sciences, 19th Ed., Gennaro, ed., Mack Publishing Co., Easton, Pa. (1995). In addition to a pharmaceutically acceptable carrier, the compositions described herein also can include binders, stabilizers, preservatives, salts, excipients, delivery vehicles and/or auxiliary agents.

In accordance with the present invention, there may be employed conventional molecular biology, microbiology, biochemical, and recombinant DNA techniques within the skill of the art. Such techniques are explained fully in the literature. The invention will be further described in the following examples, which do not limit the scope of the methods and compositions of matter described in the claims.

EXAMPLES

Example 1—Computational Design of the Artificial PRRSV-CON Genome

Full-genome sequences of 64 PRRSV isolates originating from the Midwestern states (Iowa, Nebraska and Illinois) of the U.S. were sequenced using the Roche 454-GS-FLX sequencing technology. In addition, more than 20 full-genome sequences of PRRSV isolates originating from the U.S. were collected from GenBank. After removing redundant sequences, a final set of 60 full-genome sequences of PRRSV was attained. The 60 PRRSV full-genome sequences were aligned using the MUSCLE program (Edgar RC, 2004, BMC Bioinform., 5:113). After that, a consensus genome sequence (PRRSV-CON) was generated by selecting the most common nucleotide found at each position of the viral genome, using the Jalview program. Phylogenetic analysis shows that the PRRSV-CON genome locates right at the center of the phylogenetic tree. See FIG. 1A. Consequently, the pairwise genetic distance from PRRSV-CON to the naturally occurring PRRSV strains is significantly

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shorter than the distance from any one naturally occurring PRRSV strains to each other ($p < 0.0001$). See FIG. 1B.

Example 2—Generation of an Infectious PRRSV-CON Virus

It is generally difficult to accurately determine the sequence at 5' and 3' ends of a viral genome. Thus, we realized that the sequences at the 5' and 3' untranslated regions (UTRs) of the naturally occurring PRRSV genomes analyzed in Example 1 may not be accurate. To increase the change of recovering infectious virus, we replaced the 5' and 3'UTRs of the PRRSV-CON genome with the 5' and 3' UTRs of the infectious cDNA clone FL12 (Truong et al., 2004, Virology, 325:308-19). Four DNA fragments, designated A-D, encompassing the entire PRRSV-CON genome, were chemically synthesized by Genscript (Piscataway, N.J.). Each DNA fragment was flanked by a pair of restriction enzyme sites to facilitate the cloning purposes. The T7 RNA polymerase promoter sequence was incorporated into fragment D, preceding the viral 5'end, to facilitate the in vitro transcription of the viral genome. See FIG. 2A. Individual DNA fragments were sequentially cloned into the shuttle vector that carries the corresponding restriction enzyme site, following the order from fragment A to fragment D. Once the full-length PRRSV-CON cDNA clone was generated, standard reverse genetics techniques were applied to recover viable PRRSV-CON viruses.

Briefly, the plasmid containing full-length cDNA genome of PRRSV-CON was digested with AclI for linearization. The purified, linear DNA fragment was used as the template for an in vitro transcription reaction using the mMES-SAGEmMACHINE Ultra T7 kit (Ambion, Austin, Tex.) to generate full genome viral RNA transcripts. After that, about 5 μ g of the full-genome RNA transcripts were transfected into MARC-145 cells cultured in a 6-well plate, using the TransIT[®]-mRNA Transfection Kit (Mirus Bio, Madison, Wis.). Transfected cells were cultured in DMEM containing 10% FBS at 37° C., 5% CO₂ for up to 6 days. Typically, cytopathic effect (CPE) was observed between day 4 and day 6 after transfection. When clear CPE was observed, culture supernatant containing the rescued virus was collected and stored in 0.5 mL aliquots in a 80° C. freezer. See, Truong et al. (2004, supra)

Example 3—In Vitro Characterization of the PRRSV-CON Virus

To study the reactivity with different PRRSV-specific monoclonal antibodies, MARC-145 cells were mock infected or infected with the PRRSV-CON virus or the PRRSV strain FL12. At 48 hours post-infection (p.i.), the cells were immunostained with antibodies specific to the viral nucleocapsid (N) protein or the viral nonstructural protein 1 beta (nsp1b). To study the growth kinetics of the viruses in cell culture, MARC-145 cells were infected with the PRRSV-CON or FL12 at a multiplicity of infection (MOI) of 0.01. At different time-points p.i., culture supernatant was collected and viral titers were determined by titration in MARC-145 cells.

The PRRSV-CON virus displays typical in vitro characterizations of a naturally occurring PRRSV strain. It reacts with different PRRSV-specific monoclonal antibodies including antibodies against nsp1-beta and N protein (FIG. 2B). It replicates efficiently in cell culture (FIG. 2C), and it is able to form clear and distinct plaque morphology (FIG. 3D).

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Example 4—the PRRSV-CON Virus can Infect Pigs as Efficiently as the Natural PRRSV Strain

A total of 18 PRRSV-seronegative, 3 week-old pigs were purchased from the University of Nebraska research farm. The pigs were randomly assigned into 3 experimental groups; each group was housed in a separate room in the Biosecurity Level-2 Animal Research Facilities at UNL, following the regulations established by the Institutional Animal Care and Use Committee. Pigs in group 1 were injected with PBS to act as the control. Pigs in groups 2 and 3 were inoculated intramuscularly with $10^{5.0}$ TCID₅₀ of PRRSV-CON and PRRSV strain FL12, respectively. The wild-type PRRSV strain, FL12, was included into this study for comparison purposes. The results are shown in FIG. 3. After infection, both of the PRRSV-CON and FL12-inoculated groups displayed significantly higher temperature than PBS-group (FIG. 3A), but there was no difference in temperature between PRRSV-CON-inoculated group and the FL12-inoculated group. Average daily weight gain (ADWG) was measured for each individual pig during the period of 14 days after infection. No statistical difference was observed among the three treatment groups, although pigs in the PRRSV-CON-inoculated group and the FL12-inoculated group tended to have lower ADWG than the PBS group (FIG. 3B). Viremia levels of the PRRSV-CON- and FL12-inoculated groups were almost identical (FIG. 3C). All pigs in the PRRSV-CON- and FL12-inoculated groups were seroconverted by 11 days p.i. The level of antibody response in the PRRSV-CON-inoculated group was slightly lower than that of the FL12-inoculated group (FIG. 3D). These results demonstrate that the PRRSV-CON can infect the natural host (i.e., pigs) as efficiently as the PRRSV strain, FL12.

Example 5—Evaluation of the Level of Cross-Protection Against PRRSV Strain MN-184

Materials and Methods

A total of 18 PRRSV-seronegative, 3 week-old pigs were purchased from the University of Nebraska research farm. The pigs were randomly assigned into 3 experimental groups; each group was housed in a separate room in the Biosecurity Level-2 Animal Research Facilities at UNL, following the regulations established by the Institutional Animal Care and Use Committee. Group 1 was injected with PBS and served as the non-immunization control. Group 2 was immunized by infection, intramuscularly, with PRRSV-CON at the dose of $10^{4.0}$ TCID₅₀ per pig. Group 3 was immunized by infection, intramuscularly, with the wild-type PRRSV strain, FL12, at the dose of $10^{4.0}$ TCID₅₀ per pig. See Table 1. At 53 days post-infection (p.i.), all control and immunized pigs were challenged, intramuscularly, with PRRSV strain MN-184 at a dose of $10^{5.0}$ TCID₅₀. Parameters used to evaluate protection by immunization with the PRRSV-CON virus included viremia and viral load in several different tissues as well as growth performance.

TABLE 1

Experimental Design to Evaluate Level of Cross-Protection Against PRRSV Strain MN-184

Groups	Immunized with	Challenged with
1 (n = 6)	PBS	MN-184
2 (n = 6)	PRRSV-CON	(Sub-group 2)
3 (n = 6)	PRRSV strain FL12	

To measure growth performance, each pig was weighed right before challenge infection and 15 days post-challenge.

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Body weight was recorded in pounds. Average daily weight gain (ADWG) was calculated for the period of 15 days post-challenge.

To quantitate levels of viremia after challenge infection, blood samples were taken before challenge and at days 1, 4, 7, 10, and 15 post-challenge. Serum samples were extracted from each individual blood samples and stored in a -80°C . freezer. Viremia levels were quantitated by the Animal Disease Research and Diagnostic Laboratory, South Dakota State University, using the universal RT-qPCR kit (Tetracore Inc., Rockville, Md.). Results were reported as log₁₀ copy/mL. For statistical purposes, samples that had undetected level of viral RNA were assigned a value of 0 log₁₀ copy/mL.

To quantitate levels of viral load in tissues, pigs were humanely sacrificed and necropsied on day 15 post-challenge. Samples of tonsil, lung, mediastinal lymph node and inguinal lymph node were obtained and kept individually in Whirl-Pak® bags. The samples were snap-frozen in liquid nitrogen right after collection. After that, they were stored in a -80°C . freezer. To extract RNA, tissue samples were homogenized in Trizol reagent (Life Technologies, Carlsbad, Calif.) with a ratio of 300 mg tissue in 3 mL Trizol reagent. Total RNA was extracted using the RNeasy Mini Kit (Qiagen, Valencia, Calif.) following the manufacturer's instruction. RNA concentration was quantitated by the NanoDrop® ND-1000 (NanoDrop Technologies, Inc., Wilmington, Del.) and adjusted to a final concentration of 200 ng/4.

It has been well characterized that PRRSV can colonize and persist in lymphoid tissues of infected pigs up to 150 days post-infection. In these experiments, the tissue viral load was evaluated at 15 days post-challenge, which corresponds to 67 days after the primary infection. At that time, it is likely that the pigs in the PRRSV-CON and FL12 groups still contained residual virus of the primary infection. Therefore, we used two different RT-PCR kits to quantify the viral RNA load in tissues: (i) the commercial RT-qPCR kit (Tetracore Inc., Rockville, Md.) that detects total viral RNA resulting from both the primary infection and the challenge infection, and (ii) the differential RT-PCR developed in-house that selectively detects only viral RNA from challenge infection. Five μL of each RNA sample (equivalent to 1 μg RNA) was used for each RT-qPCR reaction. Results were reported as log₁₀ copy/ μg of total RNA. For statistical

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purposes, samples that had undetected viral RNA level were assigned a value of 0 log RNA copy/1 μg of total RNA.

Results

The results of growth performance are presented in FIG. 4A. The mean ADWG of PBS-, PRRSV-CON- and FL12-immunized groups were 0.3 lbs (SD+/-0.3), 0.9 lbs (SD+/-0.6), and 1.2 lbs (SD+/-0.4), respectively. PRRSV-CON and FL12-immunized groups had greater ADWG than the PBS-immunized group. There was no statistical difference between the PRRSV-CON- and FL12-immunized groups.

The viremia levels after challenge infection are shown in FIG. 4B and Table 2. All pigs in the PBS-immunized group were viremic at all timepoints tested. The PRRSV-CON-immunized group only had 3 viremic pigs, of which 1 pig was viremic at 2 timepoints (pig #494 at 4 DPC and 7 DPC) and 2 pigs were viremic at only one timepoint (pigs #394 and 495 at 15 DPC). The remaining 3 pigs in this group (pigs #345, 410 and 459) were not viremic after challenge infection. By contrast, viremia was detected in 5 out of 6 pigs in the FL12-immunized group at two time-points or more after challenge infection. There was only 1 pig in this group (pig #440) that was not viremic at any time-point tested. Overall, the viremia level of PRRSV-CON-immunized pigs was significantly lower than that in the FL12-immunized group ($p<0.05$) and the PBS-immunized group ($p<0.0001$).

The results of total viral RNA quantitated by the universal RT-qPCR kit are shown in FIG. 4C. The PRRSV-CON- and FL12-immunized groups contained significantly lower levels of total viral RNA than the PBS-immunized group, regardless of the tissue types tested. However, there was no difference between the PRRSV-CON- and FL12-immunized groups in term of total viral RNA.

The results of MN-184 specific RNA quantitated by the differential RT-qPCR are shown in FIG. 4D. All pigs in PBS-immunized group carried MN-184 RNA in their tissues. Four pigs in the FL12-immunized group had MN-184 RNA in their tonsil and mediastinal lymph node, whereas 5 pigs in this group had MN-184 RNA in their inguinal lymph node. Remarkably, none of the pigs in the PRRSV-CON-immunized group had detectable level of MN-184 RNA in any of the tissue samples tested.

Taken together, these results clearly demonstrate that immunization of weaning pigs by infection with the non-naturally occurring PRRSV-CON resulted in significantly better cross-protection against challenge with PRRSV strain, MN-184, than did immunization with the PRRSV strain, FL12.

TABLE 2

Viremia After Challenge Infection (log ₁₀ copy/mL)							
Treatment	Pig ID	Day post-challenge infection (DPC)					
		0 DPC	1 DPC	4 DPC	7 DPC	10 DPC	15 DPC
Group 1 (Injected ("immunized") with PBS)	365	0.00	4.94	5.43	5.45	6.79	6.32
	389	0.00	6.26	6.08	5.40	7.60	6.93
	407	0.00	4.91	6.00	5.86	7.56	6.75
	416	0.00	6.20	6.04	5.20	7.18	6.78
	417	0.00	5.18	5.59	4.86	5.90	6.45
	435	0.00	5.83	5.08	5.94	5.57	5.36
	Mean	0.00	5.55	5.70	5.45	6.77	6.43
	SD	0.00	0.62	0.40	0.40	0.86	0.57

TABLE 2-continued

Viremia After Challenge Infection (log10 copy/mL)							
Treatment	Pig ID	Day post-challenge infection (DPC)					
		0 DPC	1 DPC	4 DPC	7 DPC	10 DPC	15 DPC
Group 2 (Immunized by infection with PRRSV-CON)	345	0.00	0.00	0.00	0.00	0.00	0.00
	394	0.00	0.00	0.00	0.00	0.00	2.58
	410	0.00	0.00	0.00	0.00	0.00	0.00
	459	0.00	0.00	0.00	0.00	0.00	0.00
	494	0.00	0.00	3.58	5.98	0.00	0.00
	495	0.00	0.00	0.00	0.00	0.00	2.98
	Mean	0.00	0.00	0.60	1.00	0.00	0.93
	SD	0.00	0.00	1.46	2.44	0.00	1.44
Group 3 (Immunized by infection with FL12)	349	0.00	0.00	2.81	2.92	0.00	0.00
	381	0.00	0.00	0.00	3.04	2.86	0.00
	440	0.00	0.00	0.00	0.00	0.00	0.00
	455	0.00	0.00	4.18	4.34	0.00	0.00
	487	0.00	3.59	5.28	2.40	5.60	2.68
	507	0.00	2.32	5.56	3.70	0.00	0.00
	Mean	0.00	0.99	2.97	2.73	1.41	0.45
	SD	0.00	1.58	2.50	1.50	2.35	1.09

Example 6—Evaluation of the Level of Cross-Protection Against PRRSV Strain 16244B

Materials and Methods

The experimental design was the same as described above in Example 5. A total of 18 PRRSV-seronegative, 3 week-old pigs purchased from the UNL research farm were randomly assigned into 3 experimental groups. Each group was housed in a separate room at the Biosecurity Level-2 Animal Research Facilities at UNL, following the regulations established by the Institutional Animal Care and Use Committee. Group 1 was injected with PBS and acted as the control. Group 2 was immunized, intramuscularly, by infection with PRRSV-CON at the dose of $10^{4.0}$ TCID₅₀ per pig. Group 3 was immunized, intramuscularly, by infection with the wild type PRRSV, FL12, at the dose of $10^{4.0}$ TCID₅₀ per pig. See Table 3. One pig in group 3 (pig #543) and one pig in group 2 (pig #435) were removed from this study on 14 and 23 days after primary infection, respectively, due to lameness in their legs. At day 52 post-infection (p.i.), all pigs were challenged, intramuscularly, with PRRSV strain 16244B at the challenge dose of $10^{5.0}$ TCID₅₀. Parameters used to evaluate protection by immunization with the PRRSV-CON virus, including viremia and viral load in various tissues as well as growth performance, were measured as described above in Example 5.

TABLE 3

Experimental Design to Evaluate Level of Cross-Protection Against PRRSV Strain 16244B		
Groups	Immunized with	Challenged with
1 (n = 6)	PBS	16244B
2 (n = 6)	PRRSV-CON	(sub-group 3)
3 (n = 6)	PRRSV strain FL12	

Results

The results of growth performance are shown in FIG. 5A. Mean ADWG of PBS-, PRRSV-CON-, and FL12-immunized groups were 1.1 lbs (SD+/-0.3), 1.6 lbs (SD+/-0.1), and 0.8 lbs (SD+/-0.3), respectively. The PRRSV-CON-immunized group had greater ADWG than the PBS-immu-

nized group and the FL12-immunized group; whereas the FL12-immunized group was not statistically different from the PBS-immunized group.

The results of viremia levels after challenge infection are shown in FIG. 5B and Table 4. All pigs in the PBS-immunized group were viremic at all timepoints tested. Two out of 5 pigs in the PRRSV-CON-immunized group (pigs #442 and 445) did not resolve viremia at 52 days after primary infection as viral RNA was still detected in their serum samples collected at this timepoint. After challenge infection, 3 pigs in the PRRSV-CON-immunized group were viremic at only 1 timepoint. The remaining 2 pigs in this group (pigs #436 and 438) were not viremic throughout the period of 15 days post-challenge. By contrast, all pigs in the FL12-immunized group resolved viremia by 52 days post-primary infection. After challenge infection, all pigs in this group became viremic. Overall, the viremia level of the PRRSV-CON-immunized group was significantly lower than that of the FL12-immunized group ($p < 0.0001$) or the PBS-immunized group ($p < 0.0001$).

The results of total viral RNA quantitated by the commercial RT-qPCR kit (Tetracore Inc., Rockville, Md.) are shown in FIG. 5C. Both the PRRSV-CON- and FL12-immunized groups contained significantly lower levels of total viral RNA than the PBS-immunized group, regardless of the tissue types tested. However, there was no statistical difference between the PRRSV-CON-immunized group and the FL12-immunized group in terms of total viral RNA.

The results of 16244B-specific RNA quantitated by the differential RT-qPCR are shown in FIG. 5D. All pigs in the PBS- and FL12-immunized groups carried 16244B-specific RNA in their tissues, although the levels of 16244B RNA in the FL12-immunized group was lower than those in the PBS-immunized group. By contrast, only 1 pig in the PRRSV-CON-immunized group carried 16244B-specific RNA in its inguinal lymph node, while the remaining 4 pigs in this group did not carry 16244B-specific RNA.

All together, these results clearly demonstrate that immunization of weaning pigs by infection with the non-naturally occurring PRRSV-CON resulted in significantly better cross-protection against challenge with PRRSV strain, 16244B, than did immunization with the PRRSV strain, FL12.

TABLE 4

Level of Viremia After Challenge Infection (log10 copy/mL)							
Treatment	Pig ID	Day post-challenge					
		0 DPC	1 DPC	4 DPC	7 DPC	11 DPC	14 DPC
Group 1 (Injected with PBS)	440	0.00	6.62	6.99	6.79	6.15	4.67
	441	0.00	6.61	6.93	7.11	5.79	4.81
	544	0.00	6.85	6.82	6.96	3.91	5.68
	545	0.00	7.11	7.41	7.11	6.81	5.93
	546	0.00	6.74	7.45	7.30	5.67	5.40
	547	0.00	6.77	7.51	7.36	6.73	5.52
	Mean	0.00	6.78	7.18	7.11	5.84	5.34
	SD	0.00	0.18	0.30	0.21	1.06	0.50
Group 2 (immunized by infection with PRRSV- CON)	435	Removed from experiment on day 23rd after primary infection					
	436	0.00	0.00	0.00	0.00	0.00	0.00
	437	0.00	2.48	0.00	0.00	0.00	0.00
	438	0.00	0.00	0.00	0.00	0.00	0.00
	442	2.81	0.00	0.00	0.00	0.00	2.93
	445	3.00	3.32	0.00	0.00	0.00	0.00
	Mean	1.16	1.16	0.00	0.00	0.00	0.59
	SD	1.59	1.62	0.00	0.00	0.00	1.31
Group 3 (immunized by infection with FL12)	439	0.00	4.34	6.78	3.54	2.48	0.00
	444	0.00	3.04	6.58	0.00	0.00	0.00
	446	0.00	5.26	4.84	0.00	0.00	0.00
	526	0.00	2.98	4.40	4.15	0.00	0.00
	540	0.00	3.90	4.18	5.08	3.95	0.00
	543	Removed from experiment on day 14th after primary infection					
	Mean	0.00	3.90	5.35	2.55	1.29	0.00
	SD	0.00	0.95	1.23	2.39	1.84	0.00

It is to be understood that, while the methods and compositions of matter have been described herein in conjunction with a number of different aspects, the foregoing description of the various aspects is intended to illustrate and not limit the scope of the methods and compositions of matter. Other aspects, advantages, and modifications are within the scope of the following claims.

Disclosed are methods and compositions that can be used for, can be used in conjunction with, can be used in preparation for, or are products of the disclosed methods and compositions. These and other materials are disclosed herein, and it is understood that combinations, subsets, interactions, groups, etc. of these methods and compositions

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are disclosed. That is, while specific reference to each various individual and collective combinations and permutations of these compositions and methods may not be explicitly disclosed, each is specifically contemplated and described herein. For example, if a particular composition of matter or a particular method is disclosed and discussed and a number of compositions or methods are discussed, each and every combination and permutation of the compositions and the methods are specifically contemplated unless specifically indicated to the contrary. Likewise, any subset or combination of these is also specifically contemplated and disclosed.

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SEQUENCE LISTING

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attctggtccc ctgcccacca cgttgaaagt gccgcaggct ttcattccgat tgcggcaaat 14760
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gggcgaccgt gtgggggtta agtttaattg gcgagaacca tgcggccgaa attaaaaaaa 15420
aaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaaa 15456

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<210> SEQ ID NO 2
<211> LENGTH: 540
<212> TYPE: DNA
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus
<400> SEQUENCE: 2

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gttcttgagc tcgggggtgt gggcctatct tacaggcccg aagagccact ccggtggacg 180
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atctttccaa ttgcacgaat gaccagtggg aacctgaact ttcaacaaag aatgggtcgg 300
gtcgcagctg agctttacag agccggccag ctcaccctg cagtottgaa ggctctacaa 360
gtttatgaac ggggttgccg ctgggtacccc attgttgac ctgtccctgg agtgccggtt 420
ttgcccaact cctcatatgt gagtgataaa cctttcccg gagcaactca tgtgttaacc 480
aacctgccgc tcccgagag gcccaagcct gaagactttt gccctttga gtgtgctatg 540

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<210> SEQ ID NO 3
<211> LENGTH: 180
<212> TYPE: PRT
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus
<400> SEQUENCE: 3

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

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Arg Met Val Arg Val Ala Ala Glu Leu Tyr Arg Ala Gly Gln Leu Thr
100 105 110

Pro Ala Val Leu Lys Ala Leu Gln Val Tyr Glu Arg Gly Cys Arg Trp
115 120 125

Tyr Pro Ile Val Gly Pro Val Pro Gly Val Ala Val Phe Ala Asn Ser
130 135 140

Leu His Val Ser Asp Lys Pro Phe Pro Gly Ala Thr His Val Leu Thr
145 150 155 160

Asn Leu Pro Leu Pro Gln Arg Pro Lys Pro Glu Asp Phe Cys Pro Phe
165 170 175

Glu Cys Ala Met
180

<210> SEQ ID NO 4
<211> LENGTH: 609
<212> TYPE: DNA
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 4

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tggtgccccctc gtggcgggga tgaagggaat tttgaaactg tccccgagga gttgaagttg      120
attgcgaacc gactccacat ctcttcccg ccccaaccacg cagtggacat gtctaagttt      180
gccttcatag cccctgggag tgggtgtttcc atgcggggtcg agtgccaaca cggtgcctc      240
cccgtgaca ctgtccctga aggcaactgc tgggtggcgt tgtttgactt gctccactg      300
gaagttcaga acaaagaaat tcgccatgct aaccaatttg gctatcagac caagcatggt      360
gtcgtggtgca agtacctaca gcggagggtg caagttaatg gtctccgagc agtgactgac      420
ccaaatggac ctatcgctgt acagtatttc tctgttaagg agagctggat ccgccactta      480
agactggcgg aagaacctag cctccctggg ttgaggacc tcctcagaat aagggttgag      540
cccaacacgt cgccattggc tgacaaggat gagaaaatct tccggtttgg cagtcacaag      600
tggtacggt                                     609

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<210> SEQ ID NO 5
<211> LENGTH: 203
<212> TYPE: PRT
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 5

Ala Asp Val Tyr Asp Ile Gly His Asp Ala Val Met Tyr Val Ala Glu
1 5 10 15

Gly Lys Val Ser Trp Ala Pro Arg Gly Gly Asp Glu Gly Lys Phe Glu
20 25 30

Thr Val Pro Glu Glu Leu Lys Leu Ile Ala Asn Arg Leu His Ile Ser
35 40 45

Phe Pro Pro His His Ala Val Asp Met Ser Lys Phe Ala Phe Ile Ala
50 55 60

Pro Gly Ser Gly Val Ser Met Arg Val Glu Cys Gln His Gly Cys Leu
65 70 75 80

Pro Ala Asp Thr Val Pro Glu Gly Asn Cys Trp Trp Arg Leu Phe Asp
85 90 95

Leu Leu Pro Leu Glu Val Gln Asn Lys Glu Ile Arg His Ala Asn Gln
100 105 110

Phe Gly Tyr Gln Thr Lys His Gly Val Ala Gly Lys Tyr Leu Gln Arg
115 120 125

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Arg Leu Gln Val Asn Gly Leu Arg Ala Val Thr Asp Pro Asn Gly Pro
130 135 140

Ile Val Val Gln Tyr Phe Ser Val Lys Glu Ser Trp Ile Arg His Leu
145 150 155 160

Arg Leu Ala Glu Glu Pro Ser Leu Pro Gly Phe Glu Asp Leu Leu Arg
165 170 175

Ile Arg Val Glu Pro Asn Thr Ser Pro Leu Ala Asp Lys Asp Glu Lys
180 185 190

Ile Phe Arg Phe Gly Ser His Lys Trp Tyr Gly
195 200

<210> SEQ ID NO 6

<211> LENGTH: 3588

<212> TYPE: DNA

<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 6

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gagcatctca agcactattc ccgcctgcc gacgggaact gtggttgga ctgcatttcc      180
gccatcgcca accggatggt gaattccaaa ttgaaacca cccttccga aagagtgaga      240
ccttcagatg actgggtac tgacgaggat cttgtgaata ccatccaaat cctcaggctc      300
cctgcggcct tggacaggaa cgtgtcttgt gctagcgcca agtacgtgct taagctggaa      360
ggtgagcatt ggactgtctc tgtgaccctt gggatgtccc cttctttgct ccccttgaa      420
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tccgatttg accctgctg ccttgaccga ctggctgagg tgatgcactt gcctagcagt      540
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caggtgtgct tagggaaaat catcagcctt tgtcagggtg ttgaggaatg ctgctgttcc      720
cagaacaaaa ccaaccgggt caccgccgaa gaggtcgcgg caaagattga ccagtaacct      780
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gcacaagggt acgaggttcg tcaccgtgag aggcataact ccgtgctctc taagttggag     1080
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acgaagtctg tcaagagctt gccagagaac aagcctgtcc ctgctccgag caggaaggtc     1380
agatctgatt gtggcagccc gattttaatg ggcgacaatg tcctaacag ttgggaagat     1440
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gagcctgcac ttatgccgcg gttgcaacat atttctaggc cagtgcacac ttgagtgtg     1560
ccggcccaaa ttctgcacc gcgcagagct gtgtcccgac cggtgacgcc ctcgagtga     1620
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gcagcaacgc tgacgtacca ggacgaaccc ctagatttgt ctgcatcctc acagactgaa 1740
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<210> SEQ ID NO 7

<211> LENGTH: 1196

<212> TYPE: PRT

<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 7

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Ala Gly Lys Arg Ala Arg Lys Ala Arg Ser Gly Ala Thr Ala Thr Val
1           5           10           15

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Ala His Arg Ala Leu Pro Ala Arg Glu Thr Gln Gln Ala Lys Lys His
20           25           30

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Glu	Val	Ala	Ser	Ala	Asn	Lys	Ala	Glu	His	Leu	Lys	His	Tyr	Ser	Pro
	35						40					45			
Pro	Ala	Asp	Gly	Asn	Cys	Gly	Trp	His	Cys	Ile	Ser	Ala	Ile	Ala	Asn
	50					55					60				
Arg	Met	Val	Asn	Ser	Lys	Phe	Glu	Thr	Thr	Leu	Pro	Glu	Arg	Val	Arg
65					70					75					80
Pro	Ser	Asp	Asp	Trp	Ala	Thr	Asp	Glu	Asp	Leu	Val	Asn	Thr	Ile	Gln
				85					90					95	
Ile	Leu	Arg	Leu	Pro	Ala	Ala	Leu	Asp	Arg	Asn	Gly	Ala	Cys	Ala	Ser
			100					105					110		
Ala	Lys	Tyr	Val	Leu	Lys	Leu	Glu	Gly	Glu	His	Trp	Thr	Val	Ser	Val
			115				120					125			
Thr	Pro	Gly	Met	Ser	Pro	Ser	Leu	Leu	Pro	Leu	Glu	Cys	Val	Gln	Gly
						135					140				
Cys	Cys	Glu	His	Lys	Gly	Gly	Leu	Gly	Ser	Pro	Asp	Ala	Val	Glu	Val
145					150					155					160
Ser	Gly	Phe	Asp	Pro	Ala	Cys	Leu	Asp	Arg	Leu	Ala	Glu	Val	Met	His
				165					170					175	
Leu	Pro	Ser	Ser	Ala	Ile	Pro	Ala	Ala	Leu	Ala	Glu	Met	Ser	Gly	Asp
			180					185					190		
Pro	Asn	Arg	Pro	Ala	Ser	Pro	Val	Thr	Thr	Val	Trp	Thr	Val	Ser	Gln
							200					205			
Phe	Phe	Ala	Arg	His	Arg	Gly	Gly	Glu	His	Pro	Asp	Gln	Val	Cys	Leu
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Gly	Lys	Ile	Ile	Ser	Leu	Cys	Gln	Val	Ile	Glu	Glu	Cys	Cys	Cys	Ser
225					230					235					240
Gln	Asn	Lys	Thr	Asn	Arg	Val	Thr	Pro	Glu	Glu	Val	Ala	Ala	Lys	Ile
				245					250					255	
Asp	Gln	Tyr	Leu	Arg	Gly	Ala	Thr	Ser	Leu	Glu	Glu	Cys	Leu	Ala	Arg
			260					265					270		
Leu	Glu	Arg	Ala	Arg	Pro	Pro	Ser	Ala	Met	Asp	Thr	Ser	Phe	Asp	Trp
			275				280					285			
Asn	Val	Val	Leu	Pro	Gly	Val	Glu	Ala	Ala	Thr	Gln	Thr	Thr	Lys	Gln
						295					300				
Pro	His	Val	Asn	Gln	Cys	Arg	Ala	Leu	Val	Pro	Val	Val	Thr	Gln	Glu
305					310					315					320
Ser	Leu	Asp	Lys	Asp	Ser	Val	Pro	Leu	Thr	Ala	Phe	Ser	Leu	Ser	Asn
				325					330				335		
Cys	Tyr	Tyr	Pro	Ala	Gln	Gly	Asp	Glu	Val	Arg	His	Arg	Glu	Arg	Leu
			340					345					350		
Asn	Ser	Val	Leu	Ser	Lys	Leu	Glu	Glu	Val	Val	Arg	Glu	Glu	Tyr	Gly
							360					365			
Leu	Thr	Pro	Thr	Gly	Pro	Gly	Pro	Arg	Pro	Ala	Leu	Pro	Asn	Gly	Leu
						375					380				
Asp	Glu	Leu	Lys	Asp	Gln	Met	Glu								

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450	455	460
Gly Ser Pro Ile Leu Met	Gly Asp Asn Val Pro Asn Ser Trp Glu Asp	
465	470	475 480
Leu Ala Val Gly Gly Pro Leu Asp Leu Ser Thr Pro Pro Glu Pro Met		
	485 490	495
Thr Pro Leu Ser Glu Pro Ala Leu Met Pro Ala Leu Gln His Ile Ser		
	500 505	510
Arg Pro Val Thr Pro Leu Ser Val Pro Ala Pro Ile Pro Ala Pro Arg		
	515 520	525
Arg Ala Val Ser Arg Pro Val Thr Pro Ser Ser Glu Pro Ile Ser Val		
	530 535	540
Ser Ala Pro Arg His Lys Phe Gln Gln Val Glu Glu Ala Asn Leu Ala		
545	550 555	560
Ala Ala Thr Leu Thr Tyr Gln Asp Glu Pro Leu Asp Leu Ser Ala Ser		
	565 570	575
Ser Gln Thr Glu Tyr Glu Ala Ser Pro Leu Ala Pro Leu Gln Asn Met		
	580 585	590
Gly Ile Leu Glu Val Gly Gly Gln Glu Ala Glu Glu Ile Leu Ser Glu		
	595 600	605
Ile Ser Asp Ile Pro Asn Asp Ile Asn Pro Ala Pro Val Ser Ser Ser		
	610 615	620
Ser Ser Leu Ser Ser Val Lys Ile Thr Arg Pro Lys Tyr Ser Ala Gln		
625	630 635	640
Ala Ile Ile Asp Ser Gly Gly Pro Cys Ser Gly His Leu Gln Lys Glu		
	645 650	655
Lys Glu Ala Cys Leu Ser Ile Met Arg Glu Ala Cys Asp Ala Thr Lys		
	660 665	670
Leu Gly Asp Pro Ala Thr Gln Glu Trp Leu Ser Arg Met Trp Asp Arg		
	675 680	685
Val Asp Met Leu Thr Trp Arg Asn Thr Ser Ala Tyr Gln Ala Phe Arg		
	690 695	700
Thr Leu Asp Gly Arg Phe Glu Phe Leu Pro Lys Met Ile Leu Glu Thr		
705	710 715	720
Pro Pro Pro Tyr Pro Cys Gly Phe Val Met Leu Pro His Thr Pro Ala		
	725 730	735
Pro Ser Val Gly Ala Glu Ser Asp Leu Thr Ile Gly Ser Val Ala Thr		
	740 745	750
Glu Asp Val Pro Arg Ile Leu Gly Lys Ile Glu Asn Ala Gly Glu Met		
	755 760	765
Thr Asn Gln Gly Pro Leu Ala Ser Ser Glu Glu Glu Pro Ala Asp Asp		
	770 775	780
Gln Pro Ala Lys Asp Ser Arg Ile Ser Ser Arg Gly Phe Asp Glu Ser		
785	790 795	800
Thr Ala Ala Pro Ser Ala Gly Thr Gly Gly Ala Gly Leu Phe Thr Asp		
	805 810	815
Leu Pro Pro Ser Asp Gly Val Asp Ala Asp Gly Gly Gly Pro Leu Gln		
	820 825	830
Thr Val Lys Lys Lys Ala Glu Arg Leu Phe Asp Gln Leu Ser Arg Gln		
	835 840	845
Val Phe Asn Leu Val Ser His Leu Pro Val Phe Phe Ser His Leu Phe		
	850 855	860
Lys Ser Asp Ser Gly Tyr Ser Pro Gly Asp Trp Gly Phe Ala Ala Phe		
865	870 875	880

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attgccttgg ttgttttgat ttctgtttcc atcgaggcca tggtcacag gttgagttgc   300
aaggctgata tgctgtgcgt tttacttgca atcgccagct atgtttgggt accccttacc   360
tgggttgcttt gtgtgtttcc ttgctgggtg cgctgggtct ctttgcaccc cctcaccatc   420
ctatgggttg tgtttttctt gatttctgta aatatgcctt caggaatctt ggccgtgggtg   480
ttgttggttt ctctttggct tctaggtcgt tatactaata ttgctggtct tgtaaccccc   540
tatgacattc atcattacac cagtggcccc cgcggtgttg ccgccttggc tacgcacca   600
gatgggacct acttggccgc tgccgccgc gctgcgttga ctggccgcac catgctgttt   660
accccgcttc agcttgggtc ccttcttgag                                     690

```

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<210> SEQ ID NO 9
<211> LENGTH: 230
<212> TYPE: PRT
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

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```

<400> SEQUENCE: 9

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```

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

```

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<210> SEQ ID NO 10
<211> LENGTH: 612
<212> TYPE: DNA
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

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```

<400> SEQUENCE: 10

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```

ggtgctttca gaactcaaaa gccctcactg aacaccgtca atgtggtcgg gtcctccatg   60

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ggctctggcg ggggtgttcac catcgacggg aaaattaagt gcgtaactgc cgcacatgtc 120
cttacgggta attcagctag ggtttccggg gtcgggttca atcaaatget tgactttgat 180
gtaaaagggg acttcgccat agctgattgc ccgaattggc aaggggctgc tcccaagacc 240
caattctgca aggatggatg gactggccgt gcctattggc tgacatcctc tggcgctgaa 300
cccgggtgca ttgggaatgg attcgccttc tgcttcaccg cgtgcggcga ttccgggtcc 360
ccagtgatca ccgaagccgg tgagcttgtc ggcgttcaca caggatcaaa caaacaagga 420
ggaggcattg tcacgcgccc ctacggccag ttttgtaatg tggcaccat caagctgagc 480
gaattaagtg aattctttgc tggacctaag gtcccgcctg gtgatgtgaa ggttggcagc 540
cacataatta aagacataag cgaggtgcct tcagatcttt gcgccttgc tgcgtccaaa 600
cccgaactgg aa 612

```

```

<210> SEQ ID NO 11
<211> LENGTH: 204
<212> TYPE: PRT
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

```

```

<400> SEQUENCE: 11

```

```

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

```

```

<210> SEQ ID NO 12
<211> LENGTH: 510
<212> TYPE: DNA
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

```

```

<400> SEQUENCE: 12

```

```

ggaggcctct ccaccgtcca acttctgtgt gtgtttttcc tcctgtggag aatgatggga 60
catgcctgga cgcccttggg tgctgtgggt ttttttatct tgaatgaggt tctcccagct 120

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gtcctgggtcc ggagtgtttt ctcccttgga atgtttgtgc tatcttggtc cacaccatgg 180
tctgcgcaag ttctgatgat caggcttcta acagcagctc ttaacaggaa cagatgggtca 240
cttgccctttt acagcctcgg tgcagtgacc ggttttgtcg cagatcttgc ggcaactcag 300
gggcatecgt tgcaggcagt gatgaattta agcacctatg ccttccctgcc tcggatgatg 360
gttgtgacct caccagtcct agtgattgctg tgtggtgttg tgcacctcct tgccataatt 420
ttgtacttgt ttaagtaccg ttgcctgcac aatgtccttg ttggcgatgg agtgttctct 480
gcggctttct tcttgcgata ctttgccgag 510

```

```

<210> SEQ ID NO 13
<211> LENGTH: 170
<212> TYPE: PRT
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 13

```

```

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

```

```

<210> SEQ ID NO 14
<211> LENGTH: 48
<212> TYPE: DNA
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 14

```

```

ggaaagtgtga gggaaggggt gtcgcaatcc tgcgggatga atcatgag 48

```

```

<210> SEQ ID NO 15
<211> LENGTH: 16
<212> TYPE: PRT
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 15

```

```

Gly Lys Leu Arg Glu Gly Val Ser Gln Ser Cys Gly Met Asn His Glu
1      5      10      15

```

```

<210> SEQ ID NO 16

```

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<211> LENGTH: 777
 <212> TYPE: DNA
 <213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 16

```

tcaactgactg gtgccctcgc tatgagactc aatgacgagg acttggattt ccttacgaaa      60
tggactgatt ttaagtgctt tgtttctgcg tccaacatga ggaatgcagc gggccaattc      120
atcgaggctg cctatgctaa agcacttaga gtagaacttg ccagttgggt gcaggttgat      180
aaggttcgag gtactttggc caaacttgaa gcttttctg ataccgtggc accccaactc      240
tcgcccggtg acattgttgt tgctcttggc cacacgcctg ttggcagtat cttcgacctc      300
aaggttggtg gcaccaagca taccctccaa gccattgaga ccagagtctc tgccgggtcc      360
aaaatgacgc tggcgcgcgt cgttgacca acccccacgc cccacccgc acccgtgccc      420
atccccctcc caccgaaagt tctggagaat ggccccaacg cctgggggga tgaggaccgt      480
ttgaataaga agaagaggcg caggatggaa gccgtcgga tctttgttat gggcggaag      540
aagtaccaga aattttggga caagaattcc ggtgatgtgt tttatgagga ggtccatgat      600
aacacagatg cgtgggagtg cctcagagtt ggcgaccctg ccgactttga ccctgagaag      660
ggaactctgt gtgggcatac caccattgaa gataaggctt acaatgtcta cgctcccca      720
tctggcaaga agttcctggt ccccgtaac ccagagagcg gaagagccca atgggaa      777

```

<210> SEQ ID NO 17
 <211> LENGTH: 259
 <212> TYPE: PRT
 <213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 17

```

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

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Arg Val Gly Asp Pro Ala Asp Phe Asp Pro Glu Lys Gly Thr Leu Cys
 210 215 220

Gly His Thr Thr Ile Glu Asp Lys Ala Tyr Asn Val Tyr Ala Ser Pro
 225 230 235 240

Ser Gly Lys Lys Phe Leu Val Pro Val Asn Pro Glu Ser Gly Arg Ala
 245 250 255

Gln Trp Glu

<210> SEQ ID NO 18
 <211> LENGTH: 138
 <212> TYPE: DNA
 <213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 18

gctgcaaagc tttccgtgga gcaggccctt ggcattgatga atgtcgacgg tgaactgaca 60
 gccaaagaac tggagaaact gaaaagaata attgacaaac tccagggcct gactaaggag 120
 cagtgtttaa actgctag 138

<210> SEQ ID NO 19
 <211> LENGTH: 45
 <212> TYPE: PRT
 <213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 19

Ala Ala Lys Leu Ser Val Glu Gln Ala Leu Gly Met Met Asn Val Asp
 1 5 10 15

Gly Glu Leu Thr Ala Lys Glu Leu Glu Lys Leu Lys Arg Ile Ile Asp
 20 25 30

Lys Leu Gln Gly Leu Thr Lys Glu Gln Cys Leu Asn Cys
 35 40 45

<210> SEQ ID NO 20
 <211> LENGTH: 1917
 <212> TYPE: DNA
 <213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 20

gccgccagcg gcttgaccgc ctgtggctgc gccggcttgg ttgttactga gacagcggtg 60
 aaaatagtca aatttcacaa ccggaccttc accctaggac ctgtgaattt aaaagtggcc 120
 agtgaggttg agctaaaaga cgcggtcgag cacaaccaac acccggttgc aagaccggtt 180
 gatggtggtg ttgtgctcct gcgctccgca gttccttcgc ttatagacgt cttgatctcc 240
 ggtgctgatg catctcccaa gttactcgcc cgccacgggc cgggaaacac tgggatcgat 300
 ggcacgcttt gggattttga ggccgaagcc accaaagagg aaatcgact cagtgcgcaa 360
 ataatacagg cttgtgacat taggcgcggc gacgcacctg aaattggtct cccttacaag 420
 ctgtaccctg ttatggggcaa ccttgagcgg gtaaaaggag ttttgagaa tacaagggtt 480
 ggagacatac cttacaaaac cccagtgac actggaagcc cagtgcacgc ggctgcctgc 540
 ctcacgcccc atgccactcc ggtgactgat gggcgctccg tcttggccac gacctgccc 600
 tccggttttg agttgtatgt accgaccatt ccagcgtctg tccttgatta tcttgattct 660
 aggctgact gccccaaaca gttgacagag cacggctgtg aggatgccgc attgagagac 720
 ctctccaaagt atgacttgte caccgaagc tttgttttgc ctggagttct tcgccttggtg 780
 cgtaagtacc tgtttgcccc tgtgggtaag tgcccggccc ttcacggcc ttccacttac 840
 cctgccaaga attctatggc tggaataaat gggaacaggt ttccaaccaa ggacattcag 900

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agcgtccctg aaatcgacgt tctgtgcgca cagggcgtgc gagaaaactg gcaaactgtt   960
acctcttgta ccctcaagaa acagtattgc gggaagaaga agactaggac aatactcggc   1020
accaataact tcattgcgct ggcccaccgg gcagcggtga gtggtgtcac ccagggttc   1080
atgaaaaagg cgtttaactc gcccatcgcc ctcgggaaaa acaaatttaa ggagctacag   1140
actccggtct tgggcagggt ccttgaagct gatcttgcac cctgcgatcg atccacacct   1200
gcaattgtcc gctggtttgc cgccaatctt ctttatgaac ttgcctgtgc tgaagagcat   1260
ctaccgtcgt acgtgctgaa ctgctgccac gacttactgg tcacgcagtc cggcgcagtg   1320
actaagagag gtggcctgtc gtctggcgac ccgatcactt ctgtgtccaa caccatttac   1380
agcttggtga tatatgcaca gcacatggtg ctacgttact taaaagtgg tcaccccat   1440
ggccttctgt ttctacaaga ccagctaaag tttgaggaca tgctcaaggt tcaacccctg   1500
atcgctctatt cggacgacct cgtgctgtat gccgagtcct ccaccatgcc aaactaccac   1560
tggtggggtg aacatctgaa cctgatgctg ggttttcaga cggacccaaa gaagacagcc   1620
ataacagact cgccatcatt tctaggctgt aggataataa atgggcgcca gctagtcccc   1680
aaccgtgaca ggattctcgc ggccctcgcc taccacatga aggcgagcaa tgtttctgaa   1740
tactacgcct cggcggtgtc aatactcatg gacagctgtg cttgtttgga gtatgatcct   1800
gaatgggttg aagaacttgt ggttggaata gcgcagtgcg ccgcgaagga cggctacagc   1860
tttcccggcc cgccgttctt cttgtccatg tgggaaaaac tcaggtccaa ttatgag   1917

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<210> SEQ ID NO 21

<211> LENGTH: 639

<212> TYPE: PRT

<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 21

```

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

```

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

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610	615	620	
Pro Phe Phe Leu Ser Met Trp Glu Lys Leu Arg Ser Asn Tyr Glu			
625	630	635	
 <210> SEQ ID NO 22			
<211> LENGTH: 1323			
<212> TYPE: DNA			
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus			
 <400> SEQUENCE: 22			
gggaagaagt ccagagtgtg cgggtactgc ggggccccgg ccccgtaacg cactgcctgt		60	
ggcctcgacg tctgtattta ccacaccac ttccaccagc attgtccagt cataatctgg		120	
tgtggccatc cagcgggttc tggttcttgt agtgagtgca aacccccct agggaaaggc		180	
acaagccctc tagatgaggt gttggaacaa gtcccgata agcctccacg gaccgtaatc		240	
atgcatgtgg agcaggttct caccctctt gaccagga gataccagac tcgccgga		300	
ttagtctcgg ttaggcgtgg catcagggga aatgaagtg acctaccaga cggtgattat		360	
gctagcaccg ccttgcctcc cacttgtaaa gagatcaaca tggctcgtgt cgcttctaat		420	
gtgttgcgca gcaggttcat catcggtcca cccggtgctg ggaaaacata ctggctcctt		480	
caacaggtcc aggatgtgta tgcattttac acaccaactc atcagacat gcttgacatg		540	
attaaggctt tggggacgtg ccggttcaac gtcccgagc gcacaacgct gcaattccct		600	
gccccctccc gtaccggccc gtgggttcgc atcctggccg gcggttggtg tcctggcaag		660	
aattccttcc tggatgaagc agcgtattgt aatcaccttg atgtcttgag gcttcttagc		720	
aaaactaccc tcacctgtct gggagacttc aaacaactcc acccagtggg ttttgattct		780	
cattgctatg tttttgacat catgcctcag actcaactga agaccatctg gaggtttgga		840	
cagaatatct gtgatgccat tcagccagat tacagggaca aacttggtgc catggccaac		900	
acaaccgtg taacctagc ggaacaaact gtcaagtatg ggcaagtcct cacccttac		960	
cacagggacc gagaggacgg cgccatcaca attgactcca gtcaaggcgc cacatttgat		1020	
gtggttacat tgcatttgcc cactaaagat tcaactcaaca ggcaaagagc ccttggtgct		1080	
atcaccaggg caagacatgc tatctttgtg tatgaccac acaggcaact gcagagcatg		1140	
tttgatcttc ctgcaaaagg cacaccgtc aacctcgccg tgcaccgtga cgagcagctg		1200	
atcgtgctag atagaaataa caaagaatgc acggttgctc aggcctctagg caatggggat		1260	
aaattcaggg ccacagacaa gcgcgttgta gattctctcc gcgccatttg tgcagatcta		1320	
gaa		1323	

<210> SEQ ID NO 23
 <211> LENGTH: 441
 <212> TYPE: PRT
 <213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 23

Gly Lys Lys Ser Arg Val Cys Gly Tyr Cys Gly Ala Pro Ala Pro Tyr		
1	5	10 15
Ala Thr Ala Cys Gly Leu Asp Val Cys Ile Tyr His Thr His Phe His		
	20	25 30
Gln His Cys Pro Val Ile Ile Trp Cys Gly His Pro Ala Gly Ser Gly		
	35	40 45
Ser Cys Ser Glu Cys Lys Pro Pro Leu Gly Lys Gly Thr Ser Pro Leu		
50	55	60

-continued

Asp	Glu	Val	Leu	Glu	Gln	Val	Pro	Tyr	Lys	Pro	Pro	Arg	Thr	Val	Ile	65	70	75	80
Met	His	Val	Glu	Gln	Gly	Leu	Thr	Pro	Leu	Asp	Pro	Gly	Arg	Tyr	Gln	85	90	95	
Thr	Arg	Arg	Gly	Leu	Val	Ser	Val	Arg	Arg	Gly	Ile	Arg	Gly	Asn	Glu	100	105	110	
Val	Asp	Leu	Pro	Asp	Gly	Asp	Tyr	Ala	Ser	Thr	Ala	Leu	Leu	Pro	Thr	115	120	125	
Cys	Lys	Glu	Ile	Asn	Met	Val	Ala	Val	Ala	Ser	Asn	Val	Leu	Arg	Ser	130	135	140	
Arg	Phe	Ile	Ile	Gly	Pro	Pro	Gly	Ala	Gly	Lys	Thr	Tyr	Trp	Leu	Leu	145	150	155	160
Gln	Gln	Val	Gln	Asp	Gly	Asp	Val	Ile	Tyr	Thr	Pro	Thr	His	Gln	Thr	165	170	175	
Met	Leu	Asp	Met	Ile	Lys	Ala	Leu	Gly	Thr	Cys	Arg	Phe	Asn	Val	Pro	180	185	190	
Ala	Gly	Thr	Thr	Leu	Gln	Phe	Pro	Ala	Pro	Ser	Arg	Thr	Gly	Pro	Trp	195	200	205	
Val	Arg	Ile	Leu	Ala	Gly	Gly	Trp	Cys	Pro	Gly	Lys	Asn	Ser	Phe	Leu	210	215	220	
Asp	Glu	Ala	Ala	Tyr	Cys	Asn	His	Leu	Asp	Val	Leu	Arg	Leu	Leu	Ser	225	230	235	240
Lys	Thr	Thr	Leu	Thr	Cys	Leu	Gly	Asp	Phe	Lys	Gln	Leu	His	Pro	Val	245	250	255	
Gly	Phe	Asp	Ser	His	Cys	Tyr	Val	Phe	Asp	Ile	Met	Pro	Gln	Thr	Gln	260	265	270	
Leu	Lys	Thr	Ile	Trp	Arg	Phe	Gly	Gln	Asn	Ile	Cys	Asp	Ala	Ile	Gln	275	280	285	
Pro	Asp	Tyr	Arg	Asp	Lys	Leu	Val	Ser	Met	Val	Asn	Thr	Thr	Arg	Val	290	295	300	
Thr	Tyr	Val	Glu	Lys	Pro	Val	Lys	Tyr	Gly	Gln	Val	Leu	Thr	Pro	Tyr	305	310	315	320
His	Arg	Asp	Arg	Glu	Asp	Gly	Ala	Ile	Thr	Ile	Asp	Ser	Ser	Gln	Gly	325	330	335	
Ala	Thr	Phe	Asp	Val	Val	Thr	Leu	His	Leu	Pro	Thr	Lys	Asp	Ser	Leu	340	345	350	
Asn	Arg	Gln	Arg	Ala	Leu	Val	Ala	Ile	Thr	Arg	Ala	Arg	His	Ala	Ile	355	360	365	
Phe	Val	Tyr	Asp	Pro	His	Arg	Gln	Leu	Gln	Ser	Met	Phe	Asp	Leu	Pro	370	375	380	
Ala	Lys	Gly	Thr	Pro	Val	Asn	Leu	Ala	Val	His	Arg	Asp	Glu	Gln	Leu	385	390	395	400
Ile	Val	Leu	Asp	Arg	Asn	Asn	Lys	Glu	Cys	Thr	Val	Ala	Gln	Ala	Leu	405	410	415	
Gly	Asn	Gly	Asp	Lys	Phe	Arg	Ala	Thr	Asp	Lys	Arg	Val	Val	Asp	Ser	420	425	430	
Leu	Arg	Ala	Ile	Cys	Ala	Asp	Leu	Glu								435	440		

<210> SEQ ID NO 24

<211> LENGTH: 669

<212> TYPE: DNA

<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 24

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```

gggtcgagct ctccgctccc caaggctgca cacaacttgg gattttatatt ctcacctgat      60
ttgacacagt ttgctaaact cccggtagaa cttgcacccc actggcccggt ggtgacaacc      120
cagaacaatg aaaagtggcc agaccggctg gttgccagcc ttccgccctat ccataaatat      180
agccgcgcgt gcatcggtgc cggtctatag gtggggccct cggtgtttct aggcacccct      240
gggggttgtgt catactatct cacaaaattt gttaagggcg aggctcaagt gcttccggag      300
acagtottca gcaccggcgc aattgaggtg gattgccggg agtatcttga tgatcgggag      360
cgagaagttg ctgagtcctt cccacatgcc ttcattggcg acgtcaaagg cactaccgtt      420
ggaggatgtc accatgtcac ctccaaatac cttccgcgct tccttcccaa ggaatcagtt      480
gcggtagtcg ggggtttcaag ccccgggaaa gccgcaaaag cagtttgcac attaacagat      540
gtgtacctcc cagaccttga agcttacctc caccagaga cccagtccaa gtgctggaaa      600
atgatgttgg acttcaagga agttcgactg atggtctgga aagacaaaac ggcctatatt      660
caacttgaa

```

```

<210> SEQ ID NO 25
<211> LENGTH: 223
<212> TYPE: PRT
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

```

```

<400> SEQUENCE: 25

```

```

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

```

```

<210> SEQ ID NO 26
<211> LENGTH: 462
<212> TYPE: DNA

```


-continued

<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 26

```

ggccgccatt tcacctggta tcagcttgca agctatgcct cgtacatccg agttcctgtt      60
aactctacgg tgtatttgga cccctgcatg ggccctgccc tttgcaacag aagagttgtc      120
gggtccactc attggggggc tgacctcgca gtcacccctt atgattatgg tgccaaaatc      180
attctgtcta gtgcatacca tggtgaaatg cctcctgggt acaaaatcct ggcgtgcgcg      240
gagttctcgc ttgacgatcc agtgaggtag aaacacacct ggggggtttga atcggataca      300
gcgtatctgt acgagttcac cggaaacggt gaggactggg aggattacaa tgatgcgttt      360
cgtgcgcgcc agaaggga aatttataag gccactgcca ccagcatgag gtttcatttt      420
cccccgggcc ctgtcattga accaactttg ggcctgaatt ga                          462

```

<210> SEQ ID NO 27

<211> LENGTH: 153

<212> TYPE: PRT

<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 27

```

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

```

<210> SEQ ID NO 28

<211> LENGTH: 771

<212> TYPE: DNA

<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 28

```

atgaaatggg ggctatgcaa agcctttttg acaaaattgg ccaacttttt gtggatgctt      60
tcacggaatt tttggtgtcc attgttgata tcatcatatt tttggccatt ttgtttggct      120
tcaccatcgc cggttggtcg gtggtctttt gcatcagatt ggtttgctcc gcggtactcc      180
gtgcgcgccc taccattcac cctgagcaat tacagaagat cctatgaggc ctttctttct      240
cagtgccggg tggacattcc cacctgggga actaaacatc ccttggggat gctttggcac      300
cataaggtgt caaccctgat tgatgaaatg gtgtcgcgtc gaatgtaccg catcatggaa      360
aaagcaggac aggctgcctg gaaacaggtg gtgagcgagg ctacgctgtc tcgcattagt      420

```

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```

ggtttgatg tgggtgctca ttttcagcat cttgcgcga ttgaagcga gacctgtaa 480
tatttgccct ctcggctgcc catgctacac aacctgcga tgacagggtc aaatgtaacc 540
atagtgtata atagtacttt gaatcagggtg ttgctattt ttccaacccc tggttcccgg 600
ccaaagcttc atgattttca gcaatggcta atagctgtgc attcctccat attttcctct 660
gttgacagctt cttgtactct tttgtgtgtg ctgtggttgc ggattccaat gctacgtact 720
gtttttgggtt tccactgggtt aggggcaatt tttccttcga actcacagtg a 771

```

```

<210> SEQ ID NO 29
<211> LENGTH: 256
<212> TYPE: PRT
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

```

```

<400> SEQUENCE: 29

```

```

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

```

```

<210> SEQ ID NO 30
<211> LENGTH: 765
<212> TYPE: DNA
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

```

```

<400> SEQUENCE: 30

```

```

atggctaata gctgtgcatt cctccatatt ttcctctgtt gcagcttctt gtactctttt 60

```

-continued

```

tgttgtgtg tgggtgcgga ttccaatgct acgtactgtt ttgggtttcc actgggttagg 120
ggcaattttt ccttcgaact cacagtgaat tacacgggtg gtccaccttg cctcaccg 180
caagcagcgg ctgagatcta cgaaccggc aggtctcttt ggtgcaggat agggcatgac 240
cgatgtaggg aggacgatca tgacgaacta gggttcatgg ttccgcctgg cctctccagc 300
gaaggccact tgaccagtgt ttacgcctgg ttggcggtcc tgctcttcag ctacacggcc 360
cagttccatc ccgagatatt tgggataggg aatgtgagtc aagtttatgt tgacatcaag 420
caccaattca tctgcgcoga acatgacggg cagaacgcca ccttgcctcg ccatgacaac 480
atttcagcgg tgtttcagac ctactaccaa catcaggctg acggcggcaa ttggtttcac 540
ctagaatggc tgcgcccctt cttttcctct tggttggttt taaatgttcc gtggtttctc 600
aggcgttcgc ctgcaagcca tgtttcagtt cgagtctttc agacatcaag accaacacca 660
ccgcagcagc aagctttgtt gtctctcaag acatcagctg ccttaggcat ggcgactcgt 720
cctctgaggg gattcgcaaa agctctcagt gccgcacggc gatag 765

```

```

<210> SEQ ID NO 31
<211> LENGTH: 254
<212> TYPE: PRT
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus
<400> SEQUENCE: 31

```

```

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

```

-continued

<210> SEQ ID NO 32
 <211> LENGTH: 537
 <212> TYPE: DNA
 <213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 32

```

atggctgcgc cccttctttt cctcttggtt ggttttaaat gttctgtggt ttctcaggcg      60
ttcgccctgca agccatgttt cagttcgagt ctttcagaca tcaagaccaa caccaccgca      120
gcagcaagct ttgtgtgctt ccaagacatc agctgcctta ggcatggcga ctgctcctct      180
gaggcgattc gcaaaagctc tcagtgccgc acggcgatag ggacaccctg gtacatcacc      240
atcacagcca atgtgacaga tgagaattat ttacattctt ctgatctcct catgctttct      300
tcttgcccttt tctatgcttc tgagatgagt gaaaagggat tcaagggtgt atttggaat      360
gtgtcaggca tcgtggctgt gtgtgtcaac tttaccagct acgtccaaca tgtcaaggag      420
tttaccacac gctccttggt ggctcgacat gtgcggctgc ttcatttcat gacacctgag      480
accatgaggt gggcaaccgt tttagcctgt ctttttgcca ttctgttggc aatttga      537

```

<210> SEQ ID NO 33
 <211> LENGTH: 178
 <212> TYPE: PRT
 <213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 33

```

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

Ala Ile

```

<210> SEQ ID NO 34
 <211> LENGTH: 603
 <212> TYPE: DNA
 <213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 34

```

atgttgggga aatgcttgac cgcgggctgt tgctcgcat tgctttcttt gtggtgtatc      60

```

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```

gtgccgttct gttttgtgc gtcgtcaac gccaacagca acagcagctc ccatttacag   120
ttgatttata acttgacgct atgtgagctg aatggcacag attggctggc taacaaattt   180
gattgggcag tggagacttt tgcatcttt cccgtgttga ctcacattgt ctctatggt   240
gccctcacca ccagccatct ccttgacaca gtcggtctgg tcaactgtgc taccgccggg   300
ttttatcacg ggcggtatgt cttgagtagc atctacgcgg tctgtgccct ggctgcgttg   360
atttgcttcg tcattaggtt tgcaagaac tgcagtctct ggcgctactc atgtaccaga   420
tataccaact ttcttctgga cactaagggc agactctatc gttggcggtc gcccgctc   480
atagagaaaa ggggtaaagt tgaggtcgaa ggtcatctga tcgacctcaa aagagttgtg   540
cttgatgggt cctgggcaac ccccttaacc agagtttcag cggaacaatg gggtcgtcct   600
tag                                                                    603

```

```

<210> SEQ ID NO 35
<211> LENGTH: 200
<212> TYPE: PRT
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

```

```

<400> SEQUENCE: 35

```

```

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

```

```

<210> SEQ ID NO 36
<211> LENGTH: 525
<212> TYPE: DNA
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

```

```

<400> SEQUENCE: 36

```

```

atggggctcgt ccttagacga cttctgccat gatagcacgg ctccacaaaa ggtgcttttg   60
gcgttttcta ttacctacac gccagtgatg atatatgccc taaaggtgaa tcgcggccga   120

```

-continued

```

ctgctagggc ttctgcacct ttgtattttt ctgaattgtg ctttcacott cgggtacatg 180
acattcgcg cacttcagag cacaataag gtcgcgctca ctatgggagc agtagttgca 240
ctcctttggg ggggtgtactc agccatagaa acctggaaat tcatcacctc cagatgccgt 300
ttgtgcttgc taggccgcaa gtacattctg gccctgccc accacgttga aagtgcgca 360
ggctttcatc cgattgcggc aaatgataac cagcatttg tcgtccggcg tcccggctcc 420
actacggtca acggcacatt ggtgccggg ttgaaaagcc tcgtgttggg tggcagaaaa 480
gctgttaaac agggagtggg aaacctgtc aaatatgcca aataa 525

```

```

<210> SEQ ID NO 37
<211> LENGTH: 174
<212> TYPE: PRT
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus
<400> SEQUENCE: 37

```

```

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

```

```

<210> SEQ ID NO 38
<211> LENGTH: 372
<212> TYPE: DNA
<213> ORGANISM: Porcine reproductive and respiratory syndrome virus
<400> SEQUENCE: 38

```

```

atgccaaata acaacggcaa gcagcagaag aaaaagaagg gggatggcca gccagtcaat 60
cagctgtgcc agatgtctgg taagatcatc gccagcaaa accagtccag aggcaaggga 120
ccgggaaaga aaaataagaa gaaaaacccg gagaagcccc atttctctct agcgactgaa 180
gatgacgtca gacatcactt taccctagt gagcggcaat tgtgtctgtc gtcaatccag 240
actgccttta atcaaggcgc tggaacttgt accctgtcag attcaggag gataagttac 300
actgtggagt ttagtttgcc gacgcatcat actgtgcgcc tgatccggt cacagcatca 360
ccctcagcat ga 372

```

-continued

<210> SEQ ID NO 39
 <211> LENGTH: 123
 <212> TYPE: PRT
 <213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 39

```

Met Pro Asn Asn Asn Gly Lys Gln Gln Lys Lys Lys Lys Gly Asp Gly
 1             5             10             15

Gln Pro Val Asn Gln Leu Cys Gln Met Leu Gly Lys Ile Ile Ala Gln
          20             25             30

Gln Asn Gln Ser Arg Gly Lys Gly Pro Gly Lys Lys Asn Lys Lys Lys
          35             40             45

Asn Pro Glu Lys Pro His Phe Pro Leu Ala Thr Glu Asp Asp Val Arg
          50             55             60

His His Phe Thr Pro Ser Glu Arg Gln Leu Cys Leu Ser Ser Ile Gln
          65             70             75             80

Thr Ala Phe Asn Gln Gly Ala Gly Thr Cys Thr Leu Ser Asp Ser Gly
          85             90             95

Arg Ile Ser Tyr Thr Val Glu Phe Ser Leu Pro Thr His His Thr Val
          100            105            110

Arg Leu Ile Arg Val Thr Ala Ser Pro Ser Ala
          115            120

```

<210> SEQ ID NO 40
 <211> LENGTH: 156
 <212> TYPE: DNA
 <213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 40

```

atgttcaagt atgttgggga aatgcttgac cgcgggctgt tgctcgcat tgctttcttt      60
gtggtgtatc gtgccgttct gttttgctgc gctcgtaac gccaacagca acagcagctc      120
ccatttacag ttgatttaca acttgacgct atgtga                                156

```

<210> SEQ ID NO 41
 <211> LENGTH: 51
 <212> TYPE: PRT
 <213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 41

```

Met Phe Lys Tyr Val Gly Glu Met Leu Asp Arg Gly Leu Leu Leu Ala
 1             5             10             15

Ile Ala Phe Phe Val Val Tyr Arg Ala Val Leu Phe Cys Cys Ala Arg
          20             25             30

Gln Arg Gln Gln Gln Gln Leu Pro Phe Thr Val Asp Leu Gln Leu
          35             40             45

Asp Ala Met
          50

```

<210> SEQ ID NO 42
 <211> LENGTH: 222
 <212> TYPE: DNA
 <213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 42

```

atgggggcta tgcaaagcct ttttgacaaa attggccaac tttttgtgga tgctttcacg      60
gaatttttgg tgtccattgt tgatatcacc atatttttgg ccattttgtt tggttcacc      120
atcgccggtt ggctgggtgt cttttgcacc agattggttt gctccgcggt actccgtgcg      180

```

-continued

cgccctacca ttcacctga gcaattacag aagatcctat ga

222

<210> SEQ ID NO 43

<211> LENGTH: 73

<212> TYPE: PRT

<213> ORGANISM: Porcine reproductive and respiratory syndrome virus

<400> SEQUENCE: 43

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

Asp Ala Phe Thr Glu Phe Leu Val Ser Ile Val Asp Ile Ile Ile Phe
 20 25 30

Leu Ala Ile Leu Phe Gly Phe Thr Ile Ala Gly Trp Leu Val Val Phe
 35 40 45

Cys Ile Arg Leu Val Cys Ser Ala Val Leu Arg Ala Arg Pro Thr Ile
 50 55 60

His Pro Glu Gln Leu Gln Lys Ile Leu
 65 70

What is claimed is:

1. A non-structural PRRSV-CON polypeptide having a sequence selected from the group consisting of SEQ ID NO:3, 5, and 7.

2. The polypeptide of claim 1, wherein the polypeptide is encoded by a nucleic acid, respectively, having a sequence selected from the group consisting of SEQ ID NO:2, 4, and 6.

3. A virus particle comprising the non-structural PRRSV-CON polypeptide of claim 1.

4. A composition comprising the virus particle of claim 3 and a pharmaceutically acceptable carrier.

5. The composition of claim 4, further comprising an adjuvant.

6. A composition comprising the non-structural polypeptide of claim 1 and a pharmaceutically acceptable carrier.

7. The composition of claim 6, further comprising an adjuvant.

8. A method for eliciting an immune response to PPRSV in a porcine, comprising administering, to a porcine:

- (i) an effective amount of the polypeptide of claim 1;
- (ii) an effective amount of the virus particle of claim 3; or
- (iii) an effective amount of the composition of claim 6.

9. The method of claim 8, wherein the administration is selected from the group consisting of intramuscularly, intraperitoneally, and orally.

10. A method for treating or preventing PPRS in a porcine, comprising administering, to a porcine:

- (i) an effective amount of the polypeptide of claim 1;
- (ii) an effective amount of the virus particle of claim 3; or
- (iii) an effective amount of the composition of claim 6.

11. The method of claim 10, wherein the administration is selected from the group consisting of intramuscularly, intraperitoneally, and orally.

* * * * *