

SEQUENCE LISTING

<110> Regeneron Pharmaceuticals, Inc.

<120> NON-HUMAN ANIMALS COMPRISING A HUMANIZED TTR LOCUS AND METHODS OF USE

<130> 57766-519832

<150> US 62/565,980

<151> 2017-09-29

<150> US 62/679,142

<151> 2018-06-01

<150> US 62/720,292

<151> 2018-08-21

<160> 94

<170> PatentIn version 3.5

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aaagtgatca	ccctgaagtc	caagctggtg	tccgattttc	ggaaggattt	ccagttttac	2940
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gccaagtact	tcttctacag	caacatcatg	aactttttca	agaccgaaat	caccctggcc	3180
aacggcgaga	tcagaaagcg	ccctctgatc	gagacaaacg	gcgaaaccgg	ggagatcgtg	3240
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aggaacagcg	acaagctgat	cgccagaaa	aaggactggg	acccaagaa	gtacggcggc	3420
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cagaaacagc	tgtttgtgga	acagcataag	cactacctgg	acgagatcat	cgagcagatc	3840
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tacaacaagc	acagggacaa	gcctatcaga	gagcaggccg	agaatatcat	ccacctgttc	3960

accctgacaa acctgggcg ctcctgccgcc ttcaagtact ttgacaccac catcgaccgg 4020
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<212> PRT
<213> Artificial Sequence

<220>
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<222> (4)..(10)
<223> 5' NLS

<220>
<221> MISC_FEATURE
<222> (1376)..(1391)
<223> 3' NLS

<400> 94

Met Asp Lys Pro Lys Lys Lys Arg Lys Val Lys Tyr Ser Ile Gly Leu
1 5 10 15

Asp Ile Gly Thr Asn Ser Val Gly Trp Ala Val Ile Thr Asp Glu Tyr
20 25 30

Lys Val Pro Ser Lys Lys Phe Lys Val Leu Gly Asn Thr Asp Arg His
35 40 45

Ser Ile Lys Lys Asn Leu Ile Gly Ala Leu Leu Phe Asp Ser Gly Glu
50 55 60

Thr Ala Glu Ala Thr Arg Leu Lys Arg Thr Ala Arg Arg Arg Tyr Thr
65 70 75 80

Arg Arg Lys Asn Arg Ile Cys Tyr Leu Gln Glu Ile Phe Ser Asn Glu
85 90 95

Met Ala Lys Val Asp Asp Ser Phe Phe His Arg Leu Glu Glu Ser Phe
100 105 110

Leu Val Glu Glu Asp Lys Lys His Glu Arg His Pro Ile Phe Gly Asn
115 120 125

Ile Val Asp Glu Val Ala Tyr His Glu Lys Tyr Pro Thr Ile Tyr His
130 135 140

Leu Arg Lys Lys Leu Val Asp Ser Thr Asp Lys Ala Asp Leu Arg Leu
145 150 155 160

Ile Tyr Leu Ala Leu Ala His Met Ile Lys Phe Arg Gly His Phe Leu
165 170 175

Ile Glu Gly Asp Leu Asn Pro Asp Asn Ser Asp Val Asp Lys Leu Phe
180 185 190

Ile Gln Leu Val Gln Thr Tyr Asn Gln Leu Phe Glu Glu Asn Pro Ile
195 200 205

Asn Ala Ser Gly Val Asp Ala Lys Ala Ile Leu Ser Ala Arg Leu Ser
210 215 220

Lys Ser Arg Arg Leu Glu Asn Leu Ile Ala Gln Leu Pro Gly Glu Lys
225 230 235 240

Lys Asn Gly Leu Phe Gly Asn Leu Ile Ala Leu Ser Leu Gly Leu Thr
245 250 255

Pro Asn Phe Lys Ser Asn Phe Asp Leu Ala Glu Asp Ala Lys Leu Gln
260 265 270

Leu Ser Lys Asp Thr Tyr Asp Asp Asp Leu Asp Asn Leu Leu Ala Gln
275 280 285

Ile Gly Asp Gln Tyr Ala Asp Leu Phe Leu Ala Ala Lys Asn Leu Ser
290 295 300

Asp Ala Ile Leu Leu Ser Asp Ile Leu Arg Val Asn Thr Glu Ile Thr

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

Asn Glu Leu Thr Lys Val Lys Tyr Val Thr Glu Gly Met Arg Lys Pro
530 535 540

Ala Phe Leu Ser Gly Glu Gln Lys Lys Ala Ile Val Asp Leu Leu Phe
545 550 555 560

Lys Thr Asn Arg Lys Val Thr Val Lys Gln Leu Lys Glu Asp Tyr Phe
565 570 575

Lys Lys Ile Glu Cys Phe Asp Ser Val Glu Ile Ser Gly Val Glu Asp
580 585 590

Arg Phe Asn Ala Ser Leu Gly Thr Tyr His Asp Leu Leu Lys Ile Ile
595 600 605

Lys Asp Lys Asp Phe Leu Asp Asn Glu Glu Asn Glu Asp Ile Leu Glu
610 615 620

Asp Ile Val Leu Thr Leu Thr Leu Phe Glu Asp Arg Glu Met Ile Glu
625 630 635 640

Glu Arg Leu Lys Thr Tyr Ala His Leu Phe Asp Asp Lys Val Met Lys
645 650 655

Gln Leu Lys Arg Arg Arg Tyr Thr Gly Trp Gly Arg Leu Ser Arg Lys
660 665 670

Leu Ile Asn Gly Ile Arg Asp Lys Gln Ser Gly Lys Thr Ile Leu Asp
675 680 685

Phe Leu Lys Ser Asp Gly Phe Ala Asn Arg Asn Phe Met Gln Leu Ile
690 695 700

His Asp Asp Ser Leu Thr Phe Lys Glu Asp Ile Gln Lys Ala Gln Val
705 710 715 720

Ser Gly Gln Gly Asp Ser Leu His Glu His Ile Ala Asn Leu Ala Gly
725 730 735

Ser Pro Ala Ile Lys Lys Gly Ile Leu Gln Thr Val Lys Val Val Asp

740

745

750

Glu Leu Val Lys Val Met Gly Arg His Lys Pro Glu Asn Ile Val Ile
755 760 765

Glu Met Ala Arg Glu Asn Gln Thr Thr Gln Lys Gly Gln Lys Asn Ser
770 775 780

Arg Glu Arg Met Lys Arg Ile Glu Glu Gly Ile Lys Glu Leu Gly Ser
785 790 795 800

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

Lys Leu Tyr Leu Tyr Tyr Leu Gln Asn Gly Arg Asp Met Tyr Val Asp
820 825 830

Gln Glu Leu Asp Ile Asn Arg Leu Ser Asp Tyr Asp Val Asp His Ile
835 840 845

Val Pro Gln Ser Phe Leu Lys Asp Asp Ser Ile Asp Asn Lys Val Leu
850 855 860

Thr Arg Ser Asp Lys Asn Arg Gly Lys Ser Asp Asn Val Pro Ser Glu
865 870 875 880

Glu Val Val Lys Lys Met Lys Asn Tyr Trp Arg Gln Leu Leu Asn Ala
885 890 895

Lys Leu Ile Thr Gln Arg Lys Phe Asp Asn Leu Thr Lys Ala Glu Arg
900 905 910

Gly Gly Leu Ser Glu Leu Asp Lys Ala Gly Phe Ile Lys Arg Gln Leu
915 920 925

Val Glu Thr Arg Gln Ile Thr Lys His Val Ala Gln Ile Leu Asp Ser
930 935 940

Arg Met Asn Thr Lys Tyr Asp Glu Asn Asp Lys Leu Ile Arg Glu Val
945 950 955 960

Lys Val Ile Thr Leu Lys Ser Lys Leu Val Ser Asp Phe Arg Lys Asp
965 970 975

Phe Gln Phe Tyr Lys Val Arg Glu Ile Asn Asn Tyr His His Ala His
980 985 990

Asp Ala Tyr Leu Asn Ala Val Val Gly Thr Ala Leu Ile Lys Lys Tyr
995 1000 1005

Pro Lys Leu Glu Ser Glu Phe Val Tyr Gly Asp Tyr Lys Val Tyr
1010 1015 1020

Asp Val Arg Lys Met Ile Ala Lys Ser Glu Gln Glu Ile Gly Lys
1025 1030 1035

Ala Thr Ala Lys Tyr Phe Phe Tyr Ser Asn Ile Met Asn Phe Phe
1040 1045 1050

Lys Thr Glu Ile Thr Leu Ala Asn Gly Glu Ile Arg Lys Arg Pro
1055 1060 1065

Leu Ile Glu Thr Asn Gly Glu Thr Gly Glu Ile Val Trp Asp Lys
1070 1075 1080

Gly Arg Asp Phe Ala Thr Val Arg Lys Val Leu Ser Met Pro Gln
1085 1090 1095

Val Asn Ile Val Lys Lys Thr Glu Val Gln Thr Gly Gly Phe Ser
1100 1105 1110

Lys Glu Ser Ile Leu Pro Lys Arg Asn Ser Asp Lys Leu Ile Ala
1115 1120 1125

Arg Lys Lys Asp Trp Asp Pro Lys Lys Tyr Gly Gly Phe Asp Ser
1130 1135 1140

Pro Thr Val Ala Tyr Ser Val Leu Val Val Ala Lys Val Glu Lys
1145 1150 1155

Gly Lys Ser Lys Lys Leu Lys Ser Val Lys Glu Leu Leu Gly Ile

1160

1165

1170

Thr Ile Met Glu Arg Ser Ser Phe Glu Lys Asn Pro Ile Asp Phe
 1175 1180 1185

Leu Glu Ala Lys Gly Tyr Lys Glu Val Lys Lys Asp Leu Ile Ile
 1190 1195 1200

Lys Leu Pro Lys Tyr Ser Leu Phe Glu Leu Glu Asn Gly Arg Lys
 1205 1210 1215

Arg Met Leu Ala Ser Ala Gly Glu Leu Gln Lys Gly Asn Glu Leu
 1220 1225 1230

Ala Leu Pro Ser Lys Tyr Val Asn Phe Leu Tyr Leu Ala Ser His
 1235 1240 1245

Tyr Glu Lys Leu Lys Gly Ser Pro Glu Asp Asn Glu Gln Lys Gln
 1250 1255 1260

Leu Phe Val Glu Gln His Lys His Tyr Leu Asp Glu Ile Ile Glu
 1265 1270 1275

Gln Ile Ser Glu Phe Ser Lys Arg Val Ile Leu Ala Asp Ala Asn
 1280 1285 1290

Leu Asp Lys Val Leu Ser Ala Tyr Asn Lys His Arg Asp Lys Pro
 1295 1300 1305

Ile Arg Glu Gln Ala Glu Asn Ile Ile His Leu Phe Thr Leu Thr
 1310 1315 1320

Asn Leu Gly Ala Pro Ala Ala Phe Lys Tyr Phe Asp Thr Thr Ile
 1325 1330 1335

Asp Arg Lys Arg Tyr Thr Ser Thr Lys Glu Val Leu Asp Ala Thr
 1340 1345 1350

Leu Ile His Gln Ser Ile Thr Gly Leu Tyr Glu Thr Arg Ile Asp
 1355 1360 1365

Leu	Ser	Gln	Leu	Gly	Gly	Asp	Lys	Arg	Pro	Ala	Ala	Thr	Lys	Lys
1370						1375					1380			

Ala	Gly	Gln	Ala	Lys	Lys	Lys	Lys
1385						1390	