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<110> Regeneron Pharmaceuticals, Inc.
 <120> NON-HUMAN ANIMALS COMPRISING A HUMANIZED TRKB LOCUS
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 <150> US 62/592,905
 <151> 2017-11-30
 <150> US 62/661,373
 <151> 2018-04-23
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 Met Ser Pro Trp Leu Lys Trp His Gly Pro Ala Met Ala Arg Leu Trp
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 Gly Leu Cys Leu Leu Val Leu Gly Phe Trp Arg Ala Ser Leu Ala Cys
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 Pro Thr Ser Cys Lys Cys Ser Ser Ala Arg Ile Trp Cys Thr Glu Pro
 Page 1

35

40

45

Ser Pro Gly Ile Val Ala Phe Pro Arg Leu Glu Pro Asn Ser Val Asp
 50 55 60

Pro Glu Asn Ile Thr Glu Ile Leu Ile Ala Asn Gln Lys Arg Leu Glu
 65 70 75 80

Ile Ile Asn Glu Asp Asp Val Glu Ala Tyr Val Gly Leu Arg Asn Leu
 85 90 95

Thr Ile Val Asp Ser Gly Leu Lys Phe Val Ala Tyr Lys Ala Phe Leu
 100 105 110

Lys Asn Ser Asn Leu Arg His Ile Asn Phe Thr Arg Asn Lys Leu Thr
 115 120 125

Ser Leu Ser Arg Arg His Phe Arg His Leu Asp Leu Ser Asp Leu Ile
 130 135 140

Leu Thr Gly Asn Pro Phe Thr Cys Ser Cys Asp Ile Met Trp Leu Lys
 145 150 155 160

Thr Leu Gln Glu Thr Lys Ser Ser Pro Asp Thr Gln Asp Leu Tyr Cys
 165 170 175

Leu Asn Glu Ser Ser Lys Asn Met Pro Leu Ala Asn Leu Gln Ile Pro
 180 185 190

Asn Cys Gly Leu Pro Ser Ala Arg Leu Ala Ala Pro Asn Leu Thr Val
 195 200 205

Glu Glu Gly Lys Ser Val Thr Leu Ser Cys Ser Val Gly Gly Asp Pro
 210 215 220

Leu Pro Thr Leu Tyr Trp Asp Val Gly Asn Leu Val Ser Lys His Met
 225 230 235 240

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

Ser Asp Asp Ser Gly Lys Gln Ile Ser Cys Val Ala Glu Asn Leu Val
260 265 270

Gly Glu Asp Gln Asp Ser Val Asn Leu Thr Val His Phe Ala Pro Thr
275 280 285

Ile Thr Phe Leu Glu Ser Pro Thr Ser Asp His His Trp Cys Ile Pro
290 295 300

Phe Thr Val Arg Gly Asn Pro Lys Pro Ala Leu Gln Trp Phe Tyr Asn
305 310 315 320

Gly Ala Ile Leu Asn Glu Ser Lys Tyr Ile Cys Thr Lys Ile His Val
325 330 335

Thr Asn His Thr Glu Tyr His Gly Cys Leu Gln Leu Asp Asn Pro Thr
340 345 350

His Met Asn Asn Gly Asp Tyr Thr Leu Met Ala Lys Asn Glu Tyr Gly
355 360 365

Lys Asp Glu Arg Gln Ile Ser Ala His Phe Met Gly Arg Pro Gly Val
370 375 380

Asp Tyr Glu Thr Asn Pro Asn Tyr Pro Glu Val Leu Tyr Glu Asp Trp
385 390 395 400

Thr Thr Pro Thr Asp Ile Gly Asp Thr Thr Asn Lys Ser Asn Glu Ile
405 410 415

Pro Ser Thr Asp Val Ala Asp Gln Ser Asn Arg Glu His Leu Ser Val
420 425 430

Tyr Ala Val Val Val Ile Ala Ser Val Val Gly Phe Cys Leu Leu Val
435 440 445

Met Leu Leu Leu Leu Lys Leu Ala Arg His Ser Lys Phe Gly Met Lys
450 455 460

Gly Pro Ala Ser Val Ile Ser Asn Asp Asp Asp Ser Ala Ser Pro Leu
465 470 475 480

His His Ile Ser Asn Gly Ser Asn Thr Pro Ser Ser Ser Glu Gly Gly
485 490 495

Pro Asp Ala Val Ile Ile Gly Met Thr Lys Ile Pro Val Ile Glu Asn
500 505 510

Pro Gln Tyr Phe Gly Ile Thr Asn Ser Gln Leu Lys Pro Asp Thr Phe
515 520 525

Val Gln His Ile Lys Arg His Asn Ile Val Leu Lys Arg Glu Leu Gly
530 535 540

Glu Gly Ala Phe Gly Lys Val Phe Leu Ala Glu Cys Tyr Asn Leu Cys
545 550 555 560

Pro Glu Gln Asp Lys Ile Leu Val Ala Val Lys Thr Leu Lys Asp Ala
565 570 575

Ser Asp Asn Ala Arg Lys Asp Phe His Arg Glu Ala Glu Leu Leu Thr
580 585 590

Asn Leu Gln His Glu His Ile Val Lys Phe Tyr Gly Val Cys Val Glu
595 600 605

Gly Asp Pro Leu Ile Met Val Phe Glu Tyr Met Lys His Gly Asp Leu
610 615 620

Asn Lys Phe Leu Arg Ala His Gly Pro Asp Ala Val Leu Met Ala Glu
625 630 635 640

Gly Asn Pro Pro Thr Glu Leu Thr Gln Ser Gln Met Leu His Ile Ala
645 650 655

Gln Gln Ile Ala Ala Gly Met Val Tyr Leu Ala Ser Gln His Phe Val
660 665 670

His Arg Asp Leu Ala Thr Arg Asn Cys Leu Val Gly Glu Asn Leu Leu
675 680 685

Val Lys Ile Gly Asp Phe Gly Met Ser Arg Asp Val Tyr Ser Thr Asp
690 695 700

Tyr Tyr Arg Val Gly Gly His Thr Met Leu Pro Ile Arg Trp Met Pro
705 710 715 720

Pro Glu Ser Ile Met Tyr Arg Lys Phe Thr Thr Glu Ser Asp Val Trp
725 730 735

Ser Leu Gly Val Val Leu Trp Glu Ile Phe Thr Tyr Gly Lys Gln Pro
740 745 750

Trp Tyr Gln Leu Ser Asn Asn Glu Val Ile Glu Cys Ile Thr Gln Gly
755 760 765

Arg Val Leu Gln Arg Pro Arg Thr Cys Pro Gln Glu Val Tyr Glu Leu
770 775 780

Met Leu Gly Cys Trp Gln Arg Glu Pro His Thr Arg Lys Asn Ile Lys
785 790 795 800

Ser Ile His Thr Leu Leu Gln Asn Leu Ala Lys Ala Ser Pro Val Tyr
805 810 815

Leu Asp Ile Leu Gly
820

<210> 2
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<223> Cytoplasmic Domain

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Met Ser Pro Trp Pro Arg Trp His Gly Pro Ala Met Ala Arg Leu Trp
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Gly Leu Cys Leu Leu Val Leu Gly Phe Trp Arg Ala Ser Leu Ala Cys
20 25 30

Pro Met Ser Cys Lys Cys Ser Thr Thr Arg Ile Trp Cys Thr Glu Pro
35 40 45

Ser Pro Gly Ile Val Ala Phe Pro Arg Leu Glu Pro Asn Ser Ile Asp
50 55 60

Pro Glu Asn Ile Thr Glu Ile Leu Ile Ala Asn Gln Lys Arg Leu Glu
65 70 75 80

Ile Ile Asn Glu Asp Asp Val Glu Ala Tyr Val Gly Leu Lys Asn Leu
85 90 95

Thr Ile Val Asp Ser Gly Leu Lys Phe Val Ala Tyr Lys Ala Phe Leu
100 105 110

Lys Asn Gly Asn Leu Arg His Ile Asn Phe Thr Arg Asn Lys Leu Thr
115 120 125

Ser Leu Ser Arg Arg His Phe Arg His Leu Asp Leu Ser Asp Leu Ile
130 135 140

Leu Thr Gly Asn Pro Phe Thr Cys Ser Cys Asp Ile Met Trp Leu Lys
145 150 155 160

Thr Leu Gln Glu Thr Lys Ser Ser Pro Asp Thr Gln Asp Leu Tyr Cys
165 170 175

Leu Asn Glu Ser Ser Lys Asn Thr Pro Leu Ala Asn Leu Gln Ile Pro
180 185 190

Asn Cys Gly Leu Pro Ser Ala Arg Leu Ala Ala Pro Asn Leu Thr Val
 195 200 205

Glu Glu Gly Lys Ser Val Thr Ile Ser Cys Ser Val Gly Gly Asp Pro
 210 215 220

Leu Pro Thr Leu Tyr Trp Asp Val Gly Asn Leu Val Ser Lys His Met
 225 230 235 240

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

Ser Asp Asp Ser Gly Lys Gln Ile Ser Cys Val Ala Glu Asn Leu Val
 260 265 270

Gly Glu Asp Gln Asp Ser Val Asn Leu Thr Val His Phe Ala Pro Thr
 275 280 285

Ile Thr Phe Leu Glu Ser Pro Thr Ser Asp His His Trp Cys Ile Pro
 290 295 300

Phe Thr Val Arg Gly Asn Pro Lys Pro Ala Leu Gln Trp Phe Tyr Asn
 305 310 315 320

Gly Ala Ile Leu Asn Glu Ser Lys Tyr Ile Cys Thr Lys Ile His Val
 325 330 335

Thr Asn His Thr Glu Tyr His Gly Cys Leu Gln Leu Asp Asn Pro Thr
 340 345 350

His Met Asn Asn Gly Asp Tyr Thr Leu Met Ala Lys Asn Glu Tyr Gly
 355 360 365

Lys Asp Glu Arg Gln Ile Ser Ala His Phe Met Gly Arg Pro Gly Val
 370 375 380

Asp Tyr Glu Thr Asn Pro Asn Tyr Pro Glu Val Leu Tyr Glu Asp Trp
 385 390 395 400

Thr Thr Pro Thr Asp Ile Gly Asp Thr Thr Asn Lys Ser Asn Glu Ile
 405 410 415

Pro Ser Thr Asp Val Ala Asp Gln Thr Asn Arg Glu His Leu Ser Val
420 425 430

Tyr Ala Val Val Val Ile Ala Ser Val Val Gly Phe Cys Leu Leu Val
435 440 445

Met Leu Leu Leu Leu Lys Leu Ala Arg His Ser Lys Phe Gly Met Lys
450 455 460

Gly Pro Ala Ser Val Ile Ser Asn Asp Asp Asp Ser Ala Ser Pro Leu
465 470 475 480

His His Ile Ser Asn Gly Ser Asn Thr Pro Ser Ser Ser Glu Gly Gly
485 490 495

Pro Asp Ala Val Ile Ile Gly Met Thr Lys Ile Pro Val Ile Glu Asn
500 505 510

Pro Gln Tyr Phe Gly Ile Thr Asn Ser Gln Leu Lys Pro Asp Thr Phe
515 520 525

Val Gln His Ile Lys Arg His Asn Ile Val Leu Lys Arg Glu Leu Gly
530 535 540

Glu Gly Ala Phe Gly Lys Val Phe Leu Ala Glu Cys Tyr Asn Leu Cys
545 550 555 560

Pro Glu Gln Asp Lys Ile Leu Val Ala Val Lys Thr Leu Lys Asp Ala
565 570 575

Ser Asp Asn Ala Arg Lys Asp Phe His Arg Glu Ala Glu Leu Leu Thr
580 585 590

Asn Leu Gln His Glu His Ile Val Lys Phe Tyr Gly Val Cys Val Glu
595 600 605

Gly Asp Pro Leu Ile Met Val Phe Glu Tyr Met Lys His Gly Asp Leu
610 615 620

Asn Lys Phe Leu Arg Ala His Gly Pro Asp Ala Val Leu Met Ala Glu
625 630 635 640

Gly Asn Pro Pro Thr Glu Leu Thr Gln Ser Gln Met Leu His Ile Ala
645 650 655

Gln Gln Ile Ala Ala Gly Met Val Tyr Leu Ala Ser Gln His Phe Val
660 665 670

His Arg Asp Leu Ala Thr Arg Asn Cys Leu Val Gly Glu Asn Leu Leu
675 680 685

Val Lys Ile Gly Asp Phe Gly Met Ser Arg Asp Val Tyr Ser Thr Asp
690 695 700

Tyr Tyr Arg Val Gly Gly His Thr Met Leu Pro Ile Arg Trp Met Pro
705 710 715 720

Pro Glu Ser Ile Met Tyr Arg Lys Phe Thr Thr Glu Ser Asp Val Trp
725 730 735

Ser Leu Gly Val Val Leu Trp Glu Ile Phe Thr Tyr Gly Lys Gln Pro
740 745 750

Trp Tyr Gln Leu Ser Asn Asn Glu Val Ile Glu Cys Ile Thr Gln Gly
755 760 765

Arg Val Leu Gln Arg Pro Arg Thr Cys Pro Gln Glu Val Tyr Glu Leu
770 775 780

Met Leu Gly Cys Trp Gln Arg Glu Pro His Thr Arg Lys Asn Ile Lys
785 790 795 800

Asn Ile His Thr Leu Leu Gln Asn Leu Ala Lys Ala Ser Pro Val Tyr
805 810 815

Leu Asp Ile Leu Gly
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Met Ser Ser Trp Ile Arg Trp His Gly Pro Ala Met Ala Arg Leu Trp
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Gly Phe Cys Trp Leu Val Val Gly Phe Trp Arg Ala Ala Phe Ala Cys
 20 25 30

Pro Thr Ser Cys Lys Cys Ser Ala Ser Arg Ile Trp Cys Ser Asp Pro
 35 40 45

Ser Pro Gly Ile Val Ala Phe Pro Arg Leu Glu Pro Asn Ser Val Asp
 50 55 60

Pro Glu Asn Ile Thr Glu Ile Phe Ile Ala Asn Gln Lys Arg Leu Glu
 65 70 75 80

Ile Ile Asn Glu Asp Asp Val Glu Ala Tyr Val Gly Leu Arg Asn Leu
 85 90 95

Thr Ile Val Asp Ser Gly Leu Lys Phe Val Ala His Lys Ala Phe Leu
 100 105 110

Lys Asn Ser Asn Leu Gln His Ile Asn Phe Thr Arg Asn Lys Leu Thr
 115 120 125

Ser Leu Ser Arg Lys His Phe Arg His Leu Asp Leu Ser Glu Leu Ile
130 135 140

Leu Val Gly Asn Pro Phe Thr Cys Ser Cys Asp Ile Met Trp Ile Lys
145 150 155 160

Thr Leu Gln Glu Ala Lys Ser Ser Pro Asp Thr Gln Asp Leu Tyr Cys
165 170 175

Leu Asn Glu Ser Ser Lys Asn Ile Pro Leu Ala Asn Leu Gln Ile Pro
180 185 190

Asn Cys Gly Leu Pro Ser Ala Asn Leu Ala Ala Pro Asn Leu Thr Val
195 200 205

Glu Glu Gly Lys Ser Ile Thr Leu Ser Cys Ser Val Ala Gly Asp Pro
210 215 220

Val Pro Asn Met Tyr Trp Asp Val Gly Asn Leu Val Ser Lys His Met
225 230 235 240

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

Ser Asp Asp Ser Gly Lys Gln Ile Ser Cys Val Ala Glu Asn Leu Val
260 265 270

Gly Glu Asp Gln Asp Ser Val Asn Leu Thr Val His Phe Ala Pro Thr
275 280 285

Ile Thr Phe Leu Glu Ser Pro Thr Ser Asp His His Trp Cys Ile Pro
290 295 300

Phe Thr Val Lys Gly Asn Pro Lys Pro Ala Leu Gln Trp Phe Tyr Asn
305 310 315 320

Gly Ala Ile Leu Asn Glu Ser Lys Tyr Ile Cys Thr Lys Ile His Val
325 330 335

Thr Asn His Thr Glu Tyr His Gly Cys Leu Gln Leu Asp Asn Pro Thr
340 345 350

His Met Asn Asn Gly Asp Tyr Thr Leu Ile Ala Lys Asn Glu Tyr Gly
355 360 365

Lys Asp Glu Lys Gln Ile Ser Ala His Phe Met Gly Trp Pro Gly Ile
370 375 380

Asp Asp Gly Ala Asn Pro Asn Tyr Pro Asp Val Ile Tyr Glu Asp Tyr
385 390 395 400

Gly Thr Ala Ala Asn Asp Ile Gly Asp Thr Thr Asn Arg Ser Asn Glu
405 410 415

Ile Pro Ser Thr Asp Val Thr Asp Lys Thr Gly Arg Glu His Leu Ser
420 425 430

Val Tyr Ala Val Val Val Ile Ala Ser Val Val Gly Phe Cys Leu Leu
435 440 445

Val Met Leu Phe Leu Leu Lys Leu Ala Arg His Ser Lys Phe Gly Met
450 455 460

Lys Asp Phe Ser Trp Phe Gly Phe Gly Lys Val Lys Ser Arg Gln Gly
465 470 475 480

Val Gly Pro Ala Ser Val Ile Ser Asn Asp Asp Asp Ser Ala Ser Pro
485 490 495

Leu His His Ile Ser Asn Gly Ser Asn Thr Pro Ser Ser Ser Glu Gly
500 505 510

Gly Pro Asp Ala Val Ile Ile Gly Met Thr Lys Ile Pro Val Ile Glu
515 520 525

Asn Pro Gln Tyr Phe Gly Ile Thr Asn Ser Gln Leu Lys Pro Asp Thr
530 535 540

Phe Val Gln His Ile Lys Arg His Asn Ile Val Leu Lys Arg Glu Leu
545 550 555 560

Gly Glu Gly Ala Phe Gly Lys Val Phe Leu Ala Glu Cys Tyr Asn Leu
565 570 575

Cys Pro Glu Gln Asp Lys Ile Leu Val Ala Val Lys Thr Leu Lys Asp
580 585 590

Ala Ser Asp Asn Ala Arg Lys Asp Phe His Arg Glu Ala Glu Leu Leu
595 600 605

Thr Asn Leu Gln His Glu His Ile Val Lys Phe Tyr Gly Val Cys Val
610 615 620

Glu Gly Asp Pro Leu Ile Met Val Phe Glu Tyr Met Lys His Gly Asp
625 630 635 640

Leu Asn Lys Phe Leu Arg Ala His Gly Pro Asp Ala Val Leu Met Ala
645 650 655

Glu Gly Asn Pro Pro Thr Glu Leu Thr Gln Ser Gln Met Leu His Ile
660 665 670

Ala Gln Gln Ile Ala Ala Gly Met Val Tyr Leu Ala Ser Gln His Phe
675 680 685

Val His Arg Asp Leu Ala Thr Arg Asn Cys Leu Val Gly Glu Asn Leu
690 695 700

Leu Val Lys Ile Gly Asp Phe Gly Met Ser Arg Asp Val Tyr Ser Thr
705 710 715 720

Asp Tyr Tyr Arg Val Gly Gly His Thr Met Leu Pro Ile Arg Trp Met
725 730 735

Pro Pro Glu Ser Ile Met Tyr Arg Lys Phe Thr Thr Glu Ser Asp Val
740 745 750

Trp Ser Leu Gly Val Val Leu Trp Glu Ile Phe Thr Tyr Gly Lys Gln
755 760 765

Pro Trp Tyr Gln Leu Ser Asn Asn Glu Val Ile Glu Cys Ile Thr Gln
770 775 780

Gly Arg Val Leu Gln Arg Pro Arg Thr Cys Pro Gln Glu Val Tyr Glu
Page 13

785		790		795		800
Leu Met Leu Gly Cys Trp Gln Arg Glu Pro His Met Arg Lys Asn Ile						
		805		810		815
Lys Gly Ile His Thr Leu Leu Gln Asn Leu Ala Lys Ala Ser Pro Val						
		820		825		830
Tyr Leu Asp Ile Leu Gly						
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Pro Thr Ser Cys Lys Cys Ser Ala Ser Arg Ile Trp Cys Ser Asp Pro
 35 40 45

Ser Pro Gly Ile Val Ala Phe Pro Arg Leu Glu Pro Asn Ser Val Asp
 50 55 60

Pro Glu Asn Ile Thr Glu Ile Phe Ile Ala Asn Gln Lys Arg Leu Glu
 65 70 75 80

Ile Ile Asn Glu Asp Asp Val Glu Ala Tyr Val Gly Leu Arg Asn Leu
 85 90 95

Thr Ile Val Asp Ser Gly Leu Lys Phe Val Ala His Lys Ala Phe Leu
 100 105 110

Lys Asn Ser Asn Leu Gln His Ile Asn Phe Thr Arg Asn Lys Leu Thr
 115 120 125

Ser Leu Ser Arg Lys His Phe Arg His Leu Asp Leu Ser Glu Leu Ile
 130 135 140

Leu Val Gly Asn Pro Phe Thr Cys Ser Cys Asp Ile Met Trp Ile Lys
 145 150 155 160

Thr Leu Gln Glu Ala Lys Ser Ser Pro Asp Thr Gln Asp Leu Tyr Cys
 165 170 175

Leu Asn Glu Ser Ser Lys Asn Ile Pro Leu Ala Asn Leu Gln Ile Pro
 180 185 190

Asn Cys Gly Leu Pro Ser Ala Asn Leu Ala Ala Pro Asn Leu Thr Val

195	200	205																	
Glu	Glu	Gly	Lys	Ser	Ile	Thr	Leu	Ser	Cys	Ser	Val	Ala	Gly	Asp	Pro				
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Val	Pro	Asn	Met	Tyr	Trp	Asp	Val	Gly	Asn	Leu	Val	Ser	Lys	His	Met				
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Asn	Glu	Thr	Ser	His	Thr	Gln	Gly	Ser	Leu	Arg	Ile	Thr	Asn	Ile	Ser				
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Ser	Asp	Asp	Ser	Gly	Lys	Gln	Ile	Ser	Cys	Val	Ala	Glu	Asn	Leu	Val				
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Gly	Glu	Asp	Gln	Asp	Ser	Val	Asn	Leu	Thr	Val	His	Phe	Ala	Pro	Thr				
		275					280					285							
Ile	Thr	Phe	Leu	Glu	Ser	Pro	Thr	Ser	Asp	His	His	Trp	Cys	Ile	Pro				
	290					295					300								
Phe	Thr	Val	Lys	Gly	Asn	Pro	Lys	Pro	Ala	Leu	Gln	Trp	Phe	Tyr	Asn				
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Gly	Ala	Ile	Leu	Asn	Glu	Ser	Lys	Tyr	Ile	Cys	Thr	Lys	Ile	His	Val				
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Thr	Asn	His	Thr	Glu	Tyr	His	Gly	Cys	Leu	Gln	Leu	Asp	Asn	Pro	Thr				
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His	Met	Asn	Asn	Gly	Asp	Tyr	Thr	Leu	Ile	Ala	Lys	Asn	Glu	Tyr	Gly				
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Lys	Asp	Glu	Lys	Gln	Ile	Ser	Ala	His	Phe	Met	Gly	Trp	Pro	Gly	Ile				
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Gly	Thr	Ala	Ala	Asn	Asp	Ile	Gly	Asp	Thr	Thr	Asn	Arg	Ser	Asn	Glu				
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Ile Pro Ser Thr Asp Val Thr Asp Lys Thr Gly Arg Glu His Leu Ser
420 425 430

Val Tyr Ala Val Val Val Ile Ala Ser Val Val Gly Phe Cys Leu Leu
435 440 445

Val Met Leu Leu Leu Lys Leu Ala Arg His Ser Lys Phe Gly Met
450 455 460

Lys Gly Pro Ala Ser Val Ile Ser Asn Asp Asp Asp Ser Ala Ser Pro
465 470 475 480

Leu His His Ile Ser Asn Gly Ser Asn Thr Pro Ser Ser Ser Glu Gly
485 490 495

Gly Pro Asp Ala Val Ile Ile Gly Met Thr Lys Ile Pro Val Ile Glu
500 505 510

Asn Pro Gln Tyr Phe Gly Ile Thr Asn Ser Gln Leu Lys Pro Asp Thr
515 520 525

Phe Val Gln His Ile Lys Arg His Asn Ile Val Leu Lys Arg Glu Leu
530 535 540

Gly Glu Gly Ala Phe Gly Lys Val Phe Leu Ala Glu Cys Tyr Asn Leu
545 550 555 560

Cys Pro Glu Gln Asp Lys Ile Leu Val Ala Val Lys Thr Leu Lys Asp
565 570 575

Ala Ser Asp Asn Ala Arg Lys Asp Phe His Arg Glu Ala Glu Leu Leu
580 585 590

Thr Asn Leu Gln His Glu His Ile Val Lys Phe Tyr Gly Val Cys Val
595 600 605

Glu Gly Asp Pro Leu Ile Met Val Phe Glu Tyr Met Lys His Gly Asp
610 615 620

Leu Asn Lys Phe Leu Arg Ala His Gly Pro Asp Ala Val Leu Met Ala
625 630 635 640

Glu Gly Asn Pro Pro Thr Glu Leu Thr Gln Ser Gln Met Leu His Ile
645 650 655

Ala Gln Gln Ile Ala Ala Gly Met Val Tyr Leu Ala Ser Gln His Phe
660 665 670

Val His Arg Asp Leu Ala Thr Arg Asn Cys Leu Val Gly Glu Asn Leu
675 680 685

Leu Val Lys Ile Gly Asp Phe Gly Met Ser Arg Asp Val Tyr Ser Thr
690 695 700

Asp Tyr Tyr Arg Val Gly Gly His Thr Met Leu Pro Ile Arg Trp Met
705 710 715 720

Pro Pro Glu Ser Ile Met Tyr Arg Lys Phe Thr Thr Glu Ser Asp Val
725 730 735

Trp Ser Leu Gly Val Val Leu Trp Glu Ile Phe Thr Tyr Gly Lys Gln
740 745 750

Pro Trp Tyr Gln Leu Ser Asn Asn Glu Val Ile Glu Cys Ile Thr Gln
755 760 765

Gly Arg Val Leu Gln Arg Pro Arg Thr Cys Pro Gln Glu Val Tyr Glu
770 775 780

Leu Met Leu Gly Cys Trp Gln Arg Glu Pro His Thr Arg Lys Asn Ile
785 790 795 800

Lys Ser Ile His Thr Leu Leu Gln Asn Leu Ala Lys Ala Ser Pro Val
805 810 815

Tyr Leu Asp Ile Leu Gly
820

<210> 5

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<212> PRT

<213> Artificial Sequence

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<223> Rat Sequence

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<223> Extracellular Domain

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<223> Human Sequence

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<222> (455)..(822)

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20 25 30

Pro Thr Ser Cys Lys Cys Ser Ala Ser Arg Ile Trp Cys Ser Asp Pro
35 40 45

Ser Pro Gly Ile Val Ala Phe Pro Arg Leu Glu Pro Asn Ser Val Asp
50 55 60

Pro Glu Asn Ile Thr Glu Ile Phe Ile Ala Asn Gln Lys Arg Leu Glu
65 70 75 80

Ile Ile Asn Glu Asp Asp Val Glu Ala Tyr Val Gly Leu Arg Asn Leu
85 90 95

Thr Ile Val Asp Ser Gly Leu Lys Phe Val Ala His Lys Ala Phe Leu
100 105 110

Lys Asn Ser Asn Leu Gln His Ile Asn Phe Thr Arg Asn Lys Leu Thr
115 120 125

Ser Leu Ser Arg Lys His Phe Arg His Leu Asp Leu Ser Glu Leu Ile
130 135 140

Leu Val Gly Asn Pro Phe Thr Cys Ser Cys Asp Ile Met Trp Ile Lys
145 150 155 160

Thr Leu Gln Glu Ala Lys Ser Ser Pro Asp Thr Gln Asp Leu Tyr Cys
165 170 175

Leu Asn Glu Ser Ser Lys Asn Ile Pro Leu Ala Asn Leu Gln Ile Pro
180 185 190

Asn Cys Gly Leu Pro Ser Ala Asn Leu Ala Ala Pro Asn Leu Thr Val
195 200 205

Glu Glu Gly Lys Ser Ile Thr Leu Ser Cys Ser Val Ala Gly Asp Pro
210 215 220

Val Pro Asn Met Tyr Trp Asp Val Gly Asn Leu Val Ser Lys His Met
225 230 235 240

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

Ser Asp Asp Ser Gly Lys Gln Ile Ser Cys Val Ala Glu Asn Leu Val
260 265 270

Gly Glu Asp Gln Asp Ser Val Asn Leu Thr Val His Phe Ala Pro Thr
275 280 285

Ile Thr Phe Leu Glu Ser Pro Thr Ser Asp His His Trp Cys Ile Pro
290 295 300

Phe Thr Val Lys Gly Asn Pro Lys Pro Ala Leu Gln Trp Phe Tyr Asn
305 310 315 320

Gly Ala Ile Leu Asn Glu Ser Lys Tyr Ile Cys Thr Lys Ile His Val
325 330 335

Thr Asn His Thr Glu Tyr His Gly Cys Leu Gln Leu Asp Asn Pro Thr
340 345 350

His Met Asn Asn Gly Asp Tyr Thr Leu Ile Ala Lys Asn Glu Tyr Gly
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Gly Pro Asp Ala Val Ile Ile Gly Met Thr Lys Ile Pro Val Ile Glu
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35 40 45

Glu Ile Ile Asn Glu Asp Asp Val Glu Ala Tyr Val Gly Leu Arg Asn
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Leu Thr Ile Val Asp Ser Gly Leu Lys Phe Val Ala Tyr Lys Ala Phe
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Leu Lys Asn Ser Asn Leu Arg His Ile Asn Phe Thr Arg Asn Lys Leu
85 90 95

Thr Ser Leu Ser Arg Arg His Phe Arg His Leu Asp Leu Ser Asp Leu
100 105 110

Ile Leu Thr Gly Asn Pro Phe Thr Cys Ser Cys Asp Ile Met Trp Leu
115 120 125

Lys Thr Leu Gln Glu Thr Lys Ser Ser Pro Asp Thr Gln Asp Leu Tyr
130 135 140

Cys Leu Asn Glu Ser Ser Lys Asn Met Pro Leu Ala Asn Leu Gln Ile
145 150 155 160

Pro Asn Cys Gly Leu Pro Ser Ala Arg Leu Ala Ala Pro Asn Leu Thr
165 170 175

Val Glu Glu Gly Lys Ser Val Thr Leu Ser Cys Ser Val Gly Gly Asp
180 185 190

Pro Leu Pro Thr Leu Tyr Trp Asp Val Gly Asn Leu Val Ser Lys His
195 200 205

Met Asn Glu Thr Ser His Thr Gln Gly Ser Leu Arg Ile Thr Asn Ile
210 215 220

Ser Ser Asp Asp Ser Gly Lys Gln Ile Ser Cys Val Ala Glu Asn Leu
225 230 235 240

Val Gly Glu Asp Gln Asp Ser Val Asn Leu Thr Val His Phe Ala Pro

245

250

255

Thr Ile Thr Phe Leu Glu Ser Pro Thr Ser Asp His His Trp Cys Ile
 260 265 270

Pro Phe Thr Val Arg Gly Asn Pro Lys Pro Ala Leu Gln Trp Phe Tyr
 275 280 285

Asn Gly Ala Ile Leu Asn Glu Ser Lys Tyr Ile Cys Thr Lys Ile His
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Val Thr Asn His Thr Glu Tyr His Gly Cys Leu Gln Leu Asp Asn Pro
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Thr His Met Asn Asn Gly Asp Tyr Thr Leu Met Ala Lys Asn Glu Tyr
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Gly Lys Asp Glu Arg Gln Ile Ser Ala His Phe Met Gly Arg Pro Gly
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Val Asp Tyr Glu Thr Asn Pro Asn Tyr Pro Glu Val Leu Tyr Glu Asp
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Trp Thr Thr Pro Thr Asp Ile Gly Asp Thr Thr Asn Lys Ser Asn Glu
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35 40 45

Ile Gly Met Thr Lys Ile Pro Val Ile Glu Asn Pro Gln Tyr Phe Gly
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Ile Thr Asn Ser Gln Leu Lys Pro Asp Thr Phe Val Gln His Ile Lys
65 70 75 80

Arg His Asn Ile Val Leu Lys Arg Glu Leu Gly Glu Gly Ala Phe Gly
85 90 95

Lys Val Phe Leu Ala Glu Cys Tyr Asn Leu Cys Pro Glu Gln Asp Lys
100 105 110

Ile Leu Val Ala Val Lys Thr Leu Lys Asp Ala Ser Asp Asn Ala Arg
115 120 125

Lys Asp Phe His Arg Glu Ala Glu Leu Leu Thr Asn Leu Gln His Glu
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His Ile Val Lys Phe Tyr Gly Val Cys Val Glu Gly Asp Pro Leu Ile
145 150 155 160

Met Val Phe Glu Tyr Met Lys His Gly Asp Leu Asn Lys Phe Leu Arg
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Ala His Gly Pro Asp Ala Val Leu Met Ala Glu Gly Asn Pro Pro Thr
180 185 190

Glu Leu Thr Gln Ser Gln Met Leu His Ile Ala Gln Gln Ile Ala Ala

195

200

205

Gly Met Val Tyr Leu Ala Ser Gln His Phe Val His Arg Asp Leu Ala
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Thr Arg Asn Cys Leu Val Gly Glu Asn Leu Leu Val Lys Ile Gly Asp
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Phe Gly Met Ser Arg Asp Val Tyr Ser Thr Asp Tyr Tyr Arg Val Gly
 245 250 255

Gly His Thr Met Leu Pro Ile Arg Trp Met Pro Pro Glu Ser Ile Met
 260 265 270

Tyr Arg Lys Phe Thr Thr Glu Ser Asp Val Trp Ser Leu Gly Val Val
 275 280 285

Leu Trp Glu Ile Phe Thr Tyr Gly Lys Gln Pro Trp Tyr Gln Leu Ser
 290 295 300

Asn Asn Glu Val Ile Glu Cys Ile Thr Gln Gly Arg Val Leu Gln Arg
 305 310 315 320

Pro Arg Thr Cys Pro Gln Glu Val Tyr Glu Leu Met Leu Gly Cys Trp
 325 330 335

Gln Arg Glu Pro His Thr Arg Lys Asn Ile Lys Ser Ile His Thr Leu
 340 345 350

Leu Gln Asn Leu Ala Lys Ala Ser Pro Val Tyr Leu Asp Ile Leu Gly
 355 360 365

<210> 55

<211> 31

<212> PRT

<213> Rattus norvegicus

<400> 55

Met Ser Pro Trp Pro Arg Trp His Gly Pro Ala Met Ala Arg Leu Trp
 1 5 10 15

Gly Leu Cys Leu Leu Val Leu Gly Phe Trp Arg Ala Ser Leu Ala

20

25

30

<210> 56
 <211> 398
 <212> PRT
 <213> Rattus norvegicus

<400> 56

Cys Pro Met Ser Cys Lys Cys Ser Thr Thr Arg Ile Trp Cys Thr Glu
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Pro Ser Pro Gly Ile Val Ala Phe Pro Arg Leu Glu Pro Asn Ser Ile
 20 25 30

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

Glu Ile Ile Asn Glu Asp Asp Val Glu Ala Tyr Val Gly Leu Lys Asn
 50 55 60

Leu Thr Ile Val Asp Ser Gly Leu Lys Phe Val Ala Tyr Lys Ala Phe
 65 70 75 80

Leu Lys Asn Gly Asn Leu Arg His Ile Asn Phe Thr Arg Asn Lys Leu
 85 90 95

Thr Ser Leu Ser Arg Arg His Phe Arg His Leu Asp Leu Ser Asp Leu
 100 105 110

Ile Leu Thr Gly Asn Pro Phe Thr Cys Ser Cys Asp Ile Met Trp Leu
 115 120 125

Lys Thr Leu Gln Glu Thr Lys Ser Ser Pro Asp Thr Gln Asp Leu Tyr
 130 135 140

Cys Leu Asn Glu Ser Ser Lys Asn Thr Pro Leu Ala Asn Leu Gln Ile
 145 150 155 160

Pro Asn Cys Gly Leu Pro Ser Ala Arg Leu Ala Ala Pro Asn Leu Thr
 165 170 175

Val Glu Glu Gly Lys Ser Val Thr Ile Ser Cys Ser Val Gly Gly Asp
 Page 58

180				185				190								
Pro	Leu	Pro	Thr	Leu	Tyr	Trp	Asp	Val	Gly	Asn	Leu	Val	Ser	Lys	His	
		195					200					205				
Met	Asn	Glu	Thr	Ser	His	Thr	Gln	Gly	Ser	Leu	Arg	Ile	Thr	Asn	Ile	
		210					215					220				
Ser	Ser	Asp	Asp	Ser	Gly	Lys	Gln	Ile	Ser	Cys	Val	Ala	Glu	Asn	Leu	
		225					230					235				
Val	Gly	Glu	Asp	Gln	Asp	Ser	Val	Asn	Leu	Thr	Val	His	Phe	Ala	Pro	
				245							250			255		
Thr	Ile	Thr	Phe	Leu	Glu	Ser	Pro	Thr	Ser	Asp	His	His	Trp	Cys	Ile	
				260									270			
Pro	Phe	Thr	Val	Arg	Gly	Asn	Pro	Lys	Pro	Ala	Leu	Gln	Trp	Phe	Tyr	
				275									285			
Asn	Gly	Ala	Ile	Leu	Asn	Glu	Ser	Lys	Tyr	Ile	Cys	Thr	Lys	Ile	His	
				290									300			
Val	Thr	Asn	His	Thr	Glu	Tyr	His	Gly	Cys	Leu	Gln	Leu	Asp	Asn	Pro	
				305									315			
Thr	His	Met	Asn	Asn	Gly	Asp	Tyr	Thr	Leu	Met	Ala	Lys	Asn	Glu	Tyr	
				325									330			
Gly	Lys	Asp	Glu	Arg	Gln	Ile	Ser	Ala	His	Phe	Met	Gly	Arg	Pro	Gly	
				340									350			
Val	Asp	Tyr	Glu	Thr	Asn	Pro	Asn	Tyr	Pro	Glu	Val	Leu	Tyr	Glu	Asp	
				355									365			
Trp	Thr	Thr	Pro	Thr	Asp	Ile	Gly	Asp	Thr	Thr	Asn	Lys	Ser	Asn	Glu	
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Ile	Pro	Ser	Thr	Asp	Val	Ala	Asp	Gln	Thr	Asn	Arg	Glu	His			
				385									395			

<210> 57
<211> 24
<212> PRT
<213> Rattus norvegicus

<400> 57

Leu Ser Val Tyr Ala Val Val Val Ile Ala Ser Val Val Gly Phe Cys
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Leu Leu Val Met Leu Leu Leu Leu
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<210> 58
<211> 368
<212> PRT
<213> Rattus norvegicus

<400> 58

Lys Leu Ala Arg His Ser Lys Phe Gly Met Lys Gly Pro Ala Ser Val
1 5 10 15

Ile Ser Asn Asp Asp Asp Ser Ala Ser Pro Leu His His Ile Ser Asn
20 25 30

Gly Ser Asn Thr Pro Ser Ser Ser Glu Gly Gly Pro Asp Ala Val Ile
35 40 45

Ile Gly Met Thr Lys Ile Pro Val Ile Glu Asn Pro Gln Tyr Phe Gly
50 55 60

Ile Thr Asn Ser Gln Leu Lys Pro Asp Thr Phe Val Gln His Ile Lys
65 70 75 80

Arg His Asn Ile Val Leu Lys Arg Glu Leu Gly Glu Gly Ala Phe Gly
85 90 95

Lys Val Phe Leu Ala Glu Cys Tyr Asn Leu Cys Pro Glu Gln Asp Lys
100 105 110

Ile Leu Val Ala Val Lys Thr Leu Lys Asp Ala Ser Asp Asn Ala Arg
115 120 125

Lys Asp Phe His Arg Glu Ala Glu Leu Leu Thr Asn Leu Gln His Glu
Page 60

130		135		140
His 145	Ile Val	Lys Phe Tyr 150	Gly Val Cys Val	Glu 155 Gly Asp Pro Leu Ile 160
Met 165	Val Phe Glu Tyr 165	Met Lys His Gly 170	Asp Leu Asn Lys Phe 175	Leu Arg
Ala 180	His Gly Pro Asp Ala 185	Val Leu Met Ala 190	Glu Gly Asn Pro Pro Thr	
Glu 195	Leu Thr Gln Ser Gln 200	Met Leu His Ile Ala 205	Gln Gln Ile Ala Ala	
Gly 210	Met Val Tyr Leu Ala 215	Ser Gln His Phe Val 220	His Arg Asp Leu Ala	
Thr 225	Arg Asn Cys Leu Val 230	Gly Glu Asn Leu 235	Leu Val Lys Ile Gly Asp 240	
Phe 245	Gly Met Ser Arg Asp Val 250	Tyr Ser Thr Asp Tyr Tyr Arg 255	Val Gly	
Gly 260	His Thr Met Leu Pro Ile 265	Arg Trp Met Pro Pro 270	Glu Ser Ile Met	
Tyr 275	Arg Lys Phe Thr Thr 280	Glu Ser Asp Val Trp Ser 285	Leu Gly Val Val	
Leu 290	Trp Glu Ile Phe Thr 295	Tyr Gly Lys Gln Pro 300	Trp Tyr Gln Leu Ser	
Asn 305	Asn Glu Val Ile 310	Glu Cys Ile Thr Gln 315	Gly Arg Val Leu Gln Arg 320	
Pro 325	Arg Thr Cys Pro Gln Glu Val 330	Tyr Glu Leu Met Leu 335	Gly Cys Trp	
Gln 340	Arg Glu Pro His Thr Arg 345	Lys Asn Ile Lys Asn 350	Ile His Thr Leu	

Leu Gln Asn Leu Ala Lys Ala Ser Pro Val Tyr Leu Asp Ile Leu Gly
 355 360 365

<210> 59
 <211> 31
 <212> PRT
 <213> Homo sapiens

<400> 59

Met Ser Ser Trp Ile Arg Trp His Gly Pro Ala Met Ala Arg Leu Trp
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Gly Phe Cys Trp Leu Val Val Gly Phe Trp Arg Ala Ala Phe Ala
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<210> 60
 <211> 399
 <212> PRT
 <213> Homo sapiens

<400> 60

Cys Pro Thr Ser Cys Lys Cys Ser Ala Ser Arg Ile Trp Cys Ser Asp
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Pro Ser Pro Gly Ile Val Ala Phe Pro Arg Leu Glu Pro Asn Ser Val
 20 25 30

Asp Pro Glu Asn Ile Thr Glu Ile Phe Ile Ala Asn Gln Lys Arg Leu
 35 40 45

Glu Ile Ile Asn Glu Asp Asp Val Glu Ala Tyr Val Gly Leu Arg Asn
 50 55 60

Leu Thr Ile Val Asp Ser Gly Leu Lys Phe Val Ala His Lys Ala Phe
 65 70 75 80

Leu Lys Asn Ser Asn Leu Gln His Ile Asn Phe Thr Arg Asn Lys Leu
 85 90 95

Thr Ser Leu Ser Arg Lys His Phe Arg His Leu Asp Leu Ser Glu Leu
 100 105 110

Ile Leu Val Gly Asn Pro Phe Thr Cys Ser Cys Asp Ile Met Trp Ile
 Page 62

115		120		125
Lys Thr Leu Gln Glu Ala	Lys Ser Ser Pro Asp Thr Gln Asp Leu Tyr			
130	135	140		
Cys Leu Asn Glu Ser Ser	Lys Asn Ile Pro Leu Ala Asn Leu Gln Ile			
145	150	155	160	
Pro Asn Cys Gly Leu Pro Ser Ala Asn Leu Ala Ala Pro Asn Leu Thr				
	165	170	175	
Val Glu Glu Gly Lys Ser Ile Thr Leu Ser Cys Ser Val Ala Gly Asp				
	180	185	190	
Pro Val Pro Asn Met Tyr Trp Asp Val Gly Asn Leu Val Ser Lys His				
	195	200	205	
Met Asn Glu Thr Ser His Thr Gln Gly Ser Leu Arg Ile Thr Asn Ile				
	210	215	220	
Ser Ser Asp Asp Ser Gly Lys Gln Ile Ser Cys Val Ala Glu Asn Leu				
225	230	235	240	
Val Gly Glu Asp Gln Asp Ser Val Asn Leu Thr Val His Phe Ala Pro				
	245	250	255	
Thr Ile Thr Phe Leu Glu Ser Pro Thr Ser Asp His His Trp Cys Ile				
	260	265	270	
Pro Phe Thr Val Lys Gly Asn Pro Lys Pro Ala Leu Gln Trp Phe Tyr				
	275	280	285	
Asn Gly Ala Ile Leu Asn Glu Ser Lys Tyr Ile Cys Thr Lys Ile His				
	290	295	300	
Val Thr Asn His Thr Glu Tyr His Gly Cys Leu Gln Leu Asp Asn Pro				
305	310	315	320	
Thr His Met Asn Asn Gly Asp Tyr Thr Leu Ile Ala Lys Asn Glu Tyr				
	325	330	335	

Gly Lys Asp Glu Lys Gln Ile Ser Ala His Phe Met Gly Trp Pro Gly
340 345 350

Ile Asp Asp Gly Ala Asn Pro Asn Tyr Pro Asp Val Ile Tyr Glu Asp
355 360 365

Tyr Gly Thr Ala Ala Asn Asp Ile Gly Asp Thr Thr Asn Arg Ser Asn
370 375 380

Glu Ile Pro Ser Thr Asp Val Thr Asp Lys Thr Gly Arg Glu His
385 390 395

<210> 61
<211> 24
<212> PRT
<213> Homo sapiens

<400> 61

Leu Ser Val Tyr Ala Val Val Val Ile Ala Ser Val Val Gly Phe Cys
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Leu Leu Val Met Leu Phe Leu Leu
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<210> 62
<211> 384
<212> PRT
<213> Homo sapiens

<400> 62

Lys Leu Ala Arg His Ser Lys Phe Gly Met Lys Asp Phe Ser Trp Phe
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Gly Phe Gly Lys Val Lys Ser Arg Gln Gly Val Gly Pro Ala Ser Val
20 25 30

Ile Ser Asn Asp Asp Asp Ser Ala Ser Pro Leu His His Ile Ser Asn
35 40 45

Gly Ser Asn Thr Pro Ser Ser Ser Glu Gly Gly Pro Asp Ala Val Ile
50 55 60

Ile Gly Met Thr Lys Ile Pro Val Ile Glu Asn Pro Gln Tyr Phe Gly
Page 64

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

Tyr Arg Lys Phe Thr Thr Glu Ser Asp Val Trp Ser Leu Gly Val Val
 290 295 300

Leu Trp Glu Ile Phe Thr Tyr Gly Lys Gln Pro Trp Tyr Gln Leu Ser
 305 310 315 320

Asn Asn Glu Val Ile Glu Cys Ile Thr Gln Gly Arg Val Leu Gln Arg
 325 330 335

Pro Arg Thr Cys Pro Gln Glu Val Tyr Glu Leu Met Leu Gly Cys Trp
 340 345 350

Gln Arg Glu Pro His Met Arg Lys Asn Ile Lys Gly Ile His Thr Leu
 355 360 365

Leu Gln Asn Leu Ala Lys Ala Ser Pro Val Tyr Leu Asp Ile Leu Gly
 370 375 380

<210> 63
 <211> 93
 <212> DNA
 <213> Mus musculus

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<210> 64
 <211> 1194
 <212> DNA
 <213> Mus musculus

<400> 64
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 ctcatgcaa accagaaaag gctagaaatc atcaatgaag atgacgttga agcttacgtg 180
 gggctgagaa accttacaat tgtggattcc ggcttaaagt ttgtggctta caaagcgttt 240
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 aggagacatt tccgccacct tgacttgtct gacctgatcc tgacgggtaa tccgttcacg 360
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caggatttgt actgcctcaa tgagagcagc aagaacatgc ccctggcgaa cctgcagata	480
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taccctgaag tcctctatga agactggacc acgccaactg acattgggga tactacgaac	1140
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 <211> 72
 <212> DNA
 <213> Mus musculus

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ttgctcctgc tc	72

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 <212> DNA
 <213> Mus musculus

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gagggcggtc ccgacgctgt cattattgga atgaccaaga ttcctgttat tgaaaacccc	180
cagtactttg gcatcaccaa cagtcagctc aagccagaca catttggtca gcatatcaag	240
agacacaaca tcgttctgaa gagggaactt ggggaaggag ctttcgggaa agttttcctt	300
gccgagtgtc acaacctctg cccagagcag gataagatcc tgggtggctgt gaagacgctg	360

aaggacgcca gcgacaatgc acgcaaggac tttcatcggg aagctgagct gctgaccaac	420
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atggtctttg agtacatgaa gcacggggac ctcaacaagt tccttagggc acacggggccc	540
gacgcagtgc tgatggcaga gggtaacccg cccacagagc tgacgcagtc gcagatgctg	600
cacatcgctc agcaaatcgc agcaggtatg gtctacctgg cgtcccaaca ctttgtgcac	660
cgtgacctgg ccacccggaa ctgcctggtg ggagagaacc tgctggtgaa aattggggac	720
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tatcagctat cgaacaatga ggtgatagag tgcatacccc agggaagagt cttcagcgg	960
cctcgaacgt gtccccagga ggtgtatgag ctcatgcttg gatgctggca gcgggaacca	1020
cacacccgga agaacaatca gagcatccac accctccttc agaacttggc caaggcatct	1080
cccgctctacc tggatatcct aggc	1104

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 <211> 93
 <212> DNA
 <213> Rattus norvegicus

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ctggtcttgg gcttctggag ggcttctctt gcc	93

<210> 68
 <211> 1194
 <212> DNA
 <213> Rattus norvegicus

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ctcattgcaa accagaaaag gttagaaatc atcaatgaag atgatgtcga agcttacgtg	180
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tacctgaag tcctctatga agactggacc acgccaactg acatcgggga tactacaaac	1140
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 <211> 72
 <212> DNA
 <213> Rattus norvegicus

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ctgcttctgc tc	72

<210> 70
 <211> 1104
 <212> DNA
 <213> Rattus norvegicus

<400> 70	
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gagggcgggc ccgatgctgt catcattggg atgaccaaga tccctgtcat tgaaaacccc	180
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agacacaaca tcgttctgaa gagggagctt ggagaaggag cctttgggaa agttttccta	300
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gacgtctgga gcctgggagt tgtgttgtgg gagatcttca cctacggcaa gcagccctgg	900
tatcagctat caaacaacga ggtgatagaa tgcattcccc agggcagagt cttcagcgg	960
cctcgacagt gtccccagga ggtgtacgag ctgatgctgg gatgctggca gcgggaacca	1020
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<210> 71
 <211> 93
 <212> DNA
 <213> Homo sapiens

<400> 71	
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<210> 72
 <211> 1197
 <212> DNA
 <213> Homo sapiens

<400> 72	
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ttcatcgcaa accagaaaag gttagaaatc atcaacgaag atgatgttga agcttatgtg	180

ggactgagaa atctgacaat tgtggattct ggattaaaat ttgtggctca taaagcattt	240
ctgaaaaaca gcaacctgca gcacatcaat tttacccgaa acaaactgac gagtttgtct	300
aggaaacatt tccgtcacct tgacttgtct gaactgatcc tgggtgggcaa tccatttaca	360
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caggatttgt actgcctgaa tgaaagcagc aagaatattc ccctggcaaa cctgcagata	480
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gttggttaacc tggtttccaa acatatgaat gaaacaagcc acacacaggg ctccttaagg	660
ataactaaca tttcatccga tgacagtggg aagcagatct cttgtgtggc ggaaaatctt	720
gtaggagaag atcaagattc tgtcaacctc actgtgcatt ttgcaccaac tatcacattt	780
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Lys Asn Ser Asn Leu Gln His Ile Asn Phe Thr Arg Asn Lys Leu Thr
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Lys Asp Glu Lys Gln Ile Ser Ala His Phe Met Gly Trp Pro Gly Ile
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tctgaactga tcctggtggg caatccatth acatgctcct gtgacattat gtggatcaag	480
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agcaagaata ttcccctggc aaacctgcag atacccaatt gtggtttgcc atctgcaa	600
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<220>
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<400> 79

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Ser Phe Ser Ser Phe
20 25 30

Gly Met His Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
 35 40 45

Ser Val Ile Ser Tyr Asp Gly Ile Asn Thr Tyr Tyr Thr Asp Ser Val
 50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ser Lys Asn Thr Leu Tyr
 65 70 75 80

Leu Gln Met Asn Gly Leu Arg Ala Glu Asp Thr Ala Leu Tyr Tyr Cys
 85 90 95

Val Gln Gly Ser Ile Gly Thr Val Phe Glu Tyr Trp Gly Gln Gly Thr
 100 105 110

Leu Val Thr Val Ser Ser
 115

<210> 80
 <211> 24
 <212> DNA
 <213> Artificial Sequence

<220>
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<400> 80
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24

<210> 81
 <211> 8
 <212> PRT
 <213> Artificial Sequence

<220>
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<400> 81

Gly Phe Ser Phe Ser Ser Phe Gly
 1 5

<210> 82
 <211> 24
 <212> DNA

<213> Artificial Sequence

<220>

<223> Synthetic

<400> 82

atatcatatg atggaattaa taca

24

<210> 83

<211> 8

<212> PRT

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<220>

<223> Synthetic

<400> 83

Ile Ser Tyr Asp Gly Ile Asn Thr
1 5

<210> 84

<211> 33

<212> DNA

<213> Artificial Sequence

<220>

<223> Synthetic

<400> 84

gtgcaagggt caattggaac cgtttttgaa tac

33

<210> 85

<211> 11

<212> PRT

<213> Artificial Sequence

<220>

<223> Synthetic

<400> 85

Val Gln Gly Ser Ile Gly Thr Val Phe Glu Tyr
1 5 10

<210> 86

<211> 321

<212> DNA

<213> Artificial Sequence

<220>

<223> Synthetic

<400> 86

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gggaaagttc ctaaactcct gatctatgct gcatccactt tacaatcagg ggtcccatct    180
cggttcggtg gcagtggatc tgggacagat ttcactctca ccatcagcag cctgcagcct    240
gaagatgttg caacttatta ctgtcaaaag tataaccagtg ccccatcac tttcggccct    300
gggaccaaag tggatatcaa a                                              321
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<210> 87

<211> 107

<212> PRT

<213> Artificial Sequence

<220>

<223> Synthetic

<400> 87

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1           5           10           15
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Asp Arg Val Thr Ile Thr Cys Arg Ala Ser Gln Gly Ile Ser Asn Tyr
          20           25           30
```

```
Leu Ala Trp Tyr Gln Gln Lys Pro Gly Lys Val Pro Lys Leu Leu Ile
          35           40           45
```

```
Tyr Ala Ala Ser Thr Leu Gln Ser Gly Val Pro Ser Arg Phe Gly Gly
          50           55           60
```

```
Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Ser Ser Leu Gln Pro
65           70           75           80
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```
Glu Asp Val Ala Thr Tyr Tyr Cys Gln Lys Tyr Thr Ser Ala Pro Phe
          85           90           95
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Thr Phe Gly Pro Gly Thr Lys Val Asp Ile Lys
          100          105
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<210> 88

<211> 18
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<213> Artificial Sequence

<220>
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<400> 88
cagggcatta gcaattat

18

<210> 89
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<220>
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<400> 89

Gln Gly Ile Ser Asn Tyr
1 5

<210> 90
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<220>
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<400> 90
gctgcatcc

9

<210> 91
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<400> 91

Ala Ala Ser
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<210> 92
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<220>

<223> Synthetic

<400> 92

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27

<210> 93

<211> 9

<212> PRT

<213> Artificial Sequence

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<223> Synthetic

<400> 93

Gln Lys Tyr Thr Ser Ala Pro Phe Thr

1

5

<210> 94

<211> 366

<212> DNA

<213> Artificial Sequence

<220>

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tccgggaaag ggctggagtg gattggccgt attagaaaca aggctaacag ttacgcgaca 180

gcatatgggtg cgtcgggtgac aggcagggtt accatctcca gagatgattc aaagaacacg 240

gcgtatctgc aaatgaacag cctgaaaacc gaggacacgg ccgtttatta ctgtactttc 300

ccgggtgtag tgggacgagg aggttttgac tactggggcc agggcaccct ggtcaccgtc 360

tcctca 366

<210> 95

<211> 122

<212> PRT

<213> Artificial Sequence

<220>

<223> Synthetic

<400> 95

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
 1 5 10 15

Ser Leu Lys Leu Ser Cys Thr Ala Ser Gly Phe Thr Phe Ser Gly Ser
 20 25 30

Val Ile His Trp Val Arg Gln Ala Ser Gly Lys Gly Leu Glu Trp Ile
 35 40 45

Gly Arg Ile Arg Asn Lys Ala Asn Ser Tyr Ala Thr Ala Tyr Gly Ala
 50 55 60

Ser Val Thr Gly Arg Phe Thr Ile Ser Arg Asp Asp Ser Lys Asn Thr
 65 70 75 80

Ala Tyr Leu Gln Met Asn Ser Leu Lys Thr Glu Asp Thr Ala Val Tyr
 85 90 95

Tyr Cys Thr Phe Pro Gly Val Val Gly Arg Gly Gly Phe Asp Tyr Trp
 100 105 110

Gly Gln Gly Thr Leu Val Thr Val Ser Ser
 115 120

<210> 96
 <211> 24
 <212> DNA
 <213> Artificial Sequence

<220>
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<400> 96
 gggttcacct tcagtggtc tggt

24

<210> 97
 <211> 8
 <212> PRT
 <213> Artificial Sequence

<220>
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<400> 97

Gly Phe Thr Phe Ser Gly Ser Val
1 5

<210> 98
<211> 30
<212> DNA
<213> Artificial Sequence

<220>
<223> Synthetic

<400> 98
attagaaaca aggctaacag ttacgcgaca

30

<210> 99
<211> 10
<212> PRT
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<220>
<223> Synthetic

<400> 99

Ile Arg Asn Lys Ala Asn Ser Tyr Ala Thr
1 5 10

<210> 100
<211> 39
<212> DNA
<213> Artificial Sequence

<220>
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<400> 100
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39

<210> 101
<211> 13
<212> PRT
<213> Artificial Sequence

<220>
<223> Synthetic

<400> 101

Thr Phe Pro Gly Val Val Gly Arg Gly Gly Phe Asp Tyr
1 5 10

<210> 102
<211> 339
<212> DNA
<213> Artificial Sequence

<220>
<223> Synthetic

<400> 102
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tggtaccaac agaaaccagg acagcctcct aagttgctct ttctactgggc atctaccgg 180
gaatccgggg tccctgaccg attcgggtggc agcgggtctg ggacagattt ctctctcacc 240
atcaacagcc tgcagactga agatgtggca gtttattact gtctccaata ttatagtatt 300
ccgtggacgt tcggccaagg gaccaaggtg gaaatcaaa 339

<210> 103
<211> 113
<212> PRT
<213> Artificial Sequence

<220>
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<400> 103

Asp Ile Val Met Thr Gln Ser Pro Asp Ser Leu Ala Val Ser Leu Gly
1 5 10 15

Glu Gly Ala Thr Ile Asn Cys Met Ser Ser Gln Ser Val Leu Phe Ser
20 25 30

Ser Asn Asn Lys Asn Tyr Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln
35 40 45

Pro Pro Lys Leu Leu Phe Tyr Trp Ala Ser Thr Arg Glu Ser Gly Val
50 55 60

Pro Asp Arg Phe Gly Gly Ser Gly Ser Gly Thr Asp Phe Ser Leu Thr
65 70 75 80

Ile Asn Ser Leu Gln Thr Glu Asp Val Ala Val Tyr Tyr Cys Leu Gln
85 90 95

Tyr Tyr Ser Ile Pro Trp Thr Phe Gly Gln Gly Thr Lys Val Glu Ile
100 105 110

Lys

<210> 104
<211> 36
<212> DNA
<213> Artificial Sequence

<220>
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<400> 104
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<210> 105
<211> 12
<212> PRT
<213> Artificial Sequence

<220>
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<400> 105

Gln Ser Val Leu Phe Ser Ser Asn Asn Lys Asn Tyr
1 5 10

<210> 106
<211> 9
<212> DNA
<213> Artificial Sequence

<220>
<223> Synthetic

<400> 106
tgggcatct 9

<210> 107
<211> 3
<212> PRT
<213> Artificial Sequence

<220>
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<400> 107

Trp Ala Ser
1

<210> 108

<211> 27

<212> DNA

<213> Artificial Sequence

<220>

<223> Synthetic

<400> 108

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27

<210> 109

<211> 9

<212> PRT

<213> Artificial Sequence

<220>

<223> Synthetic

<400> 109

Leu Gln Tyr Tyr Ser Ile Pro Trp Thr
1 5

<210> 110

<211> 381

<212> DNA

<213> Artificial Sequence

<220>

<223> Synthetic

<400> 110

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ccagggaagg ggctggagtg gatttcatac attagtaata gtggttatac catatactac 180

gcagactctg tgaagggccg attcaccatc tccagagaca acgccaggaa ctcaatatat 240

ctgcaagtga acagcctgag agccgaggac acggctgttt attactgttc gagacgtact 300

actatgattc ggggcattag ggcgtactac tattacggtc tggacgtctg gggccaaggg 360

accacgggtca ccgtctcctc a 381

<210> 111
 <211> 127
 <212> PRT
 <213> Artificial Sequence

<220>
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<400> 111

Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
 1 5 10 15

Ser Leu Thr Leu Ser Cys Ala Ala Ser Gly Phe Asn Phe Arg Asp Tyr
 20 25 30

Glu Met Ile Trp Val Arg Gln Thr Pro Gly Lys Gly Leu Glu Trp Ile
 35 40 45

Ser Tyr Ile Ser Asn Ser Gly Tyr Thr Ile Tyr Tyr Ala Asp Ser Val
 50 55 60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Arg Asn Ser Ile Tyr
 65 70 75 80

Leu Gln Val Asn Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys
 85 90 95

Ser Arg Arg Thr Thr Met Ile Arg Gly Ile Arg Ala Tyr Tyr Tyr Tyr
 100 105 110

Gly Leu Asp Val Trp Gly Gln Gly Thr Thr Val Thr Val Ser Ser
 115 120 125

<210> 112
 <211> 24
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<220>
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<400> 112

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24

<210> 113
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<212> PRT
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<220>
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<400> 113

Gly Phe Asn Phe Arg Asp Tyr Glu
1 5

<210> 114
<211> 24
<212> DNA
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<400> 114
attagtaata gtggttatac cata

24

<210> 115
<211> 8
<212> PRT
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<220>
<223> Synthetic

<400> 115

Ile Ser Asn Ser Gly Tyr Thr Ile
1 5

<210> 116
<211> 60
<212> DNA
<213> Artificial Sequence

<220>
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<400> 116
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<210> 117
<211> 20

<212> PRT
<213> Artificial Sequence

<220>
<223> Synthetic

<400> 117

Ser Arg Arg Thr Thr Met Ile Arg Gly Ile Arg Ala Tyr Tyr Tyr Tyr
1 5 10 15

Gly Leu Asp Val
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<210> 118
<211> 324
<212> DNA
<213> Artificial Sequence

<220>
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<400> 118
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gggaaagccc ctaagctcct gatctatgct gcatccagtt tgcaaagtgg ggtcccgtca 180
aggttcagtg gcagtggatc tgggacagat ttactctca ccatcagcag tctgcaacct 240
gaagattttg caacttacta ctgtcaacag agttacagta cccctccgat caccttcggc 300
caagggacac gactggagat taaa 324

<210> 119
<211> 108
<212> PRT
<213> Artificial Sequence

<220>
<223> Synthetic

<400> 119

Asp Ile Gln Met Thr Gln Ser Pro Ser Ser Leu Ser Ala Ser Val Gly
1 5 10 15

Asp Arg Val Thr Ile Thr Cys Arg Ala Ser Gln Ser Ile Ser Ser Tyr
20 25 30

Leu Asn Trp Tyr Gln Gln Lys Pro Gly Lys Ala Pro Lys Leu Leu Ile
 35 40 45

Tyr Ala Ala Ser Ser Leu Gln Ser Gly Val Pro Ser Arg Phe Ser Gly
 50 55 60

Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Ser Ser Leu Gln Pro
 65 70 75 80

Glu Asp Phe Ala Thr Tyr Tyr Cys Gln Gln Ser Tyr Ser Thr Pro Pro
 85 90 95

Ile Thr Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys
 100 105

<210> 120
 <211> 18
 <212> DNA
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<220>
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<400> 120
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18

<210> 121
 <211> 6
 <212> PRT
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<220>
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<400> 121

Gln Ser Ile Ser Ser Tyr
 1 5

<210> 122
 <211> 9
 <212> DNA
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<220>
 <223> Synthetic

<400> 122
gctgcatcc

9

<210> 123
<211> 3
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<220>
<223> Synthetic

<400> 123

Ala Ala Ser
1

<210> 124
<211> 30
<212> DNA
<213> Artificial Sequence

<220>
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<400> 124
caacagagtt acagtacccc tccgatacc

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<210> 125
<211> 10
<212> PRT
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<220>
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<400> 125

Gln Gln Ser Tyr Ser Thr Pro Pro Ile Thr
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