

US 20100189735A1

(19) United States (12) Patent Application Publication (10) Pub. No.: US 2010/0189735 A1

Jestin et al.

(54) CIRCOVIRUS SEQUENCES ASSOCIATED WITH PIGLET WEIGHT LOSS DISEASE (PWD)

(75) Inventors: André Jestin, Saint-Brieuc (FR); Emmanuel Albina, Tregueux (FR); Pierre Le Cann, Pledran (FR); Philippe Blanchard, Plerin (FR); Evelyne Hutet, Plerin (FR); Claire Arnauld, Saint-Brieuc (FR); Catherine Truong, Saint-Brieuc (FR); Dominique Mahe, Saint-Carreuc (FR); Roland Cariolet, Ploufragan (FR); François Madec, Saint-Brieuc (FR)

Correspondence Address: BUCHANAN, INGERSOLL & ROONEY PC POST OFFICE BOX 1404 ALEXANDRIA, VA 22313-1404 (US)

- (73) Assignee: **WYETH**, Madison, NJ (US)
- (21) Appl. No.: 12/718,183
- (22) Filed: Mar. 5, 2010

Related U.S. Application Data

(60) Continuation of application No. 11/588,237, filed on Oct. 27, 2006, now Pat. No. 7,722,883, which is a

(10) Pub. No.: US 2010/0189735 A1 (43) Pub. Date: Jul. 29, 2010

division of application No. 10/718,264, filed on Nov. 21, 2003, now Pat. No. 7,179,472, which is a division of application No. 09/514,245, filed on Feb. 28, 2000, now Pat. No. 6,703,023, which is a continuation-inpart of application No. PCT/FR98/02634, filed on Dec. 4, 1998.

(30) Foreign Application Priority Data

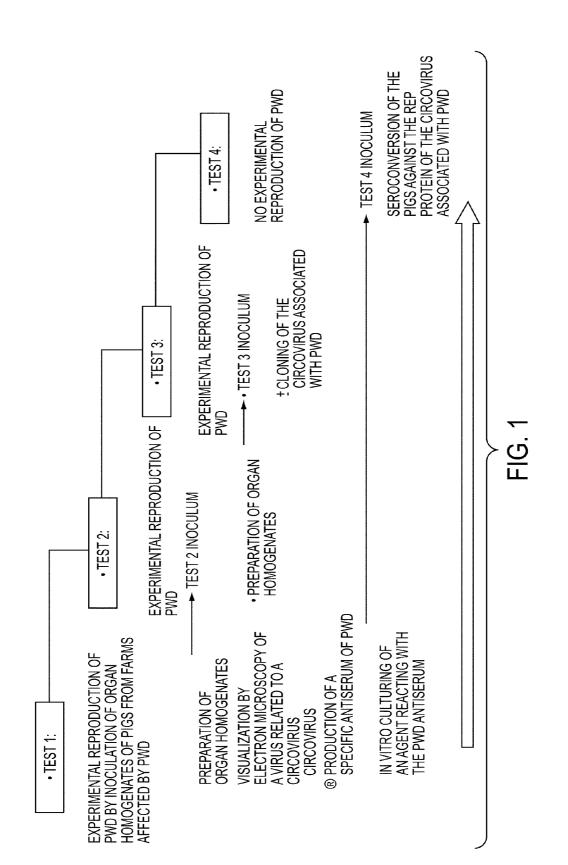
Dec. 5, 1997 (FR) 97/15396

Publication Classification

- (51) Int. Cl. *A61K 39/12* (2006.01)
- (52) U.S. Cl. 424/186.1; 424/204.1

(57) ABSTRACT

The genome sequences and the nucleotide sequences coding for the PWD circovirus polypeptides, such as the circovirus structural and non-structural polypeptides, vectors including the sequences, and cells and animals transformed by the vectors are provided. Methods for detecting the nucleic acids or polypeptides, and kits for diagnosing infection by a PWD circovirus, also are provided. Method for selecting compounds capable of modulating the viral infection are further provided. Pharmaceutical, including vaccine, compositions for preventing and/or treating viral infections caused by PWD circovirus and the use of vectors for preventing and/or treating diseases also are provided.



Jul. 29, 2010 Sheet 2 of 29

Leu Ala Ser Arg Cys Arg Cys Cys Arg Pro Leu Thr Leu Ser Phe Ala Leu Cys Trp Arg Val Glu Ala Ala Ala Ala Gly Arg Cys Arg *** His Phe His Trp Ala Gly Ala Cys Lys Pro Leu Pro Leu Val Glu Ala Ala Asp Thr Phe Ile Gly Leu GCC GTC GTG GAG CCG TCG CAG TCA CTT $\begin{array}{c} 27\\ 45\end{array}$ TGG TCG CGT GAA GCC GTC 31 TTA CGG TTC ACC AGC GCÁ CTT CGG CẮĞ CGG CAG CÃĆ CTC GGC AĞČ GTC AGT GẢĂ AAT GCC AĂĠ 51 Thr Ser Ala Leu Arg Gln Arg Gln His Leu Gly Ser Val Ser Glu Asn Ala Lys Pro Ala His Phe Gly Ser Gly Ser Thr Ser Ala Ala Ser Val Lys Met Pro Ser Gln Arg Thr Ser Ala Ala Ala Ala Pro Arg Gln Arg Gln *** Lys Cys Gln Ala Ser Phe Arg Gly Ala Val Gly Tyr Ser Thr Pro Thr *** Gly *** Tyr Asp Lys Leu Phe Ala Ala Arg Leu Gly Met Leu Pro Pro His Glu Gly Lys Ile Ile Arg Leu Phe Leu Pro Gly Cys Gly Trp Leu Leu His Thr Asn Val Arg Leu Leu Gly GTT CTT TTC GCC GGG CGT IGG GGT ATT 63 CTC CAC CCA CAA GTG GGA ATT ATT AGG 108 CCT CAA GAA AAG CGG CCC GCA ACC CCA TAA GAG GTG GGT GTT CAC TAA TAA TCC Gln Glu Lys Arg Pro Ala Thr Pro *** Glu Val Gly Val His Pro *** *** Ser Lys Lys Ser Gly Pro Gln Pro His Lys Arg Trp Val Phe Thr Leu Asn Asn Pro Arg Lys Ala Ala Arg Asn Pro Ile Arg Gly Gly Cys Ser Pro Leu Ile Ile Leu Arg Pro Pro Ser Phe Cys Phe Val Pro Ala Glu Leu Arg Gly Lys Gln Asn Asn Gly Leu Leu Leu Phe Val Phe Tyr Pro Leu Lys Trp Asp Gly Lys Lys Ile Ile Glu Ser Ser Ser Phe Phe Leu Ile Arg Ser Ser Gly Ile Glu Arg Lys Ser *** GIT TTA TGC CCT CGA AGG TTA GAG GGA AAA ACT 135 144 153 AAT 162 AAG GCT CCT CCT CTT ĒČŤ TTT TTC CGA GGÀ GGA GAA ÀÀÀ CAA AAT ÀČĞ GGA GCT TCC AAT CTC TGA TTA Phe Arg Gly Gly Glu Lys Gln Asn Thr Gly Ala Ser Asn Leu Pro Phe *** Leu Ser Glu Glu Glu Lys Asn Lys Ile Arg Glu Leu Pro Ile Ser Leu Phe Asp Tyr Pro Arg Arg Arg Lys Thr Lys Tyr Gly Ser Phe Gln Ser Pro Phe Leu Ile Ile Gln Lys His Arg Pro Leu Asn Pro Leu Pro Tyr Phe Glu Glu Gly Gly Pro Thr Lys Asn Thr Ala Leu Phe Thr Gln Phe Leu Thr Ser Ser Arg Val Glu Leu Pro Lys Thr Gln Pro Ser Ser Pro Lys Ser Ser Pro Leu Val Gly *** Arg Trp Pro TCC AAA CCT CCT 180 TCT CCC ATC 198 TTG AGG AGT CCC 216 AAA ACA AAC ACC GCT GGA GGT TTT TGT TTG TGG CGA GGA AGG TTT GGA AGA GGG TAG AAC TCC TCA CCT CCA GGG Phe Cys Leu Trp Arg Gly Arg Phe Gly Arg Gly *** Asn Ser Ser Pro Pro Gly Phe Val Cys Gly Glu Glu Gly Leu Glu Glu Gly Arg Thr Pro His Leu Gln Gly Leu Phe Val Ala Arg Lys Val Trp Lys Arg Val Glu Leu Leu Thr Ser Arg Gly Gln Ser Asn Gln *** Ser Ala Ser Lys *** Cys Pro Ser Thr Thr Asn Gln His Lys Arg Ile Lys Ser Leu Leu Leu Ser Lys Val Leu His Leu Pro Ile Lys Thr Asn Ala Phe Lys Ala Leu Phe Cys Val Lys Leu Leu Thr Phe His Tyr Lys Pro CAA ACG CTT AAA ACG ATT CIT CGT CTG AAA ATT GTT CCA CTT CAC CAT AAA ACC 225 234 243 252 261 270 GTT TGC GÃĂ TTT TGC TĂĂ GAA GCA GĂC TTT TAA CAA GGT GAA GTG CTA TTT 'I'GG Val Cys Glu Phe Cys *** Glu Ala Asp Phe *** Gln Gly Glu Val Val Phe Trp Phe Ala Asn Phe Ala Lys Lys Gln Thr Phe Asn Lys Val Lys Trp Tyr Phe Gly Leu Arg Ile Leu Leu Arg Ser Arg Leu Leu Thr Arg *** Ser Gly Ile Leu Val Gly Ser Gly Cys Arg Ser Leu Ser Leu Phe Arg Gly Ala Ser Tyr Leu Ile Ser Gly Ala Ala Val Asp Leu Phe Arg Phe Ser Gly Val Leu Leu Ile Phe Phe Val Ala Arg Gln Trp Met Ser Phe Ala Phe Pro Val Ser Trp Cys Phe Leu Ser Tyr ACG GGC GAC GGT GTA GCT CTT TCG CTT TCC TTG GCT GGT CGT CTT ATT TCT TAT 279 288 297 306 315 324 TGC CCG CTG CCA CAT CGA GAA AGC GAA AGG AAC CGA CCA GCA GAA TAA AGA ATA Cys Pro Leu Pro His Arg Glu Ser Glu Arg Asn Arg Pro Ala Glu *** Arg Ile Ala Arg Cys His Ile Glu Lys Ala Lys GIy Thr Asp Gln Gln Asn Lys Glu Tyr Pro Ala Ala Thr Ser Arg Lys Arg Lys Glu Pro Thr Ser Arg Ile Lys Asn Thr

Cys Tyr Leu Leu Gly Cys Val *** Arg Thr His Leu Glu Ala Ser Gly Pro Ser Ala Thr Phe Phe Ala Val Tyr Lys Asp Leu Thr Ser Ser Arg Pro Val Leu Pro Gln Leu Leu Ser Pro Trp Met Ser Ile Ser His Pro Ala Gly Arg Phe Trp Pro GAC GTC ATT TCT TCC GGT GTA TGA ATA GCT CAC ACC TCG AGG CGC CTT GGT CCC 333 G42 351 360 369 378 CTG CAG TAA AGA AGG CCA CAT ACT TAT CGA GTG TGG AGC TCC GCG GAA CCA GGG Leu Gln *** Arg Arg Pro His Thr Tyr Arg Val Trp Ser Ser Ala Glu Pro Gly Cys Ser Lys Glu Gly His Ile Leu Ile Glu Cys Gly Ala Pro Arg Asn Gln Gly Ala Val Lys Lys Ala Thr Tyr Leu Ser Ser Val Glu Leu Arg Gly Thr Arg Gly

Ala Cys Arg Gly Thr *** Gln Gln Ser Tyr Gly Lys Pro Ser Pro Thr Lys Pro Leu Ala Ala Val Gln Arg Ser Ser His Thr Gly Lys Gln Leu Arg Pro Arg Gln Phe Arg Leu Ser Arg Asp Val Ala Thr Leu Val Arg Lys Ser Val Pro Asp Lys CTT CGC GTC GCT GGA CAG ATG ACG ACA CTC ATG GGA AAA CCT CTG CCC CAG AAA 387 396 405 414 423 432 GAA GCG CAG CGA CCT GTC TAC TGC TGT GAG TAC CCT TTT GGA GAC GGG GTC TTT Glu Ala Gln Arg Pro Val Tyr Cys Cys Glu Tyr Pro Phe Gly Asp Gly Val Phe Lys Arg Ser Asp Leu Ser Thr Ala Val Ser Thr Leu Leu Glu Thr Gly Ser Leu Ser Ala Ala Thr Cys Leu Leu *** Val Pro Phe Trp Arg Gly Leu Trp

Ser Gln Leu Arg Ala Thr Glu Gln Leu Thr His Ser Phe Asn Gly Arg Ala Pro His Ser Tyr Gly Leu Leu Lys Arg Tyr Arg Ile His Ser Ile Glu Ala Pro Gln Thr Val Thr Ala Ser Cys Asn Gly Thr Val Tyr Thr Leu Phe Lys Arg Pro Ser CCA CTG ACA TCG GCT CGT CAA AGG ACA TTG CAT ACA CTC TTT AAA GGC GCC CGA 441 450 459 468 477 486 GGT GAC TGT AGC CGA GCA GTT TCC TGT AAC GTA TGT GAG AAA TTT CCG CGG GCT Gly Asp Cys Ser Arg Ala Val Ser Cys Asn Val Cys Glu Lys Phe Pro Arg Ala Val Thr Val Ala Glu Gln Phe Pro Val Thr Tyr Val Arg Asn Phe Arg Gly Leu *** Leu *** Pro Ser Ser Phe Leu *** Arg Met *** Glu Ile Ser Ala Gly Trp

FIG. 2b

Val Arg *** Leu Pro Gly Ala Arg Asn His Ser Ser Gly Thr Pro Gly Tyr Asn Tyr Val Asp Tyr His Ala Arg Cly Thr Thr Pro Leu Ala Leu Pro Gly Thr Ile Thr Cys Thr Met Thr Pro Gly Gly Pro Gln Pro Phe Leu Trp His Ala Arg Leu ACA TGT GCA GIA TCA CCC GGG CGG GCC AAC ACC CTT CTC GGT CAC CCG GGC ATT 549 558 567 576 585 594 TGT ACA CGT CAT AGT GGG CCC GCC CGG TTG TGG GAA GAG CCA GTG GGC CCG TAA Cvs Thr Arg His Ser Gly Pro Ala Arg Leu Trp Glu Glu Pro Val Gly Pro *** Val His Val Ile Val Gly Pro Pro Gly Cys Gly Lys Ser Gln Trp Ala Arg Asn Tyr Thr Ser *** Trp Ala Arg Pro Val Val Gly Arg Ala Ser Gly Pro Val Ile

Gln Gln Ala *** Pro Cys Arg Ser Ser Ala *** Tyr Phe Tyr Thr Thr Pro His Lys Ser Leu Arg Pro Val Gly Val Pro Leu Arg Thr Ser Ile Leu Pro Pro Ile Lys Ala Ser Gly Leu Ser Val *** Gln Phe Gly Leu Leu Phe Leu His His Ser AAA ACG ACT CGG ATC CCT GTG GAT GAC CTT CGG ATC ATC TTT ATT CAC CAC CCT 603 612 621 630 639 648 TTT TGC TGA GCC TAG GGA CAC CTA CTG GAA GCC TAG TAG AAA TAA GTG GTG GGA Phe Cys *** Ala *** Gly His Leu Leu Glu Ala *** *** Lys *** Val Val Gly Phe Ala Glu Pro Arg Asp Thr Tyr Trp Lys Pro Ser Arg Asn Lys Trp Trp Asp Leu Leu Ser Leu Gly Thr Pro Thr Gly Ser Leu Val Glu Ile Ser Gly Gly Met

Ile Asp His Leu Leu Leu Gln Gln Lys Pro His Asn Lys His Ser Thr Val Lys Ser Ile Met Ser Phe Phe Asn Asn Asn Gln Ile Ile Lys Ile Ala Pro *** Arg Pro Tyr *** Pro Ser Ser Thr Thr Thr Lys Ser Ser Lys *** Pro Gln Asn Gly ACC TAT AGT ACC TCT TCT TCA ACA ACA AAA CCT ACT AAA AAT ACC GAC CAA TGG 657 666 675 684 693 702 TGG ATA TCA TGG AGA AGA AGT TGT TGT TTT GGA TGA TTT TTA TGG CTG GTT ACC Trp Ile Ser Trp Arg Arg Ser Cys Cys Phe Gly *** Phe Leu Trp Leu Val Thr Gly Tyr His Gly Glu Glu Val Val Val Leu Asp Asp Phe Tyr Gly Trp Leu Pro Asp Ile Met Glu Lys Lys Leu Leu Phe Trp Met Ile Phe Met Ala Gly Tyr Leu

Pro His Asp Val Ser Val Thr His Gly Thr Asp Met Ser Gln Leu Ser *** Leu Pro Ile Ile *** Gln Ser Gln Thr Val Pro Ile Trp Gln Ser Tyr Leu Ser Phe Gln Ser Ser Arg Ser Leu Ser His Ser Arg Tyr Gly Asn Val Thr Ser Val Leu AAC CCT ACT AGA TGA CTC TGA CAC ACT GGC CAT AGG TAA CTG ACA TCT CTG ATT 711 720 729 738 747 756 TTG GGA TGA TCT ACT GAG ACT GTG TGA CCG GTA TCC ATT GAC TGT AGA GAC TAA Leu Gly *** Ser Thr Glu Thr Val *** Pro Val Ser Ile Asp Cys Arg Asp *** Trp Asp Asp Leu Leu Arg Leu Cys Asp Arg Tyr Pro Leu Thr Val Glu Thr Lys Gly Met Ile Tyr *** Asp Cys Val Thr Gly Ile His *** Leu *** Arg Leu Lys

Pro Tyr Gln Glu Lys Lys Pro Gly Cys Tyr Lys Ser *** Trp Cys Asp Pro Gly Pro Thr Ser Asn Arg Lys Gln Gly Ala Thr Asn Gln Asn Gly Ala Ile Leu Gly Pro Pro Val Thr Gly Lys Lys Ala Arg Leu Ile Lys Ile Val Leu Leu *** Ala TCC CCC ATG ACA AGG AAA AAA CCG GGC GTC ATA AAA CTA ATG GTC GTT AGT CCG 765 774 783 792 801 810 AGG GGG TAC TGT TCC TTT TTT GGC CCG CAG TAT TTT GAT TAC CAG CAA TCA GGC Arg Gly Tyr Cys Ser Phe Phe Gly Pro Gln Tyr Phe Asp Tyr Gln Gln Ser Gly GIy Gly Thr Val Pro Phe Leu Ala Arg Ser Ile Leu Ile Thr Ser Asn Gln Ala GIy Val Leu Phe Leu Phe Trp Pro Ala Val Phe *** Leu Pro Ala Ile Arg Pro

FIG. 2c

Gly Pro Ile Thr Ser Arg Leu Gln Gln Gly Leu Gln Leu Leu Glu Arg Asp Ser G_y Leu Phe Pro Val Gly *** Ser Ser Asp Trp Ser Tyr Phe Ser Glu Ile Pro Gly Trp Ser His Tyr Glu Glu Val Ala Thr Gly Ala Thr Ser Ala Arg *** Arg GGG GGT CCT TAC CAT GAG GAG TTG ACG ACA GGG TCG ACA TCT TCG AGA GAT AGC 819 CCC CCA GGA ATG GTA CTC CTC AAC TGC IGT CCC AGC TGT AGA AGC TCT CTA TCG Pro Pro Gly Met Val Leu Leu Asn Cys Cys Pro Ser Cys Arg Ser Ser Leu Ser Pro Gln Glu Trp Tyr Ser Ser Thr Ala Val Pro Ala Val Glu Ala Leu Tyr Arg Pro Arg Asn Gly Thr Pro Gln Leu Leu Ser Gln Leu *** Lys Leu Ser Ile Gly

Ser *** *** Lys Ala Ile Lys Ser Ser Gln Gln Leu Val Ile Trp Pro Pro Val Pro Asn Ser Ser Gln Leu Lys Pro Leu Ser Ser Ser Phe Leu Gly Arg Leu Tyr Leu Ile Val Val Lys Cys Asn Gln Phe Val Ala Pro Ser Cys Asp Val Ser Thr CTC CTA ATG ATG AAA CGT TAA AAC CTT CTG ACG ACC TCT TGT TAG GTG CCT CCA 873 GAG GAT TAC TAC TTT GCA ATT TTG GAA GAC TGC TGG AGA ACA ATC CAC GGA GGT Glu Asp Tyr Tyr Phe Ala Ile Leu Glu Asp Cys Trp Arg Thr Ile His Gly Gly Arg Ile Thr Thr Leu Gln Phe Trp Lys Thr Ala Gly Glu Gln Ser Thr Glu Val Gly Leu Leu Cys Asn Phe Gly Arg Leu Leu Glu Asn Asn Pro Arg Arg Tyr

Arg Leu Gly Ile Gln Leu Leu Pro Gly Val Arg His Gly Lys Gly Met Tyr Phe GLy Phe Ala Ser Lys Phe Cys His Val Trp Gly Thr Gly Lys Glu Trp Ile Phe Gly Ser Pro Arg Asn Ser Ala Thr Ser Gly Gly Gln Ala Arg Lys Gly Tyr Leu TGG GCT TCC GGC TAA ACT TCG TCA CCT GGG TGG GAC ACG GGA AAA GGG TAT ATT 927 936 945 954 954 963 972 ACC CGA AGG CCG ATT TGA AGC AGT GGA CCC ACC CTG TGC CCT TTT CCC ATA TAA Thr Arg Arg Pro Ile *** Ser Ser Gly Pro Thr Leu Cys Pro Phe Pro Ile *** Pro Glu Gly Arg Phe Glu Ala Val Asp Pro Pro Cys Ala Leu Phe Pro Tyr Lys Pro Lys Ala Asp Leu Lys Gln Trp Thr His Pro Val Pro Phe Ser His Ile Lys

Leu Asn Ser Leu Arg Lys Gln *** *** Met Thr Ile Thr Lys Ile Lys Ile *** Tyr Ile Val Ser Asp Lys Lys Asn Asp Cys Arg Leu Pro Lys *** Lys *** Glu Ile Phe *** Gln Thr Lys Lys Thr Ile Val Asp Tyr His Asn Lys Asn Lys Asn TTA TTT AAT GAC TCA GAA AAA ACA ATA GTG TAG CAT TAC CAA AAA TAA AAA TAA 981 990 999 1008 1017 1026 AAT AAA TTA CTG AGT CTT TTT TGT TAT CAC ATC GTA ATG GTT TTT ATT ITT ATT Asn Lys Leu Ieu Ser Leu Phe Cys Tyr His Ile Val Met Val Phe Ile Phe Ile I.e Asn Tyr *** Val Phe Phe Val Ile Thr Ser *** Trp Phe Leu Phe Leu Phe *** Ile Thr Glu Ser Phe Leu Leu Ser His Arg Asn Gly Phe Tyr Phe Tyr Ser

Lys Ser Pro Arg Glu Pro Tyr Ile Arg Gln Ile Thr Cys Leu Tyr Asp Val Lys Asn Leu Pro Asp Lys Leu Ile Phe Glu Arg Phe Gln Val Tyr Ile Thr Leu Arg Met *** Leu Thr Lys *** Ser Leu Asn Glu Ser Asn Tyr Met Phe Leu *** Gly GTA AAT CTC CCA GAA AGT CCT ATT TAA GAG ACT TAA CAT GTA TTT ATC AGT TGG 1035 1044 1053 1062 1071 1080 CAT TTA GAG GGT CTT TCA GGA TAA ATT CTC TGA ATT GTA CAT AAA TAG TCA ACC His Leu Glu Gly Leu Ser Gly *** Ile Leu *** Ile Val His Lys *** Ser Thr I e *** Arg Val Phe Gln Asp Lys Phe Ser Glu Leu Tyr Ile Asn Ser Gln Pro Phe Arg Gly Ser Phe Arg Ile Asn Ser Leu Asn Cys Thr *** Ile Val Asn Leu

FIG. 2d

Gly Cys Leu Lys Pro Ser His Asn Cys Lys Pro Ala Cys Leu Gly Pro Arg His Val Val Tyr Asn Gln Ala Thr Thr Ala Asn Gln Leu Ala Tyr Gly Leu Gly Thr *** Trp Met IIe Lys Pro Gln Pro Gln Met Lys Ser Arg Met Ala Trp Ala Gln AAT GGT GTA TTA AAA CCC GAC ACC AAC GTA AAA CCT CGC GTA TCG GGT CCG GAC 1089 1098 1098 107 1116 1125 1134 TTA CCA CAT AAT TTT GGG CTG TGG TTG CAT TTT GGA GCG CAT AGC CCA GGC CTG Leu Pro His Asn Phe Gly Leu Trp Leu His Phe Gly Ala His Ser Pro Gly Leu Tyr His IIe IIe Leu Gly Cys Gly Cys IIe Leu Glu Arg IIe Ala Gln Ala Cys Thr Thr *** Phe Trp Ala Val Val Ala Phe Trp Ser Ala *** Pro Arg Pro Val Ala Arg Cys Gln His Pro Tyr Lys Phe Pro Ala Val Ala Pro Lys Lys *** *** His Glu Val Asn Thr His Thr Ash Leu His Leu Trp Leu Gln Ash Arg Lys Asn Thr Ser Ser Met Pro Thr Pro IIe *** IIe Ser Gly Cys Ser Thr G.u Lys IIe ACA CGA GCT GTA ACC ACA CCC ATA AAT TTA CCT CGG TGT CGA CCA AAG AAA ATA 1143 1152 1161 1170 1179 1188 TCT GCT CGA CAT TGG TGT GGG TAT TTA AAT GGA GCC ACA GCT GGT TIC TTT TAT Cys Ala Arg His Trp Cys Gly Tyr Leu Asn Cly Ala Thr Ala Gly Phe Phe Tyr Val Leu Asp IIe Gly Val Gly IIe *** Met Glu Pro Gln Leu Val Ser Phe IIe Cys Ser Thr Leu Val Trp Val Phe Lys Trp Ser His Ser Trp Phe Leu Leu Leu Lys Ala Pro Val Leu *** Asn Asn Pro Arg Ala Arg Thr Gln Pro His Leu Val Asn Pro Gln Phe Trp Asp IIe Thr Gln Asp Leu Glu Pro Lys Pro Thr Phe Tyr IIe Gln Ser Ser Gly IIe Leu Grn Lys Thr *** Ser Gln Asn Pro Pro Ser Thr ATT AAC CGA CCT TGG TTA GTT AAC AAA CCA CAT CGA GAC CAA ACC CCC ACT TCA 1197 1206 1215 1224 1233 1224 TAT TTG GCT GGA ACC AAT CAA TTG TTT GGT CTA GCT CTG GTT TGG GGG TGA AGT

TAT TTG GCT GGA ACC AAT CAA TTG TTT GGT CTA GCT CTG GTT TGG GGG TGA AGT Tyr Leu Ala Gly Thr Asn Gln Leu Phe Gly Leu Ala Leu Val Trp Gly *** Ser Ile Trp Leu Glu Pro Ile Asn Cys Leu Val *** Leu Trp Phe Gly Gly Gly Val Phe Gly Trp Asn Gln Ser Ile Val Trp Ser Ser Ser Gly Leu Gly Val Lys Tyr Gln Leu Pro Leu Tyr Leu Ala Ala Lys His His Pro Pro Leu Leu Leu *** Tyr Arg Ser His Tyr Thr Phe Pro Gln Arg Ile Thr His Arg Ser Ser Tyr Asn Ile Gly Pro Thr Thr Pro Leu Pro Ser Gly *** Pro Thr Ala Pro Pro Thr Thr Leu TGG ACC TCA CCA TCC ATT TCC CGA CGG AAT ACC ACA CCG CCC TCC TCA TCA ATT 1251 1260 1269 1278 1287 1296 ACC TGG AGT GGT AGG TAA AGG GCT GCC TTA TGG TGT GGC GGG AGG AGT AGT TAA Thr Trp Ser Gly Arg *** Arg Ala Ala Leu Trp Cys Gly Gly Arg Ser Ser *** Pro Gly Val Val Gly Lys Gly Leu Pro Tyr Gly Val Ala Gly GIy Val Val Asn Leu Glu Trp *** Val Lys Gly Cys Leu Met Val Trp Arg Glu Glu *** Leu Ile

Leu Pro *** Leu Gly Leu Gln His Leu Pro Asn Cys Leu Gln Cys Gly Leu Tyr Tyr Pro Asp Tyr Ala Leu Asn Thr Ser Pro Thr Val Phe Asn Ala Asp Leu Ile Ile Pro Thr Met Pro Tro Thr Pro Pro Pro Pro *** Leu Thr Pro Met Trp Ser ATA TCC CCA GTA TCC GGT TCA ACC ACC TCC CCC AAT GTT TCA ACC GTA GGT TCT 1305 1314 1322 1341 1350 TAT AGG GGT CAT AGG CCA AGT TGG TGG AGG GGG TTA CAA AGT TGG CAT CCA AGA Tyr Arg Gly His Arg Pro Ser Trp Trp Arg Cly Leu Gln Ser Trp His Pro Arg Ile GIy Val Ile GIy Gln Val Gly Cly GIy GIy Tyr Lys Val Gly Ile Gln Asp *** Gly Ser *** Ala Lys Leu Val Glu Gly Val Thr Lys Leu Ala Ser Lys Ile

FIG. 2e

Cys Cys His Val Trp Cys Arg Lys Ser *** Leu His His Pro Arg Gln Pro Leu Val Val Thr Ser Gly Val Gly Arg Gln Asn Ser Thr Ile Pro Asp Arg Pro Tyr Leu Leu Pro Gly Leu Val Glu Lys Ile Leu Pro Ser Pro Thr Glu Pro Thr ATT GTT GTC ACC TGG GTT GTG GAG AAA CTA ATC TCC ACT ACC CCA GAG ACC CCA 1359 1368 1377 1386 1395 1404 TAA CAA CAG TGG ACC CAA CAC CTC TTT GAT TAG AGG TGA TGG GGT CTC TGG GGT ATT GTT GTC 1359 ___ ___ ___ __ Gln Gln Trp Thr Gln His Leu Phe Asp *** Arg *** Trp Gly Leu Trp Gly Asn Asn Ser Gly Pro Asn Thr Ser Leu Ile Arg Gly Asp Gly Val Ser Gly Val Thr Thr Val Asp Pro Thr Pro Leu *** Leu Glu Val Met Gly Ser Leu Gly ***

Ile *** Ile *** Gly Lys *** Tyr Pro Leu Ile Pro Phe Thr Pro Thr Pro Pro Phe Glu Tyr Lys Ala Lys Arg Ile Arg Tyr Tyr Gln Phe Pro Leu Pro Leu Pro Phe Asn Met Asn Leu Arg Glu Leu Val Thr Thr Asn Ser Leu Tyr Pro Tyr Pro TTT TAA GTA TAA ATC GGA AAG ATT AFG CCA TCA TAA CCT TTC CAT CCC CAT CCC $^{1413}_{1422}$ $^{1431}_{1431}$ aaa att cat att tag cct ttc taa tac ggt agt att gga aag gta ggg gta ggg ------_ _ _ Lys Ile His Ile *** Pro Phe *** Tyr Gly Ser Ile Gly Lys Val Gly Val Gly Lys Phe Ile Phe Ser Leu Ser Asn Thr Val Val Leu Glu Arg *** Gly *** Gly Asn Ser Tyr Leu Ala Phe Leu Ile Arg *** Tyr Trp Lys Gly Arg Gly Arg Gly

Gln His Arg Arg Leu Pro Pro Pro Val Pro Arg His Gln Ile Glu Ala Arg *** Asn Thr Gly Gly Ser Pro Pro Leu Phe Gln Gly Ile Asn Phe Arg Leu Glu Asn Thr Pro Ala Ala Gln Pro Pro Ser Ser Ser Ala Ser Thr Ser Asp *** Ser Thr CCA ACC ACG GCG GAC TCC CCC CCT CCT TGA CCG GCT ACA ACT TAG AGT CGA GCA 1467 1476 1485 1485 1494 1503 1512 GCT TGG TGC CGC CTG AGG GGG GGA GGA ACT GGC CGA TGT TGA ATC TCA GCT CGT Gly Trp Cys Arg Leu Arg Gly Gly Gly Thr Gly Arg Cys *** Ile Ser Ala Arg Val Gly Ala Ala *** Gly Gly Glu Glu Leu Ala Asp Val Glu Ser Gln Leu Val Leu Val Pro Pro Glu Gly Gly Arg Asn Trp Pro Met Leu Asn Leu Ser Ser Leu

Cys Glu Leu Ile Ala Ala Leu Thr Arg Arg Lys His His Thr Cys Ile Arg *** Val Asn Trp Ser Pro Gln Ser His Gly Gly Arg Ile Thr Leu Val Phe Glu Arg Leu Met Gly Leu His Ser Arg Thr Asp Glu Glu *** Pro Ser Tyr Leu Asn Glu ATT STA AGG TTC TAC CGA CGC TCA CAG GAG GAG AAT ACC ACT CAT GTT TAA GAG 1521 1530 1539 1548 1557 1566 TAA CAT TCC AAG ATG GCT GCG AGT GTC CTC CTC TTA TGG TGA GTA CAA ATT CTC *** His Ser Lys Met Ala Ala Ser Val Leu Leu Leu Trp *** Val Gln Ile Leu Asn Ile Pro Arg Trp Leu Arg Val Ser Ser Ser Tyr Gly Glu Tyr Lys Phe Ser Thr Phe Gln Asp Gly Cys Glu Cys Pro Pro Leu Met Val Ser Thr Asn Ser Leu

Phe Pro Pro Phe Gln Leu Tyr Gly Asp Lys Pro Ala Met Gln Leu Pro Lys Gln Ser Leu Arg Ser Asn Phe Ile G_y Thr Lys Arg Arg Trp Arg Tyr Arg Asn Arg Leu Phe Ala Pro Ile Ser Ser Va_ Arg Arg Glu Ala Gly Asp Thr Val Thr Glu ATC TTT CCG CCC TTA ACT TCT ATG GGC AGA AAG CCG CGG TAG ACA TTG CCA AAG 1575 1584 1593 1602 1611 1620 TAG AAA GGC GGG AAT TGA AGA TAC CCG TCT TTC GGC GCC ATC TGT AAC GGT TTC *** Lys Gly Gly Asn *** Arg Tyr Pro Ser Phe Gly Ala Ile Cys Asn Gly Phe Arg Lys Ala Gly Ile Glu Asp Thr Arg Leu Ser Ala Pro Ser Val Thr Val Ser Glu Arg Arg Glu Leu Lys Ile Pro Val Phe Arg Arg His Leu *** Arg Phe Leu Leu Arg Pro Thr Gly Phe Ile Thr Lys Glu Pro Pro His Lys Trp Ser Pro Gln Phe Ala Pro His Val Leu Tyr Pro Arg Arg Arg Leu Ile Asn Gly Leu His Ser Ser Pro Pro Thr Tyr Trp Ile His Asp Glu Gly Ser Ser Thr Glu Leu Ile Ala ACT TCC GCC CCA CAT GGT TTA TAC CAG AAG AGG CCT CCT ACA AAG GTT CTA CCG 1629 1638 1647 1656 1665 1674 TGA AGG CGG GGT GTA CCA AAT ATG GTC TTC TCC GGA GGA TGT TTC CAA GAT GGC *** Arg Arg Gly Val Pro Asn Met Val Phe Ser Gly Gly Cys Phe Gln Asp Gly Glu Gly Gly Val Tyr Gln Ile Trp Ser Ser Pro Glu Asp Val Ser Lys Met Ala Lys Ala Gly Cys Thr Lys Tyr Gly Leu Leu Arg Arg Met Phe Pro Arg Trp Leu

Pro Pro Arg Thr Arg Arg Arg Arg Tyr Arg Arg Arg Pro Trp Thr Met Arg Tyr Ala Pro Ala Pro Gly Asp Glu Ala Thr Val Gly Gly Gln Gly Arg *** Gly Ile ACG CCC CCG CCC AGG CAG AAG ACG CCA TTG CGC AGG AAC CGG TGC AGT AGG ATA 1683 1692 1701 1710 1719 1728 TGC GGG GGC GGG TCC GTC TTC TGC GGT AAC GCC TCC TTG GCC ACG TCA TCC TAT Cys Gly Gly Ser Val Phe Cys Gly Asn Ala Ser Leu Ala Thr Ser Ser Tyr Ala Gly Ala Gly Pro Ser Ser Ala Val Thr Pro Pro Trp Pro Arg His Prc Ile Arg Gly Arg Val Arg Leu Leu Arg *** Arg Leu Leu Gly His Val Ile Leu ***

Leu Ser Leu Leu Ala Ser Ser Tyr Tyr Phe His Phe Phe His Ala Ala Thr Thr Asn Phe Thr Phe Ser Thr Arg Gln Gln Leu Ile TTT TCA CTT TCT TCA CGC GAC GAC ATC ATA A 5' 1737 1746 1755 AAA AGT GAA AGA AGT GCG CTG CTG TAG TAT T 3' Lys Ser Glu Arg Ser Ala Leu Leu *** Tyr Lys Val Lys Glu Val Arg Cys Cys Ser Ile Lys *** _ys Lys Cys Cys Ala Ala Val Val

FIG. 2g

circopormank circopormeeh circopordfp	10 20 30 40 50 1 ACCAGCGCAC TTCGGCAGCG GCAGCACCTC GGCAGCGTCA GTGAAAATGC 1 ACCAGCGCAC TTCGGCAGCG GCAGCACCTC GGCAGCGTCA GTGAAAATGC 1 ACCAGCGCAC TTCGGCAGCG GCAGCACCTC GGCAGCGTCA GTGAAAATGC	50 50
circopormank circopormeeh circopordfp	60 70 80 90 100 51 CAAGCAACAA AAGCGGCCCG CAACCCCATA AGAGGTGGGT GTTCACCCTT 51 CAAGCAACAA AAGCGGCCCG CAACCCCATA AGAGGTGGGT GTTCACCCTT 51 CAAGCAACAA AAGCGGCCCG CAACCCCATA AGAGGTGGGT GTTCACCCTT	100 100 100
circopormank circopormeeh circopordfp	110 120 130 140 150 101 AATAATCCTT CCGAGGAGGA GAAAAACAAAI ATACGGGAGC TTCCAATCTC 101 AATAATCCTT CCGAGGAGGA GAAAAACAAAI ATACGGGAGC TTCCAATCTC 101 AATAATCCTT CCGAGGAGGA GAAAAACAAAI ATACGGGAGC TTCCAATCTC	150 150 150
circopormank circopormeeh circopordfp	160 170 180 190 200 151 [CCTTTTCAT] [TATTTTGTTT] [GCGGAGAGGA] AGGTTTGCAA] [GAGGGTAGAA 151 [CCTTTTCAT] [TATTTTGTTT] [GCGGAGAGA] AGGTTTGCAA] [GAGGGTAGAA 151 [CCTTTTCAT] [TATTTTGTTT] [GTGGCGAGGA] AGGTTTGCAA] [GAGGGTAGAA]	200 200 200
circopormank circopormeeh circopordfp	210 220 230 240 250 201 [CTGCICACCT] [CCAGGGGTTT] [GCEAATTTTG] [CTAAGAAGCA] [GACTTTTAAC] 201 [CTCCICACCT] [CCAGGGGTTT] [GCGAATTTTG] [CTAAGAAGCA] [GACTTTTAAC] 201 [CTCCICACCT] [CCAGGGGTTT] [GCGAATTTTG] [CTAAGAAGCA] [GACTTTTAAC]	250 250 250
circopormank circopormeeh circopordfp	260 270 280 290 300 251 [AAGGIGAAGT] [GGTATTTTGG] [TGCCCGCTGC] [CACATCGAGA] [AAGCGAAAGG] 251 [AAGGIGAAGT] [GGTATTTTGG] [TGCCCGCTGC] [CACATCGAGA] [AAGCGAAAGG] 251 [AAGGIGAAGT] [GGTATTTTGG] [TGCCCGCTGC] [CACATCGAGA] [AAGCGAAAGG]	300 300 300
circopormank circopormeeh circopordfp	310 320 330 340 350 301 AACCCACCAG CAGAATAAAG AATACTGCAG TAAAGAAGGC CACATACTTA 301 AACCGACCAG CAGAATAAAG AATACTGCAG TAAAGAAGGC CACATACTTA 301 AACCGACCAG CAGAATAAAG AATACTGCAG TAAAGAAGGC CACATACTTA	350 350 350
circopormank circopormeeh circopordfp	360 370 380 390 400 351 TCGACTGTGG AGCTCCCCCGG AACCAGGGGA AGCGCAGCGA CCTGTCTACT 351 TCGAGTGTGG AGCTCCCCCGG AACCAGGGGA AGCGCAGCGA CCTGTCTACT 351 TCGAGTGTGG AGCTCCCCCGG AACCAGGGGA AGCGCAGCGA CCTGTCTACT	400 400 400
circopormank circopormeeh circopordfp	410 420 430 440 450 401 GCTGIGAETA CCCTTTTGGA GACGGGGTCT TTGGTGAETG TAGCCGAGCA 401 GCTGIGAETA CCCTTTTGGA GACGGGGTCT TTGGTGAETG TAGCCGAGCA 401 GCTGIGAETA CCCTTTTGGA GACGGGGTCT TTGGTGAETG TAGCCGAGCA	450 450 450
circopormank circopormeeh circopordfp	460 470 480 490 500 451 GTTCCCTGTA ACGTATGTGA GAAATTTCCG CGGGCTGGCT GAACTTTTGA 451 GTTCCCTGTA ACGTATGTGA GAAATTTCCG CGGGCTGCT GAACTTTTGA 451 GTTTCCTGTA ACGTATGTGA GAAATTTCCG CGGGCTGCCT GAACTTTTGA	500 500 500
circopormank circopormeeh circopordfp	510 520 530 540 550 501 AAGTGAGCGG GAAGATGCAG CAGCGTGATT GGAAGACAGC TGTACACGTC 501 AAGTGAGCGG GAAGATGCAG CAGCGTGATT GGAAGACAGC TGTACACGTC 501 AAGTGAGCGG GAAGATGCAG CAGCGTGATT GGAAGACAGC TGTACACGTC	550 550 550
circopormank circopormeeh circopordfp	560 570 580 590 600 551 [ATAGIGGECC] [CGCCCGETTG] [TGGGAAGAGC] [CAGTGGGCCC] [GTAATTTTGC] 551 [ATAGIGGECC] [CGCCCGETTG] [TGGGAAGAGC] [CAGTGGGCCC] [GTAATTTTGC] 551 [ATAGIGGECC] [CGCCCGETTG] [TGGGAAGAGC] [CAGTGGGCCC] [GTAATTTTGC]	600 600 600

FIG. 3a

circopormank circopormeeh circopordfp	601 TGAGCCTAGC GACACCTACT GGAAGCCTAG TAGAAATAAC TGGTGGGATG 601 TGAGCCTAGG GACACCTACT GGAAGCCTAG TAGAAATAAC TGGTGGGATG 601 TGAGCCTAGG GACACCTACT GGAAGCCTAG TAGAAATAAC TGGTGGGATG 601 TGAGCCTAGG GACACCTACT GGAAGCCTAG TAGAAATAAC TGGTGGGATG	650 650 650
circopormank circopormech circopordfp	660 670 680 700 651 [GATATCATGG] AGAAGAAGTT] [GTTGTTTTGG] [ATGATTTTTA] [TGACTGGTTA] 651 [GATATCATGG] AGAAGAAGTT] [GTTGTTTTGG] [ATGATTTTTA] [TGACTGGTTA] 651 [GATATCATGG] AGAAGAAGTT] [GTTGTTTTGG] [ATGATTTTTA] [TGACTGGTTA]	700 700 700
circopormank circopormeeh circopordfp	710 720 730 740 750 701 [CCTTGGGATG ATCTACTGAG ACTGTGTGAC CGGTATCCAT TGACTGTAGA 701 [CCTTGGGATG ATCTACTGAG ACTGTGTGAC CGGTATCCAT TGACTGTAGA 701 [CCTTGGGATG ATCTACTGAG ACTGTGTGGAC CGGTATCCAT TGACTGTAGA	750 750 750
circopormank circopormeeh circopordfp	760 770 780 790 800 751 GACTAAAGCC GGTACTGTTC CTTTTTTGGC TCGCAGTATT TTGATTACCA 751 GACTAAAGGG GGTACTGTTC CTTTTTTGGC CCGCAGTATT TTGATTACCA 751 GACTAAAGGG GGTACTGTTC CTTTTTTGGC CCGCAGTATT TTGATTACCA	800 800 800
circopormank circopormeeh circopordfp	810 801 [GCAATCAGGC] [CCCCCAGGAA] [TGGTACTCCT] [CAACTGCTGT] [CCCAGCTGTA] 801 [GCAATCAGGC] [CCCCCAGGAA] [TGGTACTCCT] [CAACTGCTGT] [CCCAGCTGTA] 801 [GCAATCAGGC] [CCCCCAGGAA] [TGGTACTCCT] [CAACTGCTGT] [CCCAGCTGTA]	850 850 850
circopormank circopormeeh circopordfp	860 87C 880 9C0 851 GAAGCTCTCT ATCGGAGGAT TACTACTTTG CAATTTTGGA AGACTGCTGG 851 GAAGCTCICT ATCGGAGGAT TACTACTTTG CAATTTTGGA AGACTGCTGG 851 GAAGCTCTCT ATCGGAGGAT TACTACTTTG CAATTTTGGA AGACTGCTGG	900 900 900
circopormank circopormeeh circopordfp	910 901 AGAACAATCA ACGGAGGTAC CCGAAGGCCG ATTTGAAGCA GTGGACCCAC 901 AGAACAATCC ACGGAGGTAC CCGAAGGCCG ATTTGAAGCA GTGGACCCAC 901 AGAACAATCC ACGGAGGTAC CCGAAGGCCG ATTTGAAGCA GTGGACCCAC	950 950 950
circopormank circopormeeh circopordfp	960 970 980 990 1000 951 [CCTGTGCCCT] [TTTCCCATAT] [AAAATAAATT] [ACTGAGTCTT] [TTTTGTTATC] 951 [CCTGTGCCCT] [TTTCCCATAT] [AAAATAAATT] [ACTGAGTCTT] [TTTTGTTATC] 951 [CCTGTGCCCT] [TTTCCCATAT] [AAAATAAATT] [ACTGAGTCTT] [TTTTGTTATC]	$1000 \\ 1000 \\ 1000 \\ 1000$
circopormank circopormeeh circopordfp	1010 1020 1030 1040 1050 1001 ACATCGTAAT GGTTTTTATTI (TTTATTTATT) (TAGAGGGTCI) (TTTAGGATAA 1001 ACATCGTAAT GGTTTTTATTI (TTTATTTATT) (TAGAGGGTCI (TTTAGGATAA 1001 ACATCGTAAT) (GGTTTTTATT) (TTTATTCATT) (TAGAGGGTCI (TTCAGGATAA	1050 1050 1050
circopormank circopormeeh circopordfp	1060 107C 1080 1090 11C0 1051 ATTCTCTCAA TTGTACATAA ATAGTCAGCC TTACCACATA ATTTTGGGCT 1051 ATTCTCTGAA TTGTACATAA ATAGTCAGCC TTACCACATA ATTTTGGGCT 1051 ATTCTCTGAA TTGTACATAA ATAGTCAACC TTACCACATA ATTTTGGGCT	1100 1100 1100
circopormank circopormeeh circopordfp	1110 1120 1130 1140 1150 1101 GTEGCTGCAT TTTGGAGCGC ATAGCCGAGG CCTGTGTGCT CGACATTGGT 1101 GTEGCTGCAT TTTGGAGCGC ATAGCCGAGG CCTGTGTGCT CGACATTGGT 1101 GTEGTTGCAT TTTGGAGCGC ATAGCCCAGG CCTGTGTGCT CGACATTGGT	1150 1150 1150
circopormank circopormech circopordfp	1160 1170 1180 1200 1151 GTGGGTATTTI AAATGGAGCCI ACAGCTGGTTI TCTTTTATTA TTTGGGTGGA 1151 GTGGGTATTTI AAATGGAGCCI ACAGCTGGTTI TCTTTATTA TTTCGGTGGA 1151 GTGGGTATTTI AAATGGAGCCI ACAGCTGGTTI TCTTTTATTA TTTGGCTGGA	1200 1200 1200

FIG. 3b

circopormank circopormeeh circopordfp	1210 1220 1230 1240 1250 1201 ACCANTCAAT TGTTTGGTCC AGCTCAGGTT TGGGGGGTGAA CTACCTGGAG 1201 ACCAATCAAT TGTTTGGTCC AGCTCAGGTT TGGGGGGTGAA CTACCTGGAG 1201 ACCAATCAAT TGTTTGGTCT AGCTCTGGTT TGGGGGGTGAA CTACCTGGAG	1250 1250 1250
circopormank circopormeeh circopordfp	1251 TGGTAGSTAA AGGGCTGCCT TATGGTGTGG CGGGAGCAGT AGTTAATATA 1251 TGGTAGSTAA AGGGCTGCCT TATGGTGTGG CGGGAGCAGT AGTTAATATA 1251 TGGTAGSTAA AGGGCTGCCT TATGGTGTGG CGGGAGCAGT AGTTAATATA	1300 1300 1300
circopormank circopormeeh circopordfp	1310 1320 1330 1340 1350 1301 GGGGTCATAG GCCAAGTTGG TGGAGGGGGGT TACAAACTTG CATCCAAGA 1301 GGGGTCATAG GCCAAGTTGG TGGAGGGGGGT TACAAACTTG CATCCAAGA 1301 GGGGTCATAG GCCAAGTTGG TGGAGGGGGGT TACAAACTTG CATCCAAGA	1350 1350 1350
circopormank circopormeeh circopordfp	1360 1370 1380 1390 1400 1351 TAACAACAGT GGACCCAACA CCTCTTTCAT TAGAGGTGAT GGGGTCTCTG 1351 TAACAACAGT GGACCCAACA CCTCTTTGAT TAGAGGTGAT GGGGTCTCTG 1351 TAACAACAGT GGACCCAACA CCTCTTTGAT TAGAGGTGAT GGGGTCTCTG	$1400 \\ 1400 \\ 1400 \\ 1400$
circepormank circepormeeh circepordfp	1401 GGGTAAAATT CATATTTAGC CTTTCTAATA CGGTAGTATT GGAAAGGTAG 1401 GGGTAAAATT CATATTTAGC CTTTCTAATA CGGTAGTATT GGAAAGGTAG 1401 GGGTAAAATT CATATTTAGC CTTTCTAATA CGGTAGTATT GGAAAGGTAG 1401 GGGTAAAATT	$ \begin{array}{r} 1450 \\ 1450 \\ 1450 \\ 1450 \\ \end{array} $
circopormank circopormeeh circopordfp	1451 GGGTAGGGGG TTGGTGCCGC CTGAGGGGGGG GAGGAACTGG CCGATGTTGA 1451 GGGTAGGGGG TTGGTGCCGC CTGAGGGGGGG GAGGAACTGG CCGATGTTGA 1451 GGGTAGGGGG TTGGTGCCGC CTGAGGGGGGG GAGGAACTGG CCGATGTTGA 1451 GGGTAGGGGG TTGGTGCCGC CTGAGGGGGGG GAGGAACTGG CCGATGTTGA	$1500 \\ 1000 \\ 1000 \\ $
circopormank circopormeeh circopordfp	1510 1520 1530 1540 1550 1501 AFCTGAGGTG GTTAACATGC CAAGATGGCT GCGAGTATCC TCCTTTATG 1501 ATCTGAGGTA GTTAACATTC CAAGATGGCT GCGAGTATCC TCCTTTATG 1501 ATCTGAGOTC GTTAACATTC CAAGATGGCT GCGAGTCCC TCCTCTTATG	1550 1550 1550
circopormank circopormeeh circopordfp	1551 GIGAGTACAA ATTCTCTAGA AAGGCGGGGAA TCGAAGATAC CCGTCTTTCG 1551 GIGAGTACAA ATTCTCTAGA AAGGCGGGGAA TCGAAGATAC CCGTCTTTCG 1551 GIGAGTACAA ATTCTCTAGA AAGGCGGGAA TCGAAGATAC CCGICTTTCG 1551 GIGAGTACAA ATTCTCTAGA AAGGCGGGAA TCGAAGATAC CCGICTTTCG	$1600 \\ 1600 \\ 1600 \\ 1600$
circopormank circopormeeh circopordfp	1610 1620 1630 1640 1650 1601 GCGCCATCTG TAACGGTTTC TGAAGGCGGG GTGTGCCAAA TATGGTCTTC 1601 GCGCCATCTG TAACGGTTTC TGAAGGCGGG GTGTGCCAAA TATGGTCTTC 1601 GCGCCATCTG TAACGGTTTC TGAAGGCGGG GTGTACCAAA TATGGTCTTC	1650 1650 1650
circopormank circopormeeh circopordfp	1651 TCCGGASGAT GTTTCCAAGA TGGCTGCGGG GGCGGGTCCT TCTTCTGCGG 1651 TCCGGASGAT GTTTCCAAGA TGGCTGCGGG GGCGGGTCCT TCTTCTGCGG 1651 TCCGGASGAT GTTTCCAAGA TGGCTGCGGG GGCGGGTCCG TCTTCTGCGG 1651 TCCGGASGAT GTTTCCAAGA TGGCTGCGGG GGCGGGTCCG TCTTCTGCGG	$1700 \\ 1000 \\ 1000 \\ 1000 \\ 1000 \\ 1000 \\ 1000 \\ 1000 \\ 1000 \\ 1000 \\ 1000 \\ 1000 \\ 1000 \\ 1000 \\ $
circopormank circopormeeh circopordfp	1710 1720 1730 1740 1750 1701 TAACGCCTCC TTGGCCACGT CATCCTATAA AAGTGAAAGA AGTGCGCTGC 1701 TAACGCCTCC TTGGCCACGT CATCCTATAA AAGTGAAAGA AGTGCGCTGC 1701 TAACGCCTCC TTGGCCACGT CATCCTATAA AAGTGAAAGA AGTGCGCTGC	1750 1750 1750
circopormank circopormeeh circopordfp	1760 1770 1780 1790 1800 1751 [TGTAGTATT]. 1751 [TGTAGTATT]. 1751 [TGTAGTATT].	1800 1800 1800

FIG. 3c

circopormank circopormeeh circopordfp[10 20 30 40 50 1 NPSKKSSZOP HKRWVFTLNN PSEEEKNKIR ELPISLFDYF VCGEEGLEEG 1 NPSKKSSZOP HKRWVFTLNN PSEEEKNKIR ELPISLFDYF VCGEEGLEEG 1 NPSKKSSZOP HKRWVFTLNN PSEEEKNKIR ELPISLFDYF VCGEEGLEEG	50 50
circopormank circopormeeh circopordfp[60 70 80 90 100 51 RTAHLOGFAN FAKKOTENKV KWYFGARCHI EKAKGTDOON KEYCSKEGHI 51 RTPHLQGFAN FAKKOTENKV KWYFGARCHI EKAKGTDOON KEYCSKEGHI 51 RTPHLQGFAN FAKKOTENKV KWYFGARCHI EKAKGTDOON KEYCSKEGHI	100 100 100
circopormank circopormeen circopordfp[110 120 130 140 150 101 LIECGAPRNO GKRSDLSTAV STLLETGSLV TVAEOFPVTY VRNFRGLAEL 101 LIECGAPRNO GKRSDLSTAV STLLETGSLV TVAEOFPVTY VRNFRGLAEL 101 LIECGAPRNO GKRSDLSTAV STLLETGSLV TVAEOFPVTY VRNFRGLAEL	150 150 150
circopormank circopormeeh circopordfp[160 170 180 190 20C 151 [LKVSGKMÇOR DWKTAVHVIV GPPGCGKSOW ARNFAEPSOT YWKPSRNKWH 151 LKVSGKMÇOR DWKTAVHVIV GPPGCGKSOW ARNFAEPROT YWKPSRNKWH 151 LKVSGKMÇOR DWKTAVHVIV GPPGCGKSOW ARNFAEPROT YWKPSRNKWH	200 200 200
circopormank circopormeeh circopordfp[210 220 230 240 250 201 DGYHGFEVVV LDDFYGWLPW DDLLRLCDRY PLTVETKGGT VPFLARSILI 201 DGYHGFEVVV LDDFYGWLPW DDLLRLCDRY PLTVETKGGT VPFLARSILI 201 DGYHGFEVVV LDDFYGWLPW DDLLRLCDRY PLTVETKGGT VPFLARSILI	250 250 250
circopormank circopormeeh circopordfp[260 270 280 290 300 251 TSNOAPOEWY SSTAVPAVEA LYRRITTLOF WKTAGEOSTE VPEGRFEAVD 251 TSNOAPOEWY SSTAVPAVEA LYRRITTLOF WKTAGEOSTE VPEGRFEAVD 251 TSNOAPOEWY SSTAVPAVEA LYRRITTLOF WKTAGEOSTE VPEGRFEAVD	300 300 300
circopormank circopormeeh circopordfp[301310320330340350301PPCALFPYKINY301PPCALFPYKINY	350 350 350

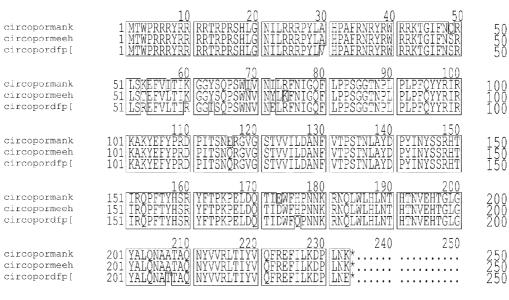
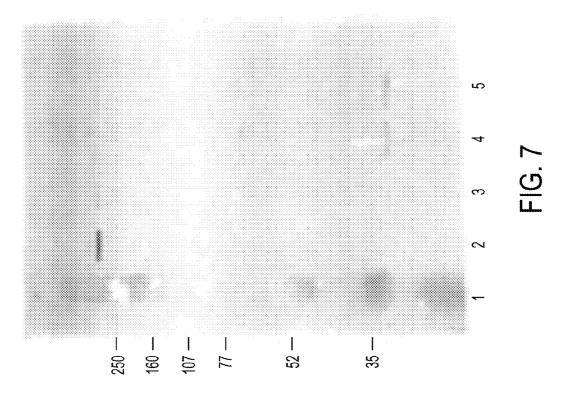


FIG. 5

circopormank circopormeeh circopordfp[1 1 1	10 MISIPPLIST MISIPPLIST MISIPPLIST	20 RLPVGVARLS RLPVGVPRLS RLPVGVPRLS	30 KITGPLALPT KITGPLALPT KITGPLALPT	40 TGRAHYDVYS TGRAHYDVYS TGRAHYDVYS	50 CLPITLLHLP CLPITLLHLP CLPITLLHLP	50 500 500
circopormank circopormeeh circopordfp[51 51 51	60 AHFOKFSOPA AHFOKFSOPA AHFOKFSOPA	70 EISHIRYREI EISHIRYREI EISHIRYRKL	80 LGYSHQRPRL LGYSHQRPRL LGYSHQRPRL	90 QKGTHSSRQV QKGTHSSRQV QKGTHSSRQV	100 AALPLVPRSS AALPLVPRSS AALPLVPRSS	100 100 100
circopormank circopormeeh circopordfp[101 101 101	110 TLDKYVAFFT TLDKYVAFFT TLDKYVAFFT	120 AVFFILLVGS AVFFILLVGS AVFFILLVGS	130 FRFLDVAAGT FRFLDVAAGT FRFLDVAAGT	140 KIPLHLVKSL KIPLHLVKSL KIPLHLVKSL	150 [LLSKISKPLE] LLSKIRKPLE] LLSKIRKPLE]	150 150 150
circopormank circopormeeh circopordfp[151 151 151	160 VSSSTLFOTF VRSSTLFOTF VRSSTLFOTF	170 LSANKIIKKG LSANKIIKKG LATNKIIKKG	180 DWKLPYFVFL DWKLPYFVFL DWKLPYFVFL	190 LLGRIIKGEH LLGRIIKGEH LLGRIIKGEH	200 PPLMGLRAAF PPLMGLRAAF PPLMGLRAAF	200 200 200
circopormank circopormeeh circopordfp[201 201 201	210 [LAWHFH* LAWHFFH LAWHFH	220	230	240	250	250 250 250



Leu Ala Ser Arg Cys Arg Cys Cys Arg Pro Leu Val Glu Ala Ala Val His Gly Trp Arg Val Glu Ala Ala Ala Ala Gly Arg Cys Cys Arg Leu Leu Leu Met Gly Gly Ala Cys Lys Pro Leu Pro Leu Val Glu Ala Ala Gly *** Cys Cys Cys Ala TGG TCG CGT GAA GCC GTC GCC GTC GTG GAG CCG TCG TGG AGT CGT CGT TGT ACG 9 ACC AGC GCA CTT CGG CĀG CGG CAG CAC CTC GGC AGC ACC TCA GCA GCA ACA TGC Thr Ser Ala Leu Arg Gln Arg Gln His Leu Gly Ser Thr Ser Ala Ala Thr Cys Pro Ala His Phe GIy Ser GIy Ser Thr Ser Ala Ala Pro Gln Gln Gln His Ala Gln Arg Thr Ser Ala Ala Ala Ala Pro Arg Gln His Leu Ser Ser Asn Met Pro

Ala Leu Leu Ile Ser Ser Ala Ser Gly Leu Gly Met Phe Pro Pro His Glu Ser Leu Leu Phe Phe Pro Leu Leu Pro Gly Trp Gly Trp Leu Leu His Thr Asn Val Trp Cys Ser Ser His Phe Phe Arg Val Gly Val Gly Tyr Phe Thr Pro Thr *** GGT CGT TCT TCT TAC CTT CTT CGC CTG GGG TTG GGG TAT FTT CCA CCC ACA AGT 63 72 81 90 99 108 CCA GCA AGA AGA ATG GAA GAA GCG GAC CCC AAC CCC ATA AAA GGT GGG TGT TCA Pro Ala Arg Arg Met Glu Glu Ala Asp Pro Asn Pro Ile Lys Gly Gly Cys Ser Gln Gln Glu Glu Trp Lys Lys Arg Thr Pro Thr Pro *** Lys Val Gly Val His Ser Lys Lys Asn Gly Arg Ser Gly Pro Gln Pro His Lys Arg Trp Val Phe Thr

Gln Ile Ile Arg Gly Phe Val Leu Ala Leu Phe Tyr Pro Ile Lys Trp Tyr Gly Arg Phe Leu Gly Glu Ser Ser Ser Arg Leu Phe Ile Arg Ser Arg Gly Ile Asp Glu Ser Tyr Asp Lys Arg Leu Arg Ala Cys Ser Phe Val Pro Asp Glu Leu Ile GAG ACT TAT TAG GAA GGC TTC TGC TCG CGT TCT TTT ATG CCC TAG AAG GTT ATA 117 126 135 144 153 162 CTC TGA ATA ATC CTT CCG AAG ACG AGC GCA AGA AAA TAC GGG ATC TTC CAA TAT Leu *** Ile Ile Leu Pro Lys Thr Ser Ala Arg Lys Tyr Gly Ile Phe Gln Tyr Ser Glu *** Ser Phe Arg Arg Arg Ala Gln Glu Asn Thr Gly Ser Ser Asn Ile Leu Asn Asn Pro Ser Glu Asp Glu Arg Lys Lys Ile Arg Asp Leu Pro Ile Ser

*** Lys Ile Ile Lys Asn Asn Ala Leu Leu Thr Ile Leu Phe Ser Ser Cys Arg Arg Asn Ser *** Lys Ile Thr Pro Ser Ser Pro Leu Ser Ser Pro Arg Val Gly Gly Ile Gln Asn Asn *** Gln Gln Arg Pro Pro Tyr His Pro Leu Val Phe Val GGG ATA AAC TAA TAA AAT AAC AAC CGC TCC TCC CAT TAC FCC TTC CTG CTT GTG 171 180 189 198 207 216 CCC TAT TTG ATT ATT TTA TTG TTG GCG AGG AGG GTA ATG AGG AAG GAC GAA CAC Pro Tyr Leu Ile Ile Leu Leu Leu Ala Arg Arg Val Met Arg Lys Asp Glu His Pro Ile *** Leu Phe Tyr Cys Trp Arg Gly Gly *** *** Gly Arg Thr Asn Thr Leu Phe Asp Tyr Phe Ile Val Gly Glu Glu Gly Asn Glu Glu Gly Arg Thr Pro

Val Glu Leu Pro Glu Ser Ile Lys His Leu Leu Leu Ser Lys Ile Phe His Leu *** Arg Trp Pro Asn Ala Leu Lys Thr Phe Phe Cys Val Lys Leu Leu Thr Phe Glu Gly Gly Pro Thr Arg *** Asn Gln Ser Ser Ala Ser Lys *** Tyr Leu Ser GAG TGG AGG TCC CCA AGC GAT TAA AAC ACT TCT TCG TCT GAA AAT TAT TTC ACT 225 234 243 252 261 270 CTC ACC TCC AGG GGT TCG CTA ATT TTG TGA AGA AGC AGA CTT TTA ATA AAG TGA Leu Thr Ser Arg Gly Ser Leu Ile Leu *** Arg Ser Arg Leu Leu Ile Lys *** Ser Pro Pro Gly Val Arg *** Phe Cys Glu Glu Ala Asp Phe *** *** Ser Glu His Leu Gln Gly Phe Ala Asn Phe Val Lys Lys Gln Thr Phe Asn Lys Val Lys

3' 5'

FIG. 8a

Pro Ile Gln Thr Gly Ala Ala Val Asp Leu Phe Arg Phe Ser Cys Ile Leu Leu His Tyr Lys Pro Ala Arg Gln Trp Met Ser Phe Ala Phe Pro Val Ser *** Cys Thr Thr Asn Pro His Gly Ser Gly Cys Arg Ser Leu Ser Leu Phe Leu Asp Ala TCA CCA TAA ACC CAC GGG CGA CGG TGT AGC TCT TTC GCT TTC CTT GTC TAG TCG 279 288 297 306 315 324 AGT GGT ATT TGG GTG CCC GCT GCC ACA TCG AGA AAG CGA AAG GAA CAG ATC AGC Ser Gly Ile Trp Val Pro Ala Ala Thr Ser Arg Lys Arg Iys Glu Gln Ile Ser Val Val Phe Gly Cys Pro Leu Pro His Arg Glu Ser Glu Arg Asn Arg Ser Ala Trp Tyr Leu Gly Ala Arg Cys His Ile Glu Lys Ala Lys GIy Thr Asp Gln Gln

Ile Phe Phe Val Ala Thr Phe Phe Ala Val *** Gln His Leu Thr Ser Ser Arg Phe Leu Ser Tyr Gln Leu Leu Ser Pro Leu Lys Ser His Ser His Pro Ala Gly Ser Tyr Leu Ile Ser Cys Tyr Leu Leu Cys Ser Val Ser Pro Thr His Leu Glu TCT TAT TTC TTA TGA CGT CAT TTC TTC CGT TGA ATG ACT ACC TCA CAC CTC GAG 333 342 351 360 369 378 AGA ATA AAG AAT ACT GCA GTA AAG AAG GCA ACT TAC TGA TGG AGT GTG GAG CTC Arg Ile Lys Asn Thr Ala Val Lys Lys Ala Thr Tyr *** Tro Ser Val Glu Leu Glu *** Arg Ile Leu Gln *** Arg Arg Gln Leu Thr Asp Gly Val Trp Ser Ser Asn Lys Glu Tyr Cys Ser Lys Glu Gly Asn Leu Leu Met Glu Cys Gly Ala Pro

Ser Arg Leu Ser Leu Pro Thr Val Gln Arg Ser Ser His Thr Gly Gln Gln Leu Leu Asp *** Pro Cys Arg Leu Ser Arg Asp Val Ala Thr Leu Val Lys Asn Ser *** Ile Glu Pro Val Val Ser His Gly Thr *** Gln Gln Ser Tyr Arg Thr Pro GAT CTA GAG TCC CTG TTG CCT CAC TGG ACA GAT GAC GAC ACT CAT GGA ACA ACC 387 396 405 414 423 432 CTA GAT CTC AGG GAC AAC GGA GTG ACC TGT CTA CTG CTG TGA GTA CCT TGT TGG Leu Asp Leu Arg Asp Asn Gly Val Thr Cys Leu Leu Leu *** Val Pro Cys Trp *** Ile Ser Gly Thr Thr Glu *** Pro Val Tyr Cys Cys Glu Tyr Leu Val Gly Arg Ser Gln Gly Gln Arg Ser Asp Leu Ser Thr Ala Val Ser Thr Leu Leu Glu

Ala Pro Thr Gln His Gly Asn Cys Leu Leu Val Arg Tyr Arg Lys Asp Ser Ile Leu Pro Leu Arg Thr Val Thr Ala Ser Cys Cys Gly Thr Val Asn Thr Leu Phe Ser Arg Ser Asp Pro Ser Arg Gln Leu Ala Ala Gly Gln Leu Thr Gln *** Phe TCT CGC CCT CAG ACC ACT GGC AAC GTC TCG TCG TGG GAC ATT GCA AAC AGT CTT 441 450 459 468 477 486 AGA GCG GGA GTC TGG TGA CCG TTG CAG AGC AGC ACC CTG TAA CGT TTG TCA GAA Arg Ala Gly Val Trp *** Pro Leu Gln Ser Ser Thr Leu *** Arg Leu Ser Glu Glu Arg Glu Ser Gly Asp Arg Cys Arg Ala Ala Pro Cys Asn Val Cys Gln Lys Ser Gly Ser Leu Val Thr Val Ala Glu Gln His Pro Val Thr Phe Val Arg Asn

Glu Ala Pro Gln Ser Phe Lys Gln Phe His Ala Pro Phe His Leu Leu Thr Ile Lys Arg Pro Ser Ala Ser Ser Lys Phe Thr Leu Pro Phe Ile Cys Phe Arg Ser Asn Gly Arg Ala Pro Gln Val Lys Ser Leu Ser Arg Ser Phe Ala Ser Ala His TAA AGG CGC CCG ACC GAC TTG AAA ACT TTC ACT CGC CCT TTT ACG TCT TCG CAC 495 504 513 522 531 540 ATT TCC GCG GGC TGG CTG AAC TTT TGA AAG TGA GCG GGA AAA TGC AGA AGC GTG Ile Ser Ala Gly Trp Leu Asn Phe *** Lys *** Ala Gly Lys Cys Arg Ser Val Phe Pro Arg Ala Gly *** Thr Phe Glu Ser Glu Arg Glu Asn Ala Glu Ala *** Phe Arg Gly Leu Ala Glu Leu Leu Lys Val Ser Gly Lys Met Gln Lys Arg Asp

FIG. 8b

Pro Leu Ser Ile Tyr Val Asp Asn His Pro Trp Arg Pro Thr Thr Phe Ala Phe Gln Phe Val Leu Thr Cys Thr Met Thr Pro Gly Gly Pro His Pro Leu Leu Leu Asn Ser Ser *** His Val Arg *** Gln Pro Ala Val Gln Thr His Tyr Phe Cys TAA CCT TCT GAT TAC ATG TGC AGT AAC ACC CCG GTG GAC CCA CAC CAT TTT CGT 549 ATT GGA AGA CTA ATG TAC ACG TCA TTG TGG GGC CAC CTG GGT GTG GTA AAA GCA Ile Gly Arg Leu Met Tyr Thr Ser Leu Trp Gly His Leu Gly Val Val Lys Ala Leu Glu Asp *** Cys Thr Arg His Cys Gly Ala Thr Trp Val Trp *** Lys Gln Trp Lys Thr Asn Val His Val Ile Val Gly Pro Pro Gly Cys Gly Lys Ser Lys

Pro Ser Ser Ile Lys Cys Val Arg Phe Gly Cys Val Pro Phe Trp Arg Ser Val His Ala Ala Leu Lys Ala Ser Gly Ser Val Val Tyr Gln Phe Gly Gly Leu Phe Ile Pro Gln *** Asn Gln Leu Gly Pro Phe Trp Met Ser Ser Val Val *** Phe TTA CCC GAC GAT TAA AAC GTC TGG GCC TTT GGT GTA TGA CCT TTG GTG GAT CTT 603 612 621 630 639 648 AAT GGG CTG CTA ATT TTG CAG ACC CGG AAA CCA CAT ACT GGA AAC CAC CTA GAA Asn Gly Leu Leu Ile Leu Gln Thr Arg Lys Pro His Thr Gly Asn His Leu Glu Met Gly Cys *** Phe Cys Arg Pro Gly Asn His Ile Leu Glu Thr Thr *** Lys Trp Ala Ala Asn Phe Ala Asp Pro Glu Thr Thr Tyr Trp Lys Pro Pro Arg Asn

Leu Pro Pro Ile Thr Val Met Thr Phe Phe His Asn Asn Asn Ile Val Lys Ile Leu His His Ser Pro *** Trp Pro Ser Ser Thr Thr Thr Ile Ser Ser Lys *** Cys Thr Thr Pro His Asn Gly His His Leu Leu Pro Gln *** Gln His Ser Lys TGT TCA CCA CCC TAC CAA TGG TAC CAC TTC TJC ACC AAC AAT AAC TAC TGA AAA 657 666 675 684 693 702 ACA AGT GGT GGG ATG GTT ACC ATG GTG AAG AAG IGG TTG TTA TTG ATG ACT TTT Thr Ser Gly Gly Met Val Thr Met Val Lys Lys Trp Leu Leu Leu Met Thr Phe Gln Val Val Gly Tro Leu Pro Trp *** Arg Ser Gly Cys Tyr *** *** Leu Leu Lys Trp Trp Asp Cly Tyr His Gly Glu Glu Val Val Val Ile Asp Asp Phe Tyr

Ala Pro Gln Gly Pro Ile Ile *** Gln Ser Gln Thr Ile Ser Ile Trp Gln Ser Pro Gln Ser Gly Gln Ser Ser Arg Ser Leu Ser His Ser Arg Tyr Gly Asn Val His Ser Ala Ala Arg Pro His Asp Val Ser Val Thr His Asp Ile Asp Met Ser TAC CGA CCG ACG GGA CCC TAC TAG ATG ACT CIG ACA CAC TAG CTA TAG GTA ACT 711 720 729 738 747 756 ATG GCT GGC IGC CCT GGG ATG ATC TAC TGA GAC IGT GTG ATC GAT ATC CAT TGA Met Ala Gly Cys Pro Gly Met Ile Tyr *** Asp Cys Val Ile Asp Ile His *** Tro Leu Ala Ala Leu Gly *** Ser Thr Glu Thr Val *** Ser Ile Ser Ile Asp Gly Trp Leu Pro Trp Asp Asp Leu Leu Arg Leu Cys Asp Arg Tyr Pro Leu Thr

Tyr Leu Ser Phe Thr Ser Ser Tyr Arg Lys Gln Gly Ala Thr Asn Gln Asn Gly Thr Ser Val Leu Pro Pro Val Thr Gly Lys Lys Ala Arg Leu Ile Arg Ile Val Gln Leu Ser *** Leu His Phe Gln Val Lys Lys Pro Gly Cys Tyr Glu Ser *** ___ ___ GAC ATC TCT GAT TTC CAC CTT GAC ATG GAA AAA ACC 765 774 774 783 792 CAT AAG ACT 801 AAT 810 GGG CGT CTG TAG AGĂ CTA AAG GTG GAA CTG TĂČ CTT TIT IĞG CCC GCA ĞŤA TTC TGA ŤTĂ ____ _ _ _ _ ____ ____ ---____ ____ ___ Leu *** Arg Leu Lys Val Glu Leu Tyr Leu Phe Trp Pro Ala Val Phe *** Leu Cys Arg Asp *** Arg Trp Asn Cys Thr Phe Phe Gly Pro Gln Tyr Ser Asp Tyr Val Glu Thr Lys Gly Gly Thr Val Pro Phe Leu Ala Arg Ser Ile Leu Ile Thr Ala Ile Leu Gly Arg Gln Phe Pro Val Gly *** Ser Ser Asp Trp Ser Tyr Phe Leu Leu *** Val Gly Asn Ser His Tyr Glu Glu Val Ala Thr Gly Ala Thr Ser Trp Cys Asp Ser Gly Thr Pro Ile Thr Ser Arg Leu Gln Gln Gly Leu Gln Leu GGT CGT TAG TCT GGG GCA ACC TTA CCA TGA GGA GTT GAC GAC AGG GTC GAC ATC 819 CCA GCA ATC AGA CCC CGT TGG AAT GGT ACT CCT CAA CTG CTG TCC CAG CTG TAG Pro Ala Ile Arg Pro Arg Trp Asn Gly Thr Pro Gln Leu Leu Ser Gln Leu *** Gln Gln Ser Asp Pro Val Gly Met Val Leu Leu Asn Cys Cys Pro Ser Cys Arg Ser Asn Gln Thr Pro Leu Glu Trp Tyr Ser Ser Thr Ala Val Pro Ala Val Glu

Ser Lys Ile Pro Pro Asn Ser Gly Gln Tyr Lys Pro Leu Ile Ser Cys Phe Leu Ala Arg *** Arg Leu Ile Val Glu Lys Thr Asn Gln Phe Phe Ala Val Ser Cys Leu Glu Lys Asp Ser Ser *** Lys Arg Pro Ile Lys Ser Ser His *** Leu Val TTC GAG AAA TAG CCT CCT AAT GAA GGA ACC ATA AAA CCT TCT TAC GAT GTC TTG 873 AAG CTC TTT ATC GGA GGA TTA CTT CCT TGG TAT TTT GGA AGA ATG CTA CAG AAC Lys Leu Phe Ile Gly Gly Leu Leu Pro Trp Tyr Phe Gly Arg Met Leu Gln Asn Ser Ser Leu Ser Glu Asp Tyr Phe Leu Gly Ile Leu Glu Glu Cys Tyr Arg Thr Ala Leu Tyr Arg Arg Ile Thr Ser Leu Val Phe Trp Lys Asn Ala Thr Glu Gln

Gly Arg Leu Phe Pro Ala Leu Glu Asp Gly Lys Gly Gly Trp Ala Arg Phe Lys Asp Val Ser Ser Pro Pro Trp Asn Thr Val Arg Glu Gly Gly His Gly Ser Asn Ile Trp Pro Pro Leu Pro Gly Thr Arg *** Gly Lys Gly Gly Met Gly Gln I TTA GGT CCC TCC TTC CCC CGG TCA AGC AGT GGG AAA GGG GGG GTA CGG GAC TTA 927 AAT CCA CGG AGG AAG GGG GCC AGT TCG TCA CCC TTT CCC CCC CAT GCC CTG AAT Asn Pro Arg Arg Lys Gly Ala Ser Ser Ser Pro Phe Pro Pro His Ala Leu Asn Ile His Gly Gly Arg Gly Pro Val Arg His Pro Phe Pro Pro Met Pro *** Ile Ser Thr Glu Glu Gly Gly Gln Phe Val Thr Leu Ser Pro Pro Cys Pro Glu Phe

Trp Ile Phe Tyr Ile Val Ser Asp Lys Lys Asp Ser Arg Leu Pro Lys *** *** Gly Tyr Ser Ile Phe *** Gln Thr Lys Lys Ile Val Glu Tyr His Asn Lys Asn Glu Met His Phe Leu Asn Ser Leu Arg Lys *** *** Lys Thr Ile Thr Lys ILe AAG GTA TAC TTT ATT TAA TGA CTC AGA AAA AAT AGT GAA GCA TTA CCA AAA ATA 981 990 999 1008 1017 1026 TTC CAT ATG AAA TAA ATT ACT GAG TCI TTT TTA TCA CTT CGT AAT GGT TTT TAT Phe His Met Lys *** Ile Thr Glu Ser Phe Leu Ser Leu Arg Asn Gly Phe Tyr Ser Ile *** Asn Lys Leu Leu Ser Leu Phe Tyr His Phe Val Met Val Phe _le Pro Tyr Glu Ile Asn Tyr *** Val Phe Phe Ile Thr Ser *** Trp Phe Leu Leu

Glu Asn Leu Thr Leu His Pro Thr Lys Leu Ile Leu Asn Glu Ser Asn Tyr Met Asn Met Leu Pro *** Thr Pro Pro Arg *** Phe *** Ile Arg Gln Ile Thr Cys Ile *** *** Pro Asn Leu Pro Pro Asp Lys Phe Asn Phe Glu Arg Phe Gln Val ATA AGT AAT TCC CAA TTC ACC CCC CAG AAA TTT TAA TTT AAG AGA CTT AAC ATG 1035 1044 1053 1062 1071 1080 TAT TCA TTA AGG GTT AAG TGG GGG GTC TTT AAA ATT AAA TTC TCT GAA TTG TAC Tyr Ser Leu Arg Val Lys Trp Gly Val Phe Lys Ile Lys Phe Ser Glu Leu Tyr Ile His *** Gly Leu Ser Gly Gly Ser Leu Lys Leu Asn Ser Leu Asn Cys Thr Phe Ile Lys Gly *** Val Gly Gly Leu *** Asn *** Ile Leu *** Ile Val His

FIG. 8d

Cys Pro *** Val Ser Ile Thr Asn Arg Thr Thr Tyr Val Thr Lys Ser Arg Leu Val His Asn Cys Pro Tyr Gln Ile Gly Pro Arg Ile Tyr Gln Lys Arg Val Cys Tyr Met Thr Val Arg Ile Asn Tyr Glu Gln Asp Tyr Ile Ser Asn Glu Phe Ala TAT GTA CCA ATG TGC CTA TAA CAT AAG GAC CAG CAT ATA TGA CAA AAG CTT GCG 1089 1098 1107 1116 1125 1134 ATA CAT GGT TAC ACG GAT ATT GTA TIC CTG GTC GTA TAT ACT GTT TTC GAA CGC Ile His Gly Tyr Thr Asp Ile Val Phe Leu Val Val Tyr Thr Val Phe Glu Arg Tyr Met Val Thr Arg Ile Leu Tyr Ser Trp Ser Tyr Ile Leu Phe Ser Asn Ala Thr Trp Leu His Gly Tyr Cys Ile Pro Gly Arg Ile Tyr Cys Phe Arg Thr Gln

Ala Ser Ala *** Thr Thr *** Met Glu Leu Leu Lys Tyr Asp *** Gly Cys Ser His Arg Pro Arg Arg Pro Arg Cys Lys Trp Cys Asn Thr Thr Glu Ala Val Ala Thr Gly Leu Gly Val His Asp Val Asn Gly Ala Thr Gln Leu Arg Leu Trp Leu TCA CGG CTC CGG ATG CAC CAG ATG TAA AGG TCG TCA AAC ATC AGA GTC GGT GTC 1143 AGT GCC GAG GCC TAC GTG GTC TAC ATT TCC AGC AGT TTG TAG TCT CAG CCA CAG Ser Ala Glu Ala Tyr Val Val Tyr Ile Ser Ser Ser Leu *** Ser Gln Pro Gln Val Pro Arg Pro Thr Trp Ser Thr Phe Pro Ala Val Cys Ser Leu Ser His Ser Cys Arg Gly Leu Arg Gly Leu His Phe Gln Gln Phe Val Val Ser Ala Thr Ala

Thr Glu Lys Thr Thr Gln Asn Ser Thr Ile Leu Leu Ser Ile *** Ser Leu Asn Pro Lys Lys Gln Gln Lys Thr Pro Leu Leu *** Tyr His Phe Arg Pro Cys Thr Gln Asn Arg Lys Asn Asn Pro Gln Phe Tyr Asp Ile Thr Phe Asp Leu Val Pro GAC CAA AGA AAA CAA CAA ACC AAC CTT CAT TAG TTA TCA CTT TAG ATC CTG TCC 1197 1206 1215 1224 1233 1242 CTG GTT TCT TTT GTT GTT TGG TTG GAA GTA ATC AAT AGT GAA ATC TAG GAC AGG Leu Val Ser Phe Val Val Trp Leu Glu Val Ile Asn Ser Glu Ile *** Asp Arg Trp Phe Leu Leu Leu Phe Gly Trp Lys *** Ser Ile Val Lys Ser Arg Thr Gly Gly Phe Phe Cys Cys Leu Val Gly Ser Asn Gln *** *** Asn Leu Gly Gln Val

Pro Pro Leu Thr Gly Pro Thr Thr Pro Ser Pro Ser Pro *** Pro Ile Ala Pro Gln Pro Tyr Leu Val Pro Leu Pro Leu Leu Leu Ala Pro Asn His Tyr Pro Pro Lys Pro Thr Phe Tyr Arg Ser His Tyr Ser Phe Pro Gln Thr Ile Thr His Arg AAA CCC CCA TTT CAT GGC CCT CAC CAT CCT CTT CCC GAC CCA ATA CCA TAC CGC 1251 1260 1269 1278 1287 1296 TTT GGG GGT AAA GTA CCG GGA GIG GIA GGA GAA GGG CTG GGT TAT GGT ATG GCG Phe Gly Gly Lys Val Pro Gly Val Val Gly Glu Gly Leu Gly Tyr Gly Met Ala Leu Gly Val Lys Tyr Arg Glu Trp *** Clu Lys Gly Trp Val Met Val Trp Arg Trp Gly *** Ser Thr Gly Ser Gly Arg Arg Arg Ala Gly Leu Trp Tyr Gly Gly

Pro Thr Thr *** Met Pro Thr Met Pro Ser Pro Gln Pro Arg Gln *** Leu Thr Leu Leu Leu Lys Cys Leu Pro *** Leu His Pro Ser His Gly Lys Asn Cys Leu Ser Ser Tyr Asn Val Tyr Pro Asp Tyr Thr Leu Ala Thr Ala Lys Thr Val Phe CCT CCT CAT CAA ATG TAT CCC CAG TAT CCA CTC CCG ACA CCG GAA ACA ATG TTT 1305 1314 1323 1332 1341 1350 GGA GGA GTA GTT TAC ATA GGG GTC ATA GGT GAG GGC TGT GGC CTT TGT TAC AAA Gly Gly Val Val Tyr Ile Gly Val Ile Gly Glu Gly Cys Gly Leu Cys Tyr Lys Glu Glu *** Phe Thr *** Gly Ser *** Val Arg Ala Val Ala Phe Val Thr Lys Arg Ser Ser Leu His Arg Gly His Arg *** Gly Leu Trp Pro Leu Leu Gln Ser

FIG. 8e

Ile Met *** Phe Leu Leu Val Pro Ala Trp Glu Gly Thr Val Arg Pro Ser Arg *** *** Arg Phe Tyr Cys Cys G_n Leu Gly Ser Gly Gln *** Gly Pro His Asp Asn Asp Asp Leu Ile Val Ala Ser Ser Gly Val Gly Arg Asp Gly Gln Thr Ile CAA TAG TAG ATT TTA TTG TCG IGA CCT CGG GTG AGG GGA CAG TGG GAC CCA CTA 1359 1368 1377 1386 1395 1404 GTT ATC ATC TAA AAT AAC AGC ACT GGA GCC CAC TCC CCT GTC ACC CTG GGT GAT Val Ile Ile *** Asn Asn Ser Thr Gly Ala His Ser Pro Val Thr Leu Gly Asp Leu Ser Ser Lys Ile Thr Ala Leu Glu Pro Thr Pro Leu Ser Pro Trp Val Ile Tyr His Leu Lys *** Gln His Trp Ser Pro Leu Pro Cys His Pro Gly *** Ser

Pro Ala Pro Gly Ser Asn Leu Arg Leu Arg Glu *** Glu Thr Thr Asn Leu Pro Pro Leu Leu Ala Leu Ile *** G_Y *** Gly Lys Lys Asn Gln Leu Ile *** Leu Pro Ser Cys Pro Trp Phe Glu Va_ Lys Val Lys Arg Ile Arg Tyr Tyr Glu Phe GCC CCT CGT CCC GGT CTT AAG TTG GAA TTG GAA AGA ATA AGA CAT CAT AAG TTT 1413 1422 1431 1440 1449 1458 CGG GGA GCA GGG CCA GAA TTC AAC CTT AAC CTT TCT TAT TCT GTA GTA TTC AAA Arg Gly Ala Gly Pro Glu Phe Asn Leu Asn Leu Ser Tyr Ser Val Val Phe Lys Gly Glu Gln Gly Gln Asn Ser Thr Leu Thr Phe Leu Ile Leu *** Tyr Ser Lys Gly Ser Arg Ala Arg Ile Gln Pro *** Pro Phe Leu Phe Cys Ser Ile Gln Arg

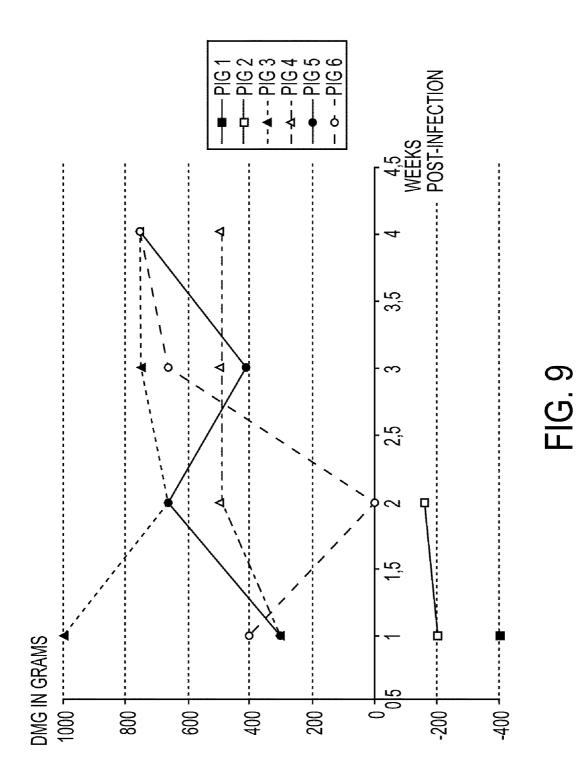
Cys Leu Ala Pro Thr Gln Gly Gly Glu Gln Pro Phe Phe Thr Met Leu Ile Ser Ala Cys Leu Pro Pro Lys Val G_Y Arg Arg Pro Ser Ser Leu *** *** Tyr Gln Pro Val Ser Arg Pro Asn Ser Gly Gly Gly Pro Pro Leu Phe Asp Asn Ile Asn CCC GTG TCT CGC CCC CAA ACT GGG GGG AGG ACC CCC TTC TTT CAG TAA TTA TAA 1467 1476 1485 1494 1503 1512 GGG CAC ACA GCG GGG GTT TGA CCC CCC TCC TGG GGG AGG AAA GTC ATT AAT ATT Gly His Arg Ala Gly Val *** Pro Pro Ser Trp Gly Lys Lys Val Ile Asn Ile Gly Thr Glu Arg Gly Phe Asp Pro Pro Pro Gly Gly Arg Lys Ser Leu Ile Leu Ala Gln Ser Gly Gly Leu Thr Pro Leu Leu Gly Glu Glu Ser His *** Tyr ***

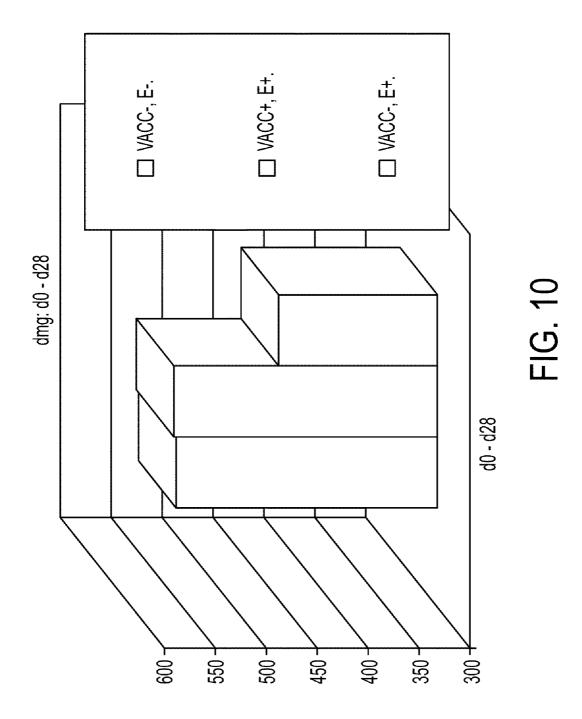
Asp *** *** Thr Trp Arg Gly Prc Pro Arg Glu Ser Gln Pro Glu Ser Ser Leu Ile Glu Asp His Gly Gly Gly Leu Leu Ala Asn Gln Ser His Asn Ala Gln Cys Phe Arg Met Met Asp Val Ala Trp Ser Pro Thr Arg Val Thr Thr Arg Lys Val CCT AGA GTA GTA CAG GTG GCG GGT CCT CCC GCA AGA CTG ACA CCA AGC GAA CTG 1521 1530 1539 1548 1557 1566 GAA TCT CAT CAT GTC CAC CGC CCA GGA GGG CGT TCT GAC TGT GGT TCG CTT GAC Glu Ser His His Val His Arg Pro Gly Gly Arg Ser Asp Cys Gly Ser Leu Asp Asn Leu Ile Met Ser Thr Ala Gin Glu Gly Val Leu Thr Val Val Arg Leu Thr Ile Ser Ser Cys Pro Pro Pro Arg Arg Ala Phe *** Leu Trp Phe Ala *** Gln

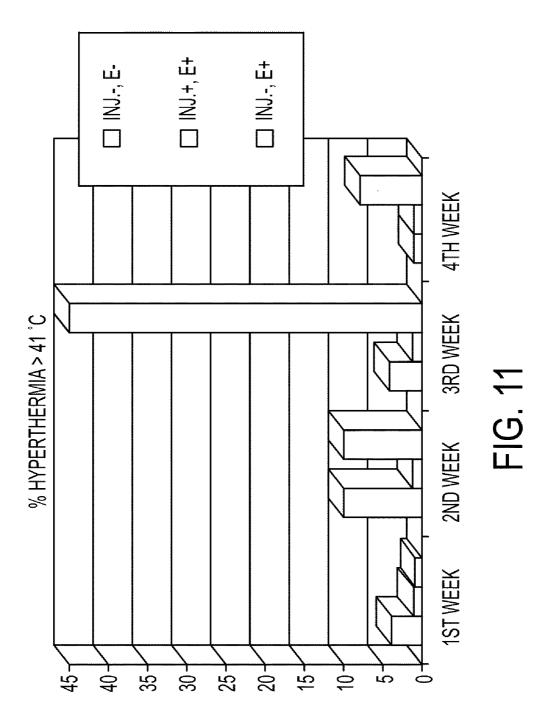
Ile Asp Ser Pro Ala Pro Ser Ala Pro Thr Ser Ser Ala Met Lys Gly Glu Gly Tyr Ile Arg Leu His Pro Leu Pro Pro His Gln Leu His Trp Lys Glu Lys Glu Thr Tyr Gly Phe Thr Arg Ser Leu Arg Thr Asn Phe Ile Gly Asn Lys Arg Arg TCA TAT AGG CTT CCA CGC CCT CTC CGC CCA CAA CTT CTA CGG TAA AAA GGA AGA 1575 1584 1593 1602 1611 1620 AGT ATA TCC GAA GGT GCG GGA GAG GCG GGT GTT GAA GAT GCC ATT TTT CTT TCT Ser Ile Ser Glu Gly Ala Gly Glu Ala Gly Val Glu Asp Ala Ile Phe Pro Ser Val Tyr Pro Lys Val Arg Glu Arg Arg Val Leu Lys Met Pro Phe Phe Leu Leu Tyr Ile Arg Arg Cys Gly Arg Gly Gly Cys *** Arg Cys His Phe Ser Phe Ser

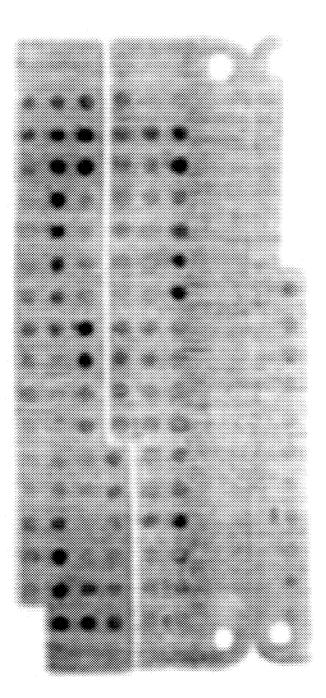
*** Ile Gln Phe Arg Phe Phe His Ala Thr Leu Ile Asp Tyr Arg Phe Val Phe Ser Thr Arg Gln Leu Tyr Thr Met Asp Ser Phe Ser Leu Leu Ala Ser Tyr Thr Asn GCA GTA TAG ACT TTT GCT TTC TTC ACG CGA CAT TCA TAA 5' 1737 1746 1755 1764 CGT CAT ATC TGA AAA CGA AAG AAG TGC GCT GTA AGT ATT 3' Arg His Ile *** Lys Arg Lys Lys Cys Ala Val Ser Ile Val Ile Ser Glu Asn Glu Arg Ser Ala Leu *** Val Ser Tyr Leu Lys Thr Lys Glu Val Arg Cys Lys Tyr

FIG. 8g





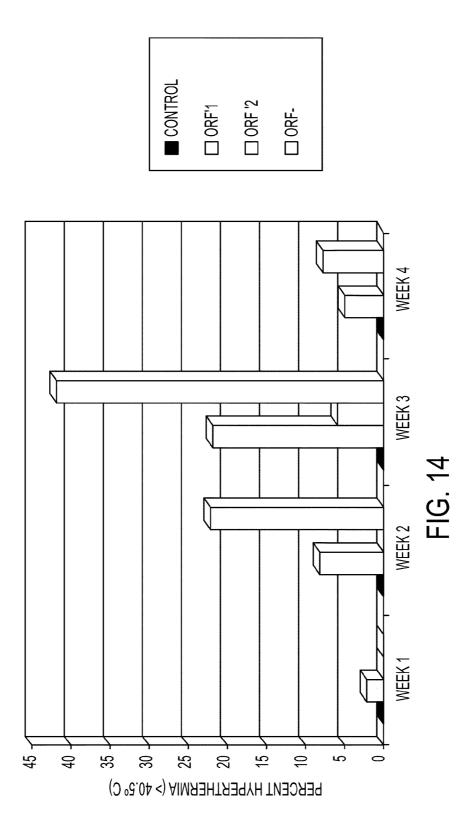


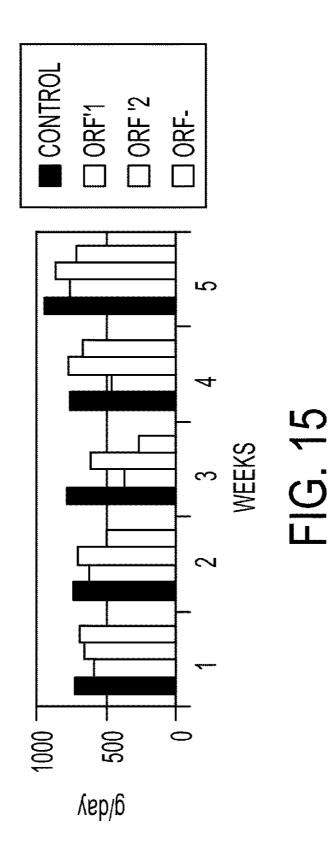


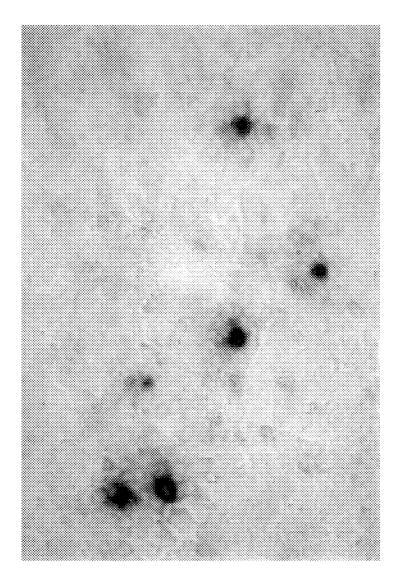
TYPE A SPOT NO. 160 TO 215

TYPE B SPOT NO. 104 TO 159

1 MTWPRRRYRR RRTRPRSHLG NILRRRPYLV HPAFRNYRW RRKTGIFNSR LSREFVLTI. RGGHSQPSWN MTYPRRRYRR RRHRPRSHLG QILRRRPWLV HPRHYRW RRKNGIFNTR LSRTFGYTVK RTTVRTPSWA	100 101 peptides 188 to 189 YYRI RKAKYEFYPR DPITSNQRGV GSTVVILDAN FVTPSTNLAY YYRI RKVKVEFWPC SPITQGDRGV GSSAVILDDN FVTKATALTY peptides 132 to 133	PELD QTIDWFQPNN KRNQLWLHLN THTNVEHTGL GYALQNATTA PVLD FTIDYFQPNN KRNQLWLRLQ TAGNVDHVGL GTAFENSIYD	peptide 152
NILRRF QILRRF	100 LPLPFQYYRI RSVPFEYYRI	RYFTPKPELD RYFTPKPVLD	235 P.LNE PPLNP
RRTRPRSHLG RRHRPRSHLG		151 TIRQPFTYHS TITQPFSYHS	VQFREFILKD VQFREFNFKD
	pcvA VNELRFNIGQ FLPPSGGTNP pcvB VDMMRFNIND FLPPGGGSNP peptide 121	150 DPYINYSSRH DPYVNYSSRH	QNYVVRLTIY QEYNIRVTMY
pcvA pcvB	pcvA_	pcvB	pcvA pcvB







CIRCOVIRUS SEQUENCES ASSOCIATED WITH PIGLET WEIGHT LOSS DISEASE (PWD)

INFORMATION ON RELATED APPLICATIONS

[0001] This application is a continuation of U.S. patent application Ser. No. 11/588,237, filed Oct. 27, 2006, now U.S. Pat. No. _____, which is a divisional of U.S. patent application Ser. No. 10/718,264, filed Nov. 21, 2003, now U.S. Pat. No. 7,179,472, which is a divisional of U.S. patent application Ser. No. 09/514,245, filed Feb. 28, 2000, now U.S. Pat. No. 6,703,023, which is a continuation-in-part of International Patent Application No. PCT/FR98/02634, filed Dec. 4, 1998, published in a non-English language, and now abandoned, which claims priority to French Application No. 97/15396, filed Dec. 5, 1997, the specifications of which are incorporated herein by reference in their entireties for all purposes.

BACKGROUND OF THE INVENTION

[0002] The invention relates to the genomic sequence and nucleotide sequences coding for polypeptides of PWD circovirus, such as the structural and nonstructural polypeptides of said circovirus, as well as vectors including said sequences and cells or animals transformed by these vectors. The invention likewise relates to methods for detecting these nucleic acids or polypeptides and kits for diagnosing infection by the PWD circovirus. The invention is also directed to a method for selecting compounds capable of modulating the viral infection. The invention further comprises pharmaceutical compositions, including vaccines, for the prevention and/or the treatment of viral infections by PWD circovirus as well as the use of a vector according to the invention for the prevention and/or the treatment of diseases by gene therapy.

[0003] Piglet weight loss disease (PWD), alternatively called fatal piglet wasting (FPW) has been widely described in North America (Harding, J. C., 1997), and authors have reported the existence of a relationship between this pathology and the presence of porcine circovirus (Daft, B. et al., 1996; Clark, E. G., 1997; Harding, J. C., 1997; Harding, J. C. and Clark, E. G., 1997; Nayar, G. P. et al., 1997). A porcine circovirus has already been demonstrated in established lines of cell cultures derived from pigs and chronically infected (Tischer, I., 1986, 1988, 1995; Dulac, G. C., 1989; Edwards, S., 1994; Allan, G. M., 1995 and McNeilly, F., 1996). This virus, during experimental infection of piglets, does not prove pathogenic for pigs (Tischer, I., 1986, Homer, G. W., 1991) and its nucleotide sequence has been determined and characterized (Tischer, I., 1982; Meehan, B. M. et al., 1997; Mankertz., A., 1997). The porcine circovirus, called PCV virus, is part of the circovirus genus of the circoviridae family (Murphy, F.A. et al., 1995) whose virion has a circular DNA of size between 1.7 and 2.3 kb, which DNA comprises three open reading frames (ORF1 to ORF3), coding for a replication protein REP involved in the initiation and termination phase of rolling circular replication (RCR) (Heyraud-Nitschke, F., et al., 1995; Harding, M. R. et al., 1993; Hanson, S. F. et al., 1995; Fontes, E. P. B. et al., 1994), coding for a capsid protein (Boulton, L. H. et al., 1997; Hackland, A. F. et al., 1994; Chu, P. W. G. et al., 1993) and coding for a nonstructural protein called a dissemination protein (Lazarowitz., S. G. et al., 1989).

[0004] The inventors of the present invention have noticed that the clinical signs perceptible in pigs and linked to infection by the PWD circovirus are very distinctive. These manifestations in general appear in pigs of 8 to 12 weeks of age, weaned for 4 to 8 weeks. The first signs are hypotonia without it being possible to speak of prostration. Rapidly (48 hours), the flanks hollow, the line of the spine becomes apparent, and the pigs "blanch." These signs are in general accompanied by hyperthermia, anorexia and most often by respiratory signs (coughing, dyspnea, polypnea). Transitory diarrhea can likewise appear. The disease state phase lasts approximately one month at the end of which the rate of mortality varies from 5 to 20%. To these mortalities, it is expedient to add a variable proportion (5-10%) of cadaveric animals which are no longer able to present an economic future. It is to be noted that outside of this critical stage of the end of post-weaning, no anomaly appears on the farms. In particular, the reproductive function is totally maintained.

[0005] On the epidemiological level, the first signs of this pathology appeared at the start of 1995 in the east of the Côtes d'Armor region in France, and the farms affected are especially confined to this area of the region. In December 1996, the number of farms concerned could not be evaluated with precision because of the absence of a specific laboratory diagnostic method or of an epidemiological surveillance system of the livestock. Based on the clinical facts as well as on results of postmortem examinations supplied by veterinarians, it is possible to estimate this number as several dozen (80-100). The contagiousness of the disease is weak to moderate. Cases are being reported outside the initial area and for the majority are following the transfer of animals coming from farms familiar with the problem. On the other hand, a characteristic of the condition is its strong remanence. Thus, farms which have been affected for a year are still affected in spite of the massive administration of therapeutics. Farms with clinical expression are drawn from various categories of specialization (breeders/fatteners, post-weaners/ fatteners) and different economic structures are concerned. In addition, the disorders appear even in farms where the rules of animal husbandry are respected.

[0006] Numerous postmortem examinations have been carried out either on farms or in the laboratory. The elements of the lesional table are disparate. The most constant macroscopic lesions are pneumonia which sometimes appears in patchy form as well as hypertrophy of the lymphatic ganglia. The other lesions above all affect the thoracic viscera including, especially, pericarditis and pleurisy. However, arthritis and gastric ulcers are also observed. The lesions revealed in the histological examination are essentially situated at the pulmonary level (interstitial pneumonia), ganglionic level (lymphoid depletion of the lymph nodes, giant cells) and renal level (glomerulonephritis, vasculitis). The infectious agents have been the subject of wide research. It has been possible to exclude the intervention of pestiviruses and Aujeszky's disease. The disorders appear in the seropositive PDRS (Porcine Dysgenic and Respiratory Syndrome, an infection linked to an arteriovirus) herds, but it has not been possible to establish the role of the latter in the genesis of the disorders (the majority of the farms in Brittany are PDRS seropositive).

[0007] The inventors of the present invention, with the aim of identifying the etiological agent responsible for PWD, have carried out "contact" tests between piglets which are obviously "ill" and SPF pigs (specific pathogen-free) from

CNEVA (Centre National d'Etudes Vétérinaires et Alimentaires, France). These tests allow the development of signs comparable to those observed on the farm to be observed in protected animal houses. The discrete signs such as moderate hyperthermia, anorexia and intermittent diarrhea appeared after one week of contact. It must be noted that the PDRS virus only diffused subsequent to the clinical signs. In addition, inoculations of organ homogenates of sick animals to healthy pigs allowed signs related to those observed on the farms to be reproduced, although with a lower incidence, linked to the favorable conditions of upkeep of the animals in the experimental installations.

[0008] Thus, the inventors of the present invention have been able to demonstrate that the pathological signs appear as a well-defined entity affecting the pig at a particular stage of its growth.

[0009] This pathology has never been described in France. However, sparse information, especially Canadian, relates to similar facts.

[0010] The disorders cannot be mastered with the existing therapeutics.

[0011] The data collected both on the farm and by experimentation have allowed the following points to be high-lighted:

- **[0012]** PWD is transmissible but its contagiousness is not very high,
- **[0013]** its etiological origin is of infectious and probably viral nature,
- [0014] PWD has a persistent character in the affected farms.

[0015] Considerable economic consequences ensue for the farms.

[0016] Thus, there is currently a significant need for a specific and sensitive diagnostic, whose production is practical and rapid, allowing the early detection of the infection.

[0017] A reliable, sensitive and practical test which allows the distinction between strains of porcine circovirus (PCV) is thus strongly desirable.

[0018] On the other hand, a need for efficient and well-tolerated treatment of infections with PWD circovirus likewise remains desirable, no vaccine currently being available against PWD circovirus.

[0019] Concerning PWD circovirus, it will probably be necessary to understand the role of the immune defense in the physiology and the pathology of the disease to develop satisfactory vaccines.

[0020] Fuller information concerning the biology of these strains, their interactions with their hosts, the associated infectivity phenomena and those of escape from the immune defenses of the host especially, and finally their implication in the development of associated pathologies, will allow a better understanding of these mechanisms. Taking into account the facts which have been mentioned above and which show in particular the limitations of combating infection by the PWD circovirus, it is thus essential today on the one hand to develop molecular tools, especially starting from a better genetic knowledge of the PWD circovirus, and likewise to perfect novel preventive and therapeutic treatments, novel methods of diagnosis and specific, efficacious and tolerated novel vaccine strategies. This is precisely the subject of the present invention.

SUMMARY OF THE INVENTION

[0021] The present invention relates to vaccines comprising a nucleotide sequence of the genome of Porcine circovirus type B, or a homologue or fragment thereof, and an acceptable pharmaceutical or veterinary vehicle. In one embodiment of the invention, the nucleotide sequence is selected from SEQ ID No. 15, SEQ ID No. 19 SEQ ID No. 23, or SEQ ID No. 25, or a homologue or fragment thereof. In another embodiment of the invention, the homologue has at least 80% sequence identity to SEQ ID No. 15, SEQ ID No. 19, SEQ ID No. 23 or SEQ ID No. 25. In yet another embodiment, the vaccines further comprising an adjuvant

[0022] The present invention also relates to vaccines comprising a polypeptide encoded by a nucleotide sequence of the genome of PCVB, or a homologue or fragment thereof, and an acceptable pharmaceutical or veterinary vehicle. In one embodiment, the homologue has at least 80% sequence identity to SEQ ID No. 15, SEQ ID No. 19, SEQ ID No. 23 or SEQ ID No. 25. In another embodiment of the invention, the nucleotide sequence is selected from SEQ ID No. 23 or SEQ ID No. 25, or a homologue or fragment thereof. In still another embodiment, the polypeptide has the amino acid sequence of SEQ ID No. 24 or SEQ ID No. 26. In yet another embodiment, the homologue has at least 80% sequence identity to SEQ ID No. 24 or SEQ ID No. 26. In another embodiment, the polypeptide has the amino acid sequence of SEQ ID No. 24 or SEQ ID No. 26. In another embodiment, the polypeptide has the amino acid sequence of SEQ ID No. 29, SEQ ID No. 30, SEQ ID No. 31, or SEQ ID No. 32.

[0023] A further aspect of the invention relates to vaccines comprising a vector and an acceptable pharmaceutical or veterinary vehicle, the vector comprising a nucleotide sequence of the genome of Porcine circovirus type B, or a homologue or fragment thereof. In one embodiment, the vaccine further comprises a gene coding for an expression product capable of inhibiting or retarding the establishment or development of a genetic or acquired disease.

[0024] The present invention also relates to vaccines comprising a cell and an acceptable pharmaceutical or veterinary vehicle, wherein the cell is transformed with a nucleotide sequence of the genome of Porcine circovirus type B, or a homologue or fragment thereof.

[0025] Still further, the present invention relates to vaccines comprising a pharmaceutically acceptable vehicle and a single polypeptide, wherein the single polypeptide consists of SEQ ID No. 26.

[0026] Additionally, the present invention relates to methods of immunizing a mammal against piglet weight loss disease comprising administering to a mammal an effective amount of the vaccines described above.

[0027] These and other aspects of the invention will become apparent to the skilled artisan in view of the teachings contained herein.

BRIEF DESCRIPTION OF THE DRAWINGS

[0028] FIG. 1: Experimental scheme which has made it possible to bring about the isolation and the identification of the circovirus associated with PWD of type A and B.

[0029] Test 1: experimental reproduction of the PWD by inoculation of pig organ homogenates from farms affected by PWD.

[0030] Test 2: experimental reproduction of PWD.

[0031] Test 3: experimental reproduction of PWD.

[0032] Test 4: no experimental reproduction of PWD.

[0033] FIG. 2: Organization of the genome of the circovirus associated with PWD of type A (PCVA)

[0034] strand of (+) polarity (SEQ ID No. 1);

[0035] strand of (-) polarity (SEQ ID No. 5, represented according to the orientation $3' \rightarrow 5'$);

[0036] sequences of amino acids of proteins encoded by the two DNA strands in the three possible reading frames SEQ ID NOS: 2-4 and 6-8 respectively.

[0037] FIG. 3: Alignment of the nucleotide sequence SEQ ID No. 1 of the PWD circovirus of type A (PCVA) and of the MEEHAN SEQ ID No. 163 strain and MANKERTZ SEQ ID No. 164 strain circoviruses of the porcine cell lines.

[0038] FIG. 4: Alignment of the sequence of amino acids SEQ ID No. 10 of a polypeptide encoded by the nucleotide sequence SEQ ID No. 9 (ORF1) of the PWD circovirus of type A (PCVA) and of corresponding nucleotide sequences of the MEEHAN SEQ ID No. 165 strain and MANKERTZ SEQ ID No. 166 strain circoviruses of the porcine cell lines.

[0039] FIG. **5**: Alignment of the sequence of amino acids SEQ ID No. 12 of a polypeptide encoded by the nucleotide sequence SEQ ID No. 11 (ORF2) of the PWD circovirus of type A (PCVA) and of corresponding nucleotide sequences of the MEEHAN SEQ ID No. 167 strain and MANKERTZ SEQ ID No. 168 strain circoviruses of the porcine cell lines.

[0040] FIG. 6: Alignment of the sequence of amino acids SEQ ID No. 14 of a polypeptide encoded by the nucleotide sequence SEQ ID No. 13 (ORF3) of the PWD circovirus of type A (PCVA) and of corresponding nucleotide sequences of the MEEHAN SEQ ID No. 169 strain and MANKERTZ SEQ ID No. 170 strain circoviruses of the porcine cell lines.

[0041] FIG. 7: Western blot analysis of recombinant proteins of the PWD circovirus of type A (PCVA).

[0042] The analyses were carried out on cell extracts of Sf9 cells obtained after infection with recombinant baculovirus PCF ORF 1.

[0043] FIG. **8**: Organization of the genome of the circovirus associated with the PWD of type B (PCVB)

[0044] strand of (+) polarity (SEQ ID No. 15);

[0045] strand of (-) polarity (SEQ ID No. 19, represented according to the orientation $3' \rightarrow 5'$);

[0046] sequence of amino acids of proteins encoded by the two DNA strands in the three possible reading frames SEQ ID NOS: 16-18 and 20-22 respectively.

[0047] FIG. **9**: Evolution of the daily mean gain (DMG) of pig farms affected by piglet weight loss disease (PWD), placed under experimental conditions.

[0048] FIG. **10**: DMG compared for the 3 batches of pigs (F1, F3 and F4) calculated over a period of 28 days, after vaccination test.

[0049] FIG. **11**: Hyperthermia greater than 41° C., expressed as a percentage compared for the 3 batches of pigs (F1, F3 and F4) calculated per week over a period of 28 days, after vaccination test.

[0050] FIG. **12**: Membranes of peptide spots corresponding to the ORF2s revealed with the aid of an infected pig serum, originating from a conventional farm.

[0051] The numbers of specific peptides of the circovirus of type B as well as their nonreactive homologs (type A) are indicated in bold.

[0052] The nonspecific immunogenic peptides are indicated in italics.

[0053] FIG. **13**: Alignment of amino acid sequences of proteins encoded by the ORF2 of the PWD circovirus of type A SEQ ID No. 12 and by the ORF'2 of the PWD circovirus of type B SEQ ID No. 26. The position of 4 peptides corresponding to specific epitopes of the PWD circovirus of type B is indicated on the corresponding sequence by a bold line, their homolog on the sequence of the PWD circovirus of type A is likewise indicated by an ordinary line.

[0054] FIG. **14**: Charts the results of experiments that demonstrate, in terms of percent hyperthermia, that vaccination with ORF'1 and ORF'2 of PCV-B enhances the level of protection in swine challenged with PCV-B (Percent hyperthermia: >40.5 C, control: not vaccinated and not challenged, ORF'1: vaccinated and challenged, ORF'2: vaccinated and challenged, ORF: not vaccinated, challenged).

[0055] FIG. **15**: Charts the results of experiments that demonstrate, in terms of animal growth, that vaccination with ORF'1 and ORF'2 of PCV-B enhances the level of protection in swine challenged with PCV-B (Control: not vaccinated, not challenged, ORF'1: vaccinated and challenged, ORF'2: vaccinated and challenged, ORF: not vaccinated, challenged).

[0056] FIG. **16**: Immunoperoxidase staining of PK15 cells at 24 h post-transfection with the pcDNA3/ORF'2 plasmid. Expression of PCVB ORF'2 was confirmed by IPMA following incubation in the presence of the swine anti-PCVB monospecific serum.

DETAILED DESCRIPTION OF THE INVENTION

[0057] The present invention relates to nucleotide sequences of the genome of PWD circovirus selected from the sequences SEQ ID No. 1, SEQ ID No. 5, SEQ ID No. 15, SEQ ID No. 19 or one of their fragments.

[0058] The nucleotide sequences of sequences SEQ ID No. 1 and SEQ ID No. 5 correspond respectively to the genome sequence of the strand of (+) polarity and of the strand of (-) polarity of the PWD circovirus of type A (or PCVA), the sequence SEQ ID No. 5 being represented according to the orientation $5'\rightarrow 3'$.

[0059] The nucleotide sequences of sequences SEQ ID No. 15 and SEQ ID No. 19 correspond respectively to the genome sequence of the strand of (+) polarity and of the strand of (-) polarity of the PWD circovirus of type B (or PCVB), the sequence SEQ ID No. 19 being represented according to the orientation $5' \rightarrow 3'$.

[0060] The present invention likewise relates to nucleotide sequences, characterized in that they are selected from:

- [0061] a) a nucleotide sequence of a specific fragment of the sequence SEQ ID No. 1, SEQ ID No. 5, SEQ ID No. 15, SEQ ID No. 19 or one of their fragments;
- **[0062]** b) a nucleotide sequence homologous to a nucleotide sequence such as defined in a);
- [0063] c) a nucleotide sequence complementary to a nucleotide sequence such as defined in a) or b), and a nucleotide sequence of their corresponding RNA;
- **[0064]** d) a nucleotide sequence capable of hybridizing under stringent conditions with a sequence such as defined in a), b) or c);
- **[0065]** e) a nucleotide sequence comprising a sequence such as defined in a), b), c) or d); and
- **[0066]** f) a nucleotide sequence modified by a nucleotide sequence such as defined in a), b), c), d) or e).

[0067] Nucleotide, polynucleotide or nucleic acid sequence will be understood according to the present invention as meaning both a double-stranded or single-stranded DNA in the monomeric and dimeric (so-called in tandem) forms and the transcription products of said DNAs.

[0068] It must be understood that the present invention does not relate to the genomic nucleotide sequences taken in their natural environment, that is to say in the natural state. It concerns sequences which it has been possible to isolate, purify or partially purify, starting from separation methods such as, for example, ion-exchange chromatography, by exclusion based on molecular size, or by affinity, or alternatively fractionation techniques based on solubility in different solvents, or starting from methods of genetic engineering such as amplification, cloning and subcloning, it being possible for the sequences of the invention to be carried by vectors.

[0069] The nucleotide sequences SEQ ID No. 1 and SEQ ID No. 15 were obtained by sequencing of the genome by the Sanger method.

[0070] Nucleotide sequence fragment according to the invention will be understood as designating any nucleotide fragment of the PWD circovirus, type A or B, of length of at least 8 nucleotides, preferably at least 12 nucleotides, and even more preferentially at least 20 consecutive nucleotides of the sequence from which it originates.

[0071] Specific fragment of a nucleotide sequence according to the invention will be understood as designating any nucleotide fragment of the PWD circovirus, type A or B, having, after alignment and comparison with the corresponding fragments of known porcine circoviruses, at least one nucleotide or base of different nature. For example, the specific nucleotide fragments of the PWD circovirus of type A can easily be determined by referring to FIG. **3** of the present invention in which the nucleotides or bases of the sequence SEQ ID No. 1 (circopordfp) are shown which are of different nature, after alignment of said sequence SEQ ID No. 1 with the other two sequences of known porcine circovirus (circopormeeh and circopormank).

[0072] Homologous nucleotide sequence in the sense of the present invention is understood as meaning a nucleotide sequence having at least a percentage identity with the bases of a nucleotide sequence according to the invention of at least 80%, preferably 90% or 95%, this percentage being purely statistical and it being possible to distribute the differences between the two nucleotide sequences at random and over the whole of their length.

[0073] Specific homologous nucleotide sequence in the sense of the present invention is understood as meaning a homologous nucleotide sequence having at least one nucleotide sequence of a specific fragment, such as defined above. Said "specific" homologous sequences can comprise, for example, the sequences corresponding to the genomic sequence or to the sequences of its fragments representative of variants of PWD circovirus of type A or B. These specific homologous sequences can thus correspond to variations linked to mutations within strains of PWD circovirus of type A and B, and especially correspond to truncations, substitutions, deletions and/or additions of at least one nucleotide. Said homologous sequences can likewise correspond to variations linked to the degeneracy of the genetic code.

[0074] The term "degree or percentage of sequence homology" refers to "degree or percentage of sequence identity between two sequences after optimal alignment" as defined in the present application.

[0075] Two amino-acids or nucleotidic sequences are said to be "identical" if the sequence of amino-acids or nucleotidic residues, in the two sequences is the same when aligned for maximum correspondence as described below. Sequence comparisons between two (or more) peptides or polynucleotides are typically performed by comparing sequences of two optimally aligned sequences over a segment or "comparison window" to identify and compare local regions of sequence similarity. Optimal alignment of sequences for comparison may be conducted by the local homology algo-

rithm of Smith and Waterman, *Ad. App. Math* 2: 482 (1981), by the homology alignment algorithm of Neddleman and Wunsch, *J. Mol. Biol.* 48: 443 (1970), by the search for similarity method of Pearson and Lipman, *Proc. Natl. Acad. Sci.* (U.S.A.) 85: 2444 (1988), by computerized implementation of these algorithms (GAP, BESTFIT, FASTA, and TFASTA in the Wisconsin Genetics Software Package, Genetics Computer Group (GCG), 575 Science Dr., Madison, Wis.), or by visual inspection.

[0076] "Percentage of sequence identity" (or degree or identity) is determined by comparing two optimally aligned sequences over a comparison window, where the portion of the peptide or polynucleotide sequence in the comparison window may comprise additions or deletions (i.e., gaps) as compared to the reference sequence (which does not comprise additions or deletions) for optimal alignment of the two sequences. The percentage is calculated by determining the number of positions at which the identical amino-acid residue or nucleic acid base occurs in both sequences to yield the number of matched positions, dividing the number of matched positions by the total number of positions in the window of comparison and multiplying the result by 100 to yield the percentage of sequence identity.

[0077] The definition of sequence identity given above is the definition that would use one of skill in the art. The definition by itself does not need the help of any algorithm, said algorithms being helpful only to achieve the optimal alignments of sequences, rather than the calculation of sequence identity.

[0078] From the definition given above, it follows that there is a well defined and only one value for the sequence identity between two compared sequences which value corresponds to the value obtained for the best or optimal alignment.

[0079] In the BLAST N or BLAST P "BLAST 2 sequence", software which is available in the web site http://www.ncbi. nlm.nih.gov/gorf/b12.html, and habitually used by the inventors and in general by the skilled man for comparing and determining the identity between two sequences, gap cost which depends on the sequence length to be compared is directly selected by the software (i.e. 11.2 for substitution matrix BLOSUM-62 for length>85).

[0080] In the present description, PWD circovirus will be understood as designating the circoviruses associated with piglet weight loss disease (PWD) of type A (PCVA) or type B (PCVB), defined below by their genomic sequence, as well as the circoviruses whose nucleic sequences are homologous to the sequences of PWD circoviruses of type A or B, such as in particular the circoviruses corresponding to variants of the type A or of the type B.

[0081] Complementary nucleotide sequence of a sequence of the invention is understood as meaning any DNA whose nucleotides are complementary to those of the sequence of the invention, and whose orientation is reversed (antiparallel sequence).

[0082] Hybridization under conditions of stringency with a nucleotide sequence according to the invention is understood as meaning a hybridization under conditions of temperature and ionic strength chosen in such a way that they allow the maintenance of the hybridization between two fragments of complementary DNA.

[0083] By way of illustration, conditions of great stringency of the hybridization step with the aim of defining the nucleotide fragments described above are advantageously the following.

5

[0084] The hybridization is carried out at a preferential temperature of 65° C. in the presence of SSC buffer, 1×SSC corresponding to 0.15 M NaCl and 0.05 M Na citrate. The washing steps, for example, can be the following:

[0085] 2×SSC, at ambient temperature followed by two washes with 2×SSC, 0.5% SDS at 65° C.; 2×0.5×SSC, 0.5% SDS; at 65° C. for 10 minutes each.

[0086] The conditions of intermediate stringency, using, for example, a temperature of 42° C. in the presence of a 2×SSC buffer, or of less stringency, for example a temperature of 37° C. in the presence of a 2×SSC buffer, respectively require a globally less significant complementarity for the hybridization between the two sequences.

[0087] The stringent hybridization conditions described above for a polynucleotide with a size of approximately 350 bases will be adapted by the person skilled in the art for oligonucleotides of greater or smaller size, according to the teaching of Sambrook et al., 1989.

[0088] Among the nucleotide sequences according to the invention, those are likewise preferred which can be used as a primer or probe in methods allowing the homologous sequences according to the invention to be obtained, these methods, such as the polymerase chain reaction (PCR), nucleic acid cloning and sequencing, being well known to the person skilled in the art.

[0089] Among said nucleotide sequences according to the invention, those are again preferred which can be used as a primer or probe in methods allowing the presence of PWD circovirus or one of its variants such as defined below to be diagnosed.

[0090] The nucleotide sequences according to the invention capable of modulating, of inhibiting or of inducing the expression of PWD circovirus gene, and/or capable of modulating the replication cycle of PWD circovirus in the host cell and/or organism are likewise preferred. Replication cycle will be understood as designating the invasion and the multiplication of PWD circovirus, and its propagation from host cell to host cell in the host organism.

[0091] Among said nucleotide sequences according to the invention, those corresponding to open reading frames, called ORF sequences, and coding for polypeptides, such as, for example, the sequences SEQ ID No. 9 (ORF1), SEQ ID No. 11 (ORF2) and SEQ ID No. 13 (ORF3) respectively corresponding to the nucleotide sequences between the positions 47 and 985 determined with respect to the position of the nucleotides on the sequence SEQ ID No. 1, the positions 1723 and 1022 and the positions 658 and 38 with respect to the position of the nucleotides on the sequence SEQ ID No. 5 (represented according to the orientation $3' \rightarrow 5'$), the ends being included, or alternatively the sequences SEQ ID No. 23 (ORF'1), SEQ ID No. 25 (ORF'2) and SEQ ID No. 27 (ORF'3), respectively corresponding to the sequences between the positions 51 and 995 determined with respect to the position of the nucleotides on the sequence SEQ ID No. 15, the positions 1734 and 1033 and the positions 670 and 357, the positions being determined with respect to the position of the nucleotides on the sequence SEQ ID No. 19 (represented according to the orientation $3' \rightarrow 5'$), the ends being included, are finally preferred.

[0092] The nucleotide sequence fragments according to the invention can be obtained, for example, by specific amplification, such as PCR, or after digestion with appropriate restriction enzymes of nucleotide sequences according to the invention, these methods in particular being described in the

work of Sambrook et al., 1989. Said representative fragments can likewise be obtained by chemical synthesis when their size is not very large and according to methods well known to persons skilled in the art.

[0093] Modified nucleotide sequence will be understood as meaning any nucleotide sequence obtained by mutagenesis according to techniques well known to the person skilled in the art, and containing modifications with respect to the normal sequences according to the invention, for example mutations in the regulatory and/or promoter sequences of polypeptide expression, especially leading to a modification of the rate of expression of said polypeptide or to a modulation of the replicative cycle.

[0094] Modified nucleotide sequence will likewise be understood as meaning any nucleotide sequence coding for a modified polypeptide such as defined below.

[0095] The present invention relates to nucleotide sequences of PWD circovirus according to the invention, characterized in that they are selected from the sequences SEQ ID No. 9, SEQ ID No. 11, SEQ ID No. 13, SEQ ID No. 23, SEQ ID No. 25, SEQ ID No. 27 or one of their fragments. **[0096]** The invention likewise relates to nucleotide sequences characterized in that they comprise a nucleotide sequence selected from:

- [0097] a) a nucleotide sequence SEQ ID No. 9, SEQ ID No. 11, SEQ ID No. 13, SEQ ID No. 23, SEQ ID No. 25, SEQ ID No. 27 or one of their fragments;
- **[0098]** b) a nucleotide sequence of a specific fragment of a sequence such as defined in a);
- [0099] c) a homologous nucleotide sequence having at least 80% identity with a sequence such as defined in a) or b);
- **[0100]** d) a complementary nucleotide sequence or sequence of RNA corresponding to a sequence such as defined in a), b) or c); and
- **[0101]** e) a nucleotide sequence modified by a sequence such as defined in a), b), c) or d).

[0102] As far as homology with the nucleotide sequences SEQ ID No. 9, SEQ ID No. 11, SEQ ID No. 13, SEQ ID No. 23, SEQ ID No. 25, SEQ ID No. 27 or one of their fragments is concerned, the homologous, especially specific, sequences having a percentage identity with one of the sequences SEQ ID No. 9, SEQ ID No. 11, SEQ ID No. 13, SEQ ID No. 23, SEQ ID No. 25, SEQ ID No. 27 or one of their fragments of at least 80%, preferably 90% or 95%, are preferred. Said specific homologous sequences can comprise, for example, the sequences corresponding to the sequences ORF1, ORF2, ORF3, ORF'1, ORF'2 and ORF'3 of PWD circovirus variants of type A or of type B. In the same manner, these specific homologous sequences can correspond to variations linked to mutations within strains of PWD circovirus of type A or of type B and especially correspond to truncations, substitutions, deletions and/or additions of at least one nucleotide.

[0103] Among nucleotide sequences according to the invention, the sequence SEQ ID No. 23 which has a homology having more than 80% identity with the sequence SEQ ID No. 9, as well as the sequence SEQ ID No. 25, are especially preferred.

[0104] Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they comprise a nucleotide sequence selected from the following sequences:

a)	SEQ	ID	No.	33	170	5'	TG <u>T</u> GG <u>C</u> GA 3';
b)	SEQ	ID	No.	34	450	5'	AG <u>T</u> TTCCT 3';
c)	SEQ	ID	No.	35	1026	5'	T <u>C</u> ATTTAGAGGGTCTTT <u>C</u> AG
d)	SEQ	ID	No.	36	1074	5'	GTCA <u>A</u> CCT 3';
e)	SEQ	ID	No.	37	1101	5'	GTGG <u>T</u> TGC 3';
f)	SEQ	ID	No.	38	1123	5'	AGCC <u>C</u> AGG 3';
g)	SEQ	ID	No.	39	1192	5'	TTGG <u>C</u> TGG 3';
h)	SEQ	ID	No.	40	1218	5'	TC <u>T</u> AGCTC <u>T</u> GGT 3';
i)	SEQ	ID	No.	41	1501	5'	ATCT <u>C</u> AG <u>C</u> TCGT 3';
j)	SEQ	ID	No.	42	1536	5'	T <u>G</u> TCCTCCT <u>C</u> TT 3';
k)	SEQ	ID	No.	43	1563	5'	TCT <u>C</u> TAGA 3';
1)	SEQ	ID	No.	44	1623	5'	TGT <u>A</u> CCAA 3';
m)	SEQ	ID	No.	45	1686	5'	TCC <u>G</u> TCTT 3';

and their complementary sequences.

[0105] In the list of nucleotide sequences a)-m) above, the underlined nucleotides are mutated with respect to the two known sequences of circovirus which are nonpathogenic to pigs. The number preceding the nucleotide sequence represents the position of the first nucleotide of said sequence in the sequence SEQ ID No. 1.

[0106] The invention comprises the polypeptides encoded by a nucleotide sequence according to the invention, preferably a polypeptide whose sequence is represented by a fragment, especially a specific fragment, of one of the six sequences of amino acids represented in FIG. 2, these six amino acid sequences corresponding to the polypeptides which can be encoded according to one of the three possible reading frames of the sequence SEQ ID No. 1 or of the sequence SEQ ID No. 5, or a polypeptide whose sequence is represented by a fragment, especially a specific fragment, of one of the six sequences of amino acids shown in FIG. 8, these six sequences of amino acids corresponding to the polypeptides which can be encoded according to one of the three possible reading frames of the sequence SEQ ID No. 15 or of the sequence SEQ ID No. 19.

[0107] The invention likewise relates to the polypeptides, characterized in that they comprise a polypeptide selected from the amino acid sequences SEQ ID No. 10, SEQ ID No. 12, SEQ ID No. 14, SEQ ID No. 24, SEQ ID No. 26, SEQ ID No. 28 or one of their fragments.

[0108] Among the polypeptides according to the invention, the polypeptide of amino acid sequence SEQ ID No. 24 which has a homology having more than 80% identity with the sequence SEQ ID No. 10, as well as the polypeptide of sequence SEQ ID No. 26, are especially preferred.

[0109] The invention also relates to the polypeptides, characterized in that they comprise a polypeptide selected from:

- [0110] a) a specific fragment of at least 5 amino acids of a polypeptide of an amino acid sequence according to the invention;
- [0111] b) a polypeptide homologous to a polypeptide such as defined in a);
- [0112] c) a specific biologically active fragment of a polypeptide such as defined in a) or b); and

[0113] d) a polypeptide modified by a polypeptide such as defined in a), b) or c).

[0114] Among the polypeptides according to the invention, the polypeptides of amino acid sequences SEQ ID No. 29, SEQ ID No. 30, SEQ ID No. 31 and SEQ ID No. 32 are also preferred, these polypeptides being especially capable of specifically recognizing the antibodies produced during infection by the PWD circovirus of type B. These polypeptides thus have epitopes specific for the PWD circovirus of type B and can thus be used in particular in the diagnostic field or as immunogenic agent to confer protection in pigs against infection by PWD circovirus, especially of type B.

[0115] In the present description, the terms polypeptide, peptide and protein are interchangeable.

[0116] It must be understood that the invention does not relate to the polypeptides in natural form, that is to say that they are not taken in their natural environment but that they can be isolated or obtained by purification from natural sources, or else obtained by genetic recombination, or alternatively by chemical synthesis and that they can thus contain unnatural amino acids, as will be described below.

[0117] Polypeptide fragment according to the invention is understood as designating a polypeptide containing at least 5 consecutive amino acids, preferably 10 consecutive amino acids or 15 consecutive amino acids.

[0118] In the present invention, specific polypeptide fragment is understood as designating the consecutive polypeptide fragment encoded by a specific fragment nucleotide sequence according to the invention.

[0119] Homologous polypeptide will be understood as designating the polypeptides having, with respect to the natural polypeptide, certain modifications such as, in particular, a deletion, addition or substitution of at least one amino acid, a truncation, a prolongation, a chimeric fusion, and/or a mutation. Among the homologous polypeptides, those are preferred whose amino acid sequence has at least 80%, preferably 90%, homology with the sequences of amino acids of polypeptides according to the invention.

[0120] Specific homologous polypeptide will be understood as designating the homologous polypeptides such as defined above and having a specific fragment of polypeptide according to the invention.

[0121] In the case of a substitution, one or more consecutive or nonconsecutive amino acids are replaced by "equivalent" amino acids. The expression "equivalent" amino acid is directed here at designating any amino acid capable of being substituted by one of the amino acids of the base structure without, however, essentially modifying the biological activities of the corresponding peptides and such that they will be defined by the following.

[0122] These equivalent amino acids can be determined either by depending on their structural homology with the amino acids which they substitute, or on results of comparative tests of biological activity between the different polypeptides, which are capable of being carried out.

[0123] By way of example, the possibilities of substitutions capable of being carried out without resulting in an extensive modification of the biological activity of the corresponding modified polypeptides will be mentioned, the replacement, for example, of leucine by valine or isoleucine, of aspartic acid by glutamic acid, of glutamine by asparagine, of arginine by lysine etc., the reverse substitutions naturally being envisageable under the same conditions.

3';

[0124] The specific homologous polypeptides likewise correspond to polypeptides encoded by the specific homologous nucleotide sequences such as defined above and thus comprise in the present definition the polypeptides which are mutated or correspond to variants which can exist in PWD circovirus, and which especially correspond to truncations, substitutions, deletions and/or additions of at least one amino acid residue.

[0125] Specific biologically active fragment of a polypeptide according to the invention will be understood in particular as designating a specific polypeptide fragment, such as defined above, having at least one of the characteristics of polypeptides according to the invention, especially in that it is:

- [0126] capable of inducing an immunogenic reaction directed against a PWD circovirus; and/or
- [0127] capable of being recognized by a specific anti-
- body of a polypeptide according to the invention; and/or [0128] capable of linking to a polypeptide or to a nucle-
- otide sequence of PWD circovirus; and/or
- **[0129]** capable of exerting a physiological activity, even partial, such as, for example, a dissemination or structural (capsid) activity; and/or
- **[0130]** capable of modulating, of inducing or of inhibiting the expression of PWD circovirus gene or one of its variants, and/or capable of modulating the replication cycle of PWD circovirus in the cell and/or the host organism.

[0131] The polypeptide fragments according to the invention can correspond to isolated or purified fragments naturally present in a PWD circovirus or correspond to fragments which can be obtained by cleavage of said polypeptide by a proteolytic enzyme, such as trypsin or chymotrypsin or collagenase, or by a chemical reagent, such as cyanogen bromide (CNBr) or alternatively by placing said polypeptide in a very acidic environment, for example at pH 2.5. Such polypeptide fragments can likewise just as easily be prepared by chemical synthesis, from hosts transformed by an expression vector according to the invention containing a nucleic acid allowing the expression of said fragments, placed under the control of appropriate regulation and/or expression elements.

[0132] "Modified polypeptide" of a polypeptide according to the invention is understood as designating a polypeptide obtained by genetic recombination or by chemical synthesis as will be described below, having at least one modification with respect to the normal sequence. These modifications will especially be able to bear on amino acids at the origin of a specificity, of pathogenicity and/or of virulence, or at the origin of the structural conformation, and of the capacity of membrane insertion of the polypeptide according to the invention. It will thus be possible to create polypeptides of equivalent, increased or decreased activity, and of equivalent, narrower, or wider specificity. Among the modified polypeptides, it is necessary to mention the polypeptides in which up to 5 amino acids can be modified, truncated at the N- or C-terminal end, or even deleted or added.

[0133] As is indicated, the modifications of the polypeptide will especially have as objective:

- **[0134]** to render it capable of modulating, of inhibiting or of inducing the expression of PWD circovirus gene and/ or capable of modulating the replication cycle of PWD circovirus in the cell and/or the host organism,
- [0135] of allowing its incorporation into vaccine compositions,
- **[0136]** of modifying its bioavailability as a compound for therapeutic use.

[0137] The methods allowing said modulations on eukaryotic or prokaryotic cells to be demonstrated are well known to the person skilled in the art. It is likewise well understood that it will be possible to use the nucleotide sequences coding for said modified polypeptides for said modulations, for example through vectors according to the invention and described below, in order, for example, to prevent or to treat the pathologies linked to the infection.

[0138] The preceding modified polypeptides can be obtained by using combinatorial chemistry, in which it is possible to systematically vary parts of the polypeptide before testing them on models, cell cultures or microorganisms for example, to select the compounds which are most active or have the properties sought.

[0139] Chemical synthesis likewise has the advantage of being able to use:

- [0140] unnatural amino acids, or
- [0141] nonpeptide bonds.

[0142] Thus, in order to improve the duration of life of the polypeptides according to the invention, it may be of interest to use unnatural amino acids, for example in D form, or else amino acid analogs, especially sulfur-containing forms, for example.

[0143] Finally, it will be possible to integrate the structure of the polypeptides according to the invention, its specific or modified homologous forms, into chemical structures of polypeptide type or others. Thus, it may be of interest to provide at the N- and C-terminal ends compounds not recognized by the proteases.

[0144] The nucleotide sequences coding for a polypeptide according to the invention are likewise part of the invention. **[0145]** The invention likewise relates to nucleotide sequences utilizable as a primer or probe, characterized in that said sequences are selected from the nucleotide sequences according to the invention.

[0146] Among the pairs of nucleotide sequences utilizable as a pair of primers according to the invention, the pairs of primers selected from the following pairs are preferred:

a) SEQ ID No. 46 5' GTG TGC TCG ACA TTG GTG TG 3', and SEQ ID No. 47 5' TGG AAT GTT AAC GAG CTG AG 3';
b) SEQ ID No. 46 5' GTG TGC TCG ACA TTG GTG TG 3', and SEQ ID No. 48 5' CTC GCA GCC ATC TTG GAA TG 3';
c) SEQ ID No. 49 5' CGC GCG TAA TAC GAC TCA CT 3', and SEQ ID No. 46 5' GTG TGC TCG ACA TTG GTG TG 3';
d) SEQ ID No. 49 5' CGC GCG TAA TAC GAC TCA CT 3', and

			- (cont	ιnu	led					
SEO ID No. 48	5'	CTC	GCA	GCC	ATC	TTG	GAA	TG	31:	and	

e) SEQ ID No. 50 5' CCT GTC TAC TGC TGT GAG TAC CTT GT 3', and SEQ ID No. 51 $\,$ 5' GCA GTA GAC AGG TCA CTC CGT TGT CC 3'.

. .

[0147] The cloning and the sequencing of the PWD circovirus, type A and B, has allowed it to be identified, after comparative analysis with the nucleotide sequences of other porcine circoviruses, that, among the sequences of fragments of these nucleic acids, were those which are strictly specific to the PWD circovirus of type A, of type B or of type A and B, and those which correspond to a consensus sequence of porcine circoviruses other than the PWD circoviruses of type A and/or B.

[0148] There is likewise a great need for nucleotide sequences utilizable as a primer or probe specific to the whole of the other known and nonpathogenic porcine circoviruses. **[0149]** Said consensus nucleotide sequences specific to all circoviruses, other than PWD circovirus of type A and B, are easily identifiable from FIG. **3** and the sequence SEQ ID No. 15, and are part of the invention.

[0150] Among said consensus nucleotide sequences, that which is characterized in that it is part of the following pair of primers is preferred:

a) SEQ ID No. 46 5' GTG TGC TCG ACA TTG GTG TG 3', and SEQ ID No. 52 5' TGG AAT GTT AAC TAC CTC AA 3'.

[0151] The invention likewise comprises a nucleotide sequence according to the invention, characterized in that said sequence is a specific consensus sequence of porcine circovirus other than PWD circovirus of type B and in that it is one of the primers of the following pairs of primers:

a)	SEQ	ID	No.	53	5'	GGC	GGC	GCC	ATC	TGT	AAC	GGT
					ΤТ	3',	and					
SEÇ) ID	No.	54		5'	GAT	GGC	GCC	GAA	AGA	CGG	GTA
					TС	3'.						

[0152] It is well understood that the present invention likewise relates to specific polypeptides of known porcine circoviruses other than PWD circovirus, encoded by said consensus nucleotide sequences, capable of being obtained by purification from natural polypeptides, by genetic recombination or by chemical synthesis by procedures well known to the person skilled in the art and such as described in particular below. In the same manner, the labeled or unlabeled mono- or polyclonal antibodies directed against said specific polypeptides encoded by said consensus nucleotide sequences are also part of the invention.

[0153] It will be possible to use said consensus nucleotide sequences, said corresponding polypeptides as well as said antibodies directed against said polypeptides in procedures or sets for detection and/or identification such as described below, in place of or in addition to nucleotide sequences, polypeptides or antibodies according to the invention, specific to PWD circovirus type A and/or B.

[0154] These protocols have been improved for the differential detection of the circular monomeric forms of specific replicative forms of the virion or of the DNA in replication and the dimeric forms found in so-called in-tandem molecular constructs. **[0155]** The invention additionally relates to the use of a nucleotide sequence according to the invention as a primer or probe for the detection and/or the amplification of nucleic acid sequences.

[0156] The nucleotide sequences according to the invention can thus be used to amplify nucleotide sequences, especially by the PCR technique (polymerase chain reaction) (Erlich, 1989; Innis et al., 1990; Rolfs et al., 1991; and White et al., 1997).

[0157] These oligodeoxyribonucleotide or oligoribonucleotide primers advantageously have a length of at least 8 nucleotides, preferably of at least 12 nucleotides, and even more preferentially at least 20 nucleotides.

[0158] Other amplification techniques of the target nucleic acid can be advantageously employed as alternatives to PCR. **[0159]** The nucleotide sequences of the invention, in particular the primers according to the invention, can likewise be employed in other procedures of amplification of a target nucleic acid, such as:

- [0160] the TAS technique (Transcription-based Amplification System), described by Kwoh et al. in 1989;
- **[0161]** the 3SR technique (Self-Sustained Sequence Replication), described by Guatelli et al. in 1990;
- **[0162]** the NASBA technique (Nucleic Acid Sequence Based Amplification), described by Kievitis et al. in 1991;
- **[0163]** the SDA technique (Strand Displacement Amplification) (Walker et al., 1992);
- **[0164]** the TMA technique (Transcription Mediated Amplification).

[0165] The polynucleotides of the invention can also be employed in techniques of amplification or of modification of the nucleic acid serving as a probe, such as:

- **[0166]** the LCR technique (Ligase Chain Reaction), described by Landegren et al. in 1988 and improved by Barany et al. in 1991, which employs a thermostable ligase;
- **[0167]** the RCR technique (Repair Chain Reaction), described by Segev in 1992;
- **[0168]** the CPR technique (Cycling Probe Reaction), described by Duck et al. in 1990;
- **[0169]** the amplification technique with Q-beta replicase, described by Miele et al. in 1983 and especially improved by Chu et al. in 1986, Lizardi et al. in 1988, then by Burg et al. as well as by Stone et al. in 1996.

[0170] In the case where the target polynucleotide to be detected is possibly an RNA, for example an mRNA, it will be possible to use, prior to the employment of an amplification reaction with the aid of at least one primer according to the invention or to the employment of a detection procedure with the aid of at least one probe of the invention, an enzyme of reverse transcriptase type in order to obtain a cDNA from the RNA contained in the biological sample. The cDNA obtained will thus serve as a target for the primer(s) or the probe(s) employed in the amplification or detection procedure according to the invention.

[0171] The detection probe will be chosen in such a manner that it hybridizes with the target sequence or the amplicon generated from the target sequence. By way of sequence, such a probe will advantageously have a sequence of at least 12 nucleotides, in particular of at least 20 nucleotides, and preferably of at least 100 nucleotides.

[0172] The invention also comprises the nucleotide sequences utilizable as a probe or primer according to the invention, characterized in that they are labeled with a radioactive compound or with a nonradioactive compound.

[0173] The unlabeled nucleotide sequences can be used directly as probes or primers, although the sequences are generally labeled with a radioactive element $(^{32}P, ^{35}S, ^{3}H, ^{125}D)$

 125 I) or with a nonradioactive molecule (biotin, acetylaminofluorene, digoxigenin, 5-bromodeoxyuridine, fluorescein) to obtain probes which are utilizable for numerous applications.

[0174] Examples of nonradioactive labeling of nucleotide sequences are described, for example, in French Patent No. 78.10975 or by Urdea et al. or by Sanchez-Pescador et al. in 1988.

[0175] In the latter case, it will also be possible to use one of the labeling methods described in patents FR-2 422 956 and FR-2 518 755.

[0176] The hybridization technique can be carried out in various manners (Matthews et al., 1988). The most general method consists in immobilizing the nucleic acid extract of cells on a support (such as nitrocellulose, nylon, polystyrene) and in incubating, under well-defined conditions, the immobilized target nucleic acid with the probe. After hybridization, the excess of probe is eliminated and the hybrid molecules formed are detected by the appropriate method (measurement of the radioactivity, of the fluorescence or of the enzymatic activity linked to the probe).

[0177] The invention likewise comprises the nucleotide sequences according to the invention, characterized in that they are immobilized on a support, covalently or noncovalently.

[0178] According to another advantageous mode of employing nucleotide sequences according to the invention, the latter can be used immobilized on a support and can thus serve to capture, by specific hybridization, the target nucleic acid obtained from the biological sample to be tested. If necessary, the solid support is separated from the sample and the hybridization complex formed between said capture probe and the target nucleic acid is then detected with the aid of a second probe, a so-called detection probe, labeled with an easily detectable element.

[0179] Another subject of the present invention is a vector for the cloning and/or expression of a sequence, characterized in that it contains a nucleotide sequence according to the invention.

[0180] The vectors according to the invention, characterized in that they contain the elements allowing the expression and/or the secretion of said nucleotide sequences in a determined host cell, are likewise part of the invention.

[0181] The vector must then contain a promoter, signals of initiation and termination of translation, as well as appropriate regions of regulation of transcription. It must be able to be maintained stably in the host cell and can optionally have particular signals specifying the secretion of the translated protein. These different elements are chosen as a function of the host cell used. To this end, the nucleotide sequences

according to the invention can be inserted into autonomous replication vectors within the chosen host, or integrated vectors of the chosen host.

[0182] Such vectors will be prepared according to the methods currently used by the person skilled in the art, and it will be possible to introduce the clones resulting therefrom into an appropriate host by standard methods, such as, for example, lipofection, electroporation and thermal shock.

[0183] The vectors according to the invention are, for example, vectors of plasmid or viral origin.

[0184] A preferred vector for the expression of polypeptides of the invention is baculovirus.

[0185] The vector pBS KS in which is inserted the intandem DNA sequence of the PWD circovirus type A (or DFP) as deposited at the CNCM on 3 Jul. 1997, under the number I-1891, is likewise preferred.

[0186] These vectors are useful for transforming host cells in order to clone or to express the nucleotide sequences of the invention.

[0187] The invention likewise comprises the host cells transformed by a vector according to the invention.

[0188] These cells can be obtained by the introduction into host cells of a nucleotide sequence inserted into a vector such as defined above, then the culturing of said cells under conditions allowing the replication and/or expression of the transfected nucleotide sequence.

[0189] The host cell can be selected from prokaryotic or eukaryotic systems, such as, for example, bacterial cells (Olins and Lee, 1993), but likewise yeast cells (Buckholz, 1993), as well as animal cells, in particular the cultures of mammalian cells (Edwards and Aruffo, 1993), and especially Chinese hamster ovary (CHO) cells, but likewise the cells of insects in which it is possible to use procedures employing baculoviruses, for example (Luckow, 1993).

[0190] A preferred host cell for the expression of the proteins of the invention is constituted by sf9 insect cells.

[0191] A more preferred host cell according to the invention is *E. coli*, such as deposited at the CNCM on 3 Jul. 1997, under the number I-1891.

[0192] The invention likewise relates to animals comprising one of said transformed cells according to the invention. [0193] The obtainment of transgenic animals according to the invention overexpres sing one or more of the genes of PWD circovirus or part of the genes will be preferably carried out in rats, mice or rabbits according to methods well known to the person skilled in the art, such as by viral or nonviral transfections. It will be possible to obtain the transgenic animals overexpressing one or more of said genes by transfection of multiple copies of said genes under the control of a strong promoter of ubiquitous nature, or selective for one type of tissue. It will likewise be possible to obtain the transgenic animals by homologous recombination in embryonic cell strains, transfer of these cell strains to embryos, selection of the affected chimeras at the level of the reproductive lines, and growth of said chimeras.

[0194] The transformed cells as well as the transgenic animals according to the invention are utilizable in procedures for preparation of recombinant polypeptides.

[0195] It is today possible to produce recombinant polypeptides in relatively large quantity by genetic engineering using the cells transformed by expression vectors according to the invention or using transgenic animals according to the invention.

[0196] The procedures for preparation of a polypeptide of the invention in recombinant form, characterized in that they employ a vector and/or a cell transformed by a vector according to the invention and/or a transgenic animal comprising one of said transformed cells according to the invention, are themselves comprised in the present invention.

[0197] Among said procedures for preparation of a polypeptide of the invention in recombinant form, the preparation procedures employing a vector, and/or a cell transformed by said vector and/or a transgenic animal comprising one of said transformed cells, containing a nucleotide sequence according to the invention coding for a polypeptide of PWD circovirus, are preferred.

[0198] The recombinant polypeptides obtained as indicated above can just as well be present in glycosylated form as in nonglycosylated form and can or cannot have the natural tertiary structure.

[0199] A preferred variant consists in producing a recombinant polypeptide used to a "carrier" protein (chimeric protein). The advantage of this system is that it allows a stabilization of and a decrease in the proteolysis of the recombinant product, an increase in the solubility in the course of renaturation in vitro and/or a simplification of the purification when the fusion partner has an affinity for a specific ligand.

[0200] More particularly, the invention relates to a procedure for preparation of a polypeptide of the invention comprising the following steps:

- **[0201]** a) culture of transformed cells under conditions allowing the expression of a recombinant polypeptide of nucleotide sequence according to the invention;
- **[0202]** b) if need be, recovery of said recombinant polypeptide.

[0203] When the procedure for preparation of a polypeptide of the invention employs a transgenic animal according to the invention, the recombinant polypeptide is then extracted from said animal.

[0204] The invention also relates to a polypeptide which is capable of being obtained by a procedure of the invention such as described previously.

[0205] The invention also comprises a procedure for preparation of a synthetic polypeptide, characterized in that it uses a sequence of amino acids of polypeptides according to the invention.

[0206] The invention likewise relates to a synthetic polypeptide obtained by a procedure according to the invention.

[0207] The polypeptides according to the invention can likewise be prepared by techniques which are conventional in the field of the synthesis of peptides. This synthesis can be carried out in homogeneous solution or in solid phase.

[0208] For example, recourse can be made to the technique of synthesis in homogeneous solution described by Houben-Weyl in 1974.

[0209] This method of synthesis consists in successively condensing, two by two, the successive amino acids in the order required, or in condensing amino acids and fragments formed previously and already containing several amino acids in the appropriate order, or alternatively several fragments previously prepared in this way, it being understood that it will be necessary to protect beforehand all the reactive functions carried by these amino acids or fragments, with the exception of amine functions of one and carboxyls of the other or vice-versa, which must normally be involved in the

formation of peptide bonds, especially after activation of the carboxyl function, according to the methods well known in the synthesis of peptides.

[0210] According to another preferred technique of the invention, recourse will be made to the technique described by Merrifield.

[0211] To make a peptide chain according to the Merrifield procedure, recourse is made to a very porous polymeric resin, on which is immobilized the first C-terminal amino acid of the chain. This amino acid is immobilized on a resin through its carboxyl group and its amine function is protected. The amino acids which are going to form the peptide chain are thus immobilized, one after the other, on the amino group, which is deprotected beforehand each time, of the portion of the peptide chain already formed, and which is attached to the resin. When the whole of the desired peptide chain has been formed, the protective groups of the different amino acids forming the peptide chain are eliminated and the peptide is detached from the resin with the aid of an acid.

[0212] The invention additionally relates to hybrid polypeptides having at least one polypeptide according to the invention, and a sequence of a polypeptide capable of inducing an immune response in man or animals.

[0213] Advantageously, the antigenic determinant is such that it is capable of inducing a humoral and/or cellular response.

[0214] It will be possible for such a determinant to comprise a polypeptide according to the invention in glycosylated form used with a view to obtaining immunogenic compositions capable of inducing the synthesis of antibodies directed against multiple epitopes. Said polypeptides or their glycosylated fragments are likewise part of the invention.

[0215] These hybrid molecules can be formed, in part, of a polypeptide carrier molecule or of fragments thereof according to the invention, associated with a possibly immunogenic part, in particular an epitope of the diphtheria toxin, the tetanus toxin, a surface antigen of the hepatitis B virus (patent FR 79 21811), the VP1 antigen of the poliomyelitis virus or any other viral or bacterial toxin or antigen.

[0216] The procedures for synthesis of hybrid molecules encompass the methods used in genetic engineering for constructing hybrid nucleotide sequences coding for the polypeptide sequences sought. It will be possible, for example, to refer advantageously to the technique for obtainment of genes coding for fusion proteins described by Minton in 1984.

[0217] Said hybrid nucleotide sequences coding for a hybrid polypeptide as well as the hybrid polypeptides according to the invention characterized in that they are recombinant polypeptides obtained by the expression of said hybrid nucleotide sequences are likewise part of the invention.

[0218] The invention likewise comprises the vectors characterized in that they contain one of said hybrid nucleotide sequences. The host cells transformed by said vectors, the transgenic animals comprising one of said transformed cells as well as the procedures for preparation of recombinant polypeptides using said vectors, said transformed cells and/or said transgenic animals are, of course, likewise part of the invention.

[0219] The polypeptides according to the invention, the antibodies according to the invention described below and the nucleotide sequences according to the invention can advantageously be employed in procedures for the detection and/or identification of PWD circovirus, or of porcine circovirus

11

other than a PWD circovirus, in a biological sample (biological tissue or fluid) capable of containing them. These procedures, according to the specificity of the polypeptides, the antibodies and the nucleotide sequences according to the invention which will be used, will in particular be able to detect and/or to identify a PWD circovirus or a porcine circovirus other than a PWD circovirus or other than the PWD circovirus of type B.

[0220] The polypeptides according to the invention can advantageously be employed in a procedure for the detection and/or the identification of PWD circovirus of type A, of type B, of type A or B, or porcine circovirus other than the PWD circovirus of type B, or of porcine circovirus other than the PWD circovirus of type A or B, in a biological sample (biological tissue or fluid) capable of containing them, characterized in that it comprises the following steps:

- **[0221]** a) contacting of this biological sample with a polypeptide or one of its fragments according to the invention (under conditions allowing an immunological reaction between said polypeptide and the antibodies possibly present in the biological sample);
- **[0222]** b) demonstration of the antigen-antibody complexes possibly formed.

[0223] In the present description, PWD circovirus, except if a particular mention is indicated, will be understood as designating a PWD circovirus of type A or of type B, and porcine circovirus other than PWD, except if a particular mention is indicated, will be understood as designating a porcine circovirus other than a PWD circovirus of type A and B.

[0224] Preferably, the biological sample is formed by a fluid, for example a pig serum, whole blood or biopsies.

[0225] Any conventional procedure can be employed for carrying out such a detection of the antigen-antibody complexes possibly formed.

[0226] By way of example, a preferred method brings into play immunoenzymatic processes according to the ELISA technique, by immunofluorescence, or radioimmunological processes (RIA) or their equivalent.

[0227] Thus, the invention likewise relates to the polypeptides according to the invention, labeled with the aid of an adequate label such as of the enzymatic, fluorescent or radioactive type.

[0228] Such methods comprise, for example, the following steps:

- **[0229]** deposition of determined quantities of a polypeptide composition according to the invention in the wells of a microtiter plate,
- **[0230]** introduction into said wells of increasing dilutions of serum, or of a biological sample other than that defined previously, having to be analyzed,
- [0231] incubation of the microplate,
- **[0232]** introduction into the wells of the microtiter plate of labeled antibodies directed against pig immunoglobulins, the labeling of these antibodies having been carried out with the aid of an enzyme selected from those which are capable of hydrolyzing a substrate by modifying the absorption of the radiation of the latter, at least at a determined wavelength, for example at 550 nm,
- **[0233]** detection, by comparison with a control test, of the quantity of hydrolyzed substrate.

[0234] The invention likewise relates to a kit or set for the detection and/or identification of PWD circovirus, of porcine circovirus other than a PWD circovirus or of porcine circovi-

rus other than the PWD circovirus of type B, characterized in that it comprises the following elements:

- [0235] a polypeptide according to the invention,
- **[0236]** if need be, the reagents for the formation of the medium favorable to the immunological or specific reaction,
- **[0237]** if need be, the reagents allowing the detection of the antigen-antibody complexes produced by the immunological reaction between the polypeptide(s) of the invention and the antibodies possibly present in the biological sample, these reagents likewise being able to carry a label, or to be recognized in their turn by a labeled reagent, more particularly in the case where the polypeptide according to the invention is not labeled,
- **[0238]** if need be, a biological reference sample (negative control) devoid of antibodies recognized by a polypeptide according to the invention,
- **[0239]** if need be, a biological reference sample (positive control) containing a predetermined quantity of antibodies recognized by a polypeptide according to the invention.

[0240] The polypeptides according to the invention allow monoclonal or polyclonal antibodies to be prepared which are characterized in that they specifically recognize the polypeptides according to the invention. It will advantageously be possible to prepare the monoclonal antibodies from hybridomas according to the technique described by Kohler and Milstein in 1975. It will be possible to prepare the polyclonal antibodies, for example, by immunization of an animal, in particular a mouse, with a polypeptide or a DNA, according to the invention, associated with an adjuvant of the immune response, and then purification of the specific antibodies contained in the serum of the immunized animals on an affinity column on which the polypeptide which has served as an antigen has previously been immobilized. The polyclonal antibodies according to the invention can also be prepared by purification, on an affinity column on which a polypeptide according to the invention has previously been immobilized, of the antibodies contained in the serum of pigs infected by a PWD circovirus.

[0241] The invention likewise relates to mono- or polyclonal antibodies or their fragments, or chimeric antibodies, characterized in that they are capable of specifically recognizing a polypeptide according to the invention.

[0242] It will likewise be possible for the antibodies of the invention to be labeled in the same manner as described previously for the nucleic probes of the invention, such as a labeling of enzymatic, fluorescent or radioactive type.

[0243] The invention is additionally directed at a procedure for the detection and/or identification of PWD circovirus, of porcine circovirus other than a PWD circovirus, or other than the PWD circovirus of type B, in a biological sample, characterized in that it comprises the following steps:

- **[0244]** a) contacting of the biological sample (biological tissue or fluid) with a mono- or polyclonal antibody according to the invention (under conditions allowing an immunological reaction between said antibodies and the polypeptides of PWD circovirus, of porcine circovirus other than a PWD circovirus, of porcine circovirus other than the PWD circovirus of type B, possibly present in the biological sample);
- **[0245]** b) demonstration of the antigen-antibody complex possibly formed.

[0246] Likewise within the scope of the invention is a kit or set for the detection and/or the identification of PWD circovirus, of porcine circovirus other than a PWD circovirus or of porcine circovirus other than the PWD circovirus of type B, characterized in that it comprises the following components:

- **[0247]** a polyclonal or monoclonal antibody according to the invention, if need be labeled;
- **[0248]** if need be, a reagent for the formation of the medium favorable to the carrying out of the immuno-logical reaction;
- **[0249]** if need be, a reagent allowing the detection of the antigen-antibody complexes produced by the immunological reaction, this reagent likewise being able to carry a label, or being capable of being recognized in its turn by a labeled reagent, more particularly in the case where said monoclonal or polyclonal antibody is not labeled;
- **[0250]** if need be, reagents for carrying out the lysis of cells of the sample tested.

[0251] The present invention likewise relates to a procedure for the detection and/or the identification of PWD, of porcine circovirus other than a PWD circovirus or of porcine circovirus other than the PWD circovirus of type B, in a biological sample, characterized in that it employs a nucleotide sequence according to the invention.

[0252] More particularly, the invention relates to a procedure for the detection and/or the identification of PWD circovirus, of porcine circovirus other than a PWD circovirus or of porcine circovirus other than the PWD circovirus of type B, in a biological sample, characterized in that it contains the following steps:

[0253] a) if need be, isolation of the DNA from the biological sample to be analyzed;

- **[0254]** b) specific amplification of the DNA of the sample with the aid of at least one primer, or a pair of primers, according to the invention;
- **[0255]** c) demonstration of the amplification products.

[0256] These can be detected, for example, by the technique of molecular hybridization utilizing a nucleic probe according to the invention. This probe will advantageously be labeled with a nonradioactive (cold probe) or radioactive element.

[0257] For the purposes of the present invention, "DNA of the biological sample" or "DNA contained in the biological sample" will be understood as meaning either the DNA present in the biological sample considered, or possibly the cDNA obtained after the action of an enzyme of reverse transcriptase type on the RNA present in said biological sample.

[0258] Another aim of the present invention consists in a procedure according to the invention, characterized in that it comprises the following steps:

- **[0259]** a) contacting of a nucleotide probe according to the invention with a biological sample, the DNA contained in the biological sample having, if need be, previously been made accessible to hybridization under conditions allowing the hybridization of the probe with the DNA of the sample;
- **[0260]** b) demonstration of the hybrid formed between the nucleotide probe and the DNA of the biological sample.

[0261] The present invention also relates to a procedure according to the invention, characterized in that it comprises the following steps:

- **[0262]** a) contacting of a nucleotide probe immobilized on a support according to the invention with a biological sample, the DNA of the sample having, if need be, previously been made accessible to hybridization, under conditions allowing the hybridization of the probe with the DNA of the sample;
- **[0263]** b) contacting of the hybrid formed between the nucleotide probe immobilized on a support and the DNA contained in the biological sample, if need be after elimination of the DNA of the biological sample which has not hybridized with the probe, with a nucleotide probe labeled according to the invention;
- **[0264]** c) demonstration of the novel hybrid formed in step b).

[0265] According to an advantageous embodiment of the procedure for detection and/or identification defined previously, this is characterized in that, prior to step a), the DNA of the biological sample is first amplified with the aid of at least one primer according to the invention.

[0266] The invention is additionally directed at a kit or set for the detection and/or the identification of PWD circovirus, of porcine circovirus other than the PWD circovirus or of porcine circovirus other than the PWD circovirus of type B, characterized in that it comprises the following elements:

- [0267] a) a nucleotide probe according to the invention;[0268] b) if need be, the reagents necessary for the car-
- rying out of a hybridization reaction; [0269] c) if need be, at least one primer according to the
- invention as well as the reagents necessary for an amplification reaction of the DNA.

[0270] The invention likewise relates to a kit or set for the detection and/or the identification of PWD circovirus, of porcine circovirus other than a PWD circovirus or of porcine circovirus other than the PWD circovirus of type B, characterized in that it comprises the following components:

- **[0271]** a) a nucleotide probe, called a capture probe, according to the invention;
- **[0272]** b) an oligonucleotide probe, called a revealing probe, according to the invention,
- [0273] c) if need be, at least one primer according to the invention, as well as the reagents necessary for an amplification reaction of the DNA.

[0274] The invention also relates to a kit or set for the detection and/or identification of PWD circovirus, of porcine circovirus other than a PWD circovirus or of porcine circovirus other than the PWD circovirus of type B, characterized in that it comprises the following elements:

- [0275] a) at least one primer according to the invention;[0276] b) if need be, the reagents necessary for carrying out a DNA amplification reaction;
- **[0277]** c) if need be, a component allowing the sequence of the amplified fragment to be verified, more particularly an oligonucleotide probe according to the invention.

[0278] The invention additionally relates to the use of a nucleotide sequence according to the invention, of a polypeptide according to the invention, of an antibody according to the invention, of a cell according to the invention, and/or of an animal transformed according to the invention, for the selection of an organic or inorganic compound capable of modulating, inducing or inhibiting the expression of genes, and/or of modifying the cellular replication of PWD circovirus or capable of inducing or of inhibiting the pathologies linked to an infection by a PWD circovirus.

[0279] The invention likewise comprises a method of selection of compounds capable of binding to a polypeptide or one of its fragments according to the invention, capable of binding to a nucleotide sequence according to the invention, or capable of recognizing an antibody according to the invention, and/or capable of modulating, inducing or inhibiting the expression of genes, and/or of modifying the cellular replication of PWD circovirus or capable of inducing or inhibiting the pathologies linked to an infection by a PWD circovirus, characterized in that it comprises the following steps:

- **[0280]** a) contacting of said compound with said polypeptide, said nucleotide sequence, or with a cell transformed according to the invention and/or administration of said compound to an animal transformed according to the invention;
- **[0281]** b) determination of the capacity of said compound to bind to said polypeptide or said nucleotide sequence, or to modulate, induce or inhibit the expression of genes, or to modulate the growth or the replication of PWD circovirus, or to induce or inhibit in said transformed animal the pathologies linked to an infection by PWD circovirus (designated activity of said compound).

[0282] The compounds capable of being selected can be organic compounds such as polypeptides or carbohydrates or any other organic or inorganic compounds already known, or novel organic compounds elaborated by molecular modeling techniques and obtained by chemical or biochemical synthesis, these techniques being known to the person skilled in the art.

[0283] It will be possible to use said selected compounds to modulate the cellular replication of PWD circovirus and thus to control infection by this virus, the methods allowing said modulations to be determined being well known to the person skilled in the art.

[0284] This modulation can be carried out, for example, by an agent capable of binding to a protein and thus of inhibiting or of potentiating its biological activity, or capable of binding to an envelope protein of the external surface of said virus and of blocking the penetration of said virus into the host cell or of favoring the action of the immune system of the infected organism directed against said virus. This modulation can likewise be carried out by an agent capable of binding to a nucleotide sequence of a DNA of said virus and of blocking, for example, the expression of a polypeptide whose biological or structural activity is necessary for the replication or for the proliferation of said virus host cells to host cells in the host animal.

[0285] The invention relates to the compounds capable of being selected by a selection method according to the invention.

[0286] The invention likewise relates to a pharmaceutical composition comprising a compound selected from the following compounds:

- **[0287]** a) a nucleotide sequence according to the invention;
- [0288] b) a polypeptide according to the invention;
- **[0289]** c) a vector, a viral particle or a cell transformed according to the invention;
- [0290] d) an antibody according to the invention;
- **[0291]** e) a compound capable of being selected by a selection method according to the invention;

possibly in combination with a pharmaceutically acceptable vehicle and, if need be, with one or more adjuvants of the appropriate immunity.

[0292] The invention also relates to an immunogenic and/or vaccine composition, characterized in that it comprises a compound selected from the following compounds:

- **[0293]** a) a nucleotide sequence according to the invention;
- [0294] b) a polypeptide according to the invention;
- **[0295]** c) a vector or a viral particle according to the invention; and
- [0296] d) a cell according to the invention.

[0297] In one embodiment, the vaccine composition according to the invention is characterized in that it comprises a mixture of at least two of said compounds a), b), c) and d) above and in that one of the two said compounds is related to the PWD circovirus of type A and the other is related to the PWD circovirus of type B.

[0298] In another embodiment of the invention, the vaccine composition is characterized in that it comprises at least one compound a), b), c), or d) above which is related to PWD circovirus of type B. In still another embodiment, the vaccine composition is characterized in that it comprises at least one compound a), b), c), or d) above which is related to PWD circovirus of type B ORF'2.

[0299] A compound related to the PWD circovirus of type A or of type B is understood here as respectively designating a compound obtained from the genomic sequence of the PWD circovirus of type A or of type B.

[0300] The invention is additionally aimed at an immunogenic and/or vaccine composition, characterized in that it comprises at least one of the following compounds:

- [0301] a nucleotide sequence SEQ ID No. 23, SEQ ID No. 25, or one of their fragments or homologues;
- **[0302]** a polypeptide of sequence SEQ ID No. 24, SEQ ID No. 26, or one of their fragments, or a modification thereof;
- **[0303]** a vector or a viral particle comprising a nucleotide sequence SEQ ID No. 23, SEQ ID No. 25, or one of their fragments or homologues;
- **[0304]** a transformed cell capable of expressing a polypeptide of sequence SEQ ID No. 24, SEQ ID No. 26, or one of their fragments, or a modification thereof; or

[0305] a mixture of at least two of said compounds.

[0306] The invention also comprises an immunogenic and/ or vaccine composition according to the invention, characterized in that it comprises said mixture of at least two of said compounds as a combination product for simultaneous, separate or protracted use for the prevention or the treatment of infection by a PWD circovirus, especially of type B.

[0307] In a preferred embodiment, the vaccine composition according to the invention comprises the mixture of the following compounds:

- [0308] a pcDNA3 plasmid containing a nucleic acid of sequence SEQ ID No. 23;
- [0309] a pcDNA3 plasmid containing a nucleic acid of sequence SEQ ID No. 25;
- [0310] a pcDNA3 plasmid containing a nucleic acid coding for the GM-CSF protein;
- **[0311]** a recombinant baculovirus containing a nucleic acid of sequence SEQ ID No. 23;
- **[0312]** a recombinant baculovirus containing a nucleic acid of sequence SEQ ID No. 25; and

[0313] if need be, an adjuvant of the appropriate immunity, especially the adjuvant AIFTM.

[0314] The invention is likewise directed at a pharmaceutical composition according to the invention, for the prevention or the treatment of an infection by a PWD circovirus.

[0315] The invention is also directed at a pharmaceutical composition according to the invention for the prevention or the treatment of an infection by the PWD circovirus of type B. **[0316]** The invention likewise concerns the use of a composition according to the invention, for the preparation of a median provide the treatment of the prevention of the prevention of the prevent of t

medicament intended for the prevention or the treatment of infection by a PWD circovirus, preferably by the PWD circovirus of type B.

[0317] Under another aspect, the invention relates to a vector, a viral particle or a cell according to the invention, for the treatment and/or the prevention of a disease by gene therapy. [0318] Finally, the invention comprises the use of a vector, of a viral particle or of a cell according to the invention for the preparation of a medicament intended for the treatment and/ or the prevention of a disease by gene therapy.

[0319] The polypeptides of the invention entering into the immunogenic or vaccine compositions according to the invention can be selected by techniques known to the person skilled in the art such as, for example, depending on the capacity of said polypeptides to stimulate the T cells, which is translated, for example, by their proliferation or the secretion of interleukins, and which leads to the production of antibodies directed against said polypeptides.

[0320] In pigs, as in mice, in which a weight dose of the vaccine composition comparable to the dose used in man is administered, the antibody reaction is tested by taking of the serum followed by a study of the formation of a complex between the antibodies present in the serum and the antigen of the vaccine composition, according to the usual techniques.

[0321] The pharmaceutical compositions according to the invention will contain an effective quantity of the compounds of the invention, that is to say in sufficient quantity of said compound(s) allowing the desired effect to be obtained, such as, for example, the modulation of the cellular replication of PWD circovirus. The person skilled in the art will know how to determine this quantity, as a function, for example, of the age and of the weight of the individual to be treated, of the state of advancement of the pathology, of the possible secondary effects and by means of a test of evaluation of the effects obtained on a population range, these tests being known in these fields of application.

[0322] According to the invention, said vaccine combinations will preferably be combined with a pharmaceutically acceptable vehicle and, if need be, with one or more adjuvants of the appropriate immunity.

[0323] Today, various types of vaccines are available for protecting animals or man against infectious diseases: attenuated living microorganisms (*M. bovis*—BCG for tuberculosis), inactivated microorganisms (influenza virus), acellular extracts (*Bordetella pertussis* for whooping cough), recombined proteins (surface antigen of the hepatitis B virus), polysaccharides (pneumococcal). Vaccines prepared from synthetic peptides or genetically modified microorganisms expressing heterologous antigens are in the course of experimentation. More recently still, recombined plasmid DNAs carrying genes coding for protective antigens have been proposed as an alternative vaccine strategy. This type of vaccination is carried out with a particular plasmid originating from a plasmid of *E. coli* which does not replicate in vivo and

which codes uniquely for the vaccinating protein. Animals have been immunized by simply injecting the naked plasmid DNA into the muscle. This technique leads to the expression of the vaccine protein in situ and to an immune response of cellular type (CTL) and of humoral type (antibody). This double induction of the immune response is one of the principal advantages of the vaccination technique with naked DNA.

[0324] The vaccine compositions comprising nucleotide sequences or vectors into which are inserted said sequences are especially described in the international application No. WO 90/11092 and likewise in the international application No. WO 95/11307.

[0325] The constitutive nucleotide sequence of the vaccine composition according to the invention can be injected into the host after having been coupled to compounds which favor the penetration of this polynucleotide into the interior of the cell or its transport to the cell nucleus. The resultant conjugates can be encapsulated in polymeric microparticles, as described in the international application No. WO 94/27238 (Medisorb Technologies International).

[0326] According to another embodiment of the vaccine composition according to the invention, the nucleotide sequence, preferably a DNA, is complexed with DEAE-dextran (Pagano et al., 1967) or with nuclear proteins (Kaneda et al., 1989), with lipids (Felgner et al., 1987) or encapsulated in liposomes (Fraley et al., 1980) or else introduced in the form of a gel facilitating its transfection into the cells (Midoux et al., 1993, Pastore et al., 1994). The polynucleotide or the vector according to the invention can also be in suspension in a buffer solution or be combined with liposomes.

[0327] Advantageously, such a vaccine will be prepared according to the technique described by Tacson et al. or Huygen et al. in 1996 or alternatively according to the technique described by Davis et al. in the international application No. WO 95/11307.

[0328] Such a vaccine can likewise be prepared in the form of a composition containing a vector according to the invention, placed under the control of regulation elements allowing its expression in man or animal. It will be possible, for example, to use, by way of in vivo expression vector of the polypeptide antigen of interest, the plasmid pcDNA3 or the plasmid pcDNA1/neo, both marketed by Invitrogen (R&D Systems, Abingdon, United Kingdom). It is also possible to use the plasmid V1Jns.tPA, described by Shiver et al. in 1995. Such a vaccine will advantageously comprise, apart from the recombinant vector, a saline solution, for example a sodium chloride solution.

[0329] Pharmaceutically acceptable vehicle is understood as designating a compound or a combination of compounds entering into a pharmaceutical composition or vaccine which does not provoke secondary reactions and which allows, for example, the facilitation of the administration of the active compound, an increase in its duration of life and/or its efficacy in the body, an increase in its solubility in solution or alternatively an improvement in its conservation. These pharmaceutically acceptable vehicles are well known and will be adapted by the person skilled in the art as a function of the nature and of the mode of administration of the chosen active compound.

[0330] As far as the vaccine formulations are concerned, these can comprise adjuvants of the appropriate immunity which are known to the person skilled in the art, such as, for example, aluminum hydroxide, a representative of the family

of muramyl peptides such as one of the peptide derivatives of N-acetyl muramyl, a bacterial lysate, or alternatively Freund's incomplete adjuvant.

[0331] These compounds can be administered by the systemic route, in particular by the intravenous route, by the intramuscular, intradermal or subcutaneous route, or by the oral route. In a more preferred manner, the vaccine composition comprising polypeptides according to the invention will be administered by the intramuscular route, through the food or by nebulization several times, staggered over time.

[0332] Their administration modes, dosages and optimum pharmaceutical forms can be determined according to the criteria generally taken into account in the establishment of a treatment adapted to an animal such as, for example, the age or the weight, the seriousness of its general condition, the tolerance to the treatment and the secondary effects noted. Preferably, the vaccine of the present invention is administered in an amount that is protective against piglet weight loss disease.

[0333] For example, in the case of a vaccine according to the present invention comprising a polypeptide encoded by a nucleotide sequence of the genome of PCV, or a homologue or fragment thereof, the polypeptide will be administered one time or several times, spread out over time, directly or by means of a transformed cell capable of expressing the polypeptide, in an amount of about 0.1 to 10 μ g per kilogram weight of the animal, preferably about 0.2 to about 5 μ g/kg, more preferably about 0.5 to about 2 μ g/kg for a dose.

[0334] The present invention likewise relates to the use of nucleotide sequences of PWD circovirus according to the invention for the construction of autoreplicative retroviral vectors and the therapeutic applications of these, especially in the field of human gene therapy in vivo.

[0335] The feasibility of gene therapy applied to man no longer needs to be demonstrated and this relates to numerous therapeutic applications like genetic diseases, infectious diseases and cancers. Numerous documents of the prior art describe the means of employing gene therapy, especially through viral vectors. Generally speaking, the vectors are obtained by deletion of at least some of the viral genes which are replaced by the genes of therapeutic interest. Such vectors can be propagated in a complementation line which supplies in trans the deleted viral functions in order to generate a defective viral vector particle for replication but capable of infecting a host cell. To date, the retroviral vectors are amongst the most widely used and their mode of infection is widely described in the literature accessible to the person skilled in the art.

[0336] The principle of gene therapy is to deliver a functional gene, called a gene of interest, of which the RNA or the corresponding protein will produce the desired biochemical effect in the targeted cells or tissues. On the one hand, the insertion of genes allows the prolonged expression of complex and unstable molecules such as RNAs or proteins which can be extremely difficult or even impossible to obtain or to administer directly. On the other hand, the controlled insertion of the desired gene into the interior of targeted specific cells allows the expression product to be regulated in defined tissues. For this, it is necessary to be able to insert the desired therapeutic gene into the interior of chosen cells and thus to have available a method of insertion capable of specifically targeting the cells or the tissues chosen. **[0337]** Among the methods of insertion of genes, such as, for example, microinjection, especially the injection of naked plasmid DNA (Derse, D. et al., 1995, and Zhao, T. M. et al., 1996), electroporation, homologous recombination, the use of viral particles, such as retroviruses, is widespread. However, applied in vivo, the gene transfer systems of recombinant retroviral type at the same time have a weak infectious power (insufficient concentration of viral particles) and a lack of specificity with regard to chosen target cells.

[0338] The production of cell-specific viral vectors, having a tissue-specific tropism, and whose gene of interest can be translated adequately by the target cells, is realizable, for example, by fusing a specific ligand of the target host cells to the N-terminal part of a surface protein of the envelope of PWD circovirus. It is possible to mention, for example, the construction of retroviral particles having the CD4 molecule on the surface of the envelope so as to target the human cells infected by the HW virus (YOUNG, J. A. T. et al., Sciences 1990, 250, 1421-1423), viral particles having a peptide hormone fused to an envelope protein to specifically infect the cells expressing the corresponding receptor (KASAHARA, N. et al., Sciences 1994, 266, 1373-1376) or else alternatively viral particles having a fused polypeptide capable of immobilizing on the receptor of the epidermal growth factor (EGF) (COSSET, F. L. et al., J. of Virology 1995, 69, 10, 6314-6322). In another approach, single-chain fragments of antibodies directed against surface antigens of the target cells are inserted by fusion with the N-terminal part of the envelope protein (VALSESIA-WITTMAN, S. et al., J. of Virology 1996, 70, 3, 2059-2064; TEARINA CHU, T. H. et al., J. of Virology 1997, 71, 1, 720-725).

[0339] For the purposes of the present invention, a gene of interest in use in the invention can be obtained from a eukaryotic or prokaryotic organism or from a virus by any conventional technique. It is, preferably, capable of producing an expression product having a therapeutic effect and it can be a product homologous to the cell host or, alternatively, heterologous. In the scope of the present invention, a gene of interest can code for an (i) intracellular or (ii) membrane product present on the surface of the host cell or (iii) secreted outside the host cell. It can therefore comprise appropriate additional elements such as, for example, a sequence coding for a secretion signal. These signals are known to the person skilled in the art.

[0340] In accordance with the aims pursued by the present invention, a gene of interest can code for a protein corresponding to all or part of a native protein as found in nature. It can likewise be a chimeric protein, for example arising from the fusion of polypeptides of various origins or from a mutant having improved and/or modified biological properties. Such a mutant can be obtained, by conventional biological techniques, by substitution, deletion and/or addition of one or more amino acid residues.

[0341] It is very particularly preferred to employ a gene of therapeutic interest coding for an expression product capable of inhibiting or retarding the establishment and/or the development of a genetic or acquired disease. A vector according to the invention is in particular intended for the prevention or for the treatment of cystic fibrosis, of hemophilia A or B, of Duchenne's or Becker's myopathy, of cancer, of AIDS and of

other bacteria or infectious diseases due to a pathogenic organism: virus, bacteria, parasite or prion. The genes of interest utilizable in the present invention are those which code, for example, for the following proteins:

- **[0342]** a cytokine and especially an interleukin, an interferon, a tissue necrosis factor and a growth factor and especially a hematopoietic growth factor (G-CSF, GM-CSF),
- **[0343]** a factor or cofactor involved in clotting and especially factor VIII, von Willebrand's factor, antithrombin III, protein C, thrombin and hirudin,
- [0344] an enzyme or an enzyme inhibitor such as the inhibitors of viral proteases,
- **[0345]** an expression product of a suicide gene such as thymidine kinase of the HSV virus (herpesvirus) of type 1,
- [0346] an activator or an inhibitor of ion channels,
- **[0347]** a protein of which the absence, the modification or the deregulation of expression is responsible for a genetic disease, such as the CFTR protein, dystrophin or minidystrophin, insulin, ADA (adenosine diaminose), glucocerebrosidase and phenylhydroxylase,
- **[0348]** a protein capable of inhibiting the initiation or the progression of cancers, such as the expression products of tumor suppressor genes, for example the P53 and Rb genes,
- **[0349]** a protein capable of stimulating an immune or an antibody response, and
- **[0350]** a protein capable of inhibiting a viral infection or its development, for example the antigenic epitopes of the virus in question or altered variants of viral proteins capable of entering into competition with the native viral proteins.

[0351] The invention thus relates to the vectors characterized in that they comprise a nucleotide sequence of PWD circovirus according to the invention, and in that they additionally comprise a gene of interest.

[0352] The present invention likewise relates to viral particles generated from said vector according to the invention. It additionally relates to methods for the preparation of viral particles according to the invention, characterized in that they employ a vector according to the invention, including viral pseudoparticles (VLP, virus-like particles).

[0353] The invention likewise relates to animal cells transfected by a vector according to the invention.

[0354] Likewise comprised in the invention are animal cells, especially mammalian, infected by a viral particle according to the invention.

[0355] The present invention likewise relates to a vector, a viral particle or a cell according to the invention, for the treatment and/or the prevention of a genetic disease or of an acquired disease such as cancer or an infectious disease. The invention is likewise directed at a pharmaceutical composition comprising, by way of therapeutic or prophylactic agent, a vector or a cell according to the invention, in combination with a vehicle acceptable from a pharmaceutical point of view.

[0356] Other characteristics and advantages of the invention appear in the examples and the figures.

[0357] The invention is described in more detail in the following illustrative examples. Although the examples may

represent only selected embodiments of the invention, it should be understood that the following examples are illustrative and not limiting.

Examples

Example 1

Cloning, Sequencing and Characterization of the PWD Circovirus of Type A (PCVA)

[0358] 1. Experimental Procedures

[0359] Experimental reproduction of the infection and its syndrome are provided (cf. FIG. 1).

[0360] A first test was carried out with pigs from a very well-kept farm, but affected by piglet weight loss disease (PWD), likewise called fatal piglet wasting (FPW). Tests carried out with SPF (specific pathogen-free) pigs showed a transfer of contaminant(s) finding expression in a complex pathology combining hyperthermia, retardation of growth, diarrhea and conjunctivitis. The PDRS (porcine dysgenic and respiratory syndrome) virus, an infectious disease due to an arteriovirus) was rapidly isolated from breeding pigs and contact pigs. It should have been possible to attribute all the clinical signs to the presence of the PDRS virus. However, two farm pigs presented signs of FPW without the PDRS virus being isolated. The histological analyses and blood formulas, however, showed that these pigs were suffering from an infectious process of viral origin.

[0361] In a second test, 8-week SPF pigs were inoculated by the intratracheal route with organ homogenates of two farm pigs suffering from FPW. The inoculated pigs exhibited hyperthermia 8 to 9 days post-infection, then their growth was retarded. Other SPF pigs, placed in contact, had similar, attenuated signs 30 days after the initial experiment. No sero-conversion with respect to a European or Canadian strain of PDRS virus was recorded in these animals.

[0362] A third test allowed the syndrome to be reproduced from samples taken from the pigs of the second test.

[0363] Conclusion

[0364] The syndrome is reproduced under the experimental conditions. It is determined by at least one infectious agent, which is transmittable by direct contact. The clinical constants are a sometimes high hyperthermia (greater than or equal to 41.5° C.) which develops 8 to 10 days after infection. Retardation of the growth can be observed. The other signs are a reversal of the blood formula (reversal of the lymphocyte/polynuclear ratio from 70/30 to 30/70) and frequent lesions on the ganglia, especially those draining the respiratory apparatus (ganglionic hypertrophy, loss of structure with necrosis and infiltration by mononucleated or plurinucleated giant cells).

2. Laboratory Studies

[0365] Various cell supports including primary pig kidney cells or cell lines, pig testicle cells, monkey kidney cells, pig lymphocytes, pig alveolar macrophages and circulating blood monocytes were used to demonstrate the possible presence of a virus. No cytopathic effect was demonstrated in these cells. On the other hand, the use of a serum of a pig sick after experimental infection allowed an intracellular antigen to be revealed in the monocytes, the macrophages and approximately 10% of pig kidney (PK) cells infected with organ homogenates. This indirect revealing was carried out kinetically at different culture times. It is evident from this that the

antigen initially appears in the nucleus of the infected cells before spreading into the cytoplasm. The successive passages in cell culture did not allow the signal to be amplified.

[0366] Under electron microscopy on organ homogenates, spherical particles labeled specifically by the serum of sick pigs, infected under the experimental conditions, were visualized. The size of these particles is estimated at 20 nm.

[0367] After two passages of these organ homogenates over pig lymphocytes and then three passages over pig kidney or testicle cells, a cytopathic effect developed and was amplified. An adenovirus was visualized in the electron microscope, which, under the experimental conditions, did not reproduce FPW (only a hyperthermia peak was noted 24 to 48 hours after infection, and then nothing more).

[0368] It has been possible to demonstrate DNA bands in certain samples of pigs infected under the experimental conditions and having exhibited signs of the disease (results not shown). A certain connection exists between the samples giving a positive result in cell culture and those having a DNA band.

[0369] Conclusion

[0370] At least two types of virus were demonstrated in the organ homogenates from pigs suffering from FPW. One is an adenovirus, but by itself alone it does not reproduce the disease. The other type of virus is a circovirus and is associated with FPW. This circovirus, of which two types have been isolated and sequenced, designated below PWD circovirus type A (or PCVA) and PWD circovirus of type B (or PCVB) have mutations with respect to the known sequences of circovirus which are nonpathogenic for the pig.

3. Cloning and Sequencing of the DNA of the PWD Circovirus of Type A

[0371] Cloning and sequencing of the DNA of PHD circovirus Type A is accomplished by extraction of the replicative form (RF) DNA, followed by cleavage by the Kpn I enzyme and amplification by a pair of primers flanking the Kpn I restriction site. The two strands of DNA are sequenced at least twice by the Sanger method.

[0372] The nucleic sequence of the strand of (+) polarity of the genome of the PWD circovirus of type A (or PCVA), strain FPW, is represented by the sequence SEQ ID No. 1 in the list of sequences, the nucleic acid sequence of the strand of (-) polarity of the genome of the PWD circovirus of type A (or PCVA) being represented by the nucleic acid sequence $3' \rightarrow 5'$ of FIG. 3 or by the sequence SEQ ID No. 5 (represented according to the orientation $5' \rightarrow 3'$) in the list of sequences.

[0373] The amino acid sequences SEQ ID No. 10, SEQ ID No. 12 and SEQ ID No. 14 of the list of sequences respectively represent the sequences of proteins encoded by the nucleic sequences of the 3 open reading frames SEQ ID No. 9 (ORF1), corresponding to the REP protein, SEQ ID No. 11 (ORF2) and SEQ ID No. 13 (ORF3), determined from the sequence SEQ ID No. 1 of the strand of (+) polarity or of the nucleic sequence SEQ ID No. 5 of the strand of (-) polarity of the genome of the PWD circovirus of type A.

4. Comparison of the Nucleotide Sequences and Amino Acids of the PWD Circovirus of Type A (or Associated with PWD) which are Obtained with the Corresponding Sequences of MEEHAN and MANKERTZ Circoviruses of Porcine Cell Lines.

[0374] DNA sequences are analyzed using, DNASIS software.

Sequences of Oligonucleotides Used as Primers or Probes in the Detection and/or Identification Procedures

1. Specific Detection of the PWD Circovirus of Type A: [0375]

SEQ ID No. 46 primer PCV 5: 5' GTG TGC TCG ACA TTG GTG TG 3'; SEQ ID No. 47 primer PCV 10: 5' TGG AAT GTT AAC GAG CTG AG 3';

2. Specific Detection of the Circovirus of the Cell Lines: **[0376]**

SEQ ID No. 46 primer PCF 5: 5' GTG TGC TCG ACA TTG GTG TG 3'; SEQ ID No. 52 primer MEE 1: 5' TGG AAT GTT AAC TAC CTC AA 3';

3. Differential Detection:

[0377] the pairs of primers used are those described, for example, in the paragraphs 1 and 2 above;

4. Detection of the Monomeric Circular Replicative Forms RF:

[0378]

SEQ ID No. 46 primer PCV 5: 5' GTG TGC TCG ACA TTG GTG TG 3'; SEQ ID No. 48 primer PCV 6: 5' CTC GCA GCC ATC TTG GAA TG 3';

5. Detection of the Vectors Carrying the Dimers in Tandem: [0379]

```
Nar dimer:

SEQ ID No. 49

primer KS 620: 5' CGC GCG TAA TAC GAC TCA CT 3';

SEQ ID No. 46

primer PCV 5: 5' GTG TGC TCG ACA TTG GTG TG 3';

Kpn dimer:

SEQ ID No. 49

primer KS 620: 5' CGC GCG TAA TAC GAC TCA CT 3';

SEQ ID No. 48
```

primer PCV 6: 5'CTC GCA GCC ATC TTG GAA TG 3';

6. Differential Detection:

[0380] The pairs of primers used are those described, for example, in paragraphs 4 and 5 above.

[0381] The procedures using the pairs or primers described in paragraphs 4 and 5 are of particular interest for differentially detecting the circular monomeric forms of specific replicative forms of the virion or of the DNA in replication and the dimeric forms found in the so-called in-tandem molecular constructs.

[0382] The in-tandem constructs of the viral genome (dimers) such as the constructs used for the preparation of the

pBS KS+tandem PCV Kpn I vector, deposited at the CNCM under the number I-1891, 3 Jul. 1997 (*E. coli* transformed by said vector) are very interesting for their use in methods of production of sufficient quantity of an inoculum formed of DNA, intended for the virus production, this in the absence of a satisfactory virus production protocol in a cell system. These said methods of production using in-tandem constructs of the viral genome will allow the virulence factors to be studied by mutation and by way of consequence will be able to be used for the production of a collection of viruses carrying the mutations indicated in the construction of vectors which will have the appropriate tropism and virulence. These vectors with autoreplicative structure have the sought gene transfer properties, especially for their applications in gene therapy, and in vaccinology.

[0383] Western-Blot Analysis of Recombinant Proteins of the PWD Circovirus of Type A

[0384] The results were obtained using a specific antiserum of the PWD circovirus produced during test 1 (cf. FIG. 1).

[0385] Type of Products Analyzed

[0386] The analyses were carried out on cell extracts of Sf9 cells obtained after infection by the recombinant baculovirus PCV ORF 1.

[0387] The culture of Sf9 cells was carried out in a 25 cm^2 Petri dish according to the standard culture methods for these cells. After centrifugation, the cell pellets are taken up with 300 µl of PBS buffer (phosphate saline buffer).

[0388] Electrophoresis (PAGE-SDS)

[0389] The electrophoresis is carried out on the cell extracts of Sf9 cells obtained previously on 5 samples (cf. Table 1 below) under the following conditions:

[0390] % polyacrylamide gel: 8%; conditions: denaturing

[0391] Voltage: 80 V; duration: 135 mn.

TABLE 1

Nature of	the s	samples	enh	iected	to	electro	horesis	
_mature or	une a	sampies	suo	lecten	w	electro	JIIOICSIS	

-	Well No.				
	1	2	3	4	5
Sample applied	PM Rainbow	Raoul 24 h	Raoul 48 h	Raoul 72 h	Raoul 96 h
µl of sample	10	15	15	15	15
μl of Laemmli 4X	0	5	5	5	5

Legends to Table 1:

Laemmli 4X: loading buffer

PM Rainbow: molecular-weight markers (35, 52, 77, 107, 160 and 250 kD)

Raoul 24 h, 48 h, 72 h and 96 h: expression products of the ORF1 of the PWD circovirus of type A.

[0392] Western Blot

[0393] After electrophoresis, the bands obtained in the different wells are transferred to nitrocellulose membrane for 1 h at 100 v in a TGM buffer (tris-glycine-methanol).

[0394] The Western blot is carried out under the following conditions:

[0395] 1) Saturation with a solution containing 5% of skimmed milk; 0.05% of Tween 20 in a TBS 1×buffer (tris buffer saline) for 30 min.

[0396] 2) 1st antibody:

[0397] 10 ml of PWD anticircovirus antibody of type A are added diluted to ¹/100, then the reaction mixture is incubated for one night at 4° C. Three washes of 10 min in TBS 1× are carried out. [0398] 3) 2nd antibody:

[0399] 10 ml of pig rabbit P164 antibody anti-immunoglobulins, coupled to peroxidase (Dakopath), are added diluted to $\frac{1}{100}$, then the reaction medium is incubated for 3 hours at 37° C. Three washes of 10 min in TBS 1× are carried out.

[0400] 4) Visualization

[0401] The substrate 4-chloro-1-naphthol in the presence of oxygenated water is used for visualization. Results

[0402] The results are shown in FIG. 7.

Kinetics of Appearance of Antibodies Specific for the REP Recombinant Protein of the PWD Circovirus of Type A Expressed in Baculovirus After Infection of Pigs by the PWD Circovirus of Type A (test 4, cf. FIG. 1)

[0403] After infection of the pigs, a sample of serum of each of the infected pigs is taken at different periods expressed in the table by the date of taking (carried out here in the same year) and is then analyzed by Western blot.

[0404] The visualization of the specific antibodies is carried out in the manner described previously.

[0405] The results obtained are shown by Table 2 below.

TABLE 2

	Ki	netics of	appearar	nce of s	pecific a	untibodi	es	
Sample	Pigs	10/6	16/06	23/06	01/07	08/07	15/07	21/07
A3 Control B2 Infec. RP+	1 2 1 2 3 4	Neg. Neg. Neg. Neg.	Neg. Neg. Neg. Neg.	Neg. Neg. Neg. Neg.	+ Neg. Neg. Neg.	+ Neg. + Neg.	Neg. Neg. ++ Neg. + Neg.	+++ Neg. + ++

Legends to Table 2:

A3 control: uninfected control animals;

B2 Infec. RP+: animals infected with pig kidney (PK) cells containing the circovirus; Neg.: negative;

+, ++, +++: intensity scale of the positive reaction;

 $10/06,\,16/06,\,23/06,\,01/07,\,08/07,\,15/07,\,21/07;$ dates expressed in day/month on which the different withdrawals of serum were carried out.

Example 2

Cloning, Sequencing and Characterization of the Type B PWD Circovirus (PCVB)

[0406] The techniques used for cloning, sequencing and characterization of the type B PWD circovirus (PCVB) are those used in Example 1 above for the type A PWD circovirus (PCVA).

[0407] The nucleic acid sequence of the strand of (+) polarity of the genome of the PWD circovirus of type B (or PCVB) is represented by the sequence SEQ ID No. 15 in the sequence listing, the nucleic acid sequence of the strand of (-) polarity of the genome of the PWD circovirus of type B (or PCVB) being represented by the nucleic acid sequence $3' \rightarrow 5'$ of FIG. **8** or by the sequence SEQ ID No. 19 (represented according to the orientation $5' \rightarrow 3'$) in the sequence listing.

[0408] The amino acid sequences, SEQ ID No. 24, SEQ ID No. 26 and SEQ ID No. 28 of the sequence listing, respectively, represent the sequences of the proteins encoded by the nucleic sequences of the 3 open reading frames SEQ ID No. 23 (ORF'1), corresponding to the REP protein, SEQ ID No. 25 (ORF'2) and SEQ ID No. 27 (ORF'3), determined from the sequence SEQ ID No. 15 of the strand of (+) polarity or from

the nucleic sequence SEQ ID No. 19 of the strand of (-) polarity of the genome of the PWD circovirus of type B.

Example 3

Comparative Analysis of Nucleotide Sequences (ORF1, ORF2 and Genomic) and Amino Acid Sequences Encoded by the ORF1 and the ORF2 of the PWD Circoviruses of Type A (PCVA) and of Type B (PCVB)

[0409] The results expressed in % of homology are shown in Tables 3 and 4 below.

TABLE 3

Compared analysis	s of the amino acid	sequences
% homology	ORF1	ORF2
PCVA/PCVB	80.4	56.2

TABLE 4	
---------	--

Co	npared analysis	of the nucleo	otide sequen	ces
% homology	Genomic	ORF1	ORF2	The remainder
PCVA/PCVB	70.4	80.4	60.1	66.1

Example 4

Observation of the Disease and Reproduction of the Disease Under Experimental Conditions

[0410] a) Test No. 1: Observation of the Disease

[0411] The objective is to take breeding animals at the start of disease and to place them under experimental conditions to follow the progression of the pathology and describe all the clinical signs thereof. This first test was carried out on 3 breeding pigs aged 10 weeks of which 2 were already ill (suffering from wasting), and on 3 other pigs aged 13 weeks, not having signs of disease. The clinical observation was spread over a period of 37 days. Two pigs of 10 weeks wasted rapidly (pigs 1 and 2, FIG. 9) and had to be painlessly killed 5 and 6 days after their arrival. A single pig exhibited hyperthermia over 5 days and diarrhea. Two other pigs exhibited dyspnea and cough, of which one additionally had hyperthermia, greater than 41° C., for the two first days of its stay. Another pig had retarded growth in the second week (pig 6, FIG. 9), without any other clinical sign being recorded. On the lesional level, 5 pigs out of 6 exhibited macroscopic lesions of gray pneumonia, the sixth exhibited cicatricial lesions on the lung.

[0412] b) Test No. 2: Reproduction of the Disease from Inocula Prepared in Farm Pigs.

[0413] The two sick pigs in test 1 served to prepare inocula which were tested in test 2 on specific-pathogen-free (SPF) pigs. The SPF pigs were aged 9 weeks at the time of inoculation. The clinical and lesional results are shown in Table 5.

TABLE 5

Summary of the measurements carried out during experimental reproduction of PWD. (The values of the control animals are reported in brackets, the underlined values indicate a difference between infected animals and control animals)

-

	Test Measurement					
	2	3	4	5	6	7
Status of the pigs	SPF CNEVA	SPF field	SPF CNEVA	SPF CNEVA	Conventional	Conventional
Age	9 weeks	6 weeks	5 weeks	5 weeks	5 weeks	6-7 weeks
Number	4	6	12	8	8	8
Inoculation route	Intratracheal route	Intratracheal route	Intratracheal + intramuscular route	Intratracheal + intramuscular route	Intratracheal + intramuscular route	Intratracheal + intramuscular route
Inoculum titer per pig	ND*	ND*	10 ^{4.53} TCID ₅₀ per ml: 1 ml IM + 5 ml IT	10 ^{4.53} TCID ₅₀ per ml: 1 ml IM + 5 ml IT	10 ^{4.53} TCID ₅₀ per ml: 1 ml IM + 5 ml IT	10 ^{4.53} TCID ₅₀ per ml: 1 ml IM + 5 ml IT
Start of hyperthermia	10 days post-infection	9-13 days post-infection	12-13 days post-infection	9-14 days post-infection	8-12 days post-infection	12 days post-infection
% of pigs in hyperthermia**	100%	83%	92%	100%	75%	88%
Number of days of hyperthermia per pig**	7	4.5	3.3	5.8	7.5	11.6
Maximum temperatures*** Hyperthermia**** % per week	40.4 to 41.7° C.	40.6 to 42.3° C.	40.2 to 41.6° C.	40.3 to 40.8° C.	40.6 to 42° C.	40.2 to 41.9° C.
W1	3.5 (3.5)	17 (36)	7 (5)	37 (17)	16 (17)	20 (28)
W2	42 (3.5)	7 (13)	13 (1)	21 (3)	52 (10)	37 (28)
W3	35 (3.5)	33 (10)	28 (7)	62 (2)	34 (12)	79 (17)
W4	21 (3.5)	28 (7)	5 (0)	6 (3)	25 (22)	55 (3)
DMG:			(-)	(-)	()	
W1	928 (1053)	417 (357)	564 (620)	650 (589)	401 (407)	509 (512)
W2	678 (1028)	428 (617)	<u>503 (718)</u>	612 (584)	294 (514)	410 (310)
W3	661 (1000)	771 (642)	381 (657)	520 (851)	375 (586)	435 (440)
W4	786 (1100)	<u>550 (657)</u>	764 (778)	641 (696)	<u>473 (610)</u>	451 (681)

	Summary of the measuremen reported in brackets, t					als are	
			Test Me	asurement			
	2	3	4	5	6	7	
Contact pigs transmission	Yes to 100%	Yes to 75%	Not tested	Not tested	Not tested	Not tested	
% of pulmonary lesions	25	75	0	25	25	12	
% of ganglionic lesions	17	33	67	25	50	12	

*ND: not determined.

**hyperthermia when the temperature is greater than 40° C.,

***range of maximum temperatures recorded at the individual level,

****the percentage corresponds to the number of temperature recordings greater than 40° C. divided by the total number of temperature recordings in the week on all of the pigs.

[0414] In this test, there was no wasting, at the very most a retardation of the growth in the second, third or fourth week after infection. These data illustrate that certain breeding conditions probably favor the expression of the disease. [0415] c) Tests No. 3 to No. 7: Reproduction of the Experi-

mental Tests

[0416] The increase in the number of the experimental tests on pigs had the mastering and better characterization of the experimental model as an objective. All of the results are presented in Table 5.

[0417] Under the experimental conditions, PWD is thus characterized by a long incubation, of 8 to 14 days, true hyperthermia over 2 to 8 days, a decrease in food consumption and a retardation of the increase in weight on the second, third or fourth week post-infection. The lesional table associated with this clinical expression includes, in the main, ganglionic hypertrophy and lesions of pneumonia.

[0418] Conclusion

[0419] The perfection of this experimental model allows the direct etiological role of the PWD circovirus in the disease to be indisputably demonstrated. In addition, this model is an indispensable tool for the understanding of pathogenic mechanisms and the study of future vaccine candidates.

Example 5

Demonstration of the Vaccine Composition Protective Efficacy Produced from Nucleic Fragments of PWD Circovirus Sequence

[0420] 1) Animals Used for the Study

[0421] Piglets having the PWD disease, reproduced under experimental conditions described in paragraph c) of Example 4, were used in a protocol for evaluating the vaccine composition efficacy, comprising nucleic fragments of PWD circovirus sequence.

[0422] 2) Tested Vaccine Composition and Vaccination Protocol

[0423] a) Components Used for the Study

[0424] The plasmids were obtained from the pcDNA3 plasmid of INVITROGENE

[0425] pcDNA3ORF-Plasmids

[0426] These plasmids are plasmids which do not carry a PWD circovirus nucleic acid insert and are used as a negative control plasmid.

[0427] pcDNA3ORF1+ Plasmid and pcDNA3ORF2+ Plasmid

[0428] The pcDNA3ORF1+ and pcDNA3ORF2+ plasmids are plasmids which carry a nucleic acid insert of the sequence of the PWD circovirus of TYPE B, and an insert comprising the nucleic acid fragment SEQ ID No. 23 (ORF'1) coding for the Rep protein of sequence SEQ ID No. 24 and an insert comprising the nucleic acid fragment SEQ ID No. 25 (ORF'2) coding for the protein of sequence SEQ ID No. 26, probably corresponding to the capsid protein, respectfully. These nucleic constructs further comprise the ATG initiation codon of the coding sequence of the corresponding protein.

[0429] GMCSF+ Plasmid

[0430] GM-CSF (granulocyte/macrophage colony stimulating factor) is a cytokine which occurs in the development, the maturation and the activation of macrophages, granulocytes and dendritic cells which present an antigen. The beneficial contribution of the GM-CSF in vaccination is considered to be a cellular activation with, especially, the recruitment and the differentiation of cells which present an antigen.

[0431] This pcDNA3-GMCSF+ plasmid carries a nucleic acid insert coding for the granulocyte/macrophage colony stimulation factor, the GM-CSF protein.

[0432] The gene coding for this GM-CSF protein was cloned and sequenced by Inumaru et al. (Immunol. Cell Biol., 1995, 73 (5), 474-476). The pcDNA3-GMCSF+ plasmid was obtained by Dr. B. Charley of NRA of Jouy-en-Josas (78, France).

[0433] Recombinant Baculoviruses

[0434] The so-called ORF-baculoviruses are viruses not carrying any insert comprising a nucleic acid fragment capable of expressing a PWD circovirus protein.

[0435] The so-called ORF1+ (BAC ORF1+) or ORF2+ (BAC ORF2+) baculoviruses are recombinant baculoviruses carrying an insert comprising a nucleic acid fragment SEQ ID No. 23 (ORF'1) and an insert comprising the nucleic acid fragment SEQ ID No. 25 (ORF'2), respectively.

[0436] Adjuvant

[0437] The adjuvant supplied by the Seppic Company, a subsidiary of AIR LIQUIDE, is the adjuvant corresponding to the reference AIF SEPPIC.

[0438] b) Vaccination Protocol

[0439] Weaned piglets aged 3 weeks are divided into four batches A, B, C and D each comprising 8 piglets.

[0440] Batches A, B and C, aged 3 weeks, each receive a first injection (injection M1) of 1 ml containing 200 micro-

grams of plasmids (naked DNA) in PBS, pH: 7.2, by the intramuscular route for each of the plasmids mentioned below for each batch, then, at the age of 5 weeks, a second injection (injection M2) comprising these same plasmids. A third injection is carried out simultaneously on the other side of the neck. This third injection comprises 1 ml of a suspension containing 5×10^6 cells infected by recombinant baculoviruses and 1 ml of AIF SEPPIC adjuvant. [0441] Batch A (F1) (Control Batch):

First Injection

[0442] pcDNA3ORF1-plasmid, pcDNA3ORF2-plasmid and GMCSF+ plasmid.

Second and Third Injection (Simultaneous)

[0443] pcDNA3ORF1-plasmid, pcDNA3ORF2-plasmid and GMCSF+ plasmid;

[0444] Cells transformed by baculoviruses not containing any nucleic acid insert coding for a PWD circovirus protein; [0445] AIF SEPPIC adjuvant.

[0446] Batch B (F2) (Control Batch):

First Injection

[0447] pcDNA3ORF1-plasmid, pcDNA3ORF2-plasmid and GMCSF+ plasmid;

Second and Third Injection (Simultaneous)

[0448] pcDNA3ORF1-plasmid, pcDNA3ORF2-plasmid and GMCSF+ plasmid;

[0449] Cells transformed by baculoviruses not containing any nucleic acid insert coding for a PWD circovirus protein; [0450] AIF SEPPIC adjuvant.

[0451] Batch C (F3):

First Injection

[0452] pcDNA3ORF1+ plasmid, pcDNA3ORF2+ plasmid and GMCSF+ plasmid;

Second and Third Injection (Simultaneous)

[0453] pcDNA3ORF1+ plasmid, pcDNA3ORF2+ plasmid and GMCSF+ plasmid;

[0454] Cells transformed by BAC ORF1+ and BAC ORF2+ recombinant baculoviruses capable of respectively expressing the Rep protein of sequence SEQ ID No. 24 and the protein of sequence SEQ ID No. 26 of the PWD circovirus of TYPE B. Batch D (F4) (control batch): no injection

[0455] The batches of piglets B, C and D are infected (tested) at the age of 6 weeks although batch A is not subjected to the test.

[0456] 3) Observation of the Batches

[0457] counting of coughing/sneezing: 15 minutes/batch/ day;

[0458] consistency of fecal matter: every day;

[0459] regular recordings: weekly taking of blood, weighing;

[0460] weighing of food refuse: 3 times per week;

[0461] calculation of the daily mean gain in weight (dmg);

[0462] The daily mean gains were calculated for each of the

batches over a period of 28 days following testing (cf. FIG. **10**), an intermediate calculation of the dmg was likewise

carried out for each of the batches over the first and second periods of 14 days. The results obtained are reported below in Table 6.

TABLE 6

	Da	ily mean gai	ns	
	F1	F2	F3	F4
d0-d14 d14-d28 d0-d28	411 g 623 g 554 g	450 g 362 g 406 g	511 g 601 g 556 g	461 g 443 g 452 g

[0463] Measurement of hyperthermia

[0464] The measurement of hyperthermia, of greater than 41° C. (cf. FIG. **11**) and greater than 40.2° C., was carried out for each of the batches over a total period of 28 days following testing. The results obtained, corresponding to the ratio expressed as a percentage between the number of temperature recordings of greater than 41° C. (or greater than 40.2° C.) and the total number of temperature recordings carried out on all of the pigs per one-week period are reported below in Tables 7 and 8, respectively, for the hyperthermia measurements of greater than 41° C. and greater than 40.2° C.

TABLE 7

	Hyper	thermia > 41	° C	
	F1	F2	F3	F4
W1	4.1	0	0	0
W2	10.7	16.	0	8.9
W3	4.7	27.	0	45.
W4	0	0	0	7.5

TABLE 8

	Hyperthermia > 40.2						
	F1	F2	F3	F4			
W1 W2 W3 W4	29.1 28.5 14.3 3.3	10.41 39.2 68.7 17.5	29.1 10.7 25.0 20.0	20.8 37.5 81.2 55			

[0465] 4) Conclusion

[0466] The recordings carried out clearly show that the animals which received the three injections of a vaccine composition comprising nucleic acid fragments of PWD circovirus according to the invention and/or capable of expressing recombinant proteins of PWD circovirus, in particular of type B, did not exhibit hyperthermia (cf. FIG. **10**). These animals additionally did not experience a decline in their growth, the dmgs being comparable to those of uninfected control animals (cf. FIG. **9**). They did not exhibit any particular clinical sign.

[0467] These results demonstrate the efficacious protection of the piglets against infection with a PWD circovirus of the invention, the primary agent responsible for PWD or FPW, provided by a vaccine composition prepared from a nucleic acid fragment of the nucleic sequence of PWD circovirus according to the invention, in particular of type B, and/or from recombinant proteins encoded by these nucleic acid fragments. **[0468]** These results in particular show that the proteins encoded by the ORF1 and ORF2 of PWD circovirus according to the invention are immunogenic proteins inducing an efficacious protective response for the prevention of infection by a PWD circovirus.

Example 6

Serological Diagnosis of PWD Circovirus by Immunodetermination Using Recombinant Proteins or Synthetic Peptides of PWD Circovirus

[0469] A. Serological Diagnosis with Recombinant Proteins

[0470] The identification and the sequencing of porcine PWD circovirus allow recombinant proteins of PWD circovirus to be produced by the techniques of genetic recombination well known to the person skilled in the art. Using these techniques, recombinant proteins encoded, in particular, by the ORF'2 of the PWD circovirus, type B, were expressed by transformed Sf9 insect cells and then isolated.

[0471] These recombinant proteins encoded by the ORF'2 are extracted, after culture of the transformed Sf9 cells, by thermal cell lysis by means of 3 cycles of freezing/thawing to -70° C./+ 37° C. Healthy Sf9 cells or nontransformed control Sf9 cells are also lysed.

[0472] Two antigenic fractions originating from nontransformed control Sf9 cells and Sf9 cells expressing the ORF'2 are precipitated at 4° C. by a 60% plus or minus 5% saturated ammonium sulfate solution. Determination of total proteins is carried out with the aid of the Biorad kit. 500 ng of control Sf9 proteins and of semipurified Sf9 proteins expressing the ORF'2, in solution in 0.05 M bicarbonate buffer pH 9.6, are passively adsorbed at the bottom of 3 different wells of a Nunc Maxisorp microplate by incubation for one night at +4° C.

[0473] The reactivity of pig sera with respect to each of these antigenic fractions is evaluated by an indirect ELISA reaction of which the experimental protocol is detailed below: [0474] Saturation step: 200μ /well of PBS1×/3% semi-

skimmed milk, 1 h 30 incubation at 37° C.

[0475] Washing: 200 µl/well of PBS1×/Tween 20: 0.05%, 3 rapid washes.

[0476] Serum incubation step: $100 \,\mu$ /well of serum diluted to $\frac{1}{100}$ in PBS1×/semi-skimmed milk, 1%/Tween 20: 0.05%, 1 h incubation at 37° C.

[0477] Washing: 200 μl/well of PBS1×/Tween 20: 0.05%, 2 rapid washes followed by 2 washes of 5 min.

- [0478] Conjugate incubation step: 50 µl/well of rabbit antipig conjugate diluted to ¹/1000 in PBS1×/semi-skimmed milk, 1%/Tween 20: 0.05%, 1 h incubation at 37° C.
- **[0479]** Washing: 200 μl/well of PBS1×/Tween 20: 0.05%, 2 rapid washes followed by 2 washes of 5 min.
- [0480] Visualization step: 100 µl/well of OPD substrate/ citrate buffer/H₂O₂, 15 min incubation at 37° C.
- [0481] Termination: 50 μ /well of 1 N H₂SO₄.
- **[0482]** Read optical density in a spectrophotometer at 490 nm.

Results

[0483] The results obtained are shown below in Table 9.

	TABLE 9	
Antigens	Reactivity of Pig Serum not inoculated with Circovirus	Reactivity of Pig Serum inoculated with Circovirus
Purified Sf9 control Sf9 expressing purified ORF'2	0.076 0.071	0.088 1.035

[0484] The results are expressed in optical density measured in a spectrophotometer at 490 nm during analysis by ELISA of the reactivity of pig sera which are or are not inoculated with the type B PWD circovirus according to the protocol indicated above.

[0485] B. Serological Diagnosis by Synthetic Peptide

[0486] The epitopic mapping of the proteins encoded, for example, by the nucleic sequences ORF1 and ORF2 of the two types of PWD circovirus (types A and B) additionally allowed immunogenic circoviral epitopes to be identified on the proteins encoded by the nucleic sequences ORF'1 and ORF'2 as well as the specific epitopes of the protein encoded by the nucleic acid sequence ORF'2 of the type B PWD circovirus. Four specific epitopes of the type B PWD circovirus and one epitope common to the two types of PWD circovirus situated on the protein encoded by the nucleic sequence ORF'2 were synthesized in peptide form. The equivalent peptides in the circovirus of type A were likewise synthesized. All peptides were evaluated as diagnostic antigens within the context of carrying out a serological test.

Results

[0487] The results obtained are shown in Table 10, below.

TABLE	10
-------	----

Туре						Infected pig serum reactivity Circovirus B		
	Peptide	PWD circovirus	PositionAA seque				Conventional 2 D0/D42	Epitopic specificity
SEQ ID NO: 29	121	В	71-85 VDMMRFN	INDFLPPG +	-/-, +++	+/-, +++	-, +++	Circovirus
SEQ ID NO: 55	177	в	70-84 NVNELRF1	IIGQFLPP +	-/-, +	+/-, +/-	+/-, -	в
SEQ ID NO: 30	132	в	115-129 QGDRGVGS	SAVILDD +	-/-, +/-	++, ++	+/-, +	Circovirus

	Results of the evaluation as a diagnostic antigen of synthetic peptides encoded by the nucleic sequences ORF2 and ORF'2 of PWD circovirus of type A and B.							
			Туре		re	Infected pig eactivity Circ		_
		Peptide	PWD circovirus	PositionAA sequence	SPF D0/D54	Conventional 1 D0/D42	Conventional 2 D0/D42	Epitopic specificity
SEQ ID NO:	56	188	A	114-127 TSNQRGVGSTVVIL	+/-, -	-, +/-	+/-, +/-	В
SEQ ID NO:	31	133	в	119-134 GVGSSAVILDDNVFTH	<-, ++	++, +++	+/-, ++	
SEQ ID NO:	57	189	А	118-132 RGVGSTVVILDANFV	+/-, -	-, +/-	+/-, +/-	
SEQ ID NO:	58	146	в	171-185 FTIDYFQPNNKRNQL	-, +/-	-, ++	-, ++	Circovirus
SEQ ID NO:	59	202	А	170-184 DQTIDWFQPNNKRNQ	+++, +++	+/-, ++	+, ++	A&B
SEQ ID NO:	32	152	в	195-209 VDHVGLGTAFENSIY	-, ++	+++, +++	+/-, +	Circovirus
SEQ ID NO:	60	208	A	194-208 NVEHTGLGYALQNAT	-, -	-, -	-, -	в

+/-, +, ++. Increasing intensities of the reactivities observed in Spot peptides on a nitrocellulose membrane. The porcine sera tested are from animals experimentally infected with the circovirus of type B within the animal houses of the CNEVA. Samples are taken from the animals before inoculation on d0 and 42 days or 54 days after inoculation, on d42, d54.

Example 7

Characterization of the Specific Epitopes of the PWD Circovirus of type B

[0488] The proteins encoded by the ORF2 of the porcine circoviruses of type A and B were chosen for this study. For each of the ORF2s (types A and B), 56 peptides of 15 amino acids which overlap every 4 amino acids were synthesized, thus covering the whole of the protein (cf. Table 11 below).

TABLE	11	1
-------	----	---

Sequence of amino acids of the 56 peptides of 15 amino acids synthesized from the nucleic sequence ORF'2 (type B) and ORF2 (type <u>A) of PWD circovirus with their corresponding spot number (cf. FIG. 12)</u>

Тур	De B ORF'2		Type A ORF2				
SI	pot No.Sequence		Spot No.Sequence				
SEQ ID NO: 171	104 MTYPRRRYRRI	RHRP SEQ ID NO:	175 160 MTWPRRRYRRRTRP				
SEQ ID NO: 172	105 RRRYRRRHRPI	RSHL SEQ ID NO:	176 161 RRRYRRRTRPRSHL				
SEQ ID NO: 173	106 RRRRHRPRSHL	GQIL SEQ ID NO:	177 162 RRRRTRPRSHLGNIL				
SEQ ID NO: 61	107 HRPRSHLGQILI	RRRP SEQ ID NO:	84 163 TRPRSHLGNILRRRP				
SEQ ID NO: 62	108 SHLGQILRRRP	WLVH SEQ ID NO:	85 164 SHLGNILRRRPYLVH				
SEQ ID NO: 63	109 QILRRRPWLVH	PRHR SEQ ID NO:	86 165 NILRRRPYLVHPAFR				
SEQ ID NO: 64	110 RRPWLVHPRHR	YRWR SEQ ID NO:	87 166 RRPYLVHPAFRNRYR				
SEQ ID NO: 65	111 LVHPRHRYRWRI	RKNG SEQ ID NO:	88 167 LVHPAFRNRYRWRRK				
SEQ ID NO: 66	112 RHRYRWRRKNG	IFNT SEQ ID NO:	89 168 AFRNRYRWRRKTGIF				
SEQ ID NO: 67	113 RWRRKNGIFNT	RLSR SEQ ID NO:	90 169 RYRWRRKTGIFNSRL				
SEQ ID NO: 68	114 KNGIFNTRLSR	IFGY SEQ ID NO:	91 170 RRKTGIFNSRLSREF				
SEQ ID NO: 69	115 FNTRLSRTFGY	IVKR SEQ ID NO:	92 171 GIFNSRLSREFVLTI				

Sequence of ami	no acids of the 56 peptide	es of 15 amino acids
synthesized from t	ne nucleic sequence ORF'2	(type B) and ORF2 (type
A) of PWD circovirus	with their corresponding	spot number (cf. FIG. 12)

	Type B ORF'2	Type A ORF2	
	Spot No.Sequence	Spot No.Sequence	
SEQ ID NO: 70	116 LSRTFGYTVKRTTVR	SEQ ID NO: 93 172 SRLSREFVLTIR	रGGH
SEQ ID NO: 71	117 FGYTVKRTTVRTPSW	SEQ ID NO: 94 173 REFVLTIRGGHS	SQPS
SEQ ID NO: 72	118 VKRTTVRTPSWAVDM	SEQ ID NO: 95 174 LTIRGGHSOPSW	VNVN
SEQ ID NO: 73	119 TVRTPSWAVDMMRFN	SEQ ID NO: 96 175 GGHSQPSWNVNE	ILRF
SEQ ID NO: 74	120 PSWAVDMMRFNINDF	SEQ ID NO: 97 176 QPSWNVNELRFN	1IGO
SEQ ID NO: 29	121 VDMMRFNINDFLPPG	SEQ ID NO: 98 177 NVNELRFNIGQF	7LPP
SEQ ID NO: 75	122 RFNINDFLPPGGGSN	SEQ ID NO: 99 178 LRFNIGQFLPPS	GGT
SEQ ID NO: 76	123 NDFLPPGGGSNPRSV	SEQ ID NO: 100 179 IGQFLPPSGGTN	1PLP
SEQ ID NO: 77	124 PPGGGSNPRSVPFEY	SEQ ID NO: 101 180 LPPSGGTNPLPL	_PFQ
SEQ ID NO: 78	125 GSNPRSVPFEYYRIR	SEQ ID NO: 102 181 GGTNPLPLPFQY	YYRI
SEQ ID NO: 79	126 RSVPFEYYRIRKVKV	SEQ ID NO: 103 182 PLPLPFQYYRIR	RKAK
SEQ ID NO: 80	127 FEYYRIRKVKVEFWP	SEQ ID NO: 104 183 PFQYYRIRKAKY	YEFY
SEQ ID NO: 81	128 RIRKVKVEFWPCSPI	SEQ ID NO: 105 184 YRIRKAKYEFYF	PRDP
SEQ ID NO: 82	129 VKVEFWPCSPITQGD	SEQ ID NO: 106 185 KAKYEFYPRDPI	ITSN
SEQ ID NO: 83	130 FWPCSPITQGDRGVG	SEQ ID NO: 107 186 EFYPRDPITSNQ	QRGV
SEQ ID NO: 174	131 SPITQGDRGVGSSAV	SEQ ID NO: 108 187 RDPITSNQRGVG	JSTV
SEQ ID NO: 30	132 QGDRGVGSSAVILDD	SEQ ID NO: 109 188 TSNQRGVGSTVV	/ILD
SEQ ID NO: 31	133 GVGSSAVILDDNFVT	SEQ ID NO: 136 189 RGVGSTVVILDA	4NFV
SEQ ID NO: 111	134 SAVILDDNFVTKATA	SEQ ID NO: 137 190 STVVILDANFVI	[PST
SEQ ID NO: 112	135 LDDNFVTKATALTYD	SEQ ID NO: 138 191 ILDANFVTPSTN	1LAY
SEQ ID NO: 113	136 FVTKATALTYDPYVN	SEQ ID NO: 139 192 NFVTPSTNLAYD	OPYI
SEQ ID NO: 114	137 ATALTYDPYVNYSSR	SEQ ID NO: 140 193 PSTNLAYDPYIN	IYSS
SEQ ID NO: 115	138 TYDPYVNYSSRIITIT	SEQ ID NO: 141 194 LAYDPYINYSSR	RHTI
SEQ ID NO: 116	139 YVNYSSRHTITQPFS	SEQ ID NO: 142 195 PYINYSSRHTIR	RQPF
SEQ ID NO: 117	140 SSRHTITQPFSYHSR	SEQ ID NO: 143 196 YSSRIITIRQPF	TYHS?
SEQ ID NO: 118	141 TITQPFSYHSRYFTP	SEQ ID NO: 144 197 HTIRQPFTYHSR	lyft
SEQ ID NO: 119	142 PFSYHSRYFTPKPVL	SEQ ID NO: 145 198 QPFTYHSRYFTF	PKPE
SEQ ID NO: 120	143 HSRYFTPKPVLDFTI	SEQ ID NO: 146 199 YHSRYFTPKPEL	DQT
SEQ ID NO: 121	144 FTPKPVLDFTIDYYFÇ) SEQ ID NO: 147 200 YFTPKPELDQTI	IDWF
SEQ ID NO: 122	145 PVLDFTIDYFQPNNK	SEQ ID NO: 148 201 KPELDQTIDWFQ) 29NN
SEQ ID NO: 123	146 FTIDYFQPNNKRNQL	SEQ ID NO: 149 202 DQTIDWFQPNNK	KRNQ
SEQ ID NO: 124	147 YFQPNNKRNQLWLRL	SEQ ID NO: 150 203 DWFQPNNKRNQL	JWLH
SEQ ID NO: 125	148 NNKRNQLWLRLQTAG	SEQ ID NO: 151 204 PNNKRNQLWLHL	INTH
SEQ ID NO: 126	149 NQLWLRLQTAGNVDH	SEQ ID NO: 152 205 RNQLWLHLNTHT	INVE

TABLE 11-continued

Sequence of amino acids of the 56 peptides of 15 amino acids synthesized from the nucleic sequence ORF'2 (type B) and ORF2 (type <u>A) of PWD circovirus with their corresponding spot number (cf. FIG. 12)</u>

	Type B ORF'2				Type A ORF2						
	Spot No.Sequence				Spot No.Sequence						
SEQ	ID	NO :	127	150	LRLQTAGNVDHVGLG	SEQ	ID	NO:	153	206	WLHLNTHTNVEHTGL
SEQ	ID	NO:	128	151	TAGNVDHVGLGTAFE	SEQ	ID	NO:	154	207	NTHTNVEHTGLGYAL
SEQ	ID	NO:	32	152	VDHVGLGTAFENSIY	SEQ	ID	NO:	155	208	NVEHTGLGYALQNAT
SEQ	ID	NO:	129	153	GLGTAFENSIYDQEY	SEQ	ID	NO :	156	209	TGLGYALQNATTAQN
SEQ	ID	NO:	130	154	AFENSIYDQEYNIRV	SEQ	ID	NO :	157	210	YALQNATTAQNYVVR
SEQ	ID	NO:	131	155	SIYDQEYNIRVTMYV	SEQ	ID	NO :	158	211	NATTAQNYVVRLTIY
SEQ	ID	NO:	132	156	QEYNIRVTMYVQFRE	SEQ	ID	NO :	159	212	AQNYVVRLTIYVQFR
SEQ	ID	NO :	133	157	IRVTMYVQFREFNFK	SEQ	ID	NO:	160	213	VVRLTIYVQFREFIL
SEQ	ID	NO:	134	158	MYVQFREFNFKDPPL	SEQ	ID	NO :	161	214	TIYVQFREFILKDPL
SEQ	ID	NO :	135	159	VQFREFNFKDPPLNP	SEQ	ID	NO :	162	215	YVQFREFILKDPLNE

[0489] These peptides were synthesized according to the "spot" method which consists of simultaneous synthesis of a large number of peptides on a cellulose solid support, each site of synthesis of a peptide constituting a spot (Synt:em, NIMES). This method involves orientation of the peptides on the plate, these being fixed covalently by the carboxy-terminal end. A spot represents approximately 50 nmol of peptide. **[0490]** The reference of the spots and corresponding peptide sequences is given in Table 11.

[0491] These membranes were used for immunoreactivity tests with respect to serum of SPF pigs which were or were not infected experimentally with the type B PWD circoviral strain as well as with respect to sera of infected pigs from conventional farms (conventional farms 1 or 2). This study allowed specific immunoreactive peptides of the circovirus of type B corresponding to the spots No. 121, No. 132, No. 133 and No. 152 (respectively of amino acid sequences SEQ ID No. 29, SEQ ID No. 30, SEQ ID No. 31 and SEQ ID No. 32) to be demonstrated. An illustration is shown in FIG. **12** where the membranes are visualized with an infected pig serum coming from a conventional farm. Nonspecific immunoreactive peptides of type [lacuna] were likewise demonstrated, among which we shall keep the peptide No. 146 SEQ ID No. 123 which is strongly immunogenic.

[0492] A comparison between the peptide sequences of circoviruses of type A and B (FIG. **13**) indicates a divergence ranging from 20 to 60% for the specific immunoreactive peptides of the type B, and a weaker divergence (13%) between the nonspecific peptides.

Example 8

Protection of Swine From Post-Weaning Multisystemic Wasting Syndrome (PMWS) Conferred by Procine Circovirus Type B (PCV-B) ORF'2 Protein.

[0493] The ORF'1-encoded protein (REP) and ORF'2-encoded putative capsid protein of PCV-B were expressed, either in insect cells by recombinant baculovirus vectors, or in

mammalian cell lines by transfection with plasmidic expression vectors. These two circovirus-derived proteins were detectable in both expression systems. As evaluated by weight gains, hyperthermia and absence of lesions following challenge, the pigs were protected against a virulent circovirus challenge after one first DNA immunization with plasmids directing ORF'2 protein and GM-CSF expression and a second injection, 15 days later, with the same plasmid preparation plus the ORF'2 recombinant protein. A lower level of protection was observed when the pigs were vaccinated with ORF'1 protein, as opposed to pigs vaccinated with ORF'2 protein.

A. Development of an Experimental Model of PMWS in Swine:

[0494] Eight 3 week-old SPF pigs were inoculated intratracheally (5 ml) and intramuscularly (1 ml).

B. Production and Control of PCV-B Plasmids:

[0495] PCV-B ORF'1 and ORF'2 genes, isolated from PCV-B challenge strain, was cloned into vector plasmid pcDNA3.1. All constructs were validated through a partial sequencing of the PCV-B genes in the final plasmids and expression control by immunoperoxidase on PK15 cells respectively transfected with each plasmid, using swine polyclonal antibodies.

[0496] Plasmid encoding GM-CSF has been co-administered.

C. Construction of Recombinant Baculoviruses:

[0497] ORF'1 and ORF'2 proteins were expressed under polyhedrin promoter control. Recombinant proteins were detected by western-blot using swine polyclonal antibodies.

D. Vaccination and Challenge:

[0498] Four groups of 7 pigs were vaccinated intramuscularly at day 0 (Do), two weeks later, they received the same plasmid preparation plus the recombinant baculovirus.

E. Monitoring:

[0499] All groups of pigs were housed in isolated experimental units with air filtration and low air pressure. Clinical observations and rectal temperatures were recorded every day. The pigs were weighed weekly.

F. Conclusions

[0500] Expression of PCV-B ORF'2 or PCV-B ORF'1 in swine resulted in a significantly enhanced level of protection as evaluated by weight evolution and body temperature evolution following challenge with PCV-B circovirus. These results are summarized in FIGS. **14** and **15**.

[0501] The invention described herein may be embodied in other specific forms without departing from the spirit or essential characteristics thereof. The specific embodiments previously described are therefore to be considered as illustrative of, and not limiting, the scope of the invention. Additionally, the disclosure of all publications and patent applications cited above and below, including International Patent Application No. PCT/FR98/02634, filed Dec. 4, 1998, and published as International Publication No. WO 99/29871 on Jun. 17, 1999, are expressly incorporated herein by reference in their entireties to the same extent as if each were incorporated by reference individually.

BIBLIOGRAPHIC REFERENCES

- [0502] Allan, G. M. et al., 1995, Vet. Microbiol., 44: 49-64.
- [0503] Barany, F., 1911, PNAS. USA, 88: 189-193.
- [0504] Boulton, L. H. et al., 1997, J. Gen. Virol., 78 (Pt 6), 1265-1270.
- [0505] Buckholz, R. G., 1993, Yeast systems for the expression of heterologous gene products. Curr. Op. Biotechnology 4: 538-542.
- [0506] Burg, J. L. et al., 1996, Mol. and Cell. Probes, 10: 257-271.
- [0507] Chu, B. C. F. et al., 1986, NAR, 14: 5591-5603.
- [0508] Chu, P. W. G. et al., 1993, Virus Research, 27: 161-171.
- [0509] Clark, E. G., 1997, American Association of Swine Practitioners, 499-501.
- **[0510]** Daft, B. et al., 1996, American Association of Veterinary Laboratory Diagnosticians, 32.
- [0511] Derse, D. et al., 1995, J. Virol., 69(3): 1907-1912.
- [0512] Duck, P. et al., 1990, Biotechniques, 9: 142-147.
- [0513] Dulac, G. C. et al., 1989, Can. J. Vet. Res., 53: 431-433.
- **[0514]** Edwards, C. P., and Aruffo, A., 1993, Current applications of COS cell based transient expression systems. Curr. Op. Biotechnology 4: 558-563.
- [0515] Edwards, S. et al., 1994, Vet. Rec., 134: 680-681.
- **[0516]** Erlich, H. A., 1989, In PCR Technology. Principles and Applications for DNA Amplification. New York: Stockton Press.
- [0517] Felgner, et al., 1987, Proc. Natl. Acad. Sci., 84: 7413.
- [0518] Fontes, E. P. B. et al., 1994, J. Biol. Chem., Vol. 269, No. 11: 8459-8465.
- [0519] Fraley et al., 1980, J. Biol. Chem., 255: 10431.
- [0520] Guateli, J. C. et al., 1990, PNAS. USA, 87: 1874-1878.
- [0521] Hackland, A. F. et al., 1994, Arch. Virol., 139: 1-22.
- [0522] Hanson, S. F. et al., 1995, Virology, 211: 1-9.

- [0523] Harding, J. C., 1997, American Association of Swine Practitioners, 503.
- [0524] Harding, R. M. et al., 1993, Journal of General Virology, 74: 323-328.
- [0525] Harding, J. C. and Clark, E. G., 1997, Swine Health and Production, Vol. 5, No. 5: 201-203.
- [0526] Heyraud-Nitschke, F. et al., 1995, Nucleic Acids Research, Vol. 23, No. 6.
- [0527] Horner, G. W., 1991, Surveillance 18(5): 23.
- [0528] Houben-Weyl, 1974, in Methode der Organischen Chemie, E. Wunsch Ed., Volume 15-I and 15-II, Thieme, Stuttgart.
- [0529] Huygen, K. et al., 1996, Nature Medicine, 2(8): 893-898.
- [0530] Innis, M. A. et al., 1990, in PCR Protocols. A guide to Methods and Applications, San Diego, Academic Press.
- **[0531]** Kaneda, et al., 1989, Science, 243: 375.
- [0532] Kievitis, T. et al., 1991, J. Virol. Methods, 35: 273-286.
- [0533] Kohler, G. et al., 1975, Nature, 256(5517): 495-497.
- [0534] Kwoh, D. Y. et al., 1989, PNAS. USA, 86: 1173-1177.
- [0535] Ladany, S. et al., 1989, J. Clin. Microbiol. 27: 2778-2783.
- [0536] Lazarowitz, S. G. et al., 1989, The EMBO Journal, Vol. 8 No. 4: 1023-1032.
- [0537] Luckow, V. A., 1993, Baculovirus systems for the expression of human gene products. Curr. Op. Biotechnology 4: 564-572.
- [0538] Mankertz, A. et al., 1997, J. Virol., 71: 2562-2566.
- **[0539]** Matthews, J. A. et al., 1988, Anal. Biochem., 169: 1-25.
- [0540] McNeilly, F. et al., 1996, Vet. Immunol. Immunopathol., 49: 295-306.
- [0541] Meehan, B. M. et al., 1997, J. Gen. Virol. 78: 221-227.
- [0542] Merrifield, R. D., 1966, J. Am. Chem. Soc., 88(21): 5051-5052.
- [0543] Midoux, 1993, Nucleic Acids Research, 21: 871-878.
- [0544] Miele, E. A. et al., 1983, J. Mol. Biol., 171: 281-295.
- [0545] Murphy, F. A. et al., 1995, Sixth Report of the International Committee on Taxonomy of Viruses. Springer-Verlag Wien New York.
- [0546] Nayar, G. P. et al., 1997, Can. Vet. J. 38(6): 385-386.
- [0547] Olins, P. O., and Lee, S. C., 1993, Recent advances in heterologous gene expression in *E. coli*. Curr. Op. Biotechnology 4: 520-525.
- [0548] Pagano et al., 1967, J. Virol., 1: 891.
- **[0549]** Rolfs, A. et al., 1991, In PCR Topics. Usage of Polymerase Chain reaction in Genetic and Infectious Disease. Berlin: Springer-Verlag.
- **[0550]** Sambrook, J. et al., 1989, In Molecular cloning: A Laboratory Manual. Cold Spring Harbor, N.Y.: Cold Spring Harbor Laboratory Press.
- [0551] Sanchez-Pescador, R., 1988, J. Clin. Microbiol., 26(10): 1934-1938.
- [0552] Segev D., 1992, in "Non-radioactive Labeling and Detection of Biomolecules". Kessler C. Springer Verlag, Berlin, N.Y.: 197-205.
- [0553] Shiver, J. W., 1995, in Vaccines 1995, eds Chanock, R.M. Brown, F. Ginsberg, H.S. & Norrby, E., pp. 95-98, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, N.Y.
- [0554] Tascon, R. E. et al., 1996, Nature Medicine, 2(8): 888-892.
- [0555] Tischer, I. et al., 1982, Nature, 295: 64-66.
- [0556] Tischer, I. et al., 1986, Arch. Virol., 91: 271-276.

- [0557] Tischer, I. et al., 1988, Zentralbl Bakteriol Mikrobiol Hyg [A] 270: 280-287.
- [0558] Tischer, I. et al., 1995, Arch. Virol., 140: 737-743.
- [0559] Urdea, M. S., 1988, Nucleic Acids Research, II: 4937-4957.
- [0560] Walker, G. T. et al., 1992, NAR 20: 1691-1696.
- [0561] Walker, G. T. et al., 1992, PNAS. USA, 89: 392-396. [0562] White, B. A. et al., 1997, Methods in Molecular
- Biology, 67, Humana Press, Towota.
- [0563] Zhao, T. M. et al., 1996, Proc. Natl. Acad. Sci., USA 93(13): 6653-6648.

SEQUENCE LISTING

<160> NUMBER OF SEQ ID NOS: 177 <210> SEQ ID NO 1 <211> LENGTH: 1759 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1)..(78) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (82)..(99) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (106)..(156) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (160)..(195) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (199)..(231) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (235)..(246) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (250)..(315) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (319)..(330) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (334)..(489) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (493)..(525) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (529)..(591) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (595)..(600) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (604)..(606) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (610)..(627) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (634)..(636) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (640)..(681) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (685)..(708) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (712)..(726) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (730)..(753) <220> FEATURE: <221> NAME/KEY: CDS

-continued

48

96

144

<222> LOCATION: (757)..(933) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (937)..(969) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (973)..(1047) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1051)..(1056) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1060)..(1071) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1075)..(1236) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1240)..(1257) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1261)..(1293) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1297)..(1350) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1354)..(1380) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1384)..(1386) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1390)..(1416) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1420)..(1425) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1429)..(1497) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1501)..(1512) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1516)..(1551) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1555)..(1566) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1570)..(1581) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1585)..(1620) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1624)..(1752) <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1756)..(1758) <400> SEQUENCE: 1 acc agc gca ctt cgg cag cgg cag cac ctc ggc agc gtc agt gaa aat Thr Ser Ala Leu Arg Gln Arg Gln His Leu Gly Ser Val Ser Glu Asn 5 10 1 15 gcc aag caa gaa aag cgg ccc gca acc cca taa gag gtg ggt gtt cac Ala Lys Gln Glu Lys Arg Pro Ala Thr Pro Glu Val Gly Val His 20 25 30 cct taataa tcc ttc cga gga gga gaa aaa caa aat acg gga gct tcc Pro Ser Phe Arg Gly Gly Glu Lys Gln Asn Thr Gly Ala Ser 35 40 45

28

-continued

_												<u> </u>	<u>ucu</u>			
	ctc Leu			tga		-	ttg Leu		-					-	192	
gjà aaa	-						gtt Val					taa	gaa Glu		240	
-	ttt Phe	taa					ttt Phe								288	
							gaa Glu				ctg Leu			aga Arg	336	
							agc Ser								384	
							tac Tyr								432	
							tgt Cys								480	
	gct Ala		-				gag Glu							tga	528	
							agt Ser								576	
	cca Pro				taa	tgc Cys	tga	gcc Ala	tag		cac His				624	
gcc Ala 195	tagi	-	aaa Lys			Gly	tgg a Trp : 200				Arg i				672	
-	ttt Phe		-				gtt Val 215						act Thr 220		720	
	gtg Val	tga					tgt Cys			taa			tac Tyr	-	768	
							gat Asp								816	
							ccc Pro 260								864	
							gaa Glu								912	
					ccg Pro		agc Ser				acc Thr 295				960	
	ccc Pro 300						agt Ser								1008	
-	-					Leu	gag Glu							ctc Leu	1056	

-continued

												con		ueu				
tga	att Ile	gta Val 330			-								999 Gly 340	-		1104		
	cat His															1152		
	tat Tyr 360															1200		
	aat Asn													acc Thr		1248		
	ggt Gly				gct Ala										taa	1296		
	agg Arg 405															1344		
	aga Arg				tgg Trp							tag	agg Arg	tga	tgg Trp	1392		
	ctc Leu							-		ttc Phe	taa		ggt Gly	-		1440		
	aag Lys															1488		
	cga Arg	-	tga		tca Ser	-	-	taa			-	-	gct Ala		-	1536		
	ctc Leu						caa Gln			tag			д1У ддд		tga	1584		
	tac Tyr												agg Arg			1632		
	cca Pro															1680		
	д1у 999		-		-			-		-	-	-				1728		
	agt Ser								tat Tyr 545	t						1759		
<21 <21	0> SH 1> LH 2> TY 3> OH	ENGTI ZPE :	I: 54 PRT	15	e A I	PWD (circo	oviru	18									
< 40	0> SI	EQUEI	ICE :	2														
Thr 1	Ser	Ala	Leu	Arg 5	Gln	Arg	Gln	His	Leu 10	Gly	Ser	Val	Ser	Glu 15	Asn			
Ala	Lys	Gln	Glu 20	Lys	Arg	Pro	Ala	Thr 25	Pro	Glu	Val	Gly	Val 30	His	Pro			
Ser	Phe	Arg 35	Gly	Gly	Glu	ГЛа	Gln 40	Asn	Thr	Gly	Ala	Ser 45	Asn	Leu	Pro			

-continued

Phe	Leu 50	Phe	Суз	Leu	Trp	Arg 55	Gly	Arg	Phe	Gly	Arg 60	Gly	Asn	Ser	Ser
Pro 65	Pro	Gly	Val	Суз	Glu 70	Phe	Суз	Glu	Ala	Asp 75	Phe	Gln	Gly	Glu	Val 80
Val	Phe	Trp	Сүз	Pro 85	Leu	Pro	His	Arg	Glu 90	Ser	Glu	Arg	Asn	Arg 95	Pro
Ala	Glu	Arg	Ile 100	Leu	Gln	Arg	Arg	Pro 105	His	Thr	Tyr	Arg	Val 110	Trp	Ser
Ser	Ala	Glu 115	Pro	Gly	Glu	Ala	Gln 120	Arg	Pro	Val	Tyr	Cys 125	Суз	Glu	Tyr
Pro	Phe 130	Gly	Asp	Gly	Val	Phe 135	Gly	Asp	Сув	Ser	Arg 140	Ala	Val	Ser	Суз
Asn 145	Val	Сув	Glu	Lys	Phe 150	Pro	Arg	Ala	Gly	Thr 155	Phe	Glu	Ser	Glu	Arg 160
Glu	Asp	Ala	Glu	Ala 165	Leu	Glu	Asp	Ser	Cys 170	Thr	Arg	His	Ser	Gly 175	Pro
Ala	Arg	Leu	Trp 180	Glu	Glu	Pro	Val	Gly 185	Pro	Phe	Сүз	Ala	Gly 190	His	Leu
Leu	Glu	Ala 195	Lys	Val	Val	Gly	Trp 200	Ile	Ser	Trp	Arg	Arg 205	Ser	Суз	Cys
Phe	Gly 210	Phe	Leu	Trp	Leu	Val 215	Thr	Leu	Gly	Ser	Thr 220	Glu	Thr	Val	Pro
Val 225	Ser	Ile	Asp	Сүз	Arg 230	Asp	Arg	Gly	Tyr	Сув 235	Ser	Phe	Phe	Gly	Pro 240
Gln	Tyr	Phe	Asp	Tyr 245	Gln	Gln	Ser	Gly	Pro 250	Pro	Gly	Met	Val	Leu 255	Leu
Asn	Сув	Суз	Pro 260	Ser	Суз	Arg	Ser	Ser 265	Leu	Ser	Glu	Asp	Tyr 270	Tyr	Phe
Ala	Ile	Leu 275	Glu	Asp	Суз	Trp	Arg 280	Thr	Ile	His	Gly	Gly 285	Thr	Arg	Arg
Pro	Ile 290	Ser	Ser	Gly	Pro	Thr 295	Leu	Суз	Pro	Phe	Pro 300	Ile	Asn	Lys	Leu
Leu 305	Ser	Leu	Phe	Сүз	Tyr 310	His	Ile	Val	Met	Val 315	Phe	Ile	Phe	Ile	His 320
Leu	Glu	Gly	Leu	Ser 325	Gly	Ile	Leu	Ile	Val 330	His	Lys	Ser	Thr	Leu 335	Pro
His	Asn	Phe	Gly 340	Leu	Trp	Leu	His	Phe 345	Gly	Ala	His	Ser	Pro 350	Gly	Leu
Сүз	Ala	Arg 355	His	Trp	Суз	Gly	Tyr 360	Leu	Asn	Gly	Ala	Thr 365	Ala	Gly	Phe
Phe	Tyr 370	Tyr	Leu	Ala	Gly	Thr 375	Asn	Gln	Leu	Phe	Gly 380	Leu	Ala	Leu	Val
Trp 385	Gly	Ser	Thr	Trp	Ser 390	Gly	Arg	Arg	Ala	Ala 395	Leu	Trp	Сүз	Gly	Gly 400
Arg	Ser	Ser	Tyr	Arg 405	Gly	His	Arg	Pro	Ser 410	Trp	Trp	Arg	Gly	Leu 415	Gln
Ser	Trp	His	Pro 420	Arg	Gln	Gln	Trp	Thr 425	Gln	His	Leu	Phe	Asp 430	Arg	Trp
Gly	Leu	Trp 435	Gly	Lys	Ile	His	Ile 440	Pro	Phe	Tyr	Gly	Ser 445	Ile	Gly	Lys

$ \begin{array}{c} Val Gly Gly Trp Cye Arg Leu Arg Gly Gly Gly Thr Gly Arg 455 (2) and 2) a$											-	con	tin	ued	
445 470 475 480 Trp Val Gln Ile Leu Lys Gly Gly Asn Arg Tyr Pro Ser Phe Gly Ala 485 480 Gly Cys Asn Gly Phe Arg Arg Gly Gly Gly Gly Ser Val Phe Cys Gly Asn Soo 500 700 810 <td></td> <td>Val</td> <td>Gly</td> <td>Gly</td> <td>Trp</td> <td></td> <td>Arg</td> <td>Leu</td> <td>Arg</td> <td>Gly</td> <td></td> <td>Gly</td> <td>Thr</td> <td>Gly</td> <td>Arg</td>		Val	Gly	Gly	Trp		Arg	Leu	Arg	Gly		Gly	Thr	Gly	Arg
$ \begin{array}{cccccccccccccccccccccccccccccccccccc$	-	Ser	Ala	Arg		Ser	ГЛа	Met	Ala		Ser	Val	Leu	Leu	
$ \begin{array}{c} 11 \ 0 \ ys \ As \ 0 \ ys \ 0 \ 0 \ ys \ 0 \ 0 \ 0 \ 0 \ 0 \ 0 \ 0 \ 0 \ 0 \ $		Gln	Ile			Gly	Gly	Asn		Tyr	Pro	Ser	Phe	-	Ala
Gly Cys Phe Glu Asp Gly Cys Gly Gly Gly Ser Val Pas Cys Gly Asp Ala Ser Leu Ala Thr Ser Ser Tyr Ser Glu Arg Ser Ala Leu Leu Tyr Ser Leu Ala Thr Ser Ser Ser Glu Arg Ser Ala Leu Leu Tyr Ser Ser Leu Arg Ser Ser Ser Ser Ser Ala Leu Leu Leu C210> SEQ ID NO 3 Ser Ser <td< td=""><td>Ile Cys</td><td>Asn</td><td></td><td></td><td>Arg</td><td>Arg</td><td>Gly</td><td></td><td></td><td>Asn</td><td>Met</td><td>Val</td><td></td><td></td><td>Gly</td></td<>	Ile Cys	Asn			Arg	Arg	Gly			Asn	Met	Val			Gly
Ala Sar Leu Ala The Sar Sar Tyr Sar S	Gly Cys	Phe		Asp	Gly	Суз	Gly		Gly	Ser	Val	Phe		Gly	Asn
530 535 540 Tyr 545 <210> SEQ ID NO 3 <211> LENGTH: 577 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 3 Pro Ala His Phe Gly Ser Gly Ser Thr Ser Ala Ala Ser Val Lys Met 1 1 10 200 25 Leu Asn Asn Pro Ser Glu Glu Glu Glu Lys Asn Lys Ile Arg Glu Leu Pro 45 11 50 Gly Arg Thr Pro His Leu Gln Gly Phe Ala Asn Phe Ala Lys Lys Gln 60 65 70 70 70 71 70 70 70 70 75 71 80 70 70 70 75 70 70 70 75 70 75 70 70 70 75 70 70 70 75 70 75 70 70 70 75 70 75 70 75 70 75 70 <			Ala	Thr	Ser	Ser		Lys	Ser	Glu	Arq		Ala	Leu	Leu
545 <210 > SEQ ID NO 3 <211 > LENGTH: 577 <212 > TYPE: PRT <212 > ORGANISM: Type A PWD circovirus $<400 > SEQUENCE: 3Pro Ala His Phe Gly Ser Gly Ser Thr Ser Ala Ala Ser Val Lys Met110$	530						1	1			-				
<pre><11> LENGTH: 577 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 3 Pro Ala His Phe Gly Ser Gly Ser Thr Ser Ala Ala Ser Val Lys Met 1 1 Pro Ser Lys Lys Ser Gly Pro Gln Pro His Lys Arg Trp Val Phe Thr 25 10 Ser Leu Asn Asn Pro Ser Glu Glu Glu Lys Asn Lys Ile Arg Glu Leu Pro 50 Gly Arg Thr Pro His Leu Gln Gly Phe Ala Asn Phe Ala Lys Lys Gln 65 Thr Phe Asn Lys Val Lys Trp Tyr Phe Gly Ala Arg Cys His Ile Glu 95 Lys Ala Lys Gly Thr Asp Gln Gln Asn Lys Glu Tyr Cys Ser Lys Glu 115 Ser Asp Leu Ser Thr Ala Val Ser Thr Leu Leu Glu Thr Gly Ser Leu 130 Cly Leu Ala Glu Lie Lys Val Ser Thr Leu Leu 140 Ser Asp Leu Ser Thr Ala Val Ser Thr Leu Leu 140 Ser Asp Leu Ala Glu Lie Lys Val Ser Thr Lie 155 Ser Gln 155 Clys Thr Ala Val His Val 110 Cly Leu Ala Glu Lus Asn Phe 155 Clys Cla Cly Cly Cly Cly Cly Cly Cly Cly Cly Cly</pre>	-														
<400> SEQUENCE: 3 Pro Ala His Phe Gly Ser Gly Ser Thr Ser Ala Ala Ser Val Lys Met 1 For Ser Lys Lys Ser Gly Pro Gln Pro His Lys Arg Try Val Phe Thr 20 Ser Lys Lys Ser Glu Glu Glu Lys Asn Lys IIe Arg Glu Leu Pro 40 For Ser Lys Lys Pro Ser Glu Glu Glu Lys Asn Lys IIe Arg Glu Leu Pro 45 Gly Arg Thr Pro His Log Gln Gln Gly Phe Ala Arg Phe Ala Lys Lys Gln 65 For Ala Lys Gly Thr Arg Asn Lys Try Try Phe Gly Ala Arg Cys His 116 Gly Ala Lys Gly Thr Arg Gln Cln Gln Gln Asn Lys Glu Glu Glu Gly Leu Glu Glu 610 Gly His 116 For Ala Lys Gly Thr Arg Gln Gln Gln Asn Lys Glu Try Cys Ser Lys Glu 610 Gly His 116 For Ala Clu Ser Thr Ala Val Ser Thr Leu Leu Glu Glu Glu Gln Gly Lys Arg 110 For Lys Thr Ala Glu Leu Lys Val Ser Thr Leu Leu Glu 110 For Ser Arg Asn Lys Ra Clu	<211> LE <212> TY	NGTH	1: 5' PRT	77	e A 1	PWD (circ	ovir	18						
1 5 10 15 Pro Ser Lys Lys Ser Gly Pro Gln Pro His Lys Arg Trp Val Phe Thr Leu Asn Asn Pro Ser Glu Glu Glu Lys Asn Lys Arg Glu Glu Pro Asn Asn Pro Ser Glu Glu Glu Lys Asn Lys Ile Arg Glu Glu Glu Asn Asn Pro Asn Try Phe Val Cys Glu Glu Glu Glu Glu Glu Fue Asn Pro Fue Ser Glu Glu Fue Asn Pro Fue Ser Try Pro Pro Pro Pro Pro Pro Ser Fue Ser Fue <td></td>															
20 25 30 Leu Asn Asn Pro Ser Glu Glu Glu Lys Asn Lys Ile Arg Glu Leu Pro Ile Ser Leu Phe Asp Tyr Phe Val Cys Glu		His	Phe		Ser	Gly	Ser	Thr		Ala	Ala	Ser	Val		Met
35 40 45 Ile Ser Leu Phe Asp Tyr 55 Val Cys Glu Glu Gly Gly Gly Gly Arg Thr Pho His Leu Gly Gly Arg Thr Pho His Leu Gly Fr Ala Asp Phe Ala Asp Pho Ala Lys Gly Gly Arg Thr Pho His Lys Tyr Tyr Tyr Pho Gly Ala Lys His Ile Gly Tyr Pho Gly Ala Arg Cys His Ile Gly Tyr Tyr Pho Gly Ala Lys Gly Tyr Ala Cys Gly Tyr Str Gly Glu Tyr Str Gly Glu Str Gly Str Gly Glu Tyr Str Glu Glu Str Glu Glu Str Glu Glu Str Glu Str Glu Str <	Pro Ser	Lys		Ser	Gly	Pro	Gln		His	Lya	Arg	Trp		Phe	Thr
50 55 60 Gly Arg Thr Pro His Leu Gln Gly Phe Ala Asn Phe Ala Lys Lys Gln $_{65}^{75}$ Kan Ser			Pro	Ser	Glu	Glu		Lys	Asn	Lya	Ile		Glu	Leu	Pro
65707580ThrPheAsnLysValLysTrpTyrPheGlyAlaArgCysHisIleGluLysAlaLysGlyThrAspGlnGlnAsnLysGluTyrCysSerLysGluGlyHisIleLeuIleGluCysGlyAlaProArgAsnGlnGlyLysArgGlyHisIleLeuIleGluCysGlyAlaProArgAsnGlnGlyLysArgSerAspLeuSerThrAlaValSerThrLeuLeuGluThrGlySerLeuValThrValAlaGluGlnPheProValThrTyrValArgAsnPheArg145ThrValAlaGluGlnPheProValThrTyrValArgAsnPheArg145ThrValAlaGluGluSerGlyLysMetGlnLysArgAspAsp145ThrValAlaGluLeuLysValSerGlyLysMetGlnLysArgAspAsp160145LysThrAlaGluSerClyLysMetGlnLysArgAsp		Leu	Phe	Asp	Tyr		Val	Сув	Gly	Glu		Gly	Leu	Glu	Glu
ThrPheAsnLysValLysTrpTyrPheGlyAlaArgCysHisJleGluLysAlaLysGlyThrAspGlnGlnAsnLysGluTyrCysSerLysGluGlyHisIleLusIleGluCysGlyAlaProArgAsnGluTyrCysSerLysArgGlyHisIleLeuIleGluCysGlyAlaProArgAsnLusGlySerLeuMaiThrValAlaGluGlnProValThrLeuGluArgAsnProMaiThrValAlaGluGlnProValThrTyrValArgAsnProMaiThrValAlaGluGlnProValThrTyrTyrArgAsnMaiThrValGluGluLeuLeuLusCluTyrTyrTyrArgMaiThrValCluLeuLysValSerGluThrTyrArgAsnArgMaiSerThrLusKaiCluLusKaiGluFinoTyrTyrTyrTyrArgMaiLusMaiMaiMaiMaiSerGluFinoFinoFinoFinoFin		Thr	Pro	His		Gln	Gly	Phe	Ala		Phe	Ala	Lys	Lys	
LysAlaLysGlyThrAspGlnGlnAspGluTyrGluTyrCysSerLysGluGlyHisIILeuIleGluCysGlyAlaProArgAsnGlnGlyLysArgSerAspLeuSerThrAlaValSerThrLeuGluThrGlySerLeuValThrValGluGluGlnGlnProValThrTyrValArgAspProGlyLeuAlaGluGluGlnProValThrTyrValArgAspProProGlyLeuAlaGluLeuLysValSerGlyLysMetGlnLysArgGlyLeuAlaGluLeuLysValSerGlyLysMetGlnLysArgGlyLeuAlaGluLeuLysValSerGlyProProGlyLysArgArgGlyLeuAlaGluLeuLysValLysGlyProProGlyClysGlyLysArgGlyLeuAlaAlaLusValLusKalGlyProProGlyClysGlyLysLysSerGlnTrgArgArgArgArgArgArg <td></td> <td>Asn</td> <td>Lys</td> <td></td> <td></td> <td>Trp</td> <td>Tyr</td> <td>Phe</td> <td></td> <td></td> <td>Arg</td> <td>Суз</td> <td>His</td> <td></td> <td></td>		Asn	Lys			Trp	Tyr	Phe			Arg	Суз	His		
Gly HisI1eLeuI1eGluCysGlyAlaProArgAsnGlnGlyLysArgSerAspLeuSerThrAlaValSerThrLeuGluThrGlySerLeuYalThrValSerThrAlaValSerThrLeuLeuGluThrGlySerLeuYalThrValAlaGluGluGlnProValThrTyrValArgAsnProArgYalLeuAlaGluLeuLeuLysValSerGlyLysMetGlnLysArgArgYalLysThrAlaValHisValSerGlyProProGlyCysGlyLysYalLysThrAlaArgAsnPheAlaGluProProProGlyCysGlyLysYalYalAsnAsnAsnPheAsnPheAlaGluProProProGlyCysGlyLysYalYalAsnAsnLysTrpAsnGlyProAsnAsnCysGlyLysYalYalYalYalYalGlyFroYalAsnAsnYalYalYalYalYalYalYalYalYalYalYalYal<	Lys Ala	Lys			Asp	Gln	Gln			Glu	Tyr	Суз			Glu
SerAspLeuSerThrAlaValSerThrLeuLeuGluThrGlySerLeuValThrValAlaGluGluGlnPheProValThrTyrValArgAspPheArg145ThrValAlaGluGluGlnFhrItoValThrTyrValArgAspPheArgGlyLeuAlaGluLeuLeuLysValSerGlyLysMetGlnLysArgAspTrpLysThrAlaValHisValIleValGlyProProGlyCysGlyLysSerGlnTrpAlaArgAsnPheAlaGluProArgAspThrLysProSerArgAsnLysTrpAspGlyTyrHisGlyGlyGlyTyrLysProSerArgAsnLysTrpTrpAspGlyTyrHisGlyGlyGlyCysTyrLysProSerArgAsnLysTrpAspGlyTyrHisGlyGlyGlyValValProSerArgAspLysTrpAspGlyTyrHisGlyGlyGlyValValProSerArgAsp	-			Ile	Glu	Суз			Pro	Arg	Asn			Гла	Arg
ValThrValAlaGluSinPhoProValThrTyrValArgArgArgGlyLeuAlaGluLeuLysValSerGlyLysMetGlnLysArgAspTrpLysThrAlaValLeuLysValSerGlyLysMetGlnLysArgAspTrpLysThrAlaValHisValIleValGlyProProGlyCysGlyLysSerGlnTrpAlaArgAsnPhoAlaAlaAspPhoAngArgAspTrpLysProSerArgAsnLysTrpAspGlyTyrHisGlyGlyGlyGluValValProSerArgAspLysTrpTrpAspGlyTyrHisGlyGluValValValLeuAspAspPhoTyrGlyTyrLeuProTypAspAspLeuLeuValLeuAspAspPhoTypGlyTypLeuProTypAspLeuLeuArg	Ser Asp		Ser	Thr	Ala			Thr	Leu	Leu			Gly	Ser	Leu
145150155160Gly Leu Ala Glu Leu Leu Lys Val Ser Gly Lys Met Gln Lys Arg Asp 165165160Trp Lys Thr Ala Val His Val Ile Val Gly Pro Pro Gly Cys Gly Lys 185185Pro Pro Gly Cys Gly Lys 190Ser Gln Trp Ala Arg Asn Phe Ala Glu Pro Arg Asp Thr Tyr Trp Lys 200195Trr Lys Gly Cys Cly Lys 205Pro Ser Arg Asn Lys Trp Trp Asp Gly Tyr His Gly Glu Glu Val Val 210165160Val Leu Asp Asp Phe Tyr Gly Trp Leu Pro Trp Asp Asp Leu Leu Arg180		Val	Ala	Glu	Gln		Pro	Val	Thr	Tyr		Arg	Asn	Phe	Arg
165170175Trp Lys Thr Ala Val His Val Ile Val Gly Pro Pro Gly Cys Gly Lys 18018590Ser Gln Trp Ala Arg Asn Phe Ala Glu Pro Arg Asp Thr Tyr Trp Lys 2009090Pro Ser Arg Asn Lys Trp Trp Asp Gly Tyr His Gly Glu Glu Val Val 21020090Val Leu Asp Asp Phe Tyr Gly Trp Leu Pro Trp Asp Asp Leu Leu Arg	145				150					155		-			160
180185190Ser Gln Trp Ala Arg Asn Phe Ala Glu Pro Arg Asp Thr Tyr Trp Lys 195200205Pro Ser Arg Asn Lys Trp Trp Asp Gly Tyr His Gly Glu Glu Val Val 210200200Val Leu Asp Asp Phe Tyr Gly Trp Leu Pro Trp Asp Asp Leu Leu Arg	-			165		-			170	-			-	175	-
195200205Pro Ser Arg Asn Lys Trp Trp Asp Gly Tyr His Gly Glu Glu Val Val 210215220Val Leu Asp Asp Phe Tyr Gly Trp Leu Pro Trp Asp Asp Leu Leu Arg	Trp Lys	Thr		Val	His	Val	Ile		Gly	Pro	Pro	Gly	-	Gly	Lys
210 215 220 Val Leu Asp Asp Phe Tyr Gly Trp Leu Pro Trp Asp Asp Leu Leu Arg			Ala	Arg	Asn	Phe		Glu	Pro	Arg	Asp		Tyr	Trp	Lys
		Arg	Asn	Lys	Trp		Asp	Gly	Tyr	His		Glu	Glu	Val	Val
		Asp	Asp	Phe			Trp	Leu	Pro		Asp	Asp	Leu	Leu	
Leu Cys Asp Arg Tyr Pro Leu Thr Val Glu Thr Lys Gly Gly Thr Val 245 250 255	Leu Cys	Aap	Arg	-	Pro	Leu	Thr	Val		Thr	ГЛа	Gly	Gly		Val

		-
-con	nu	ed

Pro	Phe	Leu	Ala 260	Arg	Ser	Ile	Leu	Ile 265	Thr	Ser	Asn	Gln	Ala 270	Pro	Gln
Glu	Trp	Tyr 275	Ser	Ser	Thr	Ala	Val 280	Pro	Ala	Val	Glu	Ala 285	Leu	Tyr	Arg
Arg	Ile 290	Thr	Thr	Leu	Gln	Phe 295	Trp	Lys	Thr	Ala	Gly 300	Glu	Gln	Ser	Thr
Glu 305	Val	Pro	Glu	Gly	Arg 310	Phe	Glu	Ala	Val	Asp 315	Pro	Pro	Сүв	Ala	Leu 320
Phe	Pro	Tyr	Lys	Ile 325	Asn	Tyr	Val	Phe	Phe 330	Val	Ile	Thr	Ser	Trp 335	Phe
Leu	Phe	Leu	Phe 340	Ile	Arg	Val	Phe	Gln 345	Asp	ГÀа	Phe	Ser	Glu 350	Leu	Tyr
Ile	Asn	Ser 355	Gln	Pro	Tyr	His	Ile 360	Ile	Leu	Gly	Сув	Gly 365	Сув	Ile	Leu
Glu	Arg 370	Ile	Ala	Gln	Ala	Cys 375	Val	Leu	Asp	Ile	Gly 380	Val	Gly	Ile	Met
Glu 385	Pro	Gln	Leu	Val	Ser 390	Phe	Ile	Ile	Trp	Leu 395	Glu	Pro	Ile	Asn	Cys 400
Leu	Val	Leu	Trp	Phe 405	Gly	Gly	Glu	Val	Pro 410	Gly	Val	Val	Gly	Lys 415	Gly
Leu	Pro	Tyr	Gly 420	Val	Ala	Gly	Gly	Val 425	Val	Asn	Ile	Gly	Val 430	Ile	Gly
Gln	Val	Gly 435	Gly	Gly	Gly	Tyr	Lys 440	Val	Gly	Ile	Gln	Asp 445	Asn	Asn	Ser
Gly	Pro 450	Asn	Thr	Ser	Leu	Ile 455	Arg	Gly	Yab	Gly	Val 460	Ser	Gly	Val	Lys
Phe 465	Ile	Phe	Ser	Leu	Ser 470	Asn	Thr	Val	Val	Leu 475	Glu	Arg	Gly	Val	Gly 480
Ala	Ala	Gly	Gly	Glu 485	Glu	Leu	Ala	Aab	Val 490	Glu	Ser	Gln	Leu	Val 495	Asn
Ile	Pro	Arg	Trp 500	Leu	Arg	Val	Ser	Ser 505	Ser	Tyr	Gly	Glu	Tyr 510	ГЛа	Phe
Ser	Arg	Lys 515	Ala	Gly	Ile	Glu	Asp 520	Thr	Arg	Leu	Ser	Ala 525	Pro	Ser	Val
Thr	Val 530	Ser	Glu	Gly	Gly	Val 535	Tyr	Gln	Ile	Trp	Ser 540	Ser	Pro	Glu	Asp
Val 545	Ser	Lys	Met	Ala	Ala 550	Gly	Ala	Gly	Pro	Ser 555	Ser	Ala	Val	Thr	Pro 560
Pro	Trp	Pro	Arg	His 565	Pro	Ile	Lys	Val	Lys 570	Glu	Val	Arg	Сүз	Cys 575	Ser
Ile															
<213 <212	0> SI L> LI 2> T 3> OH	ENGTH ZPE :	1: 5! PRT	53	ə A I	PWD (circo	oviru	18						
<400	D> SI	EQUEI	ICE :	4											
Gln 1	Arg	Thr	Ser	Ala 5	Ala	Ala	Ala	Pro	Arg 10	Gln	Arg	Gln	Lys	Cys 15	Gln
Ala	Arg	Lys	Ala 20	Ala	Arg	Asn	Pro	Ile 25	Arg	Gly	Gly	Сүз	Ser 30	Pro	Leu

33

-	COI	nt.	lr	ıu	ed

Leu	Pro	Arg 35	Arg	Arg	Гла	Thr	Lys 40	Tyr	Gly	Ser	Phe	Gln 45	Ser	Pro	Phe
Leu	Ile 50	Ile	Leu	Phe	Val	Ala 55	Arg	Lys	Val	Trp	Lys 60	Arg	Val	Glu	Leu
Leu 65	Thr	Ser	Arg	Gly	Leu 70	Arg	Ile	Leu	Leu	Arg 75	Ser	Arg	Leu	Leu	Thr 80
Arg	Ser	Gly	Ile	Leu 85	Val	Pro	Ala	Ala	Thr 90	Ser	Arg	Lys	Arg	Lys 95	Glu
Pro	Thr	Ser	Arg 100	Ile	Гла	Asn	Thr	Ala 105	Val	ГЛа	ГЛа	Ala	Thr 110	Tyr	Leu
Ser	Ser	Val 115	Glu	Leu	Arg	Gly	Thr 120	Arg	Gly	Ser	Ala	Ala 125	Thr	Сув	Leu
Leu	Leu 130	Val	Pro	Phe	Trp	Arg 135	Arg	Gly	Leu	Trp	Leu 140	Pro	Ser	Ser	Phe
Leu 145	Arg	Met	Glu	Ile	Ser 150	Ala	Gly	Trp	Leu	Asn 155	Phe	Lys	Ala	Gly	Arg 160
Сүз	Arg	Ser	Val	Ile 165	Gly	Arg	Gln	Leu	Tyr 170	Thr	Ser	Trp	Ala	Arg 175	Pro
Val	Val	Gly	Arg 180	Ala	Ser	Gly	Pro	Val 185	Ile	Leu	Leu	Ser	Leu 190	Gly	Thr
Pro	Thr	Gly 195	Ser	Leu	Val	Glu	Ile 200	Ser	Gly	Gly	Met	Asp 205	Ile	Met	Glu
ГÀа	Lys 210	Leu	Leu	Phe	Trp	Met 215	Ile	Phe	Met	Ala	Gly 220	Tyr	Leu	Gly	Met
Ile 225	Tyr	Asp	Сүз	Val	Thr 230	Gly	Ile	His	Leu	Arg 235	Leu	ГÀа	Gly	Val	Leu 240
Phe	Leu	Phe	Trp	Pro 245	Ala	Val	Phe	Leu	Pro 250	Ala	Ile	Arg	Pro	Pro 255	Arg
Asn	Gly	Thr	Pro 260	Gln	Leu	Leu	Ser	Gln 265	Leu	ГÀа	Leu	Ser	Ile 270	Gly	Gly
Leu	Leu	Leu 275	Cys	Asn	Phe	Gly	Arg 280	Leu	Leu	Glu	Asn	Asn 285	Pro	Arg	Arg
Tyr	Pro 290	Lys	Ala	Asp	Leu	Lys 295	Gln	Trp	Thr	His	Pro 300	Val	Pro	Phe	Ser
His 305	Ile	Lys	Ile	Thr	Glu 310	Ser	Phe	Leu	Leu	Ser 315	His	Arg	Asn	Gly	Phe 320
Tyr	Phe	Tyr	Ser	Phe 325	Arg	Gly	Ser	Phe	Arg 330	Ile	Asn	Ser	Leu	Asn 335	Сув
Thr	Ile	Val	Asn 340	Leu	Thr	Thr	Phe	Trp 345	Ala	Val	Val	Ala	Phe 350	Trp	Ser
Ala	Pro	Arg 355	Pro	Val	Сүз	Ser	Thr 360	Leu	Val	Trp	Val	Phe 365	Lys	Trp	Ser
His	Ser 370	Trp	Phe	Leu	Leu	Leu 375	Phe	Gly	Trp	Asn	Gln 380	Ser	Ile	Val	Trp
Ser 385	Ser	Ser	Gly	Leu	Gly 390	Val	Гла	Tyr	Leu	Glu 395	Trp	Val	Гла	Gly	Cys 400
Leu	Met	Val	Trp	Arg 405	Glu	Glu	Leu	Ile	Gly 410	Ser	Ala	Lys	Leu	Val 415	Glu
Gly	Val	Thr	Lys 420	Leu	Ala	Ser	Lys	Ile 425	Thr	Thr	Val	Asp	Pro 430	Thr	Pro

-continued	
Leu Leu Glu Val Met Gly Ser Leu Gly Asn Ser Tyr Leu Ala Phe Leu 435 440 445	
Ile Arg Tyr Trp Lys Gly Arg Gly Arg Gly Leu Val Pro Pro Glu Gly450455460	
Gly Arg Asn Trp Pro Met Leu Asn Leu Ser Ser Leu Thr Phe Gln Asp465470475480	
Gly Cys Glu Cys Pro Pro Leu Met Val Ser Thr Asn Ser Leu Glu Arg 485 490 495	
Arg Glu Leu Lys Ile Pro Val Phe Arg Arg His Leu Arg Phe Leu Lys 500 505 510	
Ala Gly Cys Thr Lys Tyr Gly Leu Leu Arg Arg Met Phe Pro Arg Trp 515 520 525	
Leu Arg Gly Arg Val Arg Leu Leu Arg Arg Leu Leu Gly His Val Ile	
530 535 540 Leu Lys Lys Cys Ala Ala Val	
545 550	
<210> SEQ ID NO 5 <211> LENGTH: 1759 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus <400> SEOUENCE: 5	
aatactacaq caqcqcactt ctttcacttt tataqqatqa cqtqqccaaq qaqqcqttac	60
cgcagaagac ggacccgccc ccgcagccat cttggaaacg tcctccggag aagaccatat	120
ttggtacace cegeetteag aaacegttae agatggegee gaaagaeggg tatetteaat	180
teeegeettt etagagaatt tgtaeteace ataagaggag gacaetegea geeatettgg	240
aatgttaacg agctgagatt caacatcggc cagtteetee eeecteagg eggeaceaac	300
cccctacccc tacctttcca atactaccgt attagaaagg ctaaatatga attttacccc	360
agagacccca tcacctctaa tcaaagaggt gttgggtcca ctgttgttat cttggatgcc	420
aactttgtaa ccccctccac caacttggcc tatgacccct atattaacta ctcctcccgc	480
cacaccataa ggcagccott tacctaccac tocaggtact toaccoocaa accagagota	540
gaccaaacaa ttgattggtt ccagccaaat aataaaagaa accagctgtg gctccattta	600
aatacccaca ccaatgtcga gcacacaggc ctgggctatg cgctccaaaa tgcaaccaca	660
gcccaaaatt atgtggtaag gttgactatt tatgtacaat tcagagaatt tatcctgaaa	720
gaccetetaa atgaataaaa ataaaaacea ttaegatgtg ataacaaaaa agacteagta	780 840
atttatttta tatgggaaaa gggcacaggg tgggtccact gcttcaaatc ggccttcggg tacctccgtg gattgttctc cagcagtctt ccaaaattgc aaagtagtaa tcctccgata	900
gagagettet acagetggga cageagttga ggagtaceat teetgggggg eetgattget	960
ggtaatcaaa atactgoggg ccaaaaaagg aacagtacce cetttagtet etacagtcaa	1020
tggataccgg tcacacagtc tcagtagatc atcccaaggt aaccagccat aaaaatcatc	1080
caaaacaaca acttcttctc catgatatcc atcccaccac ttatttctac taggcttcca	1140
gtaggtgtcc ctaggctcag caaaattacg ggcccactgg ctcttcccac aaccgggcgg	1200
gcccactatg acgtgtacag ctgtcttcca atcacgctgc tgcatcttcc cgctcacttt	1260
caaaagttca gccagcccgc ggaaatttct cacatacgtt acaggaaact gctcggctac	1320

												con	tin	ued		
agto	acca	aaa q	gacco	ccgt	ct c	caaaa	agggt	c act	caca	agca	gta	gaca	ggt (cgct	geget	tt 1380
cccc	tggt	tc d	cgcg	gaget	tc ca	acact	tcgat	c aaq	gtato	gtgg	ccti	tctti	cac 1	tgca	gtati	tc 1440
ttta	ttt	cgc t	tggt	cggti	tc ci	tttcç	gctt	c cto	gato	gtgg	cag	cggg	cac (caaa	ataco	ca 1500
ctto	acct	tg t	taaa	aagt	ct g	cttct	ttago	c aaa	aatto	cgca	aac	ccct	gga g	ggtga	agga	gt 1560
tcta	iccct	cct t	tccaa	aacci	tt c	ctcgo	ccaca	a aad	caaaa	ataa	tca	aaaa	add 4	agati	tggaa	ag 1620
ctcc	cgta	att t	tgti	tttt	ct c	ctcct	togga	a ago	gatta	atta	agg	gtga	aca (ccca	cctct	tt 1680
atgg	ggtt	gc g	gggco	cgcti	tt t	cttgo	cttg	g cat	ttt	cact	gac	gctg	ccg a	aggt	gctgo	cc 1740
gctg	lccda	aag t	tgcg	ctggi	t											1759
<211 <212 <213	.> LH :> T) :> OF	EQ II ENGTH YPE : RGANI EQUEI	H: 50 PRT ISM:	67 Тур	e A 1	PWD (circo	oviru	15							
					I.011	Pro	Ι.011	Val	Glu	۸la	۵la	Aen	Thr	Dha	TIO	
1 1	лта	CYS	цүз	5 5	ысц	Pro	ьeu	var	10	лıd	лта	чар	1111	рпе 15	116	
Gly	Leu	Leu	Phe 20	Leu	Pro	Gly	Сүз	Gly 25	Trp	Leu	Leu	His	Thr 30	Asn	Val	
Arg	Leu	Leu 35	Gly	Glu	Ser	Ser	Ser 40	Phe	Phe	Leu	Ile	Arg 45	Ser	Ser	Gly	
Ile	Glu 50	Arg	Lys	Ser	Lys	Thr 55	Gln	Pro	Ser	Ser	Pro 60	Lys	Ser	Ser	Pro	
Leu 65	Val	Gly	Arg	Trp	Pro 70	Asn	Ala	Phe	Lys	Ala 75	Leu	Phe	Сүз	Val	Lys 80	
Leu	Leu	Thr	Phe	His 85	Tyr	Гла	Pro	Ala	Arg 90	Gln	Trp	Met	Ser	Phe 95	Ala	
Phe	Pro	Val	Ser 100	Trp	Суз	Phe	Leu	Ser 105	Tyr	Gln	Leu	Leu	Ser 110	Pro	Trp	
Met	Ser	Ile 115	Ser	His	Pro	Ala	Gly 120	Arg	Phe	Trp	Pro	Phe 125	Arg	Leu	Ser	
Arg	Asp 130	Val	Ala	Thr	Leu	Val 135	Arg	Lys	Ser	Val	Pro 140	Asp	Lys	Thr	Val	
Thr 145	Ala	Ser	Суз	Asn	Gly 150	Thr	Val	Tyr	Thr	Leu 155	Phe	Гла	Arg	Pro	Ser 160	
Ala	Ser	Ser	Lys	Phe 165		Leu	Pro	Phe	Ile 170	Сув	Суа	Arg	Ser	Gln 175	Phe	
Val	Ala	Thr	Cys 180		Met	Thr	Pro	Gly 185	Gly	Pro	Gln	Pro	Phe 190	Leu	Trp	
His	Ala	Arg 195	Leu	Lys	Ala	Ser	Gly 200		Ser	Val	Gln	Phe 205	Gly	Leu	Leu	
Phe	Leu 210	His	His	Ser	Pro	Tyr 215	Pro	Ser	Ser	Thr	Thr 220	Thr	Гла	Ser	Ser	
Lys 225	Pro	Gln	Asn	Gly	Gln 230	Ser	Ser	Arg	Ser	Leu 235	Ser	His	Ser	Arg	Tyr 240	
Gly	Asn	Val	Thr	Ser 245	Val	Leu	Pro	Pro	Val 250	Thr	Gly	Гла	Гла	Ala 255	Arg	
Leu	Ile	Lys	Ile 260	Val	Leu	Leu	Ala	Gly 265	Trp	Ser	His	Tyr	Glu 270	Glu	Val	

-continued

											-	con	LIN	uea		
		275					280					285				
Jln	Phe 290	Val	Ala	Pro	Ser	Сув 295	Asp	Val	Ser	Thr	Gly 300	Ser	Pro	Arg	Asn	
Ser 305	Ala	Thr	Ser	Gly	Gly 310		Ala	Arg	Lys	Gly 315	Tyr	Leu	Ile	Phe	Gln 320	
ſhr	Lys	Lys	Thr	Ile 325	Val	Asp	Tyr	His	Asn 330	ГЛа	Asn	ГЛа	Asn	Met 335	Leu	
ſhr	Lys	Ser	Leu 340	Asn	Glu	Ser	Asn	Tyr 345	Met	Phe	Leu	Gly	Trp 350	Met	Ile	
jàr	Pro	Gln 355	Pro	Gln	Met	ГАЗ	Ser 360	Arg	Met	Ala	Trp	Ala 365	Gln	Thr	Ser	
3er	Met 370	Pro	Thr	Pro	Ile	Ile 375	Ser	Gly	Суз	Ser	Thr 380	Glu	Lys	Ile	Ile	
Gln 385	Ser	Ser	Gly	Ile	Leu 390	Gln	Lys	Thr	Ser	Gln 395	Asn	Pro	Pro	Ser	Thr 400	
-	Pro			405				-	410					415		
	Ile		420					425					430			
	Ser	435					440					445				
	Glu 450					455					460					
465					470					475					480	
	Ser Glu			485					490					495		
	Arg		500	-				505					510			
-	His	515		-			520					525		-	_	
	530 Asp	-		-		535					540					
545	_				550		-		-	555	-				560	
				565												
<21: <21:	0> SH 1> LH 2> TY	ENGTH PE :	H: 5 PRT	80												
	3> OF 0> SE				e A 1	PMD (circo	oviru	ıs							
	J> SH Arg				Ala	Ala	Ala	Glv	Ara	Cvs	Ara	His	Phe	His	Trp	
1	Leu			5					10					15		
	Ile		20		-		-	25					30	-	-	
	Lys	35					40					45				
-1	50	2-				55					60					

-continued

set Set Arg Val Gu Leu He He low Arg 11e Lye Set Leu Leu Leu Set Set Leu Leu Keu Set Set Leu Leu Keu Set Set Leu Leu Keu Set Set Leu Feu Set Set Gu Yu Val Leu Leu IIe Phe 90 Val Ala Thr Phe Phe Ala Val Var Leu Ku Val Leu Leu IIe Phe Phe Val Ala Thr Phe Phe Ala 100 Val Leu Leu IIe Phe Phe Val Ala Thr Phe Phe Ala Val Tyr Lye Aeg Leu Thr Set Set Arg Pro Val Leu Pro Leu Ala Ala 111 Val Gin Arg Set Set His Thr Gil Yug Gin Leu Arg Pro Arg Gin His 135 Set Yr Gily Leu Leu Lyg Arg Tyr Arg IIe His Set IIe Glu Ala Pro 150 143 Tyr Gily Leu Leu Yug Arg Tyr Arg IIe His Set IIe Glu Ala Pro 150 145 Fet Yug Gin Phe His Ala Pro 160 145 Fet Yug Gin Phe His Ala Pro 160 145 Fet Yug Gin Phe His Ala Pro 160 146 Fet Yug Gin Phe His Ala Pro 160 147 Fet Yug Gin Phe His Ala Pro 160 148 Fet Yug Gin Phe His Ala Pro 160 149 Fet Yug Gin Phe His Ala Pro 160 140 Fet Yug Gin Phe His Ala Pro 160 141 Fet Yug Gin Phe His Ala Pro 170 145 Fet Yug Gin Phe His Ala Pro 170 145 Fet Yug Gin Phe His Ala Pro 1120 145 Fet Yug Ala Pro 1120 145 Fet Yug Ala Pro 1120 145 Fet Yug Ala Pro 1120
859095Arg Phe Ser Giy Val Leu Leu IIe Phe Phe Val Ala Thr Phe Phe AlaVal Tyr Lyg Asp Leu Thr Ser Ser Arg Pro Val Leu Pro Leu Ala Ala130Val Gin Arg Ser Ser Hie Thr Gly Lye Cin Leu Arg Pro Arg Cin Hie130Ser Tyr Qly Leu Leu Lye Arg Tyr Arg IIe His145Val Cy Ser Tyr Val Aep Tyr Hie Ala Arg Cin Hie150Cin Ser Phe Lye Cin Phe Hie Ala Pro161Cin Ser Phe Lye Cin Phe Hie Ala Pro162Cin Ser Tyr Val Aep Tyr Hie Ala Arg Cin Hie Leu Leu Thr IIe Pro175Leu Cye Ser Tyr Val Aep Tyr Hie Ala Ser Gin Thr Pro Leu Arg19619719819819919919919019019019119119219319419519519619719819819919919919919019019119119219319419519519519619719819819819919919919919919919919919919919919919919919919919919
100 105 110 Val Tyr Hy Ap Leu Th's Ser Ser Arg Pro Val Leu Pro Leu Pro Leu Pro Leu Pro Leu Pro Leu Pro Leu Arg Glu Ala Val Gin Arg For Ser For Th's Glu Ya Glu Pro Arg Glu Ala Ala Val Gin Arg For For Int Glu Ala Pro Leu Luu Th's For
lif 1 10 1 10 1 10 1 120 1 125 Val Gin Arg Ser Ser His Thr Gly Lys Gin Leu Arg Pro Arg Gin His 130 1 15 1 15 1 16 17 17 18 18 18 19 19 18 18 18 19 19 19 11 19 10 115 Ser Tyr Gly Leu Leu Lys Arg Tyr Arg I le His Ser I le Glu Ala Pro 145 11 19 11 19 10 115 Leu Cys Ser Tyr Val Arg Tyr His Ala Pro Leu His Leu Leu Thr I le Pro 175 19 19 19 19 19 19 19 19 19 19 19 19 19
130135140Ser Tyr GUy Leu Leu Lyw Arg Tyr Arg Ile His Ser Ile Glu Ala Pro 155160Gln Ser Phe Lyw Gln Phe His Ala Pro 160Leu His Leu Leu Thr Ile Pro 170Leu Cyw Ser Tyr Val Asp Tyr His Ala Arg Gly Thr Thr Pro Leu Ala 190Leu Pro Gly Thr Ile Lyw Ser Leu Arg Pro Val Gly Val Pro Leu Arg 200Thr Ser Ile Leu Pro Pro Ile Ser Ile Met Ser Phe Phe An Arn Arn 215216Gln Ala Thr Ang Gly Ala Ile Leu Gly Gly Val Pro Leu Arg 200Thr Ser Ile Leu Pro Pro Ile Ser Ile Met Ser Phe Phe An Arn Arn 215217218Pro Ile Trp Gln Ser Tyr Leu Ser Phe Pro Thr Ser Asp Arg Lyw Gln 260219210Pro Ile Trp Gln Ser Tyr Leu Ser Phe Ser Glu Ile Pro Pro Ana Ser Ser 275210211212212213214215216217218219219210210211212212213214215216217218218219219210211212213213214215216217218218219219219219219219219210211211212 <t< td=""></t<>
145150155160Gin Ser Phe Lyg Gin Phe His Ala Pro 165Leu His Leu Leu Thi The Pro 175110Leu Yg Ser Tyr Val Asp Tyr His Ala Arg Gly Thr Thr Pro 185Leu Arg 200Leu Yg Gir Thr Hie Lyg Ser Leu Arg Pro 211Val Gly Val Pro 200Leu Arg 2205Leu Pro 211Pro 211Pro 211Val Gly Val Pro 211Leu Pro 221Pro 211He Ker Phe 220Pro 2205Thr Sar The Leu Pro 211Pro 211Pro 220Pro 2205Pro 212He Ker Phe 230Pro 250He Ker Phe 230Pro 213He Ker Phe 245Pro 250He Ker Gin Thr Val 230Pro 214He Ker Phe 245Pro 250He Ker Gin Thr Val 230Pro 215He Ker Gin Val Gly Gly Leu Phe 260Pro Val 270Cly Ala Thr Asn Gin Asn Gly Ala The Leu Gly Gly Leu Phe 260Pro Val 270Cly Ser Ser Asp Trp Ser Tyr Phe Ser Glu He Pro Pro Asn Ser Ser 2300Clu Leu Lys Pro Leu Ser Ser Pre 230Leu Gly Ang Nag Lyg Glu Trp He Pro 300Ala Ser Lyg Phe Cyg His Va Ann Asp Cyg Arg Leu Pro Lyg Lyg Glu 320Tyr He Val Ser Ang Lyg Lyu Ann Asp Cyg Arg Pro 346Pro Asp Lyg Lyu Lieu He Phe Gin Val Tyr His Fin 346Pro Asp Lyg Lyu Lieu He Phe Glu His Leu Trp 350Leu Pro Asp Lyg Lyu Ann Ann Col Hen Trp Asp He His Leu Trp 366Sar Lyg Val Gly Thr His Glu Val Ann Thr Ala Ann Gln Leu Ala Tyr 366Sar Lyg Val Gly Thr His Glu Val Ann Thr Kan Leu His Leu Trp 365Leu Glu Pro Lyg Pro Thr Hen Tyr
163 170 175 Leu Va Ser Tyr Va Asp Tyr Ha Als Ya Ua Va Asp Tyr Ha Als Ya Ua Asp Tyr Ha Als Ya Ua Asp Tyr Ha Als Ya Ua Guy Tyr Ha Asp Tyr Leu Asp Tyr Leu Asp Tyr Leu Asp Tyr Clu Asp Tyr Ser
180185190Leu Pro Gly Thr 11eLys Ser Leu Arg Pro Val Gly Val Pro Leu Arg 205Thr ser 11eLeu Pro Pro 11e Ser 11eNet Ser Phe Phe Asn Asn Asn 210Gln 11e11eLys 11eAla Pro Arg Pro 11e11e22511eIte I and Pro Arg Pro 11e11eGln Ser Gln Thr Val 250Pro 11eTrp Gln Ser TyrLeu Ser Phe Pro Thr Ser Asn Arg Lyg Gln 26022611eThr Asn Gln Asn Gly Ala11eLeu Gly Gly Leu Phe Pro Val 260Gly Ser Ser Asp Trp Ser Tyr Phe Ser Glu 11ePro Pro Asn Ser Ser 280Gln Leu Lys Pro Leu Ser Ser Ser Phe Leu Gly Arg Leu Tyr Gly Phe 300Ala Ser Lys Phe Cys Hia Val Trp Gly Thr Gly Kag Leu Pro Lys Lys Glu 325Tyr 11e Val Ser Asp Lys Lys Lys Asn Asp Cys Arg Leu Pro Lys Lys Glu 326Asn Leu Pro Ash Ser Ser 320Leu Arg Val Val Tyr Asn Gln Ala Thr Thr Ala Asn Gln Leu Ala Tyr 336129Thr His Glu Val 335129Tyr Asn Gln Asn Pro Gln Phe Tyr Asp Lie Thr Ash 340120Tyr Asn Gln Asn Pro Gln Phe Tyr Asp Lie Thr Ash 340120121121121122122122123123123124124125125125125126129127120128129129129129129129129129129129129129129129129129129<
195200205Thr SerIleLeuProProIleSerIleMetSerPheAsnAsnAsn210IleLaProIleSerIleIleIleIleAsnAsnAsn240225IleIleLaProIleAsnArnAsnAsn240226IleIleAsnAsnAsnAsnAsn240ProIleTryGlnSerTyrLeuSerProThSerGlyAlaThrAsnGlyGlyLeuProProVal240ClySerSerAsnGlyAlaIleProProVal240ClyAsnAsnGlyGlyLeuProProVal240ClySerSerAsnGlyAlaIleProProVal240ClySerSerAsnTyrProProNanSerSerSerGlySerSerAsnTyrProProNanSerSerSerSerAsnSerSerAsnTyrProProProProNanSerSerAsnSerSerAsnAsnSerSerSerSerSerSerAsnLeuProAsnAsnGlnVal
210 215 220 G1 10 10 10 230 10 11 110 230 10 10 10 10 20 10 20 10 10 20 10 20 10 10 20 20 20 10 20 10 20 10 20 10 20 10 20 10 20 10 20 10 20 10 20 10 20 <t< td=""></t<>
225 230 235 240 Pro Ite Tr G1n Ser Ty Lue Ser Asn Arg Lys G1n G1y Ala Tr Ser Aso Y Lue Pro Tr Ser Aso Arg Lys G1n G1y Ala Tr Ser Aso Tr Ser Aso Ser Pro Val G1y Ser Ser Tr Ser Tr Ser Ser Ser Tr Ser
G1y Ala Th As G1n Frage Ser Ser Frage Frage Ser Frage Ser Tr Ser Ser Frage Ser Ser Ser Ser Ser Ser Ser Ser Frage Ser Frage Ser Ser </td
310 260 265 270 Gly Ser Asp Ty Ser Tyr Pho Ser Glu Ile Pro Pro Asn Ser Ser Gln Leu Lys Pro Leu Ser Ser Ser Pho Leu Gly Asp Ser Ser Ala Ser Lys Pro Leu Ser Ser Pho Leu Gly Tyr Gly Pho Ala Ser Lys Pro Leu Ser Asp Cys Arg Cys Arg Leu Tyr Gly Pho Jaco Tyr Jaco Tyr Gly Pho Jaco Tyr Jaco Jaco<
275 280 285 GIn Leu Lys Pro Leu Ser Ser Ser Phe Leu Glu Tyr Glu Tyr Glu Phe Ala Ser Lys Pho Cys His Val Tyr Glu Tyr Glu Tyr He Phe Ser Ser Ass Val Tyr Glu Tyr He Phe Ser Ser Ass Ser Lys Lys Ass Ser Ass Lys Lus Ass Cys Ass Ser Ser Mas Ser Pho Glu Asr Tyr Ser Ser Mas Asr Pho Glu Tyr Ser Ser Ser Mas Ser Cys Arg Pho Glu Ser Ser Ser Ser Fin Glu Ser
GIn Leu Leu Ser S
Ala Ser Lys Phe Cys His Val Trp Gly Trp Gly Lys Gly Trp Jus Jus Jus Jus Tyr I Val Ser Asp Lys Ass Asp Lys Asp Lys Asp Asp Lys Asp Asp Lys Asp Asp Asp Lys Asp Asp Lys Asp Asp Asp Lys Asp
Tyr Ile Val Sar Ass Lys Ass A
Asn Leu Pro Asp Leu Ile Pro Glu Arg Pro Stap Val Stap Leu Asp Val Type Leu Arg Yas Val Type Stap Lue Asp Val Type Stap Lue Type Glu Leu Stap Type The Asp Stap Lue Type Glu Leu Stap The Stap The The The Asp Asp Stap Lue Type Glu Stap The Stap Stap The Stap The Asp Asp Lue Type Leu Glu Asp Asp Asp Stap The Stap The Stap Stap The Stap
LeuArgYalValYuAsnGlnAlaThrAlaAsnGlnAlaTyrGlyLeuGlyThrHisHisValYalYalYalYalYalYalYalYalGlyLeuThiHisYalYalYalYalYalYalYalYalYalYalYalYalYalYalGlyLeuArgYalYalYalYalYalYalYalYalYalYalYalYalYalYalYalYalGlyLeuYalYalYalYalYalYalYalYalYalYalYalYalYalYalYalYalYalGlyYal
G1y Leu G1y Th His G1n Ais G1n Ais Th Ass Leu His Leu Trp Leu G1n Ass Arg Ass Ass Ass Ile His Leu His Leu Trp Leu G1n Ass Arg Ass Ass Ass Ile His Leu Asp Leu G1n Ass Arg Ass Ass Ass Ile Ass Asp Leu G1n Ass Arg Arg Fro Asp
LauGlnAsnArgLysAsnAsnAsnProGlnProNgAsnProNgSerNgNgNgNgAsnAsnAsnAsnSerNg <t< td=""></t<>
LeuGluProLysProProProProProGluArgIleMarsArgArgArgSerMarsIleMarsArgArgArgMarsIleArgA
Arg IleThrHisArg SerSerTyrAsnIleHisProAspTyrAlaLeuAsnThrSerProThrValPheAsnAlaAspLeuIleValThrSerAsnThrSerProThrValPheAsnAlaAspLeuIleValValThrSerGlyValGlyArgGlnAsnSerThrIleProAspArgProTyrPheGlu450ForForForForForForForForForForFor
Asn Thr Ser Pro Thr Val Phe Asn Ala Asp Leu Ile Val Val Thr Ser 435 440 445 Gly Val Gly Arg Gln Asn Ser Thr Ile Pro Asp Arg Pro Tyr Phe Glu 450 455 460
Gly Val Gly Arg Gln Asn Ser Thr Ile Pro Asp Arg Pro Tyr Phe Glu 450 455 460

-continued

												COIL		ueu	
465					470					475					480
Asn	Thr	Gly	Gly	Ser 485	Pro	Pro	Leu	Phe	Gln 490	-	Ile	Asn	Phe	Arg 495	Leu
Glu .	Asn	Val	Asn 500	Trp	Ser	Pro	Gln	Ser 505	His	Gly	Gly	Arg	Ile 510	Thr	Leu
Val	Phe	Glu 515	Arg	Ser	Leu	Arg	Ser 520	Asn	Phe	Ile	Gly	Thr 525	Lys	Arg	Arg
Trp .	Arg 530	Tyr	Arg	Asn	Arg	Phe 535		Pro	His	Val	Leu 540	Tyr	Pro	Arg	Arg
Arg 545		Ile	Asn	Gly	Leu 550	His		Arg	Pro	Arg 555		Arg	Arg	Arg	Arg 560
Tyr .	Arg	Arg	Arg				Met	Arg			His	Phe	Phe		
Ala '	Thr	Thr	Asn	565					570					575	
			580												
<210	> SF	11 O	омо	8											
<211	> LE	ENGTH	ł: 5!												
<212 <213				Тур	e A i	PWD ·	circ	ovir	ıs						
<400							2	·							
Leu . 1	Ala	Ser	Arg	Cys 5	Arg	Суз	Суз	Arg	Pro 10	Leu	Thr	Leu	Ser	Phe 15	Ala
Leu	Cya	Ser	Phe 20	Arg	Gly	Ala	Val	Gly 25	Tyr	Ser	Thr	Pro	Thr 30	Gly	Tyr
Asp	Lys	Arg 35	Pro	Pro	Ser	Phe	Cys 40	Phe	Val	Pro	Ala	Glu 45	Leu	Arg	Gly
ГЛа	Gln 50	Asn	Asn	Gln	Lys	His 55	Arg	Pro	Leu	Asn	Pro 60	Leu	Pro	Tyr	Phe
Glu 65	Glu	Gly	Gly	Pro	Thr 70	Gln	Ser	Asn	Gln	Ser 75	Ala	Ser	Lys	Суз	Pro 80
Ser	Thr	Thr	Asn	Gly 85	His	Gly	Ser	Gly	Суз 90	Arg	Ser	Leu	Ser	Leu 95	Phe
Arg	Gly	Ala	Ser 100		Leu	Ile	Ser	Cys 105		Leu	Leu	Gly	Cys 110		Arg
Thr 3	His	Leu 115		Ala	Ser	Gly	Pro 120		Ala	Сув	Arg	Gly 125		Gln	Gln
Ser			Lys	Pro	Ser		Thr	Lys	Pro	Ser			Arg	Ala	Thr
Glu	130 Gln	Leu	Thr	His				Gly	Arg		140 Pro	Gln	Val	Lys	
145		_			150				_	155		_			160
Leu	Ser	Arg	Ser	Ser 165	Ala	Ala	Ala	His	Asn 170	Ser	Ser	Leu	Gln	Val 175	Arg
Leu	Pro	Gly	Ala 180	Arg	Asn	His	Ser	Ser 185	Gly	Thr	Pro	Gly	Tyr 190	Asn	Gln
Gln .	Ala	Pro 195	Суз	Arg	Ser	Ser	Ala 200	-	Phe	Tyr	Thr	Thr 205	Pro	His	Ile
Asp :	His 210	Leu	Leu	Leu	Gln	Gln 215	-	Pro	His	Asn	Lys 220	His	Ser	Thr	Val
Lys 225	Pro	His	Asp	Val	Ser 230		Thr	His	Gly	Thr 235	Asp	Met	Ser	Gln	Leu 240

											-	con	tın	uea			
Ser	Leu	Pro	Tyr	Gln 245	Glu	Lys	Гла	Pro	Gly 250	Суз	Tyr	Lys	Ser	Trp 255	Суз		
Asp	Pro	Gly	Gly 260	Pro	Ile	Thr	Ser	Arg 265	Leu	Gln	Gln	Gly	Leu 270	Gln	Leu		
Leu	Glu	Arg 275	Asp	Ser	Ser	Lys	Ala 280	Ile	Lys	Ser	Ser	Gln 285	Gln	Leu	Val		
Ile	Trp 290	Pro	Pro	Val	Arg	Leu 295	Gly	Ile	Gln	Leu	Leu 300	Pro	Gly	Val	Arg		
His 305	Gly	Lys	Gly	Met	Tyr 310	Phe	Leu	Asn	Ser	Leu 315	Arg	Lys	Gln	Met	Thr 320		
Ile	Thr	Lys	Ile	Lys 325	Ile	Lys	Ser	Pro	Arg 330	Glu	Pro	Tyr	Ile	Arg 335	Gln		
Ile	Thr	Суз	Leu 340		Asp	Val	Lys	Gly 345		Leu	Lys	Pro	Ser 350		Asn		
Cys	Lys	Pro 355		Суз	Leu	Gly	Pro 360		His	Ala	Arg	Сув 365		His	Pro		
Tyr	Lys 370		Pro	Ala	Val	Val 375	Pro	Lys	Lys	Lys	Ala 380		Val	Leu	Asn		
Asn 385		Arg	Ala	Arg	Thr 390		Pro	His	Leu	Val 395		Leu	Pro	Leu	Tyr 400		
	Ala	Ala	Lys			Pro	Pro	Leu			Tyr	Leu	Pro				
Leu	Gln	His		405 Pro	Asn	Суз	Leu		410 Cys	Gly	Leu	Tyr		415 Cys	His		
Val	Trp	-	420 Arg	Гла	Ser	Leu	His	425 His	Pro	Arg	Gln		430 Leu	Ile	Ile		
Gly	Lys	435 Tyr	Pro	Leu	Ile	Pro	440 Phe	Thr	Pro	Thr	Pro	445 Pro	Gln	His	Arg		
Arg	450 Leu	Pro	Pro	Pro	Val	455 Pro	Arg	His	Gln	Ile	460 Glu	Ala	Arg	Cys	Glu		
465 Leu	Ile	Ala	Ala	Leu	470 Thr	Arq	Arg	Lvs	His	475 His	Thr	Cvs	Ile	Arq	480 Phe		
				485		-	Asp	-	490			-		495			
			500				Ile	505					510				
		515			-		520		-			525		-	-		
	530					535	Thr				540		GIU	ГЛЗ	AIa		
Val 545		Asp	Leu	Leu	Ser 550		Leu	Ala	Ser	Ser 555	Tyr	Tyr					
<211 <212 <213 <220 <221 <222	L> LH 2> TY 3> OF 0> FH L> NZ 2> LC	EQ II ENGTH YPE: RGANI EATUH AME/I DCATI	H: 9: DNA ISM: RE: KEY: ION:	39 Type CDS (1)			circo	oviru	15								
							ccg Pro									48	
	ctt	aat	aat	cct	tcc	gag	gag	gag		aac	aaa	ata	cgg		ctt	96	

-continued

Thr	Leu	Asn	Asn 20	Pro	Ser	Glu	Glu	Glu 25	Lys	Asn	Lys	Ile	Arg 30	Glu	Leu		
								gtt Val								144	
								д1У ддд								192	
								tat Tyr								240	
								cag Gln								288	
								gga Gly 105								336	
		-					~ ~	agt Ser				~ ~		000		384	
								cct Pro								432	
								gtg Val								480	
								ata Ile								528	
-	-	-		-	-			gct Ala 185				-				576	
								gat Asp								624	
								tgg Trp								672	
	~	~	~					act Thr	-	~ ~			~~~	~~		720	
Val	Pro	Phe	Leu	Ala 245	Arg	Ser	Ile	ttg Leu	Ile 250	Thr	Ser	Asn	Gln	Ala 255	Pro	768	
Gln	Glu	Trp	Tyr 260	Ser	Ser	Thr	Ala	gtc Val 265	Pro	Ala	Val	Glu	Ala 270	Leu	Tyr	816	
Arg	Arg	Ile 275	Thr	Thr	Leu	Gln	Phe 280	tgg Trp	ГЛа	Thr	Ala	Gly 285	Ğlu	Gln	Ser	864	
Thr	Glu 290	Val	Pro	Ğlu	Gly	Arg 295	Phe	gaa Glu								912	
						aat Asn		tga								939	

<210> SEQ ID NO 10 <211> LENGTH: 312 <212> TYPE: PRT														
<213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 10														
Met P: 1	ro Ser	Lys	Lys 5	Ser	Gly	Pro	Gln	Pro 10	His	Гла	Arg	Trp	Val 15	Phe
Thr Le	eu Asn	Asn 20	Pro	Ser	Glu	Glu	Glu 25	Lys	Asn	Гла	Ile	Arg 30	Glu	Leu
Pro I	le Ser 35	Leu	Phe	Asp	Tyr	Phe 40	Val	Суз	Gly	Glu	Glu 45	Gly	Leu	Glu
Glu G 50	ly Arg 0	Thr	Pro	His	Leu 55	Gln	Gly	Phe	Ala	Asn 60	Phe	Ala	Lys	Гла
Gln Tì 65	hr Phe	Asn	Lys	Val 70	Lys	Trp	Tyr	Phe	Gly 75	Ala	Arg	Сүз	His	Ile 80
Glu Ly	ys Ala	Lys	Gly 85	Thr	Asp	Gln	Gln	Asn 90	ГÀа	Glu	Tyr	Суз	Ser 95	Lys
Glu G	ly His	Ile 100	Leu	Ile	Glu	Суз	Gly 105	Ala	Pro	Arg	Asn	Gln 110	Gly	Lys
Arg Se	er Asp 115	Leu	Ser	Thr	Ala	Val 120	Ser	Thr	Leu	Leu	Glu 125	Thr	Gly	Ser
	al Thr 30	Val	Ala	Glu	Gln 135	Phe	Pro	Val	Thr	Tyr 140	Val	Arg	Asn	Phe
Arg G 145	ly Leu	Ala	Glu	Leu 150	Leu	ГÀа	Val	Ser	Gly 155	ГÀа	Met	Gln	Gln	Arg 160
Asp T:	rp Lys	Thr	Ala 165	Val	His	Val	Ile	Val 170	Gly	Pro	Pro	Gly	Cys 175	Gly
Lys Se	er Gln	Trp 180	Ala	Arg	Asn	Phe	Ala 185	Glu	Pro	Arg	Asp	Thr 190	Tyr	Trp
Lys P:	ro Ser 195		Asn	ГЛа	Trp	Trp 200	Asp	Gly	Tyr	His	Gly 205	Glu	Glu	Val
	al Leu 10	Asp	Asp	Phe	Tyr 215	Gly	Trp	Leu	Pro	Trp 220	Asp	Asp	Leu	Leu
Arg Le 225	eu Cys	Asp	Arg	Tyr 230	Pro	Leu	Thr	Val	Glu 235	Thr	ГАЗ	Gly	Gly	Thr 240
Val P:	ro Phe	Leu	Ala 245	Arg	Ser	Ile	Leu	Ile 250	Thr	Ser	Asn	Gln	Ala 255	Pro
Gln G	lu Trp	Tyr 260	Ser	Ser	Thr	Ala	Val 265	Pro	Ala	Val	Glu	Ala 270	Leu	Tyr
Arg A	rg Ile 275		Thr	Leu	Gln	Phe 280	Trp	Lys	Thr	Ala	Gly 285	Glu	Gln	Ser
	lu Val 90	Pro	Glu	Gly	Arg 295	Phe	Glu	Ala	Val	Asp 300	Pro	Pro	Сүз	Ala
Leu Pl 305	he Pro	Tyr	Lys	Ile 310	Asn	Tyr								
<211> <212> <213> <220> <221>	<pre><210> SEQ ID NO 11 <211> LENGTH: 702 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1)(699)</pre>													

-continued

age cat ctt gga ac at ctc ctc ogg aga aga cc at tt tg gta cac ccc 2596gec tt caga ac cgt tac aga tgg ogc cg aga gag gg ggt at ct tc at ta pe arg aga tgg tt gg tr cac ac tgg ggg ta ct ac tag ag ggg dg cac ac tgg sec cg ctt tc aga aga tt gta ctc ac a ta aga gg gg ag cac tgg sec cg ct tc aga agg gg ac ca ac ccc ct ac ac tag gg gg cac ac tgg sec cg cc tt ct agg gg gg ac ca ac ccc ct ac ccc tt cac at a ag gg gg cac tag sec ccc ccc ccc ccc tag gg gg ac cac ac ccc ct ac ccc tt ccc at at ac sec ccc ccc ccc ccc ccc ccc ccc ccc ccc												COII	CIII	ucu		
Met Thi Tip Pro Arig arig Arig Tyr Arig Arig Arig Arig Thr Arig Pro Arig 15 age cat ct: Gup and at c ct: Gup age agin cat tat tag gta Cat cc: 96 96 96 96 96 96 96 96 96 96	<400> S	equei	NCE :	11												
Ser His Lev Gly Am 11e Lev Arg Arg Arg Pro Tyr Lev Val His Pro gcc ttc ags acc gt tac aga tgg cgc cgs aag acg ggt atc ttc aat 144 Ala The Arg Ann Arg Tyr Arg Trp Arg Arg Lyo Thr Gly 11e Fhe Ann 144 Ala The Arg Ann Arg Tyr Arg Trp Arg Arg Lyo Thr Gly 11e Fhe Ann 144 Ala The Arg Ann Arg Tyr Arg Trp Arg Arg Lyo Thr Gly 11e Fhe Ann 142 Ala The Arg Ann Arg Tyr Arg Trp Arg Arg Gly Gly Gly Gly Gly Gly Gly Gly Gly Glo Ser 192 Gcd Ct Ct agg aga tt gt acc acc ct acc at a ag ggg gga cca tac 192 Gd Ct Ct gg at ggt tacc acc cta ccc tt cc ca tac ag gg gg cc ca 200 Gln Fro Ser Trp Ann Vil Ann Olu Lev Arg The Ann Fro Lev Pro Phe Gln Tyr 200 Set cc cc ccc ta ggc ggc acc aac cc ct ac cct tt cc ca tac acc 288 Lew Fro Pro Ser Gly Gly Gly Thr Ann Pro Lev Pro Phe Gln Tyr 201 100 105 110 110 105 110 111 101 110 112 102 324 113 114 120 114 105 117 115 110 123 126 127 123 127 128 126 128 129 120 </td <td></td> <td></td> <td></td> <td></td> <td></td> <td></td> <td></td> <td></td> <td>Arg</td> <td></td> <td></td> <td></td> <td></td> <td>Pro</td> <td></td> <td>48</td>									Arg					Pro		48
hla he kig kan Aig Tyr Aig Trp Arg Arg Lye Trr Giy He Phe Am 45 tor oge ett tot aga gaa ttt gta ote ace ata aga gga gga cae teg 192 see Arg Leu Ser Arg Glu Phe Val Leu Thr He Arg Gly Gly Hs ser 240 song oca tot teg aa ggt ag ta cae og otg ag tte aae ata og ag og ag cae teg 240 sin ro ser Trp Am Val Am Olu Leu Arg Phe Am Tie Oly Gly Hs ser 240 song oca tot teg ag og ag ac aae cec ce ta cee ot a cet tet caa tae 288 set to Pro Pro Ser Gly Gly Thr Am Pro Leu Pro Leu Pro Phe Gln Tyr 95 set at aga ag ot aaa tat gaa ttt ate tug ag pro 11e 336 100 101 105 110 ace ot at ag og og tt ggt tug og to cat ggt ce ta at ace 334 110 100 101 105 ace ot at ace on ce ce ca cae ag og tt ggt tug the try tro Arg App Pro 11e 334 110 110 125 ace ot ta at ace aga ggt dt ggt caa ect tig get cet at at ace 334 115 126 127 128 ace tot ace ce ce ce ace ata agg cag to the set ta gat cea ace at agg cag 384 126 127 128 125 ace tot ace ce ce ce ace ace ata agg cag cee tt ace tae cae agg 480 127 126			Gly					Arg					Val			96
See Arg Gueu Ser Arg Gu phe val Leu Thr The Arg GUy GUy His ser 50 60 60 60 60 60 60 60 60 60 6	-	Arg		-		-	Trp	-	-	-	-	Gly				144
Sin Pro Ser Trip Ann Val Ann Ölu Leu Arg PPhe Ann Ile Öly Gln Phe 55 56 57 57 57 57 57 58 58 54 55 55 55 55 55 55 55 55 55	Ser Arg					Phe					Arg					192
Heau Pro Pro Ser Gi y Gi y Thr Asm Pro Leu Pro Leu Pro Phe Gin Tyr 95 20 20 20 20 20 20 20 20 20 20 20 20 20 2	Gln Pro				Val			-	-	Phe				-	Phe	240
Fyr Arg Ile Arg Lyø Ala Lyø Tyr Glu Phe Tyr Pro Arg Arg Pro 11e 110 111 110 110 110 111 110 110 111 110 110 111 110 111 110 111 110 111 110 110 110 110 111 110 <t< td=""><td></td><td></td><td></td><td>Gly</td><td></td><td></td><td></td><td></td><td>Leu</td><td></td><td></td><td></td><td></td><td>Gln</td><td></td><td>288</td></t<>				Gly					Leu					Gln		288
Thr Ser Aem Gln Arg Gly Val Gly Ser Thr Val Val IIe Leu Asp Ala 125 115 120 125 aca tht gta acc coc toc acc acc thg gcc tat gac coc tat att acc 432 aca tht gta acc coc toc acc aca thg gcc tat gac coc tat att acc 432 aca tht gta acc coc toc acc acc ata agg cag coc th acc tac cac toc agg 480 rst prive for the tar the tar			Arg					Glu					Asp			336
Asn Phe Val Thr Pro Ser Thr Asn Leu Ala Tyr Asp Pro Tyr Ile Asn 140 Ha Tyr Asp Pro Tyr Ile Asn 140 480 tac too too cogo cac acc ata agg cag coc ttt acc tac cac too agg 150 480 Tyr Ser Ser Arg His Thr Ile Arg Gln Pro Phe Thr Tyr His Ser Arg 160 528 tac tto acc coc aaa coa gag cta gac caa aca att gat tgg tto cag 175 528 Tyr Phe Thr Pro Lys Pro Glu Leu Asp Gln Thr Ile Asp Trp Phe Gln 165 576 cca aat aat aaa aga aac cag ctg tgg cto cat tta aat acc cac acc cac acc 267 576 pro Asn Asn Lys Arg Asn Gln Leu Trp Leu His Leu Asn Thr His Thr 180 576 aat gto gag cac aca ggc ctg gg ttg at got ct caa att ga tag acc aca aca aca att aga gaa acc aca gag ctg leg cto caa att aga gaa acc aca aca 624 624 Asn Val Glu His Thr Gly Leu Gly Tyr Ala Leu Gln Ann Ala Thr Thr 200 672 gcc caa aat tat gtg gta agg ttg act att tat gta caa tto aga gaa acc aca aca aca att aga gac cot cta aat gaa taa 220 702 ctt atc ctg aaa gac cot cta aat gaa taa 230 702 c210 > SEQ ID NO 12 230 c211 > SEQ ID NO 12 230 c212 > SEQ ID NO 12 213 c213 > ORGANISM: Type A FWD circovirus c400 > SEQUENCE: 12 10 Wet Thr Trp Pro Arg Arg Arg Trr Arg Arg Arg Arg Trr Arg Pro Arg 15 Ser His Leu Gly Asn Ile Leu Arg Arg Arg Pro Tyr Leu Val His Pro<		Asn					Gly					Ile				384
Tyr Ser Ser Arg His Thr Ile Arg Gln Pro Pho Pho Thr Tyr His Ser Arg 150 Arg 160 tac ttc acc ccc aaa cca gag cta gac caa aca att gat tgg ttc cag Tyr Phe Thr Pro Ser Pro Glu Leu Ap Gln Thr Ile App Trp Phe Gln 170 528 cca aat aat aaa aga aac cag ctg tgg ctc cat tta aat acc cac acc Pro Asn Asn Lys Arg Asn Gln Leu Trp Leu His Leu Asn Thr His Thr 180 576 aat gtc gag cac aca ggc ctg gg cta gcg ctc caa aat gca acc aca Asn Val Glu His Thr Gly Leu Gly Tyr Ala Leu Gln Asn Ala Thr Thr 195 624 gcc caa aat att gtg gta agg ttg act att tat gta caa ttc aga gaa Ala Gln Asn Tyr Val Val Arg Leu Thr Ile Tyr Val Gln Phe Arg Glu 210 672 ctt atc ctg aaa gac cct cta aat gaa taa Phe Ile Leu Lys Asp Pro Leu Asn Glu 210 702 ctl att ctrops Pro Leu Asn Glu 210 230 cta ttr Trp Pro Arg Arg Arg Tyr Arg Arg Arg Arg Tr Arg Pro Arg 10 15 cta thr Trp Pro Arg Arg Arg Arg Arg Arg Pro Tyr Leu Val His Pro	Asn Phe	Val				Thr		-	-		Āsp					432
Tyr Phe Thr Pro Lys Pro Glu Leu Àsp Gln Thr Ile Asp Trp Phe Gln 170 175 175 1 cca aat aat aaa aga aac cag ctg tgg ctc cat tta aat acc cac acc Pro Asn Asn Lys Arg Asn Gln Leu Trp Leu His Leu Asn Thr His Thr 180 190 6 aat gtc gag cac aca ggc ctg ggc tat gcg ctc caa aat gca acc aca Asn Val Glu His Thr Gly Leu Gly Tyr Ala Leu Gln Asn Ala Thr Thr 195 200 20 215 20 20 20 20 20 20 20 20 20 20 20 20 20	Tyr Ser				Thr					Phe					Arg	480
Pro Asn Asn Lys Arg Asn Gln Leu Trp Leu His Leu Asn Thr His Thr 180 aat gtc gag cac aca ggc ctg ggc tat gcg ctc caa aat gca acc aca Asn Val Glu His Thr Gly Leu Gly Tyr Ala Leu Gln Asn Ala Thr Thr 195 gcc caa aat tat gtg gta agg ttg act att tat gta caa ttc aga gaa Ala Gln Asn Tyr Val Val Arg Leu Thr Ile Tyr Val Gln Phe Arg Glu 210 210 215 215 220 220 220 220 220 220 220 220 220 22				Lys					Gln					Phe		528
Asm Val Glu His Thr Gly Leu Gly Tyr Ala Leu Gln Asm Ala Thr Thr 195 200 205 205 205 205 200 672 205 200 205 200 205 200 205 200 205 200 205 200 205 200 205 200 205 200 205 200 200			Lys	-		-	-	Trp					Thr			576
Ala Gln Asn Tyr Val Val Arg Leu Thr Ile Tyr Val Gln Phe Arg Glu 210 215 220 ttt atc ctg aaa gac cct cta aat gaa taa 702 Phe Ile Leu Lys Asp Pro Leu Asn Glu 225 230 <210> SEQ ID NO 12 <211> LENGTH: 233 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 12 Met Thr Trp Pro Arg Arg Arg Tyr Arg Arg Arg Arg Thr Arg Pro Arg 1 5 10 15 Ser His Leu Gly Asn Ile Leu Arg Arg Arg Pro Tyr Leu Val His Pro		Glu					Gly					Asn				624
Phe Ile Leu Lys Asp Pro Leu Asn Glu 225 230 <210> SEQ ID NO 12 <211> LENGTH: 233 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 12 Met Thr Trp Pro Arg Arg Arg Tyr Arg Arg Arg Arg Thr Arg Pro Arg 1 5 10 15 Ser His Leu Gly Asn Ile Leu Arg Arg Arg Pro Tyr Leu Val His Pro	Ala Gln	Asn		~ ~	~	Arg	~				Val			-	~	672
<pre><211> LENGTH: 233 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 12 Met Thr Trp Pro Arg Arg Arg Tyr Arg Arg Arg Arg Thr Arg Pro Arg 1 5 10 15 Ser His Leu Gly Asn Ile Leu Arg Arg Arg Pro Tyr Leu Val His Pro</pre>	Phe Ile				Pro				taa							702
Met Thr Trp Pro Arg Arg Arg Tyr Arg Arg Arg Arg Thr Arg Pro Arg 1 5 10 15 Ser His Leu Gly Asn Ile Leu Arg Arg Arg Pro Tyr Leu Val His Pro	<211> L <212> T	ENGTI YPE :	H: 23 PRT	33	e A 1	PWD (circo	oviru	18							
1 5 10 15 Ser His Leu Gly Asn Ile Leu Arg Arg Pro Tyr Leu Val His Pro	<400> S	equei	NCE :	12												
	Met Thr 1	Trp	Pro		Arg	Arg	Tyr	Arg		Arg	Arg	Thr	Arg		Arg	
	Ser His	Leu		Asn	Ile	Leu	Arg		Arg	Pro	Tyr	Leu		His	Pro	

Ala Phe Arg Aem Arg Tyr Arg Try Arg Try Arg Lye Thr GUy He Phe Aem 45 Set Arg Cue Ser Arg Cue Phe Val Leu Thr He Arg Gly CUy His Ser 55 So The Pro Ser Try Aem Yal Aem Clu Leu Arg Phe Aem 11e CUy Clm Phe Glm Pro 60 Jeel Pro For Ser Try Aem Yal Aem Clu Leu Arg Phe Aem 11e CUy Clm Phe Glm Pro 65 Try Arg He Arg Lye Ala Lye Tyr Clu Phe Tyr Pro Arg Aep Pro 11e 100 The For Aem Clu Arg Cly Val Cly Ser Thr Val Val Ite Leu Arp Ala 125 Try Ser Arg Glu Phe Val Leu Arg Tyr App Pro Tyr 11e Aem 100 Try Ser Ser Arg His Thr 11e Arg Cln Pro Phe Thr Tyr His Ser Arg 100 Tyr Marg I Pro Lye Pro Clu Leu Aup Cln Thr The Arg Trp Phe Cln 100 Tyr Arg Glu His Thr 10e Arg Cln Pro Phe Thr Thr His Ser Arg 100 Tyr Phe Thr Pro Lye Pro Clu Leu Aup Cln Thr The Arg Trp Phe Cln 176 Tyr Arg Glu His Thr Cly Leu Cly Tyr Ala Leu Cln Aem Thr His 180 Aam Val Clu His Thr Cly Leu Cly Tyr Ala Leu Cln Aem Ala Thr Thr 180 Aal An Arg Ya Am Clu Leu Arg Clu Pro Viz Clu Pro Yal Clu Pro Arg Glu 190 Phe Tel Leu Lye Arg Pro Leu Aem Clu 220 Aal Ala Rin Tyr Val Val Arg Leu Thr 11e Tyr Val Clu Pro Arg Glu Yal Clu Pro Yal Clu Yal															CIII	uou			
505560Clin Pro Ser Try Any Val Aen Glu Leu Arg Phe Aam 11e Gly Gln Phe 65667778Leu Pro Pro Ser Gly Gly Thr Ann Pro Leu Pro Leu Pro Pro Gln Tyr 95797979797870707071727273747575757676767677787878787979707070717273747475757575767677767776777677767776777677777677777677777877787877787879797979797979797979797979797979797979 <td>А</td> <td>la</td> <td>Phe</td> <td></td> <td>Asn</td> <td>Arg</td> <td>Tyr</td> <td>Arg</td> <td></td> <td>Arg</td> <td>Arg</td> <td>Lys</td> <td>Thr</td> <td></td> <td>Ile</td> <td>Phe</td> <td>Asn</td> <td></td> <td></td>	А	la	Phe		Asn	Arg	Tyr	Arg		Arg	Arg	Lys	Thr		Ile	Phe	Asn		
65 70 75 80 Lew Pro For Ser Gily GLY The Ann Pro Lew Pro Lew Pro Pro Gin Tyr 85 97 Tyr Arg Ile Arg Lye Ala Lye Tyr Glu Phe Tyr Pro Arg Asp Pro Ile 100 110 Thr Ser Ann Glu Arg Gly Val Gly Ser Thr Val Val The Lew Asp Ala 115 110 Ann Phe Val Thr Pro Ser The Ann Lew Ala Tyr Asp Pro Tyr Ile Aan 110 120 Tyr Ser Ser Arg His Thr Ile Arg Gln Pro Phe Thr Tyr His Ser Arg 116 120 Tyr Phe Thr Pro Lye Pro Glu Lew Asp Gln Thr Ile Asp Trr Phe Gln 165 120 Tyr Phe Thr Pro Lye Pro Glu Lew App Gln Thr Ile Asp Trr Phe Gln 165 120 Ann Val Glu His Thr Gly Lew Gly Tyr Ala Lew Gln Asn Ala Thr Thr 185 120 Ana Tyr Val Val Yar Jue Thr Ile Tyr Val Gln Phe Arg Glu 210 215 Pro Han Tyr Val Val Arg Lew Thr Ile Tyr Val Gln Phe Arg Glu 210 210 Pro His Ewi Lye App Pro Lew Ann Glu 220 220 Pro His Ewi Lye App Pro Lew Ang Glu 210 210 Callos ERQUENCE: 13 3 Arg Ata Aca at acg go cca act at the tot act agg ctt cca gta agg gg gt c Net His Ser Lye He Gly Pro Lew Ala Lew Pro His Her Min Gly Val 30 Callos Er Lye Her The Gly Exc Law Ala Lew Pro Val Gly Val 1 48 Callos Er Lye Her The Gly Pro Lew Ala Ewi Pro Val Gly Val 30 48 Callos Thr His Ser Lie Ser Glo Can Acc agg gg cca gr gg gg cat to ca cac	S	er	-	Leu	Ser	Arg	Glu		Val	Leu	Thr	Ile	-	Gly	Gly	His	Ser		
959095Tyr Arg 11e Arg Luy Ala Lys Tyr Glu Phe Tyr Pro Arg Asp Pro 11e110Thr Ser Ann Gln Arg Gly Val Gly Ser Thr Val Val Tle Leu Asp Ala115Ann Phe Val Thr Pro Ser Thr Ann Leu Ala Tyr Ap Pro Tyr Ile Asn130Tyr Ber Ser Arg His Thr 11e Arg Gln Pro Phe Thr Tyr His Ser Arg145Tyr Phe Thr Pro Lys Pro Glu Leu Asp Gln Thr 11e Asp Trp Phe Gln170171Pro Asn Asn Lys Ag Gln Clu Leu Asp Gln Thr 11e Asp Trp Phe Gln185186Tyr Phe Thr Pro Lys Pro Clu Leu Gly Tyr Ala Leu Cln Asn Thr Thr180Ann Val Glu Hie Thr Gly Leu Gly Tyr Ala Leu Cln Asn Ala Thr Thr180210211211211212211212212213214214215210210211211211212212213214214215215216217218218219219219210210211211211212213214215215216217218218219219219210211211212213 </td <td></td> <td></td> <td>Pro</td> <td>Ser</td> <td>Trp</td> <td>Asn</td> <td></td> <td>Asn</td> <td>Glu</td> <td>Leu</td> <td>Arg</td> <td></td> <td>Asn</td> <td>Ile</td> <td>Gly</td> <td>Gln</td> <td></td> <td></td> <td></td>			Pro	Ser	Trp	Asn		Asn	Glu	Leu	Arg		Asn	Ile	Gly	Gln			
100105110110Thr Ser Asn Gin Arg Giy Val Giy Ser Thr Val Val Ile Leu Asp Ala 125Asn Phe Val Thr Pro Ser Thr Asn Leu Ala Tyr Asp Pro Tyr Ile Asn 135JivoTyr Ser Ser Arg Hie Thr Ile Arg Gin Pro Phe Thr Tyr Hie Ser Arg 160Tyr Phe Thr Pro Lyg Pro Giu Leu Asp Gin Thr Ile Asp Trp Phe Gin 165Tyr Phe Thr Pro Lyg Pro Giu Leu Asp Gin Thr Ile Asp Trp Phe Gin 165Tyr Phe Thr Pro Lyg Pro Giu Leu Asp Gin Thr Ile Asp Trp Phe Gin 185Asn Val Giu His Thr Gly Leu Gly Tyr Ala Leu Gin Ann Ala Thr Thr 180Ann Val Giu His Thr Gly Leu Gly Tyr Ala Leu Gin Ann Ala Thr Thr 200210221222Phe Ile Leu Lys Asp Pro Leu Asn Glu 2252210SEQ (ID NO 13 c211> LENGTH: s21 223022202220Phe The Ser The Phe PhO chreovirus c220> FEATURE: c220> FEATURE: 	L	eu	Pro	Pro	Ser	-	Gly	Thr	Asn	Pro		Pro	Leu	Pro	Phe		Tyr		
Thr Ser Aan Gin Arg Giy Val Giy Ser Thr Val Val Lie Leu Aep Ala 115 Am Phe Val Thr Pro Ser Thr Aan Leu Ala Tyr Aep Pro Tyr Lie Aan 130 Tyr Ser Ser Arg His Thr 11e Arg Gin Pro Phe Thr Tyr His Ser Arg 145 Tyr Phe Thr Pro Lyr Pro Glu Leu Ap Gin Thr 11e Aep Tyr Phe Gin 170 181 181 182 182 183 184 185 185 185 185 186 186 186	Т	yr	Arg	Ile	-	Lys	Ala	Lys	Tyr		Phe	Tyr	Pro	Arg	_	Pro	Ile		
Ann Phe Val Thr Pro Ser Thr Asn Leu Ala Tyr Asp Pro Tyr He Asn 130Tyr Ser Ser Arg His Thr He Arg Gin Pro Phe Thr Tyr His Ser Arg 145Tyr Phe Thr Pro Lyp Pro Glu Leu Asp Gin Thr He Asp Trp Phe Gln 170Tyr Phe Thr Pro Lyp Arg Asn Gin Leu Trp Leu His Leu Asn Thr His Thr 180Asn Val Glu His Thr Gly Leu Gly Tyr Ala Leu Gln Asn Ala Thr Thr 200Ala Gin Asn Tyr Val Val Arg Leu Thr He Tyr Val Gln Phe Arg Glu 210Phe He Leu Lys Asp Pro Leu Ann Glu 225***********************************	Т	'hr	Ser			Arg	Gly	Val	-		Thr	Val	Val			Asp	Ala		
Tyr Ser Ser Arg His Thr 11e Arg Gln Pro Phe Thr Tyr His Ser Arg 155Tyr Phe Thr Pro Lye Pro Glu Leu Ang Gln Thr 11e Ang Try Phe Gln 170Pro Aen Aen Lye Arg Aen Gln Leu Try Leu His Leu Aen Thr His Thr 180Aen Val Glu His Thr Gly Leu Gly Tyr Ala Leu Gln Aen Ala Thr Thr 210Ala Gln Aen Tyr Val Val Arg Leu Thr 11e Tyr Val Gln Phe Arg Glu 210210225Ala Gln Aen Tyr Val Val Arg Leu Thr 11e Tyr Val Gln Phe Arg Glu 21022623124102411242242243243244244244245246246247248248249249240250241241241242243244244244244245246246247248248248248244244245245246246247248248244245245245246246247248248248244244245245246246246247248248244244<	A	sn			Thr	Pro	Ser			Leu	Ala	Tyr	-		Tyr	Ile	Asn		
Tyr Phe Thr Pro Lye Pro Glu Leu App Gln Thr 11e Aep Trp Phe Gln 170 Pro Aan Aen Lye Arg Asn Gln Leu Trp Leu His Leu Aan Thr His Thr 180 Aan Val Glu Hie Thr Gly Leu Gly Tyr Ala Leu Gln Aan Ala Thr Thr 195 Ala Gln Aan Tyr Val Val Arg Leu Thr 11e Tyr Val Gln Phe Arg Glu 210 Phe 11e Leu Lye App Pro Leu Asn Glu 225 <pre>color SEG ID NO 13 <cli>clio SEG ID NO 14 10 10 10 10 10 10 10 10 10 10</cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></cli></pre>		-		Ser	Arg	His			Arg	Gln	Pro			Tyr	His	Ser	-		
Pro Asn Asn Lys Arg Asn Gh Leu Trp Leu His Leu Asn Thr His Thr 190Asn Val Glu His Thr Gly Leu Gly Tyr Ala Leu Gln Asn Ala Thr Thr 200Ala Gln Asn Tyr Val Val Arg Leu Thr Ile Tyr Val Gln Phe Arg Glu 210Phe Ile Leu Lys Asp Pro Leu Asn Glu225<210 SEQ ID NO 13 <211 Line Tyr Val Val Arg Pro Leu Asn Glu 230<211 Strength: 521 <222 Strength			Phe	Thr	Pro	-		Glu	Leu	Asp			Ile	Asp	Trp				
Am Val Giu His Thr Gly Leu Gly Tyr Ala Leu Gln Am Ala Thr Thr 200 Ala Gln Am Tyr Val Val Arg Leu Thr Ile Tyr Val Gln Phe Arg Glu 210 Phe Ile Leu Lyr Amp Pro Leu Am Glu 225 (210) SEQ ID NO 13 (211) LENGTH: 621 (212) TYPE: DNA (212) SEQANISM: Type A PND circovirus (220) FEATURE: (212) TYPE: CDS (222) FEATURE: (212) SEQUENCE: 13 atg ata tot ato cat cat att tot att agg ott cas gta ggt gtc 1 5 Cot agg ctc age aaa att acg gge cca ctg gct ctt cca acc ggg Pro Arg Leu Ser Lyr Ile Thr Gly Pro Leu Ala Leu Pro Thr Thr Gly 20 Crag gcc cac tat gac gtg tac age tgt ctt cca atc acg ctg ctg cat Arg Ala His Tyr Any Val Tyr Ser Cys Leu Pro Ile Thr Leu Leu His 35 Ctt ccc gct cact ta cas agt tac age cag cca ctg gct ctt cca 46 Arg Ala His Tyr Any Val Tyr Ser Cys Leu Pro Ala Glu Ile Ser His 50 Ctt ccc gt cac ttt caa aag ttc age cag cca ctg gct ctt cca 47 Arg Ala His Tyr Any Val Tyr Ser Cys Leu Pro Ala Glu Ile Ser His 50 Ctt ccc gt cac tt cas aga tt acg gcc ca ctg gct ct cca att acg ccc cgt ctc 240 Arg Leu Ser Lys Ile Thr Gly Pro Leu Ala Leu Pro Thr Thr Gly 20 Ct agg ct ca ct ta gac gtg tac age tgt ctt cca att acg ctg ctg ct ct Arg Ala His Tyr Any Val Tyr Ser Cys Leu Pro Ile Thr Leu Leu His 45 Ctt ccc gct cac ttt caa aag ttc age cag cca ctg gcg gaa att tot cac 240 16 17 18 20 20 20 20 20 24 25 24 25 24 24 25 24 25 24 25 24 24 25 24 25 24 24 25 24 24 25 24 24 25 24 24 25 24 24 25 24 25 24 24 25 24 25 24 24 24 25 24 24 25 24 24 24 25 24 24 24 25 24 24 25 24 24 24 25 24 24 24 25 24 24 24 24 24 25 24 24 24 24 25 24 24 25 24 24 24 25 24 24 24 25 24 24 24 25 24 24 24 25 24 24 24 25 24 24 24 25 24 24 25 24 24 25 24 24 25 24 25 24 24 25 24 24 25 2	Ρ	ro	Asn	Asn	-		Asn	Gln	Leu	-		His	Leu	Asn			Thr		
Ala Gin Am Tyr Val Val Arg Leu Thr Ile Tyr Val Gin Phe Arg Glu 210 220 Phe Ile Leu Lys Amp Pro Leu Am Glu 230 	A	sn	Val	Glu		Thr	Gly	Leu	Gly		Ala	Leu	Gln	Asn		Thr	Thr		
210215220Phe Ile Leu Lys Asp Pro Leu Asn Glu 2252210c210> SEQ ID NO 13 c211> LENOTH: 621 c212> TYPE: DNA c212> TYPE: DNA c220> PEATURE: c220> PEATURE: c221> NAME/KEY: CDS c222> LOCATION: (1)(618)c400> SEQUENCE: 13atg ata toc atc cca cca ctt att tot act agg ctt cca gta ggt gtc 1048 10cct agg ctc agc aaa att acg ggc cca ctg gct ctt ccc act a acc ggg 2096 30cct agg ctc agc aaa att acg ggc cca ctg gct ctt ccc act a acc ggg 2096 30cct agg ctc agc aga gtg tac agc tgt ctt cca atc acg ctg ctg cat 35144 45atg ath is Tyr Asp Val Tyr Ser Cys Leu Pro 11e Thr Leu Leu His 35144 45cct cc gct cac ttt caa agg ttc agc ccg ccg gg aa att tct cac 50192 60cct agg tac agc tgt ct gct cac at agg ctg ctc ccc gct cac 46192 26cct acg gt tac agc tgg ctc agt cac caa aga ccc cgt ctc 30240cct agc gg cac cat tg acg tgt ct gct agt cac caa aga ccc cgt ctc 50192 80cct acg tt cac agg aaa ctg ctc ggc tac agt cac caa aga ccc cgt ctc 75240 80caa aag gg tac tac acg acg tg ac agt cac caa aga ccc cgt ctc 70280 75caa aag gg tact cac ag cag tag ag cag gt gct ccc ccc tcc ccc tcc 75281 80caa aag gg tact cac ag cag tag ag cag ctg ctg cg ct ccc ccc cgt ccc 75281 80caa aag gg tact cac ag cag tag ag cag gt cg ct ccc ccc cgt ccc 75281 80caa aag gg tact cac ag cag tag cag cg cg ct ccc ccc cgt ccc 75281 76caa aag gg tact cac ag cag tag cag ctg cg cg cg ccc ccc ccc cgt ccc 75281 75	A	la	Gln		Tyr	Val	Val	Arg		Thr	Ile	Tyr	Val		Phe	Arg	Glu		
225 230 $(210) \leq SEQ ID NO 13$ $(211) \leq LENGTH: 621$ $(212) \leq TTFE: DNA$ $(212) \leq TTFE: DNA$ $(212) \leq TTFE: CDS$ $(222) \in PEATURE: (222) \leq PEATURE:(222) \leq PEATURE:(22) \leq PEATURE:(2$			210		-			215				-				5			
<pre><11> LENGTH: 621 <212> TYPE: DNA <213> ORCANISM: Type A PWD circovirus <220> FEATURE: <221> NAME/KEY: CDS <222> LOCATION: (1)(618) <400> SEQUENCE: 13 atg ata tcc atc cca cca ctt att tct act agg ctt cca gta ggt gtc Met Ile Ser Ile Pro Pro Leu Ile Ser Thr Arg Leu Pro Val Gly Val 1 5 0 15 96 cct agg ctc agc aaa att acg ggc cca ctg gct ctt ccc aca acc ggg 96 Pro Arg Leu Ser Lys Ile Thr Gly Pro Leu Ala Leu Pro Thr Thr Gly 30 cgg gcc cac tat gac gtg tac agc tgt ctt ccc atc acg ctg ctg cat Arg Ala His Tyr Asp Val Tyr Ser Cys Leu Pro Ile Thr Leu Leu His 35 60 ctt ccc gct cac ttt caa aag ttc agc cag ccc gcg gaa att tct cac 30 5 60 ctt ccc gct cac ttt caa aag ttc agc cag ccc gcg gaa att tct cac 45 60 ctt ccc gct cac ttt caa aag ttc agc cag ccc gcg gaa att tct cac 55 60 ata cgt tac agg aaa ctg ct ggt ca agt cac cac aa aga ccc cgt ctc 60 caa aag ggt act cac agc agt aga cag tcg cgt gcg ctt ccc ctg gtt 61 Lys Gly Thr His Ser Ser Arg Gln Val Ala Ala Leu Pro Leu Val 85 90 95</pre>	2	25					230												
atg ata tcc atc cca cca ctt att tct act agg ctt cca gta ggt ggt ggt gtc48Met Ile Ser Ile Pro Pro Leu Ile Ser Thr Arg Leu Pro Val Gly Val15cct agg ctc agc aaa att acg ggc cca ctg gct ctt ccc aca acc ggg96Pro Arg Leu Ser Lys Ile Thr Gly Pro Leu Ala Leu Pro Thr Thr Gly96cgg gcc cac tat gac gtg tac agc tgt ctt cca atc acg ctg ctg ctg cat144Arg Ala His Tyr Asp Val Tyr Ser Cys Leu Pro Ile Thr Leu Leu His144ctt ccc gct cac ttt caa aag ttc agc cag cag ccc gcg gaa att tct cac192Leu Pro Ala His Phe Gln Lys Phe Ser Gln Pro Ala Glu Ile Ser His192ata cgt tac agg aaa ctg ctc ggc tac agt cag cac cac caa aga ccc cgt ctc ro2401e Arg Tyr Arg Lys Leu Leu Gly Tyr Ser 75160 Arg Pro Arg Leu Val2065909095	< < < < <	211 212 213 220	L> LE 2> TY	ENGTH	H: 62 DNA														
MetIleSerIleProLeuIleSerThrArgLeuProValGlyVal115151101010101596cctaggccaaggagaattacgggccaacaacggg96ProArgLeuSerLysIleThrGlyProLeuAlaLeuProThrGly96cgggcccaatatgaggtgtataggtgtcttccaatcaggctgfdfArgAlaHisTyrAspValTyrSerCysLeuProIleThrHisArgAlaHisTyrAspValTyrSerCysLeuProIleThrLeuHisArgAlaHisTyrAspValTyrSerCysLeuProIleThrLeuHisArgAlaHisTyrAspValTyrSerCysLeuProIleThrLeuHisHisArgAlaHisPhoGlnLysPhoSerGlnProAlaGluIleSerHisIleCttccaggcaacagcagcagcagcagcagcagcagcag240IleLeu <t< td=""><td></td><td></td><td>L> NA</td><td>EATUF AME/F</td><td>RE: KEY:</td><td>CDS</td><td></td><td></td><td>circo</td><td>viru</td><td>ıs</td><td></td><td></td><td></td><td></td><td></td><td></td><td></td><td></td></t<>			L> NA	EATUF AME/F	RE: KEY:	CDS			circo	viru	ıs								
ProArgLeuSerLysIleThrGlyProLeuAlaLeuProThrGlyCgggcccactatgacgtgtacagctgftacaccatcctgctgcat144ArgAlaHisTyrAspValTyrSerCysLeuProIleArgLeuHis144Cttcccgctcactttcaaaagttcagccagcagcaggcdcat144LeuProAlaHisProCadaagcccgcggaaatttctcaccac192Cttcccgcdtacaggaaactgctcggccaccadagialu1leSerHis50NSrSrSrSrSrSrSrSrSrAlaAlaSerHisSer192atacggtacaggaaactggcdtacaggcaccaaaggccccgftctalu65NYrArgLysLeuGlnYrSerHisGlnArgProArgLeu24061LysSrYrSerArgGlnValAlaAlaLeuProLeuYal24061LysSerSerArgSerSer </td <td></td> <td>222 400</td> <td>l> N7 2> LC D> SE</td> <td>EATUF AME/F DCATI EQUEN</td> <td>RE: KEY: ION: ICE:</td> <td>CDS (1) 13</td> <td>(6:</td> <td>18)</td> <td></td>		222 400	l> N7 2> LC D> SE	EATUF AME/F DCATI EQUEN	RE: KEY: ION: ICE:	CDS (1) 13	(6:	18)											
ArgAlaHisTyrAspValTyrSerCysLeuProIleThrLeuHis3535353540354545192cttcccgctcactttcaaaagttcagccagcccggggaaattttccac192LeuProAlaHisPheGlnLysPheSerGlnProAlaGlu11eSerHis5055555657606011eSer192atacgttacaggaaactgctgggccaccaaagaccc240atacgttacaggagacagggcggccaccagagacag24065707075SerHisGlnArgProArgLeu24065707075SerGlnArgProArgLeu24061LysGlyThrHisSerSerArgGlnArgProArgLeu70707075Serggctccccctggtt288GlnLysGlyThrHisSerSerArgGlnValAlaAlaLeuVal95caaaagggggg5gggggggg	a M	222 400 tg let	l> NA 2> LC D> SE ata	EATUF AME/F DCATI EQUEN tcc	RE: KEY: ION: NCE: atc	CDS (1) 13 cca Pro	(6: cca	18) ctt	att	tct	act Thr					Gly		48	
Leu Pro Ala His Phe Gln Lys Phe Ser Gln Pro Ala Glu Ile Ser His 50 ata cgt tac agg aaa ctg ctc ggc tac agt cac caa aga ccc cgt ctc 11e Arg Tyr Arg Lys Leu Leu Gly Tyr Ser His Gln Arg Pro Arg Leu 65 caa aag ggt act cac agc agt aga cag gtc gct gcg ctt ccc ctg gtt Gln Lys Gly Thr His Ser Ser Arg Gln Val Ala Ala Leu Pro Leu Val 85 90 95	a M 1 c	400 tg let	l> NZ 2> LC D> SE ata Ile agg	EATUF AME/H DCATI EQUEN tcc Ser ctc	RE: KEY: ION: NCE: atc Ile agc Ser	CDS (1) 13 cca Pro 5 aaa	(6: cca Pro att	18) ctt Leu acg	att Ile ggc	tct Ser cca Pro	act Thr 10 ctg	Arg	Leu ctt	Pro ccc	Val aca Thr	Gly 15 acc	Val 999		
Ile Arg Tyr Arg Lys Leu Leu Gly Tyr Ser His Gln Arg Pro Arg Leu 80 65 70 75 80 caa aag ggt act cac agc agt aga cag gtc gct gcg ctt ccc ctg gtt 288 Gln Lys Gly Thr His Ser Ser Arg Gln Val Ala Ala Leu Pro Leu Val 90 95	a M 1 P C	222 400 let ct ro	l> NA 2> LO D> SE ata Ile Arg gcc	EATUR AME/H DCATI EQUEN tcc Ser ctc Leu cac His	RE: (CEY: ION: NCE: atc Ile agc Ser 20 tat	CDS (1) 13 cca Pro 5 aaa Lys gac	(6: cca Pro att Ile gtg	18) ctt Leu acg Thr tac	att Ile Gly agc Ser	tct Ser cca Pro 25 tgt	act Thr 10 ctg Leu ctt	Arg gct Ala cca	Leu ctt Leu atc	Pro ccc Pro acg Thr	Val aca Thr 30 ctg	Gly 15 acc Thr ctg	Val 999 Gly cat	96	
Gln Lys Gly Thr His Ser Ser Arg Gln Val Ala Ala Leu Pro Leu Val 85 90 95	a M I C P C A	222 400 Itg let Pro Sgg arg	l> NZ 2> LC D> SE ata Ile Arg gcc Ala ccc Pro	EATUH AME/H DCATI EQUEN tcc Ser ctc Leu cac His 35 gct	RE: KEY: ION: NCE: atc Ile agc Ser 20 tat Tyr cac	CDS (1) 13 cca Pro 5 aaa Lys gac Asp ttt	(6: cca Pro att Ile gtg Val caa	18) ctt Leu Thr tac Tyr aag Lys	att Ile ggc Gly agc Ser 40 ttc	tct Ser Cca Pro 25 tgt Cys agc	act Thr 10 ctg Leu ctt Leu cag	Arg gct Ala cca Pro ccc	Leu ctt Leu atc Ile gcg Ala	Pro ccc Pro acg Thr 45 gaa	Val aca Thr 30 ctg Leu att	Gly 15 acc Thr ctg Leu tct	Val ggg Gly cat His cac	96 144	
con one and too aca ote dat and tat did doo too to tot aca dia 336	a M I C P C C A C C L I I I I	2222 400 let cct 270 egg arg ett eu ta	L> NA 2> LC D> SE ata Ile Arg gcc Ala ccc Pro 50 cgt	EATUH AME/H DCATI EQUEN tcc Ser ctc Leu cac His 35 gct Ala tac	RE: KEY: ION: NCE: atc Ile agc Ser 20 tat Tyr cac His agg	CDS (1). 13 cca Pro 5 aaa Lys gac Asp ttt Phe aaa	(6) ccaa Pro att Ile gtg Val caa Gln ctg Leu	18) ctt Leu acg Thr tac Tyr tac Tyr 25 55 ctc	att Ile ggc Gly agc Ser 40 ttc Phe ggc	tct Ser 25 tgt Cys agc Ser tac	act Thr 10 ctg Leu ctt Leu cag Gln agt	Arg gct Ala cca Pro ccc Pro cac	Leu ctt Leu atc Ile gcg Ala 60 caa	Pro ccc Pro acg Thr 45 gaa Glu aga	Val aca Thr 30 ctg Leu att Ile ccc	Gly 15 acc Thr ctg Leu tct Ser cgt	Val ggg Gly cat His cac His ctc Leu	96 144 192	
Pro Arg Ser Ser Thr Leu Asp Lys Tyr Val Ala Phe Phe Thr Ala Val	a M P C A C C A L L I I 6 C C	2222 400 let ggg arg stt eu tta le 5 aa	L> NA 2> LC D> SF ata Ile Arg gcc Ala ccc Pro 50 cgt Arg aag	EATUH ME/I DCATJ EQUEN tcc Ser ctc Leu cac His 35 gct Ala tac Tyr ggt	RE: KEY: ION: ACE: atc Ile agc Ser 20 tat Tyr cacc His agg Arg act	CDS (1) 13 cca Pro 5 aaaa Lys gac Asp ttt Phe aaaa Lys cac His	cca Pro att Ile gtg Val caa Gln ctg Leu 70 agc	18) ctt Leu Thr tac Tyr aagg Lys 55 ctc Leu agt	att Ile ggc Gly agc Ser 40 ttc Phe ggc Gly aga	tct Ser 25 tgt Cys agc Ser tac Tyr cag	act Thr 10 ctg Leu ctt Leu cag Gln agt Ser ytc Val	Arg gct Ala cca Pro ccc Pro cac His 75 gct	Leu ctt Leu atc Ile gcg Ala 60 caa Gln gcg	Pro ccc Pro acg Thr 45 gaa Glu aga Arg ctt	Val aca Thr 30 ctg Leu att Ile ccc Pro ccc	Gly 15 acc Thr ctg Leu tct Ser cgt Arg ctg Leu	Val ggg Gly cat His cac His ctc Leu 80 gtt	96 144 192 240	

-continued

											-	con	tin	ued					
			100					105					110						
												gat Asp 125				384			
								-		~	-	ctt Leu		-		432			
												ttc Phe				480			
												aag Lys				528			
	-									-		gaa Glu				576			
												ttt Phe 205		tga		621			
<211 <212 <213	L> LE 2> TY 3> OF	EQ II ENGTH (PE : RGANI EQUEN	H: 2 PRT ISM:	26 Тур	϶ΑΙ	PWD (circo	oviru	18										
					Pro	Leu	Ile	Ser	Thr 10	Arg	Leu	Pro	Val	Gly 15	Val				
Pro	Arg	Leu		Lys	Ile	Thr	Gly		Leu	Ala	Leu	Pro		Thr	Gly				
			20					25					30						
Arg	Ala	His 35		Asp	Val	Tyr	Ser 40		Leu	Pro	Ile	Thr 45		Leu	His				
_		35	Tyr	-		-	40	Суз					Leu						
Leu	Pro 50	35 Ala	Tyr His	Phe	Gln	Lуя 55	40 Phe	Cys Ser	Gln	Pro	Ala 60	45	Leu Ile	Ser	His				
Leu Ile 65	Pro 50 Arg	35 Ala Tyr	Tyr His Arg	Phe Lys	Gln Leu 70	Lys 55 Leu	40 Phe Gly	Cys Ser Tyr	Gln Ser	Pro His 75	Ala 60 Gln	45 Glu	Leu Ile Pro	Ser Arg	His Leu 80				
Leu Ile 65 Gln	Pro 50 Arg Lys	35 Ala Tyr Gly	Tyr His Arg Thr	Phe Lys His 85 Thr	Gln Leu 70 Ser	Lys 55 Leu Ser	40 Phe Gly Arg	Cys Ser Tyr Gln	Gln Ser Val 90	Pro His 75 Ala	Ala 60 Gln Ala	45 Glu Arg	Leu Ile Pro Pro	Ser Arg Leu 95	His Leu 80 Val				
Leu Ile 65 Gln Pro	Pro 50 Arg Lys Arg	35 Ala Tyr Gly Ser	Tyr His Arg Thr Ser 100	Phe Lys His 85 Thr	Gln Leu 70 Ser Leu	Lys 55 Leu Ser Asp Gly	40 Phe Gly Arg Lys	Cys Ser Tyr Gln Tyr 105	Gln Ser Val 90 Val	Pro His 75 Ala Ala	Ala 60 Gln Ala Phe	45 Glu Arg Leu	Leu Ile Pro Pro Thr 110	Ser Arg Leu 95 Ala	His Leu 80 Val Val				
Leu Ile 65 Gln Pro Phe	Pro 50 Arg Lys Arg Phe	35 Ala Tyr Gly Ser Ile 115 Lys	Tyr His Arg Thr Ser 100 Leu	Phe Lys His 85 Thr Leu	Gln Leu 70 Ser Leu Val	Lys 55 Leu Ser Asp Gly	40 Phe Gly Arg Lys Ser 120 Leu	Cys Ser Tyr Gln Tyr 105 Phe	Gln Ser Val 90 Val Arg	Pro His 75 Ala Ala Phe	Ala 60 Gln Ala Phe Leu	45 Glu Arg Leu Phe Asp	Leu Ile Pro Pro Thr 110 Val	Ser Arg Leu 95 Ala Ala	His Leu 80 Val Val Ala				
Leu Ile 65 Gln Pro Phe Gly	Pro 50 Arg Lys Arg Phe Thr 130	35 Ala Tyr Gly Ser Ile 115 Lys	Tyr His Arg Thr Ser 100 Leu Ile	Phe Lys His 85 Thr Leu Pro	Gln Leu 70 Ser Leu Val Leu	Lys 55 Leu Ser Asp Gly His 135 Val	40 Phe Gly Arg Lys Ser 120 Leu	Cys Ser Tyr Gln Tyr 105 Phe Val	Gln Ser Val 90 Val Arg Lys	Pro His 75 Ala Ala Phe Ser	Ala 60 Gln Ala Phe Leu 140	45 Glu Arg Leu Phe Asp 125	Leu Ile Pro Pro Thr 110 Val Leu	Ser Arg Leu 95 Ala Ala Ser	His Leu 80 Val Val Ala Lys				
Leu Ile 65 Gln Pro Phe Gly Ile 145	Pro 50 Arg Lys Arg Phe Thr 130 Arg	35 Ala Tyr Gly Ser Ile 115 Lys Lys	Tyr His Arg Thr Ser 100 Leu Ile Pro	Phe Lys His 85 Thr Leu Pro Leu	Gln Leu Ser Leu Val Leu Glu 150	Lys 55 Leu Ser Asp Gly His 135 Val	40 Phe Gly Arg Lys Ser 120 Leu Arg	Cys Ser Tyr Gln Tyr 105 Phe Val Ser	Gln Ser Val 90 Val Arg Lys Ser	Pro His 75 Ala Ala Ala Ser Thr 155 Asp	Ala 60 Gln Ala Phe Leu Leu Leu	45 Glu Arg Leu Phe Asp 125 Leu	Leu Ile Pro Pro Thr 110 Val Leu Gln	Ser Arg Leu 95 Ala Ala Ser Thr	His Leu 80 Val Val Ala Lys Phe 160				
Leu Ile 65 Gln Pro Phe Gly Ile 145 Leu Phe	Pro 50 Arg Lys Arg Phe Thr 130 Arg Ala Val	35 Ala Tyr Gly Ser Ile 115 Lys Lys Thr Phe	Tyr His Arg Thr Ser 100 Leu Ile Pro Asn Leu 180	Phe Lys 85 Thr Leu Pro Leu Lys 165 Leu	Gln Leu 70 Ser Leu Val Leu Glu 150 Ile Leu	Lys 55 Leu Ser Asp Gly His 135 Val Ile Gly	40 Phe Gly Arg Lys Ser 120 Leu Arg Arg	Cys Ser Tyr Gln Tyr 105 Phe Val Ser Lys Ile 185	Gln Ser Val Arg Lys Ser Gly 170 Ile	Pro His 75 Ala Ala Phe Ser Thr 155 Asp Lys	Ala 60 Gln Ala Phe Leu Leu Leu Trp Gly	45 Glu Arg Leu Phe Asp 125 Leu Phe	Leu Ile Pro Thr 110 Val Leu Gln Leu His 190	Ser Arg 55 Ala Ala Ser Thr Pro 175	His Leu 80 Val Val Lys Phe 160 Tyr				

<210> SEQ ID NO 15 <211> LENGTH: 1767

48

96

		00110111000
<212> TYPE: DNA	4	
	- Type B PWD circovirus	
<220> FEATURE:		
<221> NAME/KEY:	CDS	
<222> LOCATION:	(1)(111)	
<220> FEATURE:		
<221> NAME/KEY:		
<222> LOCATION:	(115)(243)	
<220> FEATURE:	675 G	
<221> NAME/KEY:		
<222> LOCATION: <220> FEATURE:	(247)(267)	
<221> NAME/KEY:	CDS	
<222> LOCATION:		
<220> FEATURE:	(2,2,),,(3,3,3)	
<221> NAME/KEY:	CDS	
<222> LOCATION:		
<220> FEATURE:		
<221> NAME/KEY:	CDS	
<222> LOCATION:	(421)(447)	
<220> FEATURE:		
<221> NAME/KEY:	CDS	
<222> LOCATION:	(451)(471)	
<220> FEATURE:		
<221> NAME/KEY:		
<222> LOCATION:	(475)(510)	
<220> FEATURE:	ana	
<221> NAME/KEY:		
<222> LOCATION: <220> FEATURE:	(514)(516)	
<2220> FEATORE: <221> NAME/KEY:	CDS	
<2221> NAME/REL: <222> LOCATION:		
<222> ECCHIICK.	(320) (725)	
<221> NAME/KEY:	CDS	
<222> LOCATION:		
<220> FEATURE:		
<221> NAME/KEY:	CDS	
<222> LOCATION:	(757)(759)	
<220> FEATURE:		
<221> NAME/KEY:		
<222> LOCATION:	(763)(804)	
<220> FEATURE:		
<221> NAME/KEY:		
<222> LOCATION:	(808)(861)	
<220> FEATURE: <221> NAME/KEY:	CDC	
<2221> NAME/REI: <222> LOCATION:		
<2223 LOCATION: <220> FEATURE:	(865)(984)	
<221> NAME/KEY:	CDS	
<222> LOCATION:		
<220> FEATURE:	()	
<221> NAME/KEY:	CDS	
<222> LOCATION:	: (1177)(1233)	
<220> FEATURE:		
<221> NAME/KEY:	CDS	
<222> LOCATION:	: (1237)(1359)	
<220> FEATURE:		
<221> NAME/KEY:		
	: (1363)(1476)	
<220> FEATURE:		
<221> NAME/KEY:		
	: (1480)(1737)	
<220> FEATURE:	CDC	
<221> NAME/KEY:	: (1741)(1767)	
<400> SEQUENCE:		
	: ID : cgg cag cgg cag cac ctc c	ade ade ace tea dea dea
	1 Arg Gln Arg Gln His Leu (
1	5 10	15
	a aga aga atg gaa gaa gcg g	
-	a Arg Arg Met Glu Glu Ala A	
20	25	30

-continued

												COIL		ucu		
	glà aaa											agc Ser				144
	glà aaa															192
	gta Val 65															240
ttg Leu 80	tga			aga Arg								att Ile 90				288
	gcc Ala 95															336
	gca Ala											gag Glu				384
	agg Arg	-					-		-	-	-	gta Val		-		432
-	gcg Ala		-		-							ctg Leu	taa	cgt Arg	-	480
	gaa Glu 155													gga Gly		528
-	aga Arg	-				-		-		-		-				576
	ggt Gly 185															624
	cat His															672
	aag Lys															720
	atc Ile		tga		tgt Cys							ctg Leu	tag	aga Arg		768
	gtg Val													cca Pro	<u> </u>	816
	aga Arg														tag	864
-	ctc Leu												-	-		912
	aac Asn															960
	gcc Ala							taa				tct Ser				1008

-continued

cgt and ggt but tat bat ba tag agg ggt agg ggt cut. 1056 att aas tot tu gaa tig tac ata ggt aag tag ggt ag tut gt. 1104 1300 335 att aas tot tu gaa tig tac ata ggt aag gg gat att gt. 1104 1300 345 cig gdt gta tat att gt tac ata ggt aag gg gg gdt att gt. 1152 1300 345 cig gdt gta tat att gt tu tu gaa gg ggt get tut. 1152 141 aas tot tu cu ag agt tig tag tu cag ca cag cig gg gdt at gt. 1152 155 345 345 156 345 346 171 Tip ted Glu Yal Tip and set at an agt gaa atc tag ggt ad ggt gdt tut. 1248 340 345 346 171 Tip ted Glu Yal Tip and set du lig dlu Gly Cys Gly tut. 1344 341 Tip ted Glu Yal Tip and set du lig dlu Gly Cys Gly tut. 1344 342 410 410 440 342 410 410 440 342 410 410 440 410 410 410 410 410 410 410 410 410 410 410 410 410 410
<pre>lie Lyop Phe Ser Gu Lem Tyr IIe km Gly Tyr The Amp IIe Val 340 345 345 345 345 345 345 345 345 345 345</pre>
Leu Val Val Tyr Thr Val Phe Glu Arg Ser Ala Glu Ala Tyr Val 365 tao att tee age agt tig tag tet cag cea cag etg git tet titt Tyr Ile Ser Ser Ser Leu Ser Glu Pro Glu Leu Val Ser Phe 375 ser Glu Pro Glu Val Ser Glu Pro Glu Leu Val Ser Phe 375 geg gg gg gg gg gg gg gg ag ag ag ag gg ctg gg tat gg tag gg ag gg ct dig tag ag rag gr yal Trp Leu Glu Val Ile Arm Ser Glu Ile Gly Tyr Gly Met Ala 410 410 410 411 412 413 414 415 415 416 417 417 418 419 419 419 419 419 419 419 419
Tyr lle Ser Ser Ser Leu Ser Cln Pro Cln heu Val Ser Phe 375 gtt tgg ttg gaa gta atc aat agt gaa atc tag gac agg ttt ggg val Trp Leu Glu Val 11e Aan Ser Glu 11e Aap Arg Phe Gly 390 aaa gta cog gga gtg gta gga gaa ggg ctg ggt tat ggt atg gcg Lyw Val Pro Cly Val Val Gly Glu Gly Leu Gly Tyr Gly Met Ala 405 405 407 408 409 409 409 409 409 409 409 409
Val Trp Leu Glu Val ILe Aem Ser Glu Ile 390 Amp Arg Phe Gly 395 Amp Arg Phe Gly 396 1296 aaa gta cog gag gc gt t agga gag ot u ggt tat ggt at ggt ag go 405 410 1296 ye Val Pro Gly Val Val Gly Glu Gly Leu Gly Tyr Gly Met Ala 415 415 1344 gga gt gt tac ata ggg gt cat agt aggt ag
Lye Val Pro Gly Val Val Gly Glu Gly Leu Gly Tyr Gly Met Ala 410Alsgga gta gtt tac ata ggg gtc ata ggt gad ggc tgt ggc ctt tgt 4201344 430gaa gtt atc ata aat aac agc acg gga gco cac cc cc ct gtc 1ye Val Ile Ile 4351344 430aaa gtt atc ata aat aac agc acg ggg gco cag gaa ttc aac ctt acc 4451392Lye Val Ile Ile 435Asn Asn Ser Thr Gly Ala His Ser Pro Val 4451392Lye Ugy Agr Arg Gly Ala Gly Pro Glu Phe Asn Leu Asn Leu Ser 4501440Leu Gly Agr Arg Gly Ala Gly Cac gaa ttc aac ctt acc cc tc cc 4501440Let gta gta ttc aaa ggg cac agg ggg ggg ggg ggg ggg gg
GIY Val Tyr The GIY <
LyoVal11e11eAnn Ann SerThrGiyAla HisSerProVal 445ctgggtggtcgggggggaggaggggga<
Leu Gly App Arg Gly Ala Gly Pro Glu Phe Asn Leu Asn Leu Ser 460 tct gta gta ttc aaa ggg cac aga ggg gg tt tg 465 val val Phe Lys Gly Hia Arg Ala Gly Val v
Ser Val Val Phe Lys Oly His Arg Ala Giy Val 475 476 477 477 477 477 477 477 477
GIY Lys Lys Val Ile Asn Ile Glu Ser His His Wal His Arg Pro 480 485 485 490 490 490 1584 490 ggg cgt tct gac tgg cgt 10 Gly Arg 1632 gag gcg ggg dgg gad gad gad gad acg 1632 gu Ala Glu Ala Alp Pro Ser Pro Ala Na Thr 515 500 Gig ggg ggg ggd acg gad Gad 1632 gcd gag gcg ggg dgg ggd gad caa 1680 Ala Glu Ala Ala Glu Asp Glu Ala Ala Glu Ala Ala Glu Ala Ala Glu Ala Ala Ala Ala Ala Glu Ala Ala Ala Ala Ala Ala Ala Ala Ala
ggg ogt tet gae tgt ggt teg ett gae agt ata tee gaa ggt ggg1584d1y Arg Ser Asp CysG1y Ser Leu Asp Ser Ile Ser Glu Gly Ala1632gag geg ggt gtt Glu Ala Gly Val Glu Asp Ala IlePhe Pro Ser Pro Ala Val Thr 5151632geg ggg ggt ggt gae gag cea ggg geg geg geg gag gat etg gee asp 5301680Ala Gly Val Asp Glu Pro Gly Ala Ala AlaAla Ser Val Thr Pro Stor Try Ile1680get geg ggg ggg ggt gt tet tet tet tet t
gag gcg ggt gtt gaa gat gcc att ttt cct tct cca gcg gta acg Glu Ala Gly Val glu Asp Ala le the pro 520 ser Pro Ala Val Thr 520 ser Pro Ala Val Thr 5251630gcg ggg ggt ggt gac gag cca ggg gcg gcg gcg gcg gcg gag ga
gcg ggg gtg gac gag cca ggg gcg gcg gcg gcg gac gac
gct gcg ggg gcg gtg tct tct tct tcg gta acg cct cct tgg ata 1728 Ala Ala Gly Ala Val Ser Ser Ser Val Thr Pro Pro Trp Ile 545 550 cat atc tga aaa cga aag aag tgc gct gta agt att 1767 His Ile Lys Arg Lys Lys Cys Ala Val Ser Ile 1767 > SEQ ID NO 16 565 > SEQ UD NO 16 565 > TYPE: PRT > ORGANISM: Type B PWD circovirus >> SEQUENCE: 16 Ser Ala Leu Arg Gln Arg Gln His Leu Gly Ser Thr Ser Ala Ala 10 15
cat atc tga aaa cga aag aag tgc gct gta agt att 1767 His Ile Lys Arg Lys Lys Cys Ala Val Ser Ile 560 565 > SEQ ID NO 16 > LENGTH: 569 > TYPE: PRT >> ORGANISM: Type B PWD circovirus >> SEQUENCE: 16 Ser Ala Leu Arg Gln Arg Gln His Leu Gly Ser Thr Ser Ala Ala 15
<pre>>> SEQ ID NO 16 >> LENGTH: 569 >> TYPE: PRT >> ORGANISM: Type B PWD circovirus >> SEQUENCE: 16 Ser Ala Leu Arg Gln Arg Gln His Leu Gly Ser Thr Ser Ala Ala 5 10 15</pre>
Ser Ala Leu Arg Gln Arg Gln His Leu Gly Ser Thr Ser Ala Ala 5 10 15
5 10 15
Cys Pro Ala Arg Arg Met Glu Glu Ala Asp Pro Asn Pro Ile Lys

-continued

											-	con	tin	ued	
			20					25					30		
Gly	Gly	Суз 35	Ser	Leu	Ile	Ile	Leu 40	Pro	Lys	Thr	Ser	Ala 45	Arg	Lys	Tyr
Gly	Ile 50	Phe	Gln	Tyr	Pro	Tyr 55	Leu	Ile	Ile	Leu	Leu 60	Leu	Ala	Arg	Arg
Val 65	Met	Arg	Lys	Aap	Glu 70	His	Leu	Thr	Ser	Arg 75	Gly	Ser	Leu	Ile	Leu 80
Arg	Ser	Arg	Leu	Leu 85	Ile	Lys	Ser	Gly	Ile 90	Trp	Val	Pro	Ala	Ala 95	Thr
Ser	Arg	Lys	Arg 100	-	Glu	Gln	Ile	Ser 105	Arg	Ile	Lys	Asn	Thr 110	Ala	Val
Lys	Lys	Ala 115	Thr	Tyr	Trp	Ser	Val 120	Glu	Leu	Leu	Asp	Leu 125	Arg	Asp	Asn
Gly	Val 130		Сүз	Leu	Leu	Leu 135	Val	Pro	Cys	Trp	Arg 140	Ala	Gly	Val	Trp
Pro 145	Leu	Gln	Ser	Ser	Thr 150		Arg	Leu	Ser	Glu 155	Ile	Ser	Ala	Gly	Trp 160
Leu	Asn	Phe	Lys	Ala 165	Gly	Lys	Суз	Arg	Ser 170		Ile	Gly	Arg	Leu 175	Met
Tyr	Thr	Ser	Leu 180	Trp	Gly	His	Leu	Gly 185	Val	Val	ГЛа	Ala	Asn 190	Gly	Leu
Leu	Ile	Leu 195	Gln	Thr	Arg	Lys	Pro 200		Thr	Gly	Asn	His 205	Leu	Glu	Thr
Ser	Gly 210		Met	Val	Thr	Met 215	Val	Lys	Lys	Trp	Leu 220	Leu	Leu	Met	Thr
Phe 225	Met	Ala	Gly	Суа	Pro 230	_	Met	Ile	Tyr	Asp 235	-	Val	Ile	Asp	Ile 240
His	Leu	Arg	Leu	Lys 245	Val	Glu	Leu	Tyr	Leu 250		Trp	Pro	Ala	Val 255	Phe
Leu	Pro	Ala	Ile 260	Arg	Pro	Arg	Trp	Asn 265	-	Thr	Pro	Gln	Leu 270	Leu	Ser
Gln	Leu	Lys 275	Leu	Phe	Ile	Gly	Gly 280		Leu	Pro	Trp	Tyr 285	Phe	Gly	Arg
Met	Leu 290	Gln	Asn	Asn	Pro	Arg 295		Lys	Gly	Ala	Ser 300	Ser	Ser	Pro	Phe
Pro 305		His	Ala		Asn 310		His		-	Ile 315		Glu	Ser	Phe	Leu 320
		Arg	Asn				Tyr			Arg		Lys	Trp	Gly 335	Val
Phe	Lys	Ile	Lys 340		Ser	Glu	Leu	Tyr 345			Gly	Tyr	Thr 350	Asp	Ile
Val	Phe	Leu 355		Val	Tyr	Thr	Val 360		Glu	Arg	Ser	Ala 365	Glu	Ala	Tyr
Val	Val 370		Ile	Ser	Ser	Ser 375		Ser	Gln	Pro	Gln 380	Leu	Val	Ser	Phe
Val 385		Trp	Leu	Glu	Val 390	Ile	Asn	Ser	Glu	Ile 395		Arg	Phe	Gly	Gly 400
	Val	Pro	Gly	Val 405			Glu	Gly	Leu 410	Gly	Tyr	Gly	Met	Ala 415	
Gly	Val	Val	Tyr 420		Gly	Val	Ile	Gly 425			Суз	Gly	Leu 430	Cys	Tyr
			120					.2.3					100		

```
-continued
```

Lys Val Ile Ile Asn Asn Ser Thr Gly Ala His Ser Pro Val Thr Leu Gly Asp Arg Gly Ala Gly Pro Glu Phe Asn Leu Asn Leu Ser Tyr Ser Val Val Phe Lys Gly His Arg Ala Gly Val Pro Pro Ser Trp Gly Lys Lys Val Ile Asn Ile Glu Ser His His Val His Arg Pro Gly Gly Arg Ser Asp Cys Gly Ser Leu Asp Ser Ile Ser Glu Gly Ala Gly Glu Ala Gly Val Glu Asp Ala Ile Phe Pro Ser Pro Ala Val Thr Val Ala Gly Val Asp Glu Pro Gly Ala Ala Ala Glu Asp Leu Ala Lys Met Ala Ala Gly Ala Val Ser Ser Ser Ser Val Thr Pro Pro Trp Ile Arg His Ile Lys Arg Lys Lys Cys Ala Val Ser Ile <210> SEQ ID NO 17 <211> LENGTH: 542 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 17 Pro Ala His Phe Gly Ser Gly Ser Thr Ser Ala Ala Pro Gln Gln Gln His Ala Gln Gln Glu Glu Trp Lys Lys Arg Thr Pro Thr Pro Lys Val Gly Val His Ser Glu Ser Phe Arg Arg Arg Ala Gln Glu Asn Thr Gly Ser Ser Asn Ile Pro Ile Leu Phe Tyr Cys Trp Arg Gly Gly Arg Thr Asn Thr Ser Pro Pro Gly Val Arg Phe Cys Glu Glu Ala Asp Phe 65 70 75 80 Ser Glu Val Val Phe Gly Cys Pro Leu Pro His Arg Glu Ser Glu Arg Asn Arg Ser Ala Glu Arg Ile Leu Gln Arg Arg Gln Leu Thr Asp Gly Val Trp Ser Ser Ile Ser Gly Thr Thr Glu Pro Val Tyr Cys Cys Glu Tyr Leu Val Gly Glu Arg Glu Ser Gly Asp Arg Cys Arg Ala Ala Pro Cys As
n Val Cys Gl
n Lys Phe Pro Arg Ala Gly Thr \mbox{Phe} Glu
 Ser Glu Arg Glu Asn Ala Glu Ala Cys Thr Arg His Cys Gly Ala Thr Trp Val Trp Lys Gln Met Gly Cys Phe Cys Arg Pro Gly Asn His Ile Leu Glu 180 185 190 Thr Thr Lys Gln Val Val Gly Trp Leu Pro Trp Arg Ser Gly Cys Tyr Leu Leu Trp Leu Ala Ala Leu Gly Ser Thr Glu Thr Val Ser Ile Ser

-continued

	210					215					220				
Ile 225	Aab	Cys	Arg	Asp	Arg 230	Trp	Asn	Сув	Thr	Phe 235	Phe	Gly	Pro	Gln	Tyr 240
Ser	Asp	Tyr	Gln	Gln 245	Ser	Asp	Pro	Val	Gly 250	Met	Val	Leu	Leu	Asn 255	Сув
Cya	Pro	Ser	Cys 260	Arg	Ser	Ser	Leu	Ser 265	Glu	Aab	Tyr	Phe	Leu 270	Gly	Ile
Leu	Glu	Glu 275	Суз	Tyr	Arg	Thr	Ile 280	His	Gly	Gly	Arg	Gly 285	Pro	Val	Arg
His	Pro 290	Phe	Pro	Pro	Met	Pro 295	Asn	Lys	Leu	Leu	Ser 300	Leu	Phe	Tyr	His
Phe 305	Val	Met	Val	Phe	Ile 310	Ile	His	Gly	Leu	Ser 315	Gly	Gly	Ser	Leu	Lys 320
Leu	Asn	Ser	Leu	Asn 325	Сүз	Thr	Tyr	Met	Val 330	Thr	Arg	Ile	Leu	Tyr 335	Ser
Trp	Ser	Tyr	Ile 340	Leu	Phe	Ser	Asn	Ala 345	Val	Pro	Arg	Pro	Thr 350	Trp	Ser
Thr	Phe	Pro 355	Ala	Val	Сүз	Ser	Leu 360	Ser	His	Ser	Trp	Phe 365	Leu	Leu	Leu
Phe	Gly 370	Trp	Lys	Ser	Ile	Val 375	Lys	Ser	Arg	Thr	Gly 380	Leu	Gly	Val	Lys
Tyr 385	Arg	Glu	Trp	Glu	Lys 390	Gly	Trp	Val	Met	Val 395	Trp	Arg	Glu	Glu	Val 400
Arg	Ala	Val	Ala	Phe 405	Val	Thr	Lys	Leu	Ser 410	Ser	Lys	Ile	Thr	Ala 415	Leu
Glu	Pro	Thr	Pro 420	Leu	Ser	Pro	Trp	Val 425	Ile	Gly	Glu	Gln	Gly 430	Gln	Asn
Ser	Thr	Leu 435	Thr	Phe	Leu	Ile	Leu 440	Tyr	Ser	ГÀа	Gly	Thr 445	Glu	Arg	Gly
Phe	Asp 450	Pro	Pro	Pro	Gly	Gly 455	Arg	Lys	Ser	Leu	Ile 460	Leu	Asn	Leu	Ile
Met 465	Ser	Thr	Ala	Gln	Glu 470	Gly	Val	Leu	Thr	Val 475	Val	Arg	Leu	Thr	Val 480
Tyr	Pro	Lys	Val	Arg 485	Glu	Arg	Arg	Val	Leu 490	ГÀа	Met	Pro	Phe	Phe 495	Leu
Leu	Gln	Arg	Arg 500	Trp	Arg	Gly	Trp	Thr 505	Ser	Gln	Gly	Arg	Arg 510	Arg	Arg
Ile	Trp	Pro 515	Arg	Trp	Leu	Arg	Gly 520	Arg	Суз	Leu	Leu	Leu 525	Arg	Arg	Leu
Leu	Gly 530	Tyr	Val	Ile	Ser	Glu 535	Asn	Glu	Arg	Ser	Ala 540	Leu	Val		
<211 <212)> SH L> LH 2> TY 3> OH	ENGTH 7PE :	H: 50 PRT	56	e B 1	PWD (circo	oviru	15						
<400)> SI	EQUEI	ICE :	18											
Gln 1	Arg	Thr	Ser	Ala 5	Ala	Ala	Ala	Pro	Arg 10	Gln	His	Leu	Ser	Ser 15	Asn
Met	Pro	Ser	Lуз 20	ГЛа	Asn	Gly	Arg	Ser 25	Gly	Pro	Gln	Pro	His 30	ГЛа	Arg

-continued

												COIL		ucu	
Trp	Val	Phe 35	Thr	Leu	Asn	Asn	Pro 40	Ser	Glu	Asp	Glu	Arg 45	Lys	Lys	11
Arg	Asp 50	Leu	Pro	Ile	Ser	Leu 55	Phe	Asp	Tyr	Phe	Ile 60	Val	Gly	Glu	G
Gly 65	Asn	Glu	Glu	Gly	Arg 70	Thr	Pro	His	Leu	Gln 75	Gly	Phe	Ala	Asn	Ph 80
Val	Lys	Lys	Gln	Thr 85	Phe	Asn	Гла	Val	Lys 90	Trp	Tyr	Leu	Gly	Ala 95	Arg
СЛа	His	Ile	Glu 100	Lys	Ala	ГÀа	Gly	Thr 105		Gln	Gln	Asn	Lys 110	Glu	Tyr
Сув	Ser	Lys 115	Glu	Gly	Asn	Leu	Leu 120	Met	Glu	Сув	Gly	Ala 125	Pro	Arg	Ser
Gln	Gly 130	Gln	Arg	Ser	Asp	Leu 135	Ser	Thr	Ala	Val	Ser 140	Thr	Leu	Leu	Glu
Ser 145	Gly	Ser	Leu	Val	Thr 150	Val	Ala	Glu	Gln	His 155	Pro	Val	Thr	Phe	Val 160
Arg	Asn	Phe	Arg	Gly 165	Leu	Ala	Glu	Leu	Leu 170	Lys	Val	Ser	Gly	Lys 175	Met
Gln	Lys	Arg	Asp 180		Lys	Thr	Asn	Val 185	His	Val	Ile	Val	Gly 190	Pro	Pro
Gly	Cys	Gly 195	Lys	Ser	ГЛа	Trp	Ala 200		Asn	Phe	Ala	Asp 205		Glu	Thr
Thr	Tyr 210			Pro	Pro	Arg 215		Lys	Trp	Trp	Asp 220		Tyr	His	Gly
Glu 225	Glu	Val	Val	Val	Ile 230		Asp	Phe	Tyr	Gly 235		Leu	Pro	Trp	Asp 240
	Leu	Leu	Arg	Leu 245		Asp	Arg	Tyr	Pro 250		Thr	Val	Glu	Thr 255	
Gly	Gly	Thr	Val 260		Phe	Leu	Ala	Arg 265		Ile	Leu	Ile	Thr 270		Asn
Gln	Thr			Glu	Trp	Tyr			Thr	Ala	Val			Val	Glu
Ala	Leu	275 Tyr	Arg	Arg	Ile		280 Ser	Leu	Val	Phe		285 Lys	Asn	Ala	Thr
	290 Gln	Ser	Thr	Glu		295 Gly	Gly	Gln	Phe		300 Thr	Leu	Ser	Pro	
305 Сув	Pro	Glu	Phe	Pro	310 Tyr	Glu	Ile	Asn	Tyr	315 Val	Phe	Phe	Ile	Thr	320 Ser
Trp	Phe	Leu	Leu	325 Phe	Ile	Lys	Gly	Val	330 Gly	Gly	Leu	Ile	Val	335 His	Thr
Trp	Leu	His	340 Gly	Tyr	Суз	Ile	Pro	345 Gly	Arg	Ile	Tyr	Суз	350 Phe	Arg	Thr
	Cys	355					360					365			
	370 Thr	-	-		-	375					380				
385	Gln		-		390	-	-			395					400
				405					410					415	
_	Tyr	-	420	-				425	-	-		_	430		-
Pro	Leu	Leu	Gln	Ser	Tyr	His	Leu	ГÀа	Gln	His	Trp	Ser	Pro	Leu	Pro

-continued

-continued	
435 440 445	
Cys His Pro Gly Ser Gly Ser Arg Ala Arg Ile Gln Pro Pro Phe Leu 450 455 460	
Phe Cys Ser Ile Gln Arg Ala Gln Ser Gly Gly Leu Thr Pro Leu Leu	
465 470 475 480	
Gly Glu Glu Ser His Ile Ser Ser Cys Pro Pro Pro Arg Ala Phe 485 490 495	
Leu Trp Phe Ala Gln Tyr Ile Arg Arg Cys Gly Arg Gly Gly Cys Arg	
500 505 510	
Cys His Phe Ser Phe Ser Ser Gly Asn Gly Gly Gly Gly Arg Ala 515 520 525	
Arg Gly Gly Gly Gly Gly Ser Gly Gln Asp Gly Cys Gly Gly Gly Val	
530 535 540	
Phe Phe Gly Asn Ala Ser Leu Asp Thr Ser Tyr Leu Lys Thr Lys 545 550 555 560	
Glu Val Arq Cys Lys Tyr	
565	
<210> SEQ ID NO 19	
2215 LENGTH: 1767 2212 TYPE: DNA	
213> ORGANISM: Type B PWD circovirus	
400> SEQUENCE: 19	
atacttaca gogcacttot ttogttttoa gatatgaogt atocaaggag gogttacog	ga 60
agaagaagac accgcccccg cagccatctt ggccagatcc tccgccgccg cccctggct	tc 120
ytecaceee gecacegtta eegetggaga aggaaaaatg geatetteaa eaceegeet	tc 180
teeegeacet teggatatae tgteaagega aceaeagtea gaaegeeete etgggeggt	tg 240
gacatgatga gattcaatat taatgacttt cttcccccag gagggggggtc aaaccccc	gc 300
tetgtgeeet ttgaataeta cagaataaga aaggttaagg ttgaattetg geeetgete	cc 360
ccgatcaccc agggtgacag gggagtgggc tccagtgctg ttattttaga tgataactt	tt 420
gtaacaaagg ccacageeet caeetatgae eeetatgtaa aetaeteete eegeeatae	cc 480
ataacccagc cetteteeta ceacteeegg taetttaeee ecaaacetgt eetagattt	tc 540
actattgatt acttccaacc aaacaacaaa agaaaccagc tgtggctgag actacaaac	ct 600
gctggaaatg tagaccacgt aggcctcggc actgcgttcg aaaacagtat atacgacca	ag 660
gaatacaata teegtgtaae catgtatgta caatteagag aatttaattt taaagaeee	cc 720
ccacttaacc cttaatgaat aataaaaacc attacgaagt gataaaaaag actcagtaa	at 780
ttatttcata tggaaattca gggcatgggg gggaaagggt gacgaactgg cccccttco	ct 840
ccgtggattg ttctgtagca ttcttccaaa ataccaagga agtaatcctc cgataaaga	ag 900
cttctacagc tgggacagca gttgaggagt accattccaa cggggtctga ttgctggta	aa 960
tcagaatact gcgggccaaa aaaggtacag ttccaccttt agtctctaca gtcaatgga	at 1020
atcgatcaca cagteteagt agateateee agggeageea geeataaaag teateaata	aa 1080
caaccacttc ttcaccatgg taaccatccc accacttgtt tctaggtggt ttccagtat	tg 1140
tggtttccgg gtctgcaaaa ttagcagccc atttgctttt accacaccca ggtggcccc	ca 1200
caatgacgtg tacattagtc ttccaatcac gcttctgcat tttcccgctc actttcaaa	aa 1260

-continued	
yttcagccag cccgcggaaa tttctgacaa acgttacagg gtgctgctct gcaacggtca	1320
ccagacteee getetecaae aaggtactea cageagtaga caggteaete egttgteeet	1380
gagatetagg ageteeacae teeateagta agttgeette tttaetgeag tattetttat	1440
cetgetgate tgtteettte getttetega tgtggeageg ggeaceeaaa taceaettea	1500
ctttattaaa agtetgette tteacaaaat tagegaacee etggaggtga ggtgttegte	1560
ctteeteatt acceteeteg eeaacaataa aataateaaa tagggatatt ggaagateee	1620
gtattttett gegetegtet teggaaggat tatteagagt gaacaeeeae etttatggg	1680
gttggggtcc gcttcttcca ttcttcttgc tgggcatgtt gctgctgagg tgctgccgag	1740
gtgetgeege tgeegaagtg egetggt	1767
<210> SEQ ID NO 20 <211> LENGTH: 567 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 20	
Gly Ala Cys Lys Pro Leu Pro Leu Val Glu Ala Ala Gly Cys Cys	
$\begin{array}{cccccccccccccccccccccccccccccccccccc$	
Ala Trp Cys Ser Ser His Phe Phe Arg Val Gly Val Gly Tyr Phe Thr 20 25 30	
Pro Thr Glu Ser Tyr Asp Lys Arg Leu Arg Ala Cys Ser Phe Val Pro 35 40 45	
Asp Glu Leu Ile Gly Ile Gln Asn Asn Gln Gln Arg Pro Pro Tyr His 50 55 60	
Pro Leu Val Phe Val Glu Gly Gly Pro Thr Arg Asn Gln Ser Ser Ala 65 70 75 80	
Ser Lys Tyr Leu Ser Thr Thr Asn Pro His Gly Ser Gly Cys Arg Ser 85 90 95	
Leu Ser Leu Phe Leu Asp Ala Ser Tyr Leu Ile Ser Cys Tyr Leu Leu 100 105 110	
Cys Ser Val Ser Pro Thr His Leu Glu Ile Glu Pro Val Val Ser His 115 120 125	
Gly Thr Gln Gln Ser Tyr Arg Thr Pro Ser Arg Ser Asp Pro Ser Arg 130 135 140	
Gln Leu Ala Ala Gly Gln Leu Thr Gln Phe Asn Gly Arg Ala Pro Gln 145 150 155 160	
Val Lys Ser Leu Ser Arg Ser Phe Ala Ser Ala His Asn Ser Ser His 165 170 175	
Val Arg Gln Pro Ala Val Gln Thr His Tyr Phe Cys Ile Pro Gln Asn 180 185 190	
Gln Leu Gly Pro Phe Trp Met Ser Ser Val Val Phe Cys Thr Thr Pro 195 200 205	
His Asn Gly His His Leu Leu Pro Gln Gln His Ser Lys His Ser Ala 210 215 220	
Ala Arg Pro His Asp Val Ser Val Thr His Asp Ile Asp Met Ser Gln 225 230 235 240	
Leu Ser Leu His Phe Gln Val Lys Lys Pro Gly Cys Tyr Glu Ser Trp 245 250 255	

-cont	п	n	11	ρ	С

Leu	Leu	Glu 275	Гла	Asp	Ser	Ser	Lys 280	Arg	Pro	Ile	Гла	Ser 285	Ser	His	Leu
Val	Ile 290		Pro	Pro	Leu	Pro 295	Gly	Thr	Arg	Gly	Lys 300	Gly	Gly	Met	Gly
Gln 305	Ile	Glu	Met	His	Phe 310	Leu	Asn	Ser	Leu	Arg 315	Lys	Гла	Thr	Ile	Thr 320
Lys	Ile	Ile	Pro	Asn 325	Leu	Pro	Pro	Asp	Lys 330	Phe	Asn	Phe	Glu	Arg 335	Phe
Gln	Val	Tyr	Met 340	Thr	Val	Arg	Ile	Asn 345	Tyr	Glu	Gln	Asp	Tyr 350	Ile	Ser
Asn	Glu	Phe 355	Ala	Thr	Gly	Leu	Gly 360	Val	His	Asp	Val	Asn 365	Gly	Ala	Thr
Gln	Leu 370	Arg	Leu	Trp	Leu	Gln 375	Asn	Arg	Lys	Asn	Asn 380	Pro	Gln	Phe	Tyr
Asp 385	Ile	Thr	Phe	Asp	Leu 390	Val	Pro	Lys	Pro	Thr 395	Phe	Tyr	Arg	Ser	His 400
Tyr	Ser	Phe	Pro	Gln 405	Thr	Ile	Thr	His	Arg 410	Ser	Ser	Tyr	Asn	Val 415	Tyr
Pro	Asp	Tyr	Thr 420	Leu	Ala	Thr	Ala	Lys 425	Thr	Val	Pro	Asn	Asp 430	Asp	Leu
Ile	Val	Ala 435	Ser	Ser	Gly	Val	Gly 440	Arg	Asp	Gly	Gln	Thr 445	Ile	Pro	Ser
СЛа	Pro 450	Trp	Phe	Glu	Val	Lys 455	Val	Lys	Arg	Ile	Arg 460	Tyr	Tyr	Glu	Phe
Pro 465	Val	Ser	Arg	Pro	Asn 470	Ser	Gly	Gly	Gly	Pro 475	Pro	Leu	Phe	Aab	Asn 480
Ile	Asn	Phe	Arg	Met 485	Met	Asp	Val	Ala	Trp 490	Ser	Pro	Thr	Arg	Val 495	Thr
Thr	Arg	Lys	Val 500	Thr	Tyr	Gly	Phe	Thr 505	Arg	Ser	Leu	Arg	Thr 510	Asn	Phe
Ile	Gly	Asn 515	Lys	Arg	Arg	Trp	Arg 520	Tyr	Arg	His	Arg	Pro 525	His	Val	Leu
Trp	Pro 530	Arg	Arg	Arg	Leu	Ile 535	Gln	Gly	Leu	His	Ser 540	Arg	Pro	Arg	His
Arg 545	Arg	Arg	Arg	Tyr	Arg 550	Arg	Arg	Pro	Tyr	Thr 555	Met	Asp	Ser	Phe	Ser 560
Leu	Leu	Ala		Tyr 565		Asn									
<213 <212	D> SH L> LH 2> TY 3> OH	ENGTH ZPE :	H: 50 PRT		∋ B I	PWD (circo	oviru	15						
<400)> SH	EQUEI	ICE :	21											
Trp 1	Arg	Val	Glu	Ala 5	Ala	Ala	Ala	Gly	Arg 10	Суз	Суз	Arg	Leu	Leu 15	Leu
Met	Gly	Leu	Leu 20	Phe	Phe	Pro	Leu	Leu 25	Pro	Gly	Trp	Gly	Trp 30	Leu	Leu
His	Thr	Asn 35	Val	Arg	Phe	Leu	Gly 40	Glu	Ser	Ser	Ser	Arg 45	Leu	Phe	Ile
Arg	Ser	Arg	Gly	Ile	Asp	Arg	Asn	Ser	Lys	Ile	Thr	Pro	Ser	Ser	Pro

-continued

											-	con	tin	ued	
	50					55					60				
Leu 65	Ser	Ser	Pro	Arg	Val 70	Gly	Arg	Trp	Pro	Asn 75	Ala	Leu	Lys	Thr	Phe 80
Phe	Cya	Val	Lys	Leu 85	Leu	Thr	Phe	His	Tyr 90	ГЛа	Pro	Ala	Arg	Gln 95	Trp
Met	Ser	Phe	Ala 100	Phe	Pro	Val	Ser	Cys 105		Leu	Ser	Tyr	Gln 110	Leu	Leu
Ser	Pro	Leu 115	Lys	Ser	Ile	Ser	His 120	Pro	Ala	Gly	Leu	Asp 125	Pro	Cys	Arg
Leu	Ser 130	Arg	Asp	Val	Ala	Thr 135	Leu	Val	Lys	Asn	Ser 140	Leu	Pro	Leu	Arg
Thr 145	Val	Thr	Ala	Ser	Cys 150	Суз	Gly	Thr	Val	Asn 155	Thr	Leu	Phe	Lys	Arg 160
Pro	Ser	Ala	Ser	Ser 165	-	Phe	Thr	Leu	Pro 170	Phe	Ile	Суз	Phe	Arg 175	Ser
Gln	Phe	Val	Leu 180	Thr	Сүв	Thr	Met	Thr 185	Pro	Gly	Gly	Pro	His 190	Pro	Leu
Leu	Leu	His 195	Ala	Ala	Leu	Гла	Ala 200	Ser	Gly	Ser	Val	Val 205	Tyr	Gln	Phe
Gly	Gly 210		Phe	Leu	His	His 215	Ser	Pro	Trp	Pro	Ser 220	Ser	Thr	Thr	Thr
Ile 225	Ser	Ser	Lys	Pro	Gln 230	Ser	Gly	Gln	Ser	Ser 235	Arg	Ser	Leu	Ser	His 240
Ser	Arg	Tyr	Gly	Asn 245	Val	Thr	Ser	Val	Leu 250	Pro	Pro	Val	Thr	Gly 255	Lys
Lys	Ala	Arg	Leu 260	Ile	Arg	Ile	Val	Leu 265	Leu	Val	Gly	Asn	Ser 270	His	Tyr
Glu	Glu	Val 275	Ala	Thr	Gly	Ala	Thr 280	Ser	Ala	Arg	Arg	Leu 285	Ile	Val	Glu
Lys	Thr 290	Asn	Gln	Phe	Phe	Ala 295	Val	Ser	Сув	Asp	Val 300	Ser	Ser	Pro	Pro
Trp 305	Asn	Thr	Val	Arg	Glu 310	Gly	Gly	His	Gly	Ser 315	Asn	Gly	Tyr	Ser	Ile 320
Phe	Gln	Thr	Lys	Lys 325	Ile	Val	Glu	Tyr	His 330	Asn	Lys	Asn	Asn	Met 335	Leu
Pro	Thr		Pro 340		Phe		Arg			Thr	Суз		His 350		Cys
Pro	Tyr	Gln 355	Ile	Gly	Pro	Arg	Ile 360	Tyr	Gln	Lys	Arg	Val 365	Сув	His	Arg
Pro	Arg 370	Arg	Pro	Arg	Суз	Lys 375	Trp	Суз	Asn	Thr	Thr 380	Glu	Ala	Val	Ala
Pro 385	Lys	Lys	Gln	Gln	Lys 390	Thr	Pro	Leu	Leu	Tyr 395	His	Phe	Arg	Pro	Cys 400
Thr	Gln	Pro	Tyr	Leu 405	Val	Pro	Leu	Pro	Leu 410	Leu	Leu	Ala	Pro	Asn 415	His
Tyr	Pro	Pro	Leu 420	Leu	Leu	ГЛа	Суз	Leu 425	Pro	Leu	His	Pro	Ser 430	His	Gly
Lys	Asn	Cys 435	Leu	Arg	Phe	Tyr	Cys 440	Суз	Gln	Leu	Gly	Ser 445	Gly	Gln	Gly

Ile 465	Leu	Ala	Сүз	Leu	Pro 470	Pro	Lys	Val	Gly	Arg 475	Arg	Pro	Ser	Ser	Leu 480
Tyr	Gln	Ile	Glu	Asp 485	His	Gly	Gly	Gly	Leu 490	Leu	Ala	Asn	Gln	Ser 495	His
Asn	Ala	Gln	Сув 500	Tyr	Ile	Arg	Leu	His 505	Pro	Leu	Pro	Pro	His 510	Gln	Leu
His	Trp	Lys 515	Glu	Lys	Glu	Leu	Pro 520	Leu	Pro	Pro	Pro	Pro 525	Pro	Arg	Ala
Leu	Pro 530	Pro	Pro	Pro	Pro	Asp 535	Pro	Trp	Ser	Pro	Gln 540	Pro	Pro	Pro	Thr
Lys 545	Lys	Lys	Pro	Leu	Ala 550	Glu	Lys	Ser	Val	Asp 555	Tyr	Arg	Phe	Val	Phe 560
Ser	Thr	Arg	Gln	Leu 565	Tyr										
<21: <21: <21:	L> LI 2> TY 3> OF		H: 50 PRT ISM:	59 Туре	∋ B I	PWD (circo	oviru	18						
		EQUEI													
Leu 1	Ala	Ser	Arg	Суя 5	Arg	Суз	Суз	Arg	Pro 10	Leu	Val	Glu	Ala	Ala 15	Val
His	Gly	Ala	Leu 20	Leu	Ile	Ser	Ser	Ala 25	Ser	Gly	Leu	Gly	Met 30	Phe	Pro
Pro	His	Glu 35	Ser	Gln	Ile	Ile	Arg 40	Gly	Phe	Val	Leu	Ala 45	Leu	Phe	Tyr
Pro	Ile 50	Lys	Trp	Tyr	Gly	Lys 55	Ile	Ile	Lys	Asn	Asn 60	Ala	Leu	Leu	Thr
Ile 65	Leu	Phe	Ser	Ser	Суз 70	Arg	Val	Glu	Leu	Pro 75	Glu	Ser	Ile	Lys	His 80
Leu	Leu	Leu	Ser	Lys 85	Ile	Phe	His	Leu	Pro 90	Ile	Gln	Thr	Gly	Ala 95	Ala
Val	Asp	Leu	Phe 100	Arg	Phe	Ser	Сув	Ile 105	Leu	Leu	Ile	Phe	Phe 110	Val	Ala
Thr	Phe	Phe 115	Ala	Val	Gln	His	Leu 120	Thr	Ser	Ser	Arg	Ser 125	Arg	Leu	Ser
Leu	Pro 130	Thr			-	Ser 135		His		Gly		Gln	Leu	Ala	Pro
Thr 145	Gln	His	Gly	Asn	Cys 150	Leu	Leu	Val	Arg	Tyr 155	Arg	Lys	Asp	Ser	Ile 160
Glu	Ala	Pro	Gln	Ser 165	Phe	ГЛа	Gln	Phe	His 170	Ala	Pro	Phe	His	Leu 175	Leu
Thr	Ile	Pro	Leu 180	Ser	Ile	Tyr	Val	Asp 185	Asn	His	Pro	Trp	Arg 190	Pro	Thr
Thr	Phe	Ala 195	Phe	Pro	Ser	Ser	Ile 200	Lys	Суз	Val	Arg	Phe 205	Gly	Суз	Val
Pro	Phe 210	Trp	Arg	Ser	Val	Leu 215	Pro	Pro	Ile	Thr	Val 220	Met	Thr	Phe	Phe
His 225	Asn	Asn	Asn	Ile	Val 230	Гла	Ile	Ala	Pro	Gln 235	Gly	Pro	Ile	Ile	Gln 240
Ser	Gln	Thr	Ser	Ser	Ile	Trp	Gln	Ser	Tyr	Leu	Ser	Phe	Thr	Ser	Ser

-continued

											_	COII	tın	ueu		
				245					250					255		
Tyr	Arg	Lys	Gln 260		Ala	Thr	Asn	Gln 265	Asn	Gly	Ala	Ile	Leu 270	Gly	Arg	
Gln	Phe	Pro 275	Val	Gly	Ser	Ser	Asp 280		Ser	Tyr	Phe	Ser 285	Lys	Ile	Pro	
Pro	Asn 290	Ser	Gly	Gln	Tyr	Lys 295		Leu	Ile	Ser	Суя 300	Phe	Leu	Gly	Arg	
Leu 305		Pro	Ala	Leu	Glu 310	Asp	Gly	Lys	Gly	Gly 315	Trp	Ala	Arg	Phe	Lys 320	
Trp	Ile	Phe	Trp	Ile 325	Val	Ser	Asp	Lys	Lys 330	Asp	Ser	Arg	Leu	Pro 335	Lys	
Glu	Asn	Leu	Thr 340	Leu	His	Pro	Thr	Lys 345	Leu	Ile	Leu	Asn	Glu 350	Ser	Asn	
Tyr	Met	Сув 355	Pro	Val	Ser	Ile	Thr 360	Asn	Arg	Thr	Thr	Tyr 365	Val	Thr	Lys	
Ser	Arg 370		Ala	Ser	Ala	Thr 375	Thr	Met	Glu	Leu	Leu 380	Lys	Tyr	Asp	Gly	
Суз 385		Thr	Glu	ГЛа	Thr 390	Thr	Gln	Asn	Ser	Thr 395	Ile	Leu	Leu	Ser	Ile 400	
Ser	Leu	Asn	Pro	Pro 405	Leu	Thr	Gly	Pro	Thr 410	Thr	Pro	Ser	Pro	Ser 415	Pro	
Pro	Ile	Ala	Pro 420	Pro	Thr	Thr	Met	Pro 425	Thr	Met	Pro	Ser	Pro 430	Gln	Pro	
Arg	Gln	Leu 435	Thr	Ile	Met	Phe	Leu 440	Leu	Val	Pro	Ala	Trp 445	Glu	Gly	Thr	
Val	Arg 450		Ser	Arg	Pro	Ala 455	Pro	Gly	Ser	Asn	Leu 460	Arg	Leu	Arg	Glu	
Glu 465		Thr	Asn	Leu	Pro 470	-	Leu	Ala	Pro	Thr 475	Gln	Gly	Gly	Glu	Gln 480	
Pro	Phe	Phe	Thr	Met 485	Leu	Ile	Ser	Asp	Thr 490	Trp	Arg	Gly	Pro	Pro 495	Arg	
Glu	Ser	Gln	Pro 500	Glu	Ser	Ser	Leu	Ile 505	Asp	Ser	Pro	Ala	Pro 510	Ser	Ala	
Pro	Thr	Ser 515	Ser	Ala	Met	Lys	Gly 520	Glu	Gly	Ala	Thr	Val 525	Thr	Ala	Pro	
Thr	Ser 530				Ala				Ser	-			Ile	Ala	Ala	
Pro 545	Ala	Thr	Asp	Glu	Glu 550	Glu	Thr	Val	Gly	Gly 555	Gln	Ile	Arg	Ile	Gln 560	
Phe	Arg	Phe	Phe	His 565	Ala	Thr	Leu	Ile								
<21 <21 <22 <22 <22 <22	1 > L; 2 > T'; 3 > O; 0 > F; 1 > N; 2 > L()	EATUH AME/H	H: 9 DNA ISM: RE: RE: KEY: ION:	45 Type CDS (1)	e B 1		circ	ovir	15							
					aat Asn											48

-continued

	gtg Val			-					-	-		-	-			96
	gat Asp															144
	aat Asn 50															192
	aag Lys															240
	cac His		~ ~				~ ~							<u> </u>		288
-	agt Ser		-				-	-		-		-		-		336
-	gga Gly			-	-	-			-		-		-	-		384
	999 Gly 130															432
	aat Asn															480
-	aag Lys	-	-		-			-		-						528
	tgt Cys															576
	tac Tyr															624
	gaa Glu 210															672
	cta Leu					Asp		Tyr	Pro		Thr					720
	gga Gly		-			-	-	-	-		-			-		768
	acc Thr	<u> </u>		<u> </u>						<u> </u>			<u> </u>	<u> </u>	<u> </u>	816
-	ctt Leu							-	-			-		-		864
	caa Gln 290															912
	cct Pro									tga						945

<210> SEQ ID NO 24 <211> LENGTH: 314 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 24 Met Pro Ser Lys Lys Asn Gly Arg Ser Gly Pro Gln Pro His Lys Arg 5 10 15 1 Trp Val Phe Thr Leu Asn Asn Pro Ser Glu Asp Glu Arg Lys Lys Ile 25 20 Arg Asp Leu Pro Ile Ser Leu Phe Asp Tyr Phe Ile Val Gly Glu Glu 40 35 Gly Asn Glu Glu Gly Arg Thr Pro His Leu Gln Gly Phe Ala Asn Phe - - 55 50 60 Val Lys Lys Gln Thr Phe Asn Lys Val Lys Trp Tyr Leu Gly Ala Arg 75 70 Cys His Ile Glu Lys Ala Lys Gly Thr Asp Gln Gln Asn Lys Glu Tyr 90 Cys Ser Lys Glu Gly Asn Leu Leu Met Glu Cys Gly Ala Pro Arg Ser 100 105 110 Gln Gly Gln Arg Ser Asp Leu Ser Thr Ala Val Ser Thr Leu Leu Glu 120 125 115 Ser Gly Ser Leu Val Thr Val Ala Glu Gln His Pro Val Thr Phe Val 130 135 140 Arg Asn Phe Arg Gly Leu Ala Glu Leu Leu Lys Val Ser Gly Lys Met145150155160 Gln Lys Arg Asp Trp Lys Thr Asn Val His Val Ile Val Gly Pro Pro 170 165 175 Gly Cys Gly Lys Ser Lys Trp Ala Ala Asn Phe Ala Asp Pro Glu Thr 180 185 190 Thr Tyr Trp Lys Pro Pro Arg Asn Lys Trp Trp Asp Gly Tyr His Gly 195 200 205 Glu Glu Val Val Val Ile Asp Asp Phe Tyr Gly Trp Leu Pro Trp Asp 210 215 220 Asp Leu Leu Arg Leu Cys Asp Arg Tyr Pro Leu Thr Val Glu Thr Lys 225 230 235 240 Gly Gly Thr Val Pro Phe Leu Ala Arg Ser Ile Leu Ile Thr Ser Asn 255 245 250 Gln Thr Pro Leu Glu Trp Tyr Ser Ser Thr Ala Val Pro Ala Val Glu 265 260 270 Ala Leu Tyr Arg Arg Ile Thr Ser Leu Val Phe Trp Lys Asn Ala Thr 275 280 285 Glu Gln Ser Thr Glu Glu Gly Gly Gln Phe Val Thr Leu Ser Pro Pro 290 295 300 Cys Pro Glu Phe Pro Tyr Glu Ile Asn Tyr 310 305 <210> SEQ ID NO 25 <211> LENGTH: 702 <212> TYPE: DNA <213> ORGANISM: Type B PWD circovirus <220> FEATURE: <221> NAME/KEY: CDS

								-
-	С	01	nt	1	n	u	е	a

												con	tin	ued			 	
<222>	LO	CATI	ON :	(1)	(6	99)												
<400>	SE	QUEI	ICE :	25														
atg ac Met Th 1																48		
agc ca Ser Hi				-			-	-	-				-			96		
cgc ca Arg Hi	is															144		
ctc to Leu Se 5(er															192		
ccc to Pro Se 65																240		
ccc co Pro Pi								-					-			288		
aga at Arg Il																336		
cag gg Gln Gl	ly															384		
ttt gt Phe Va 13			-	-		-				-			-			432		
tcc to Ser Se 145																480		
ttt ac Phe Th																528		
aac aa Asn As																576		
gta ga Val As	ab		~					~ ~				· ·				624		
cag ga Gln Gl 21																672		
aat tt Asn Pf 225			-						taa							702		
<210> <211> <212> <213>	LE TY	NGTH PE :	I: 23 PRT	33	e B :	PWD -	circ	ovir	18									
<400>	SE	QUEI	ICE :	26														
Met Tł 1	hr	Tyr	Pro	Arg 5	Arg	Arg	Tyr	Arg	Arg 10	Arg	Arg	His	Arg	Pro 15	Arg			
Ser Hi	is	Leu	Gly	Gln	Ile	Leu	Arg	Arg	Arg	Pro	Trp	Leu	Val	His	Pro			

											-	con	tin	ued	
			20					25					30		
Arg	His	Arg 35	Tyr	Arg	Trp	Arg	Arg 40	ГЛа	Asn	Gly	Ile	Phe 45	Asn	Thr	Arg
Leu	Ser 50	Arg	Thr	Phe	Gly	Tyr 55	Thr	Val	Lys	Arg	Thr 60	Thr	Val	Arg	Thr
Pro 65	Ser	Trp	Ala	Val	Asp 70	Met	Met	Arg	Phe	Asn 75	Ile	Asn	Asp	Phe	Leu 80
Pro	Pro	Gly	Gly	Gly 85	Ser	Asn	Pro	Arg	Ser 90	Val	Pro	Phe	Glu	Tyr 95	Tyr
Arg	Ile	Arg	Lys 100	Val	Lys	Val	Glu	Phe 105	Trp	Pro	Суа	Ser	Pro 110	Ile	Thr
Gln	Gly	Asp 115		Gly	Val	Gly	Ser 120		Ala	Val	Ile	Leu 125		Asp	Asn
Phe			Lys	Ala	Thr			Thr	Tyr	Asp			Val	Asn	Tyr
	130 Ser	Arg	His	Thr	Ile	135 Thr	Gln	Pro	Phe		140 Tyr	His	Ser	Arg	Tyr
145 Phe	Thr	Pro	Lys	Pro	150 Val	Leu	Asp	Phe	Thr	155 Ile	Asp	Tyr	Phe	Gln	160 Pro
Asn	Asn	Lvs	- Ara	165 Asn	Gln	Leu	- Trp	Leu	170 Arg	Leu	- Gln	- Thr	Ala	175 Glv	Asn
			180					185					190		
Val	Asb	His 195	Val	Gly	Leu	Gly	Thr 200	Ala	Phe	Glu	Asn	Ser 205	Ile	Tyr	Asp
Gln	Glu 210	Tyr	Asn	Ile	Arg	Val 215	Thr	Met	Tyr	Val	Gln 220	Phe	Arg	Glu	Phe
Asn 225	Phe	Lys	Asp	Pro	Pro 230	Leu	Asn	Pro							
		EQ II													
<212	2 > T?	ENGTI ZPE : RGANI	DNA		e B :	PWD -	circ	oviru	15						
<220 <223)> FI L> N2	EATUI AME/I	RE: KEY:	CDS											
		EQUE		(1) 27	(3										
				cca Pro 5											
ttc				aaa Lys					ttt					ccc	
	-		20	-				25					30		-
				gac Asp											
				ttt Phe											
	-			gtg Val	-		-				-			-	
				cac His 85											

```
-continued
```

315

tet agg age tee aca ete cat cag taa Ser Arg Ser Ser Thr Leu His Gln 100 <210> SEQ ID NO 28 <211> LENGTH: 104 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 28 Met Val Thr Ile Pro Pro Leu Val Ser Arg Trp Phe Pro Val Cys Gly 1 5 10 15 Phe Arg Val Cys Lys Ile Ser Ser Pro Phe Ala Phe Thr Thr Pro Arg 25 20 30 Trp Pro His Asn Asp Val Tyr Ile Ser Leu Pro Ile Thr Leu Leu His 40 35 45 Phe Pro Ala His Phe Gln Lys Phe Ser Gln Pro Ala Glu Ile Ser Asp 55 50 60 Lys Arg Tyr Arg Val Leu Leu Cys Asn Gly His Gln Thr Pro Ala Leu 70 65 75 80 Gln Gln Gly Thr His Ser Ser Arg Gln Val Thr Pro Leu Ser Leu Arg 85 90 95 Ser Arg Ser Ser Thr Leu His Gln 100 <210> SEQ ID NO 29 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 29 Val Asp Met Met Arg Phe Asn Ile Asn Asp Phe Leu Pro Pro Gly 5 1 10 15 <210> SEQ ID NO 30 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 30 Gln Gly Asp Arg Gly Val Gly Ser Ser Ala Val Ile Leu Asp Asp 1 5 10 15 <210> SEQ ID NO 31 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 31 Gly Val Gly Ser Ser Ala Val Ile Leu Asp Asp Asn Phe Val Thr 1 5 10 15 <210> SEQ ID NO 32 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 32 Val Asp His Val Gly Leu Gly Thr Ala Phe Glu Asn Ser Ile Tyr 5 10 1 15

<210> SEQ ID NO 33 <211> LENGTH: 8 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 33 8 tgtggcga <210> SEQ ID NO 34 <211> LENGTH: 8 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 34 agtttcct 8 <210> SEQ ID NO 35 <211> LENGTH: 20 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 35 tcatttagag ggtctttcag 20 <210> SEQ ID NO 36 <211> LENGTH: 8 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 36 gtcaacct 8 <210> SEQ ID NO 37 <211> LENGTH: 8 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 37 gtggttgc 8 <210> SEQ ID NO 38 <211> LENGTH: 8 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 38 agcccagg 8 <210> SEQ ID NO 39 <211> LENGTH: 8 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 39 ttggctgg 8 <210> SEQ ID NO 40 <211> LENGTH: 12 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus

-continued

	-concinued
<400> SEQUENCE: 40	
tetagetetg gt	12
<210> SEQ ID NO 41 <211> LENGTH: 12 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus	
<400> SEQUENCE: 41	
ateteagete gt	12
<210> SEQ ID NO 42 <211> LENGTH: 12 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus	
<400> SEQUENCE: 42	
tgtcctcctc tt	12
<210> SEQ ID NO 43 <211> LENGTH: 8 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus	
<400> SEQUENCE: 43	
tctctaga	8
<210> SEQ ID NO 44 <211> LENGTH: 8 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus	
<400> SEQUENCE: 44	
tgtaccaa	8
<210> SEQ ID NO 45 <211> LENGTH: 8 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus	
<400> SEQUENCE: 45 tccgtctt	8
<210> SEQ ID NO 46 <211> LENGTH: 20 <212> TYPE: DNA <213> ORGANISM: Artificial Sequence <220> FEATURE: <223> OTHER INFORMATION: Primer	
<400> SEQUENCE: 46	
gtgtgctcga cattggtgtg	20
<210> SEQ ID NO 47 <211> LENGTH: 20 <212> TYPE: DNA <213> ORGANISM: Artificial Sequence <220> FEATURE: <223> OTHER INFORMATION: Primer <400> SEQUENCE: 47	

-	C	$\neg n$	t 1	n	ue	\sim
	0		L 7		uC	сı.

tggaatgtta acgagctgag	20
<210> SEQ ID NO 48 <211> LENGTH: 20 <212> TYPE: DNA <213> ORGANISM: Artificial Sequence <220> FEATURE: <223> OTHER INFORMATION: Primer	
<400> SEQUENCE: 48	
ctcgcagcca tcttggaatg	20
<pre><210> SEQ ID NO 49 <211> LENGTH: 20 <212> TYPE: DNA <213> ORGANISM: Artificial Sequence <220> FEATURE: <223> OTHER INFORMATION: Primer</pre>	
<400> SEQUENCE: 49	
cgcgcgtaat acgactcact	20
<210> SEQ ID NO 50 <211> LENGTH: 26 <212> TYPE: DNA <213> ORGANISM: Artificial Sequence <220> FEATURE: <223> OTHER INFORMATION: Primer	
<400> SEQUENCE: 50	
cctgtctact gctgtgagta ccttgt	26
<210> SEQ ID NO 51 <211> LENGTH: 26 <212> TYPE: DNA <213> ORGANISM: Artificial Sequence <220> FEATURE: <223> OTHER INFORMATION: Primer	
<400> SEQUENCE: 51	
gcagtagaca ggtcactccg ttgtcc	26
<210> SEQ ID NO 52 <211> LENGTH: 20 <212> TYPE: DNA <213> ORGANISM: Artificial Sequence <220> FEATURE: <223> OTHER INFORMATION: Primer	
<400> SEQUENCE: 52	
tggaatgtta actacctcaa	20
<210> SEQ ID NO 53 <211> LENGTH: 23 <212> TYPE: DNA <213> ORGANISM: Artificial Sequence <220> FEATURE: <223> OTHER INFORMATION: Primer	
<400> SEQUENCE: 53	
ggcggcgcca tctgtaacgg ttt	23

```
-continued
```

<210> SEQ ID NO 54 <211> LENGTH: 23 <212> TYPE: DNA <213> ORGANISM: Artificial Sequence <220> FEATURE: <223> OTHER INFORMATION: Primer <400> SEQUENCE: 54 23 gatggcgccg aaagacgggt atc <210> SEQ ID NO 55 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 55 Asn Val Asn Glu Leu Arg Phe Asn Ile Gly Gln Phe Leu Pro Pro 5 10 15 1 <210> SEQ ID NO 56 <211> LENGTH: 14 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 56 Thr Ser Asn Gln Arg Gly Val Gly Ser Thr Val Val Ile Leu 1 5 10 <210> SEQ ID NO 57 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 57 Arg Gly Val Gly Ser Thr Val Val Ile Leu Asp Ala Asn Phe Val 5 10 15 1 <210> SEQ ID NO 58 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 58 Phe Thr Ile Asp Tyr Phe Gln Pro Asn Asn Lys Arg Asn Gln Leu 1 5 10 15 <210> SEQ ID NO 59 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 59 Asp Gln Thr Ile Asp Trp Phe Gln Pro Asn Asn Lys Arg Asn Gln 1 5 10 15 <210> SEQ ID NO 60 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 60 Asn Val Glu His Thr Gly Leu Gly Tyr Ala Leu Gln Asn Ala Thr 5 10 15 1

68

<210> SEQ ID NO 61 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 61 His Arg Pro Arg Ser His Leu Gly Gln Ile Leu Arg Arg Arg Pro 15 1 5 10 <210> SEQ ID NO 62 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 62 Ser His Leu Gly Gln Ile Leu Arg Arg Arg Pro Trp Leu Val His 1 5 10 15 <210> SEQ ID NO 63 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 63 Gln Ile Leu Arg Arg Arg Pro Trp Leu Val His Pro Arg His Arg 1 5 10 15 <210> SEQ ID NO 64 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 64 Arg Arg Pro Trp Leu Val His Pro Arg His Arg Tyr Arg Trp Arg 10 1 5 15 <210> SEQ ID NO 65 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEOUENCE: 65 Leu Val His Pro Arg His Arg Tyr Arg Trp Arg Arg Lys Asn Gly 1 5 10 15 <210> SEQ ID NO 66 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 66 Arg His Arg Tyr Arg Trp Arg Arg Lys Asn Gly Ile Phe Asn Thr 1 5 10 15 <210> SEQ ID NO 67 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 67 Arg Trp Arg Arg Lys Asn Gly Ile Phe Asn Thr Arg Leu Ser Arg

-continued 1 5 10 15 <210> SEQ ID NO 68 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEOUENCE: 68 Lys Asn Gly Ile Phe Asn Thr Arg Leu Ser Arg Thr Phe Gly Tyr 1 5 10 15 <210> SEQ ID NO 69 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 69 Phe Asn Thr Arg Leu Ser Arg Thr Phe Gly Tyr Thr Val Lys Arg 1 5 10 15 <210> SEQ ID NO 70 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 70 Leu Ser Arg Thr Phe Gly Tyr Thr Val Lys Arg Thr Thr Val Arg 5 10 15 1 <210> SEQ ID NO 71 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 71 Phe Gly Tyr Thr Val Lys Arg Thr Thr Val Arg Thr Pro Ser Trp 1 5 10 15 <210> SEQ ID NO 72 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 72 Val Lys Arg Thr Thr Val Arg Thr Pro Ser Trp Ala Val Asp Met 1 5 10 15 <210> SEQ ID NO 73 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 73 Thr Val Arg Thr Pro Ser Trp Ala Val Asp Met Met Arg Phe Asn 5 10 1 15 <210> SEQ ID NO 74 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 74

70

Pro Ser Trp Ala Val Asp Met Met Arg Phe Asn Ile Asn Asp Phe 5 1 10 15 <210> SEQ ID NO 75 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEOUENCE: 75 Arg Phe As
n Ile As
n Asp Phe Leu Pro \mbox{Pro} Gly Gly S
er Asn 1 5 10 15 <210> SEQ ID NO 76 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 76 Asn Asp Phe Leu Pro Pro Gly Gly Gly Ser Asn Pro Arg Ser Val 1 5 10 15 <210> SEQ ID NO 77 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 77 Pro Pro Gly Gly Gly Ser Asn Pro Arg Ser Val Pro Phe Glu Tyr 5 10 1 15 <210> SEQ ID NO 78 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 78 Gly Ser Asn Pro Arg Ser Val Pro Phe Glu Tyr Tyr Arg Ile Arg 5 10 1 15 <210> SEQ ID NO 79 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEOUENCE: 79 Arg Ser Val Pro Phe Glu Tyr Tyr Arg Ile Arg Lys Val Lys Val 1 5 10 15 <210> SEQ ID NO 80 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 80 Phe Glu Tyr Tyr Arg Ile Arg Lys Val Lys Val Glu Phe Trp Pro 5 10 1 15 <210> SEQ ID NO 81 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 81

Arg Ile Arg Lys Val Lys Val Glu Phe Trp Pro Cys Ser Pro Ile 1 5 10 15 <210> SEQ ID NO 82 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 82 Val Lys Val Glu Phe Trp Pro Cys Ser Pro Ile Thr Gln Gly Asp 1 5 10 15 <210> SEQ ID NO 83 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 83 Phe Trp Pro Cys Ser Pro Ile Thr Gln Gly Asp Arg Gly Val Gly 5 10 1 15 <210> SEQ ID NO 84 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 84 Thr Arg Pro Arg Ser His Leu Gly Asn Ile Leu Arg Arg Arg Pro 5 10 15 1 <210> SEQ ID NO 85 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEOUENCE: 85 Ser His Leu Gly Asn Ile Leu Arg Arg Arg Pro Tyr Leu Val His 15 1 5 10 <210> SEQ ID NO 86 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 86 Asn Ile Leu Arg Arg Arg Pro Tyr Leu Val His Pro Ala Phe Arg 1 5 10 15 <210> SEQ ID NO 87 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 87 Arg Arg Pro Tyr Leu Val His Pro Ala Phe Arg Asn Arg Tyr Arg 5 10 15 1 <210> SEQ ID NO 88 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus

<400> SEQUENCE: 88 Leu Val His Pro Ala Phe Arg Asn Arg Tyr Arg Trp Arg Arg Lys 1 5 10 15 <210> SEQ ID NO 89 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 89 Ala Phe Arg Asn Arg Tyr Arg Trp Arg Arg Lys Thr Gly Ile Phe 1 5 10 15 <210> SEQ ID NO 90 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 90 Arg Tyr Arg Trp Arg Arg Lys Thr Gly Ile Phe Asn Ser Arg Leu 5 10 1 15 <210> SEQ ID NO 91 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 91 Arg Arg Lys Thr Gly Ile Phe Asn Ser Arg Leu Ser Arg Glu Phe 1 5 10 15 <210> SEQ ID NO 92 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 92 Gly Ile Phe Asn Ser Arg Leu Ser Arg Glu Phe Val Leu Thr Ile 5 1 10 15 <210> SEQ ID NO 93 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 93 Ser Arg Leu Ser Arg Glu Phe Val Leu Thr Ile Arg Gly Gly His 1 5 10 15 <210> SEQ ID NO 94 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 94 Arg Glu Phe Val Leu Thr Ile Arg Gly Gly His Ser Gln Pro Ser 1 5 10 15 <210> SEQ ID NO 95 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus

<400> SEOUENCE: 95 Leu Thr Ile Arg Gly Gly His Ser Gln Pro Ser Trp Asn Val Asn 1 5 10 15 <210> SEQ ID NO 96 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 96 Gly Gly His Ser Gln Pro Ser Trp Asn Val Asn Glu Leu Arg Phe 1 5 10 15 <210> SEQ ID NO 97 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 97 Gln Pro Ser Trp Asn Val Asn Glu Leu Arg Phe Asn Ile Gly Gln 1 5 10 15 <210> SEQ ID NO 98 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 98 Asn Val Asn Glu Leu Arg Phe Asn Ile Gly Gln Phe Leu Pro Pro 5 10 1 15 <210> SEQ ID NO 99 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 99 Leu Arg Phe Asn Ile Gly Gln Phe Leu Pro Pro Ser Gly Gly Thr 1 5 10 15 <210> SEQ ID NO 100 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 100 Ile Gly Gln Phe Leu Pro Pro Ser Gly Gly Thr Asn Pro Leu Pro 15 1 5 10 <210> SEQ ID NO 101 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 101 Leu Pro Pro Ser Gly Gly Thr Asn Pro Leu Pro Leu Pro Phe Gln 1 5 10 15 <210> SEQ ID NO 102 <211> LENGTH: 15 <212> TYPE: PRT

```
-continued
```

<213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 102 Gly Gly Thr Asn Pro Leu Pro Leu Pro Phe Gln Tyr Tyr Arg Ile 10 1 5 15 <210> SEQ ID NO 103 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 103 Pro Leu Pro Leu Pro Phe Gln Tyr Tyr Arg Ile Arg Lys Ala Lys 1 5 10 15 <210> SEQ ID NO 104 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 104 Pro Phe Gln Tyr Tyr Arg Ile Arg Lys Ala Lys Tyr Glu Phe Tyr 10 1 5 15 <210> SEQ ID NO 105 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 105 Tyr Arg Ile Arg Lys Ala Lys Tyr Glu Phe Tyr Pro Arg Asp Pro 1 5 10 15 <210> SEQ ID NO 106 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 106 Lys Ala Lys Tyr Glu Phe Tyr Pro Arg Asp Pro Ile Thr Ser Asn 1 5 10 15 <210> SEQ ID NO 107 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 107 Glu Phe Tyr Pro Arg Asp Pro Ile Thr Ser Asn Gln Arg Gly Val 1 5 10 15 <210> SEQ ID NO 108 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 108 Arg Asp Pro Ile Thr Ser Asn Gln Arg Gly Val Gly Ser Thr Val 1 5 10 15 <210> SEQ ID NO 109 <211> LENGTH: 15

```
-continued
```

<212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 109 Thr Ser Asn Gln Arg Gly Val Gly Ser Thr Val Val Ile Leu Asp 1 5 10 15 <210> SEQ ID NO 110 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 110 Gly Val Gly Ser Ser Ala Val Ile Leu Asp Asp Asn Phe Val Thr 5 10 15 1 <210> SEQ ID NO 111 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 111 Ser Ala Val Ile Leu Asp Asp Asn Phe Val Thr Lys Ala Thr Ala 1 5 10 15 <210> SEQ ID NO 112 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 112 Leu Asp Asp Asn Phe Val Thr Lys Ala Thr Ala Leu Thr Tyr Asp 1 5 10 15 <210> SEQ ID NO 113 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 113 Phe Val Thr Lys Ala Thr Ala Leu Thr Tyr Asp Pro Tyr Val Asn 1 5 10 15 <210> SEQ ID NO 114 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 114 Ala Thr Ala Leu Thr Tyr Asp Pro Tyr Val Asn Tyr Ser Ser Arg 1 5 10 15 <210> SEQ ID NO 115 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 115 Thr Tyr Asp Pro Tyr Val Asn Tyr Ser Ser Arg His Thr Ile Thr 10 1 5 15 <210> SEQ ID NO 116

```
-continued
```

<211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 116 Tyr Val Asn Tyr Ser Ser Arg His Thr Ile Thr Gln Pro Phe Ser 1 5 10 15 <210> SEQ ID NO 117 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 117 Ser Ser Arg His Thr Ile Thr Gln Pro Phe Ser Tyr His Ser Arg 1 5 10 15 <210> SEQ ID NO 118 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 118 Thr Ile Thr Gln Pro Phe Ser Tyr His Ser Arg Tyr Phe Thr Pro 1 5 10 15 <210> SEQ ID NO 119 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 119 Pro Phe Ser Tyr His Ser Arg Tyr Phe Thr Pro Lys Pro Val Leu 1 10 15 5 <210> SEQ ID NO 120 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 120 His Ser Arg Tyr Phe Thr Pro Lys Pro Val Leu Asp Phe Thr Ile 1 5 10 15 <210> SEQ ID NO 121 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 121 Phe Thr Pro Lys Pro Val Leu Asp Phe Thr Ile Asp Tyr Phe Gln 1 5 10 15 <210> SEQ ID NO 122 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 122 Pro Val Leu Asp Phe Thr Ile Asp Tyr Phe Gln Pro Asn Asn Lys 1 5 10 15

```
-continued
```

<210> SEQ ID NO 123 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 123 Phe Thr Ile Asp Tyr Phe Gln Pro Asn Asn Lys Arg Asn Gln Leu 1 5 10 15 <210> SEQ ID NO 124 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 124 Tyr Phe Gln Pro Asn Asn Lys Arg Asn Gln Leu Trp Leu Arg Leu 5 10 1 15 <210> SEQ ID NO 125 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 125 Asn Asn Lys Arg Asn Gln Leu Trp Leu Arg Leu Gln Thr Ala Gly 1 - 5 10 15 <210> SEQ ID NO 126 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 126 Asn Gln Leu Trp Leu Arg Leu Gln Thr Ala Gly Asn Val Asp His 5 1 10 15 <210> SEQ ID NO 127 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 127 Leu Arg Leu Gln Thr Ala Gly Asn Val Asp His Val Gly Leu Gly 1 5 10 15 <210> SEQ ID NO 128 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 128 Thr Ala Gly Asn Val Asp His Val Gly Leu Gly Thr Ala Phe Glu 1 5 10 15 <210> SEQ ID NO 129 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 129 Gly Leu Gly Thr Ala Phe Glu Asn Ser Ile Tyr Asp Gln Glu Tyr 1 5 10 15

```
-continued
```

<210> SEO ID NO 130 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 130 Ala Phe Glu Asn Ser Ile Tyr Asp Gln Glu Tyr Asn Ile Arg Val 1 5 10 15 <210> SEQ ID NO 131 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 131 Ser Ile Tyr Asp Gln Glu Tyr Asn Ile Arg Val Thr Met Tyr Val 1 5 10 15 <210> SEQ ID NO 132 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 132 Gln Glu Tyr Asn Ile Arg Val Thr Met Tyr Val Gln Phe Arg Glu 1 5 10 15 <210> SEQ ID NO 133 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 133 Ile Arg Val Thr Met Tyr Val Gln Phe Arg Glu Phe Asn Phe Lys 5 1 10 15 <210> SEQ ID NO 134 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 134 Met Tyr Val Gln Phe Arg Glu Phe Asn Phe Lys Asp Pro Pro Leu 1 5 10 15 <210> SEQ ID NO 135 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 135 Val Gln Phe Arg Glu Phe Asn Phe Lys Asp Pro Pro Leu Asn Pro 1 5 10 15 <210> SEQ ID NO 136 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 136 Arg Gly Val Gly Ser Thr Val Val Ile Leu Asp Ala Asn Phe Val 5 10 1 15

<210> SEQ ID NO 137 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 137 Ser Thr Val Val Ile Leu Asp Ala Asn Phe Val Thr Pro Ser Thr 10 1 5 15 <210> SEQ ID NO 138 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 138 Ile Leu Asp Ala Asn Phe Val Thr Pro Ser Thr Asn Leu Ala Tyr 1 5 10 15 <210> SEQ ID NO 139 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 139 Asn Phe Val Thr Pro Ser Thr Asn Leu Ala Tyr Asp Pro Tyr Ile 1 5 10 15 <210> SEQ ID NO 140 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 140 Pro Ser Thr Asn Leu Ala Tyr Asp Pro Tyr Ile Asn Tyr Ser Ser 1 5 10 15 <210> SEQ ID NO 141 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEOUENCE: 141 Leu Ala Tyr Asp Pro Tyr Ile As
n Tyr Ser Ser Arg His Thr Ile 1 5 10 15 <210> SEQ ID NO 142 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 142 Pro Tyr Ile Asn Tyr Ser Ser Arg His Thr Ile Arg Gln Pro Phe 1 5 10 15 <210> SEQ ID NO 143 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 143 Tyr Ser Ser Arg His Thr Ile Arg Gln Pro Phe Thr Tyr His Ser

-continued 1 5 10 15 <210> SEQ ID NO 144 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 144 His Thr Ile Arg Gln Pro Phe Thr Tyr His Ser Arg Tyr Phe Thr 1 5 10 15 <210> SEQ ID NO 145 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 145 Gln Pro Phe Thr Tyr His Ser Arg Tyr Phe Thr Pro Lys Pro Glu 1 5 10 15 <210> SEQ ID NO 146 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 146 Tyr His Ser Arg Tyr Phe Thr Pro Lys Pro Glu Leu Asp Gln Thr 5 10 15 1 <210> SEQ ID NO 147 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 147 Tyr Phe Thr Pro Lys Pro Glu Leu Asp Gln Thr Ile Asp Trp Phe 1 5 10 15 <210> SEQ ID NO 148 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 148 Lys Pro Glu Leu Asp Gln Thr Ile Asp Trp Phe Gln Pro Asn Asn 1 5 10 15 <210> SEQ ID NO 149 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 149 Asp Gln Thr Ile Asp Trp Phe Gln Pro Asn Asn Lys Arg Asn Gln 5 10 1 15 <210> SEQ ID NO 150 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 150

Asp Trp Phe Gln Pro Asn Asn Lys Arg Asn Gln Leu Trp Leu His 1 5 10 15 <210> SEQ ID NO 151 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 151 Pro Asn Asn Lys Arg Asn Gln Leu Trp Leu His Leu Asn Thr His 1 5 10 15 <210> SEQ ID NO 152 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 152 Arg Asn Gln Leu Trp Leu His Leu Asn Thr His Thr Asn Val Glu 1 5 10 15 <210> SEQ ID NO 153 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 153 Trp Leu His Leu Asn Thr His Thr Asn Val Glu His Thr Gly Leu 5 10 15 1 <210> SEQ ID NO 154 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 154 Asn Thr His Thr Asn Val Glu His Thr Gly Leu Gly Tyr Ala Leu 1 5 10 15 <210> SEQ ID NO 155 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 155 Asn Val Glu His Thr Gly Leu Gly Tyr Ala Leu Gln Asn Ala Thr 1 5 10 15 <210> SEQ ID NO 156 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 156 Thr Gly Leu Gly Tyr Ala Leu Gln Asn Ala Thr Thr Ala Gln Asn 5 10 1 15 <210> SEQ ID NO 157 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 157

60

120

180

240 300

Tyr Ala Leu Gln Asn Ala Thr Thr Ala Gln Asn Tyr Val Val Arg 5 10 1 15 <210> SEQ ID NO 158 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEOUENCE: 158 Asn Ala Thr Thr Ala Gln Asn Tyr Val Val Arg Leu Thr Ile Tyr 1 5 10 15 <210> SEQ ID NO 159 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 159 Ala Gln Asn Tyr Val Val Arg Leu Thr Ile Tyr Val Gln Phe Arg 10 1 5 15 <210> SEQ ID NO 160 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 160 Val Val Arg Leu Thr Ile Tyr Val Gln Phe Arg Glu Phe Ile Leu 10 15 1 5 <210> SEQ ID NO 161 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEOUENCE: 161 Thr Ile Tyr Val Gln Phe Arg Glu Phe Ile Leu Lys Asp Pro Leu 1 5 10 15 <210> SEQ ID NO 162 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 162 Tyr Val Gln Phe Arg Glu Phe Ile Leu Lys Asp Pro Leu Asn Glu 1 5 10 15 <210> SEQ ID NO 163 <211> LENGTH: 1759 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 163 accagegeae tteggeageg geageacete ggeagegtea gtgaaaatge eaageaagaa aageggeeeg caaceecata agaggtgggt gtteaceett aataateett eegaggagga gaaaaacaaa atacgggagc ttccaatctc cctttttgat tattttgttt gcggagagga aggtttggaa gagggtagaa ctcctcacct ccaggggttt gcgaattttg ctaagaagca qacttttaac aaqqtqaaqt qqtattttqq tqcccqctqc cacatcqaqa aaqcqaaaqq

-continued

aaccgaccag cagaataaag aatactgcag taaagaaggc cacatactta tcgagtgtgg	360
ageteegegg aaccagggga agegeagega eetgtetaet getgtgagta eeettttgga	420
gacggggtct ttggtgactg tagccgagca gttccctgta acgtatgtga gaaatttccg	480
cgggctggct gaacttttga aagtgagcgg gaagatgcag aagcgtgatt ggaagacagc	540
tgtacacgtc atagtgggcc cgcccggttg tgggaagagc cagtgggccc gtaattttgc	600
tgagcctagg gacacctact ggaagcctag tagaaataag tggtgggatg gatatcatgg	660
agaagaagtt gttgttttgg atgattttta tggctggtta ccttgggatg atctactgag	720
actgtgtgac cggtatccat tgactgtaga gactaaaggg ggtactgttc cttttttggc	780
ccgcagtatt ttgattacca gcaatcaggc cccccaggaa tggtactcct caactgctgt	840
cccagctgta gaagctctct atcggaggat tactactttg caattttgga agactgctgg	900
agaacaatcc acggaggtac ccgaaggccg atttgaagca gtggacccac cctgtgccct	960
tttcccatat aaaataaatt actgagtett ttttgttate acategtaat ggtttttatt	1020
tttatttatt tagagggtct tttaggataa attctctgaa ttgtacataa atagtcagcc	1080
ttaccacata attttgggct gtggttgcat tttggagcgc atagcccagg cctgtgtgct	1140
cgacattggt gtgggtattt aaatggagcc acagctggtt tcttttatta tttgggtgga	1200
accaatcaat tgtttggtcc agctcaggtt tgggggtgaa gtacctggag tggtaggtaa	1260
agggctgcct tatggtgtgg cgggaggagt agttaatata ggggtcatag gccaagttgg	1320
tggagggggt tacaaagttg gcatccaaga taacaacagt ggacccaaca cctctttgat	1380
tagaggtgat ggggtctctg gggtaaaatt catatttagc ctttctaata cggtagtatt	1440
ggaaaggtag gggtaggggg ttggtgccgc ctgagggggg gaggaactgg ccgatgttga	1500
atttcagcta gttaacattc caagatggct gcgagtatcc tccttttatg gtgagtacaa	1560
attetgtaga aaggegggaa ttgaagatae eegtettteg gegeeatetg taaeggttte	1620
tgaaggeggg gtgtgeeaaa tatggtette teeggaggat gttteeaaga tggetgeggg	1680
ggegggteet tettetgegg taacgeetee ttggeeaegt cateetataa aagtgaaaga	1740
agtgcgctgc tgtagtatt	1759
<210> SEQ ID NO 164 <211> LENGTH: 1759 <212> TYPE: DNA <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 164	
accagogoac ttoggoagog goagoacoto ggoagogtoa gtgaaaatgo caagoaagaa	60
aageggeeeg caaceecata agaggtgggt gttcaceett aataateett eegaggagga	120
gaaaaacaaa atacgggagc ttccaatctc cctttttgat tattttgttt gcggagagga	180
aggtttggaa gagggtagaa ctcctcacct ccaggggttt gctaattttg ctaagaagca	240
gacttttaac aaggtgaagt ggtattttgg tgcccgctgc cacatcgaga aagcgaaagg	300
aaccgaccag cagaataaag aatactgcag taaagaaggc cacatactta tcgagtgtgg	360
ageteegegg aaceagggga agegeagega eetgtetaet getgtgagta eeettttgga	420
gacggggtct ttggtgactg tagccgagca gttccctgta acgtatgtga gaaatttccg	480
cgggctggct gaacttttga aagtgagcgg gaagatgcag aagcgtgatt ggaagacagc	540

tgtacacgtc atagtgggcc cgcccggttg tgggaagagc cagtgggccc gtaattttgc	600
tgagcctagc gacacctact ggaagcctag tagaaataag tggtgggatg gatatcatgg	660
agaagaagtt gttgttttgg atgattttta tggctggtta ccttgggatg atctactgag	720
actgtgtgac cggtatccat tgactgtaga gactaaaggc ggtactgttc cttttttggc	780
tcgcagtatt ttgattacca gcaatcaggc cccccaggaa tggtactcct caactgctgt	840
cccagctgta gaagctctct atcggaggat tactactttg caattttgga agactgctgg	900
agaacaatca acggaggtac ccgaaggccg atttgaagca gtggacccac cctgtgccct	960
tttcccatat aaaataaatt actgagtett ttttgttate acategtaat ggtttttatt	1020
tttatttatt tagagggtct tttaggataa attctctgaa ttgtacataa atagtcagcc	1080
ttaccacata attttgggct gtggttgcat tttggagcgc atagcccagg cctgtgtgct	1140
cgacattggt gtgggtattt aaatggagcc acagctggtt tcttttatta tttgggtgga	1200
accattcaat tgtttggtcc agctcaggtt tgggggtgaa gtacctggag tggtaggtaa	1260
agggctgcct tatggtgtgg cgggaggagt agttaatata ggggtcatag gccaagttgg	1320
tggagggggt tacaaagttg gcatccaaga taacaacagt ggacccaaca cctctttcat	1380
tagaggtgat ggggtctctg gggtaaaatt catatttagc ctttctaata cggtagtatt	1440
ggaaaggtag gggtaggggg ttggtgccgc ctgagggggg gaggaactgg ccgatgttga	1500
atctgaggtg gttaacatgc caagatggct gcgagtatcc tccttttatg gtgattacaa	1560
attetttaga aaggeggeaa ttgaagatae eegtettteg gegeeatetg taaeggttte	1620
tgaaggeggg gtgtgeeaaa tatggtette teeggaggat gttteeaaga tggetgeggg	1680
ggcgggtcct tcttctgcgg taacgcctcc ttggccacgt catcctataa aagtgaaaga	1740
agtgcgctgc tgtagtatt	1759
<210> SEQ ID NO 165 <211> LENGTH: 312 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus	
<400> SEQUENCE: 165	
<400> SEQUENCE: 165 Met Pro Ser Lys Lys Ser Gly Pro Gln Pro His Lys Arg Trp Val Phe 1 5 10 15	
~ Met Pro Ser Lys Lys Ser Gly Pro Gln Pro His Lys Arg Trp Val Phe	
Met Pro Ser Lys Lys Ser Gly Pro Gln Pro His Lys Arg Trp Val Phe 1 5 10 15 Thr Leu Asn Asn Pro Ser Gly Gly Gly Lys Asn Lys Ile Arg Gly Leu	
Met Pro Ser Lys Lys Ser Gly Pro Gln Pro His Lys Arg Trp Val Phe 1 5 10 15 15 Thr Leu Asn Asn Pro Ser Gly Gly Gly Lys Asn Lys Ile Arg Gly Leu 20 25 30 Pro Ile Ser Leu Phe Asp Tyr Phe Val Cys Gly Gly Gly Gly Leu Gly	
Met Pro Ser Lys Ser Gly Pro Gln Pro His Lys Arg Trp Val Phe 1 No Asn Asn Pro Ser Gly Gly Gly Lys Asn Lys Arg Gly Leu Asn Leu Asn Asn Pro Ser Leu Phe Val Cys Gly Gly Gly Leu Gly Asn Phe Val Ya Asn Asn Asn Asn Ser Gly Gly Leu Gly Gly Gly Gly Gly Gly Gly Leu Gly Asn Phe Ala Asn Phe Ala Leu Leu Gly Asn Phe Ala Asn Phe Ala Leu Leu Ser Leu Gly Asn Phe Ala Leu Leu </td <td></td>	
Met Pro Ser Lys Lys Ser Gly Pro Gln Pro His Lys Arg Trp Val Phe 1 1^{1} Pro Ser Lys Lys Ser Gly Gly Gly Lys Asn Lys IIe Arg Gly Leu 20 1^{1} Cly Cly Cly Lys Asn Lys IIe Arg Gly Leu 20 1^{1} Cly Cly Cly Cly Cly Cly Cly Gly Gly Cly Leu Gly Pro IIe Ser Leu Phe Asp Tyr Phe Val Cys Cly Cly Gly Gly Gly Leu Gly 30 1^{1} Cly Cly Arg Thr Pro His Leu Cln Cly Phe Ala Asn Phe Ala Lys Lys 50 1^{1} Cln Thr Phe Asn Lys Val Lys Trp Tyr Phe Gly Ala Arg Cys His IIe	
Met Pro Ser Lys Lys Ser Gly Pro Gln Pro His Lys Arg Trp Val Phe $\begin{array}{c} 1\\ 1\\ 1\end{array}$ Pro Ser Lys Lys Ser Gly Gly Gly Lys Asn Lys IIe Arg Gly Leu $\begin{array}{c} 1\\ 15\\ 15\end{array}$ Pro Leu Asn Asn Pro Ser Gly Gly Gly Lys Asn Lys IIe Arg Gly Leu $\begin{array}{c} 1\\ 20\\ 30\end{array}$ Pro IIe Ser Leu Phe Asp Tyr Phe Val Cys Gly Gly Gly Gly Gly Leu Gly $\begin{array}{c} 1\\ 45\end{array}$ Gly Gly Arg Thr Pro His Leu Gln Gly Phe Ala Asn Phe Ala Lys Lys $\begin{array}{c} 5\\ 5\\ 5\\ 6\end{array}$ Pro His Asn Lys Val Lys Trp Tyr Phe Gly Ala Arg Cys His IIe $\begin{array}{c} 8\\ 80\end{array}$ Gly Lys Ala Lys Gly Thr Asp Gln Gln Asn Lys Gly Tyr Cys Ser Lys	
MetProSerLysLysSerGlyProGlnProHisLysArgTrpValPhe1 1^{1} <td></td>	

-continued

												COII		ueu	
	130					135					140				
Arg 145	Gly	Leu	Ala	Gly	Leu 150	Leu	ГЛЗ	Val	Ser	Gly 155	Lys	Met	Gln	Gln	Arg 160
Asp	Trp	Lys	Thr	Ala 165	Val	His	Val	Ile	Val 170	Gly	Pro	Pro	Gly	Cys 175	Gly
Lys	Ser	Gln	Trp 180	Ala	Arg	Asn	Phe	Ala 185	Gly	Pro	Arg	Asp	Thr 190	Tyr	Trp
ГÀа	Pro	Ser 195	Arg	Asn	Гла	Trp	Trp 200	Asp	Gly	Tyr	His	Gly 205	Gly	Gly	Val
Val	Val 210	Leu	Asp	Asp	Phe	Tyr 215	Gly	Trp	Leu	Pro	Trp 220	Asp	Asp	Leu	Leu
Arg 225	Leu	Сув	Asp	Arg	Tyr 230	Pro	Leu	Thr	Val	Gly 235	Thr	Lys	Gly	Gly	Thr 240
Val	Pro	Phe	Leu	Ala 245	Arg	Ser	Ile	Leu	Ile 250	Thr	Ser	Asn	Gln	Ala 255	Pro
Gln	Gly	Trp	Tyr 260		Ser	Thr	Ala	Val 265		Ala	Val	Gly	Ala 270		Tyr
Arg	Arg	Ile 275		Thr	Leu	Gln	Phe 280		Lys	Thr	Ala	Gly 285		Gln	Ser
Thr			Pro	Gly	Gly	Arg		Gly	Ala	Val	-		Pro	Cys	Ala
Leu 305	290 Phe	Pro	Tyr	Гла	Ile 310	295 Asn	Tyr				300				
<212 <213	2> T) 3> OF	ENGTI (PE : RGANI EQUEI	PRT [SM:	Тур	e A 1	PWD	circ	ovir	us						
Met 1	Pro	Ser	Lys	Lys 5	Ser	Gly	Pro	Gln	Pro 10	His	LÀa	Arg	Trp	Val 15	Phe
Thr	Leu	Asn	Asn 20	Pro	Ser	Gly	Gly	Gly 25	Lys	Asn	Lys	Ile	Arg 30	Gly	Leu
Pro	Ile	Ser 35	Leu	Phe	Asp	Tyr	Phe 40	Val	Суз	Gly	Gly	Gly 45	Gly	Leu	Gly
Gly	Gly 50	Arg	Thr	Ala	His	Leu 55	Gln	Gly	Phe	Ala	Asn 60	Phe	Ala	Lys	Lys
Gln 65	Thr	Phe	Asn	Lys	Val 70	Lys	Trp	Tyr	Phe	Gly 75	Ala	Arg	Сүз	His	Ile 80
Gly	Lys	Ala	Lys	Gly 85	Thr	Asp	Gln	Gln	Asn 90	ГÀа	Gly	Tyr	Сүз	Ser 95	Lys
Gly	Gly	His	Ile 100	Leu	Ile	Gly	Суз	Gly 105	Ala	Pro	Arg	Asn	Gln 110	Gly	Lys
Arg	Ser	Asp 115	Leu	Ser	Thr	Ala	Val 120	Ser	Thr	Leu	Leu	Gly 125	Thr	Gly	Ser
Leu	Val 130	Thr	Val	Ala	Gly	Gln 135	Phe	Pro	Val	Thr	Tyr 140	Val	Arg	Asn	Phe
Arg 145	Gly	Leu	Ala	Gly	Leu 150	Leu	Lys	Val	Ser	Gly 155	Lys	Met	Gln	Gln	Arg 160
Asp	Trp	Lys	Thr	Ala 165	Val	His	Val	Ile	Val 170	Gly	Pro	Pro	Gly	Cys 175	Gly

Lys	Ser	Gln	Trp 180	Ala	Arg	Asn	Phe	Ala 185	Gly	Pro	Ser	Asp	Thr 190	Tyr	Trp
Lys	Pro	Ser 195	Arg	Asn	Lys	Trp	Trp 200	Asp	Gly	Tyr	His	Gly 205	Gly	Gly	Val
Val	Val 210	Leu	Asp	Asp	Phe	Tyr 215	Gly	Trp	Leu	Pro	Trp 220	Asp	Asp	Leu	Leu
Arg 225	Leu	Cys	Asp	Arg	Tyr 230	Pro	Leu	Thr	Val	Gly 235		Lys	Gly	Gly	Thr 240
Val	Pro	Phe	Leu	Ala 245	Arg	Ser	Ile	Leu	Ile 250	Thr	Ser	Asn	Gln	Ala 255	Pro
Gln	Gly	Trp	Tyr 260	Ser	Ser	Thr	Ala	Val 265	Pro	Ala	Val	Gly	Ala 270	Leu	Tyr
Arg	Arg	Ile 275	Thr	Thr	Leu	Gln	Phe 280	Trp	Lys	Thr	Ala	Gly 285	Gly	Gln	Ser
Thr	Gly 290	Val	Pro	Gly	Gly	Arg 295	Phe	Gly	Ala	Val	Asp 300	Pro	Pro	Суз	Ala
Leu 305	Phe	Pro	Tyr	Lys	Ile 310	Asn	Tyr								
) NO H: 23												
<212	2 > T?	ZPE:	PRT		e A 1	PWD	circo	ovir	us						
<400)> SI	EQUEI	NCE:	167											
Met 1	Thr	Trp	Pro	Arg 5	Arg	Arg	Tyr	Arg	Arg 10	Arg	Arg	Thr	Arg	Pro 15	Arg
Ser	His	Leu	Gly 20	Asn	Ile	Leu	Arg	Arg 25	Arg	Pro	Tyr	Leu	Ala 30	His	Pro
Ala	Phe	Arg 35	Asn	Arg	Tyr	Arg	Trp 40	Arg	Arg	Lys	Thr	Gly 45	Ile	Phe	Asn
Ser	Arg 50	Leu	Ser	Thr	Glu	Phe 55	Val	Leu	Thr	Ile	Arg 60	Gly	Gly	His	Ser
Gln 65	Pro	Ser	Trp	Asn	Val 70	Asn	Tyr	Leu	Lys	Phe 75	Asn	Ile	Gly	Gln	Phe 80
Leu	Pro	Pro	Ser	Gly 85	Gly	Thr	Asn	Pro	Leu 90	Pro	Leu	Pro	Phe	Gln 95	Tyr
Tyr	Arg	Ile	Arg 100	Lys	Ala	Lys	Tyr	Glu 105	Phe	Tyr	Pro	Arg	Asp 110	Pro	Ile
Thr	Ser	Asn 115	Gln	Arg	Gly	Val	Gly 120		Thr	Val	Val	Ile 125	Leu	Asp	Ala
Asn	Phe 130		Thr	Pro	Ser	Thr 135	Asn	Leu	Ala	Tyr	Asp 140	Pro	Tyr	Ile	Asn
Tyr 145	Ser	Ser	Arg	His	Thr 150		Arg	Gln	Pro	Phe 155		Tyr	His	Ser	Arg 160
Tyr	Phe	Thr	Pro	Lys 165	Pro	Glu	Leu	Asp	Gln 170	Thr	Ile	Asp	Trp	Phe 175	His
Pro	Asn	Asn	Lys 180	Arg	Asn	Gln	Leu	Trp 185	Leu	His	Leu	Asn	Thr 190	His	Thr
Asn	Val	Glu 195	His	Thr	Gly	Leu	Gly 200		Ala	Leu	Gln	Asn 205	Ala	Ala	Thr
Ala	Gln 210		Tyr	Val	Val	Arg 215			Ile	Tyr	Val 220		Phe	Arg	Glu
	2 9														

```
-continued
```

Phe Ile Leu Lys Asp Pro Leu Asn Lys <210> SEQ ID NO 168 <211> LENGTH: 233 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 168 Met Thr Trp Pro Arg Arg Arg Tyr Arg Arg Arg Arg Thr Arg Pro Arg Ser His Leu Gly Asn Ile Leu Arg Arg Arg Pro Tyr Leu Val His Pro Ala Phe Arg Asn Arg Tyr Arg Trp Arg Arg Lys Thr Gly Ile Phe Asn Cys Arg Leu Ser Lys Glu Phe Val Ile Thr Ile Arg Gly Gly His Ser Gln Pro Ser Trp Ile Val Asn Ile Leu Arg Phe Asn Ile Gly Gln Phe 65 70 75 80 Leu Pro $\ensuremath{\operatorname{Pro}}$ Ser Gly Gly Thr Asn $\ensuremath{\operatorname{Pro}}$ Leu Pro $\ensuremath{\operatorname{Pro}}$ Phe Gln $\ensuremath{\operatorname{Tyr}}$ Tyr Arg Ile Arg Lys Ala Lys Tyr Glu Phe Tyr Pro Arg Asp Pro Ile Thr Ser Asn Glu Arg Gly Val Gly Ser Thr Val Val Ile Leu Asp Ala Asn Phe Val Thr Pro Ser Thr Asn Leu Ala Tyr Asp Pro Tyr Ile Asn 130 135 140 Tyr Ser Ser Arg His Thr Ile Arg Gln Pro Phe Thr Tyr His Ser Arg Tyr Phe Thr Pro Lys Pro Glu Leu Asp Gln Thr Ile Glu Trp Phe His Pro Asn Asn Lys Arg Asn Gln Leu Trp Leu His Leu Asn Thr His Thr Asn Val Glu His Thr Gly Leu Gly Tyr Ala Leu Gln Asn Ala Ala Thr Ala Gln Asn Tyr Val Val Arg Leu Thr Ile Tyr Val Gln Phe Arg Glu Phe Ile Leu Lys Asp Pro Leu Asn Lys <210> SEQ ID NO 169 <211> LENGTH: 206 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 169 Met Ile Ser Ile Pro Pro Leu Ile Ser Thr Arg Leu Pro Val Gly Val Pro Arg Leu Ser Lys Ile Thr Gly Pro Leu Ala Leu Pro Thr Thr Gly Arg Ala His Tyr Asp Val Tyr Ser Cys Leu Pro Ile Thr Leu Leu His 35 40 45 Leu Pro Ala His Phe Gln Lys Phe Ser Gln Pro Ala Glu Ile Ser His

```
-continued
```

Ile Arg Tyr Arg Glu Leu Leu Gly Tyr Ser His Gln Arg Pro Arg Leu Gln Lys Gly Thr His Ser Ser Arg Gln Val Ala Ala Leu Pro Leu Val Pro Arg Ser Ser Thr Leu Asp Lys Tyr Val Ala Phe Phe Thr Ala Val Phe Phe Ile Leu Leu Val Gly Ser Phe Arg Phe Leu Asp Val Ala Ala Gly Thr Lys Ile Pro Leu His Leu Val Lys Ser Leu Leu Leu Ser Lys Ile Arg Lys Pro Leu Glu Val Arg Ser Ser Thr Leu Phe Gln Thr Phe Leu Ser Ala Asn Lys Ile Ile Lys Lys Gly Asp Trp Lys Leu Pro Tyr Phe Val Phe Leu Leu Gly Arg Ile Ile Lys Gly Glu His Pro Pro Leu Met Gly Leu Arg Ala Ala Phe Leu Ala Trp His Phe His <210> SEQ ID NO 170 <211> LENGTH: 206 <212> TYPE: PRT <213> ORGANISM: Type A PWD circovirus <400> SEQUENCE: 170 Met Ile Ser Ile Pro Pro Leu Ile Ser Thr Arg Leu Pro Val Gly Val Ala Arg Leu Ser Lys Ile Thr Gly Pro Leu Ala Leu Pro Thr Thr Gly Arg Ala His Tyr Asp Val Tyr Ser Cys Leu Pro Ile Thr Leu Leu His Leu Pro Ala His Phe Gln Lys Phe Ser Gln Pro Ala Glu Ile Ser His Ile Arg Tyr Arg Glu Leu Leu Gly Tyr Ser His Gln Arg Pro Arg Leu Gln Lys Gly Thr His Ser Ser Arg Gln Val Ala Ala Leu Pro Leu Val Pro Arg Ser Ser Thr Leu Asp Lys Tyr Val Ala Phe Phe Thr Ala Val Phe Phe Ile Leu Leu Val Gly Ser Phe Arg Phe Leu Asp Val Ala Ala Gly Thr Lys Ile Pro Leu His Leu Val Lys Ser Leu Leu Leu Ser Lys Ile Arg Lys Pro Leu Glu Val Ser Ser Ser Thr Leu Phe Gln Thr Phe Leu Ser Ala Asn Lys Ile Ile Lys Lys Gly Asp Trp Lys Leu Pro Tyr Phe Val Phe Leu Leu Gly Arg Ile Ile Lys Gly Glu His Pro Pro Leu Met Gly Leu Arg Ala Ala Phe Leu Ala Trp His Phe His

-continued <210> SEQ ID NO 171 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 171 Met Thr Tyr Pro Arg Arg Arg Tyr Arg Arg Arg Arg His Arg Pro 1 5 10 15 <210> SEQ ID NO 172 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 172 Arg Arg Arg Tyr Arg Arg Arg Arg His Arg Pro Arg Ser His Leu 1 5 10 15 <210> SEQ ID NO 173 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 173 Arg Arg Arg Arg His Arg Pro Arg Ser His Leu Gly Gln Ile Leu 1 - 5 10 15 <210> SEQ ID NO 174 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 174 Ser Pro Ile Thr Gln Gly Asp Arg Gly Val Gly Ser Ser Ala Val 5 10 15 1 <210> SEQ ID NO 175 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 175 Met Thr Trp Pro Arg Arg Arg Tyr Arg Arg Arg Arg Thr Arg Pro 1 5 10 15 <210> SEQ ID NO 176 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 176 Arg Arg Arg Tyr Arg Arg Arg Arg Thr Arg Pro Arg Ser His Leu 1 5 10 15 <210> SEQ ID NO 177 <211> LENGTH: 15 <212> TYPE: PRT <213> ORGANISM: Type B PWD circovirus <400> SEQUENCE: 177 Arg Arg Arg Arg Thr Arg Pro Arg Ser His Leu Gly Asn Ile Leu 1 5 10 15

We claim:

1. A vaccine comprising a nucleotide sequence of the genome of Porcine circovirus type B, or a homologue or fragment thereof, and an acceptable pharmaceutical or veterinary vehicle, wherein the nucleotide sequence is selected from SEQ ID No. 15 or SEQ ID No. 19.

2. The vaccine of claim **1**, wherein the homologue has at least 80% sequence identity to SEQ ID No. 15 or SEQ ID No. 19.

3. A vaccine comprising a polypeptide encoded by a nucleotide sequence of the genome of Porcine circovirus type B, or a homologue or fragment thereof, and an acceptable pharmaceutical or veterinary vehicle.

4. The vaccine of claim **3**, wherein the homologue has at least 80% sequence identity to SEQ ID No. 15 or SEQ ID No. 19.

5. The vaccine of claim **3**, wherein the nucleotide sequence is selected from SEQ ID No. 23 or SEQ ID No. 25, or a homologue or fragment thereof.

6. The vaccine of claim **5**, wherein the homologue has at least 80% sequence identity to SEQ ID No. 23 or SEQ ID No. 25.

7. The vaccine of claim 5, wherein the nucleotide sequence is SEQ ID No. 25.

8. The vaccine of claim **3**, wherein the polypeptide has the amino acid sequence of SEQ ID No. 24 or SEQ ID No. 26.

9. The vaccine of claim 8, wherein the polypeptide has the amino acid sequence of SEQ ID No. 26.

10. The vaccine of claim 3, wherein the homologue has at least 80% sequence identity to SEQ ID No. 24 or SEQ ID No. 26.

11. The vaccine of claim 10, wherein the homologue has at least 80% sequence identity to SEQ ID No. 26.

12. The vaccine of claim **3**, wherein the polypeptide has the amino acid sequence of SEQ ID No. 29, SEQ ID No. 30, SEQ ID No. 31, or SEQ ID No. 32.

13. A vaccine comprising a vector and an acceptable pharmaceutical or veterinary vehicle, the vector comprising a nucleotide sequence of the genome of Porcine circovirus type B, or a homologue or fragment thereof.

14. A vaccine according to claim 13, further comprising a gene coding for an expression product capable of inhibiting or retarding the establishment or development of a genetic or acquired disease.

15. A vaccine comprising a cell and an acceptable pharmaceutical or veterinary vehicle, wherein the cell is transformed with a nucleotide sequence of the genome of Porcine circovirus type B, or a homologue or fragment thereof.

16. A vaccine comprising a pharmaceutically acceptable vehicle and a single polypetide, wherein the single polypeptide consists of SEQ ID No. 26.

* * * * *