

SEQUENCE LISTING

<110> The Florey Institute of Neuroscience and Mental Health

<120> COMPOSITIONS AND METHODS FOR TREATING DISORDERS ASSOCIATED WITH LOSS-OF-FUNCTION MUTATIONS IN SYNGAP1

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<140> PCT/AU2021/050436

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<150> 2020901507

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Leu Ala Gly Arg His Phe Thr Glu Gln Trp Tyr Pro Val Thr Leu Pro
355 360 365

Thr Gly Ser Gly Gly Ser Gly Gly Met Gly Ser Gly Gly Gly Gly Gly
370 375 380

Ser Gly Gly Gly Ser Gly Gly Lys Gly Lys Gly Gly Cys Pro Ala Val
385 390 395 400

Arg Leu Lys Ala Arg Tyr Gln Thr Met Ser Ile Leu Pro Met Glu Leu
405 410 415

Tyr Lys Glu Phe Ala Glu Tyr Val Thr Asn His Tyr Arg Met Leu Cys
420 425 430

Ala Val Leu Glu Pro Ala Leu Asn Val Lys Gly Lys Glu Glu Val Ala
435 440 445

Ser Ala Leu Val His Ile Leu Gln Ser Thr Gly Lys Ala Lys Asp Phe
450 455 460

Leu Ser Asp Met Ala Met Ser Glu Val Asp Arg Phe Met Glu Arg Glu
465 470 475 480

His Leu Ile Phe Arg Glu Asn Thr Leu Ala Thr Lys Ala Ile Glu Glu
485 490 495

Tyr Met Arg Leu Ile Gly Gln Lys Tyr Leu Lys Asp Ala Ile Gly Glu
500 505 510

Phe Ile Arg Ala Leu Tyr Glu Ser Glu Glu Asn Cys Glu Val Asp Pro
515 520 525

Ile Lys Cys Thr Ala Ser Ser Leu Ala Glu His Gln Ala Asn Leu Arg
530 535 540

Met Cys Cys Glu Leu Ala Leu Cys Lys Val Val Asn Ser His Cys Val
545 550 555 560

Phe Pro Arg Glu Leu Lys Glu Val Phe Ala Ser Trp Arg Leu Arg Cys
565 570 575

Ala Glu Arg Gly Arg Glu Asp Ile Ala Asp Arg Leu Ile Ser Ala Ser
580 585 590

Leu Phe Leu Arg Phe Leu Cys Pro Ala Ile Met Ser Pro Ser Leu Phe
595 600 605

Gly Leu Met Gln Glu Tyr Pro Asp Glu Gln Thr Ser Arg Thr Leu Thr
610 615 620

Leu Ile Ala Lys Val Ile Gln Asn Leu Ala Asn Phe Ser Lys Phe Thr
625 630 635 640

Ser Lys Glu Asp Phe Leu Gly Phe Met Asn Glu Phe Leu Glu Leu Glu
645 650 655

Trp Gly Ser Met Gln Gln Phe Leu Tyr Glu Ile Ser Asn Leu Asp Thr
660 665 670

Leu Thr Asn Ser Ser Ser Phe Glu Gly Tyr Ile Asp Leu Gly Arg Glu
675 680 685

Leu Ser Thr Leu His Ala Leu Leu Trp Glu Val Leu Pro Gln Leu Ser
690 695 700

Lys Glu Ala Leu Leu Lys Leu Gly Pro Leu Pro Arg Leu Leu Asn Asp
705 710 715 720

Ile Ser Thr Ala Leu Arg Asn Pro Asn Ile Gln Arg Gln Pro Ser Arg
725 730 735

Gln Ser Glu Arg Pro Arg Pro Gln Pro Val Val Leu Arg Gly Pro Ser
740 745 750

Ala Glu Met Gln Gly Tyr Met Met Arg Asp Leu Asn Ser Ser Ile Asp
755 760 765

Leu Gln Ser Phe Met Ala Arg Gly Leu Asn Ser Ser Met Asp Met Ala
770 775 780

Arg Leu Pro Ser Pro Thr Lys Glu Lys Pro Pro Pro Pro Pro Gly
785 790 795 800

Gly Gly Lys Asp Leu Phe Tyr Val Ser Arg Pro Pro Leu Ala Arg Ser
805 810 815

Ser Pro Ala Tyr Cys Thr Ser Ser Ser Asp Ile Thr Glu Pro Glu Gln
820 825 830

Lys Met Leu Ser Val Asn Lys Ser Val Ser Met Leu Asp Leu Gln Gly
835 840 845

Asp Gly Pro Gly Gly Arg Leu Asn Ser Ser Ser Val Ser Asn Leu Ala
850 855 860

Ala Val Gly Asp Leu Leu His Ser Ser Gln Ala Ser Leu Thr Ala Ala
865 870 875 880

Leu Gly Leu Arg Pro Ala Pro Ala Gly Arg Leu Ser Gln Gly Ser Gly
885 890 895

Ser Ser Ile Thr Ala Ala Gly Met Arg Leu Ser Gln Met Gly Val Thr
900 905 910

Thr Asp Gly Val Pro Ala Gln Gln Leu Arg Ile Pro Leu Ser Phe Gln
915 920 925

Asn Pro Leu Phe His Met Ala Ala Asp Gly Pro Gly Pro Pro Gly Gly
930 935 940

His Gly Gly Gly Gly Gly His Gly Pro Pro Ser Ser His His His His
945 950 955 960

His His His His His His Arg Gly Gly Glu Pro Pro Gly Asp Thr Phe
965 970 975

Ala Pro Phe His Gly Tyr Ser Lys Ser Glu Asp Leu Ser Ser Gly Val
980 985 990

Pro Lys Pro Pro Ala Ala Ser Ile Leu His Ser His Ser Tyr Ser Asp
995 1000 1005

Glu Phe Gly Pro Ser Gly Thr Asp Phe Thr Arg Arg Gln Leu Ser
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Leu Gln Asp Asn Leu Gln His Met Leu Ser Pro Pro Gln Ile Thr
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Ile Gly Pro Gln Arg Pro Ala Pro Ser Gly Pro Gly Gly Gly Ser
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Gly Gly Gly Ser Gly Gly Gly Gly Gly Gly Gln Pro Pro Pro Leu
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Gln Arg Gly Lys Ser Gln Gln Leu Thr Val Ser Ala Ala Gln Lys
1070 1075 1080

Pro Arg Pro Ser Ser Gly Asn Leu Leu Gln Ser Pro Glu Pro Ser
1085 1090 1095

Tyr Gly Pro Ala Arg Pro Arg Gln Gln Ser Leu Ser Lys Glu Gly
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Ser Ile Gly Gly Ser Gly Gly Ser Gly Gly Gly Gly Gly Gly Gly
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Leu Lys Pro Ser Ile Thr Lys Gln His Ser Gln Thr Pro Ser Thr
1130 1135 1140

Leu Asn Pro Thr Met Pro Ala Ser Glu Arg Thr Val Ala Trp Val
1145 1150 1155

Ser Asn Met Pro His Leu Ser Ala Asp Ile Glu Ser Ala His Ile
1160 1165 1170

Glu Arg Glu Glu Tyr Lys Leu Lys Glu Tyr Ser Lys Ser Met Asp
1175 1180 1185

Glu Ser Arg Leu Asp Arg Val Lys Glu Tyr Glu Glu Glu Ile His
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Tyr Glu Arg Arg Leu Leu Ser Gln Glu Glu Gln Thr Ser Lys Ile
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Leu Met Gln Tyr Gln Ala Arg Leu Glu Gln Ser Glu Lys Arg Leu
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Arg Gln Gln Gln Ala Glu Lys Asp Ser Gln Ile Lys Ser Ile Ile
1250 1255 1260

Gly Arg Leu Met Leu Val Glu Glu Glu Leu Arg Arg Asp His Pro
1265 1270 1275

Ala Met Ala Glu Pro Leu Pro Glu Pro Lys Lys Arg Leu Leu Asp
1280 1285 1290

Ala Gln Glu Arg Gln Leu Pro Pro Leu Gly Pro Thr Asn Pro Arg
1295 1300 1305

Val Thr Leu Ala Pro Pro Trp Asn Gly Leu Ala Pro Pro Ala Pro
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Asn Thr Ala Asp His
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Gln Tyr Val His Ser Pro Tyr Asp Arg Pro Gly Trp Asn Pro Arg Phe
35 40 45

Cys Ile Ile Ser Gly Asn Gln Leu Leu Met Leu Asp Glu Asp Glu Ile
50 55 60

His Pro Leu Leu Ile Arg Asp Arg Arg Ser Glu Ser Ser Arg Asn Lys
65 70 75 80

Leu Leu Arg Arg Thr Val Ser Val Pro Val Glu Gly Arg Pro His Gly
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Glu His Glu Tyr His Leu Gly Arg Ser Arg Arg Lys Ser Val Pro Gly
100 105 110

Gly Lys Gln Tyr Ser Met Glu Gly Ala Pro Ala Ala Pro Phe Arg Pro
115 120 125

Ser Gln Gly Phe Leu Ser Arg Arg Leu Lys Ser Ser Ile Lys Arg Thr
130 135 140

Lys Ser Gln Pro Lys Leu Asp Arg Thr Ser Ser Phe Arg Gln Ile Leu
145 150 155 160

Pro Arg Phe Arg Ser Ala Asp His Asp Arg Ala Arg Leu Met Gln Ser
165 170 175

Phe Lys Glu Ser His Ser His Glu Ser Leu Leu Ser Pro Ser Ser Ala
180 185 190

Ala Glu Ala Leu Glu Leu Asn Leu Asp Glu Asp Ser Ile Ile Lys Pro
195 200 205

Val His Ser Ser Ile Leu Gly Gln Glu Phe Cys Phe Glu Val Thr Thr
210 215 220

Ser Ser Gly Thr Lys Cys Phe Ala Cys Arg Ser Ala Ala Glu Arg Asp

225		230		235		240
Lys Trp Ile Glu Asn Leu Gln Arg Ala Val Lys Pro Asn Lys Asp Asn						
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Ser Arg Arg Val Asp Asn Val Leu Lys Leu Trp Ile Ile Glu Ala Arg						
		260		265		270
Glu Leu Pro Pro Lys Lys Arg Tyr Tyr Cys Glu Leu Cys Leu Asp Asp						
		275		280		285
Met Leu Tyr Ala Arg Thr Thr Ser Lys Pro Arg Ser Ala Ser Gly Asp						
		290		295		300
Thr Val Phe Trp Gly Glu His Phe Glu Phe Asn Asn Leu Pro Ala Val						
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Arg Ala Leu Arg Leu His Leu Tyr Arg Asp Ser Asp Lys Lys Arg Lys						
		325		330		335
Lys Asp Lys Ala Gly Tyr Val Gly Leu Val Thr Val Pro Val Ala Thr						
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Leu Ala Gly Arg His Phe Thr Glu Gln Trp Tyr Pro Val Thr Leu Pro						
		355		360		365
Thr Gly Ser Gly Gly Ser Gly Gly Met Gly Ser Gly Gly Gly Gly Gly						
		370		375		380
Ser Gly Gly Gly Ser Gly Gly Lys Gly Lys Gly Gly Cys Pro Ala Val						
385		390		395		400
Arg Leu Lys Ala Arg Tyr Gln Thr Met Ser Ile Leu Pro Met Glu Leu						
		405		410		415
Tyr Lys Glu Phe Ala Glu Tyr Val Thr Asn His Tyr Arg Met Leu Cys						
		420		425		430

Ala Val Leu Glu Pro Ala Leu Asn Val Lys Gly Lys Glu Glu Val Ala
435 440 445

Ser Ala Leu Val His Ile Leu Gln Ser Thr Gly Lys Ala Lys Asp Phe
450 455 460

Leu Ser Asp Met Ala Met Ser Glu Val Asp Arg Phe Met Glu Arg Glu
465 470 475 480

His Leu Ile Phe Arg Glu Asn Thr Leu Ala Thr Lys Ala Ile Glu Glu
485 490 495

Tyr Met Arg Leu Ile Gly Gln Lys Tyr Leu Lys Asp Ala Ile Gly Glu
500 505 510

Phe Ile Arg Ala Leu Tyr Glu Ser Glu Glu Asn Cys Glu Val Asp Pro
515 520 525

Ile Lys Cys Thr Ala Ser Ser Leu Ala Glu His Gln Ala Asn Leu Arg
530 535 540

Met Cys Cys Glu Leu Ala Leu Cys Lys Val Val Asn Ser His Cys Val
545 550 555 560

Phe Pro Arg Glu Leu Lys Glu Val Phe Ala Ser Trp Arg Leu Arg Cys
565 570 575

Ala Glu Arg Gly Arg Glu Asp Ile Ala Asp Arg Leu Ile Ser Ala Ser
580 585 590

Leu Phe Leu Arg Phe Leu Cys Pro Ala Ile Met Ser Pro Ser Leu Phe
595 600 605

Gly Leu Met Gln Glu Tyr Pro Asp Glu Gln Thr Ser Arg Thr Leu Thr
610 615 620

Leu Ile Ala Lys Val Ile Gln Asn Leu Ala Asn Phe Ser Lys Phe Thr

625		630		635		640									
Ser	Lys	Glu	Asp	Phe	Leu	Gly	Phe	Met	Asn	Glu	Phe	Leu	Glu	Leu	Glu
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Trp	Gly	Ser	Met	Gln	Gln	Phe	Leu	Tyr	Glu	Ile	Ser	Asn	Leu	Asp	Thr
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Leu	Thr	Asn	Ser	Ser	Ser	Phe	Glu	Gly	Tyr	Ile	Asp	Leu	Gly	Arg	Glu
		675					680					685			
Leu	Ser	Thr	Leu	His	Ala	Leu	Leu	Trp	Glu	Val	Leu	Pro	Gln	Leu	Ser
	690					695					700				
Lys	Glu	Ala	Leu	Leu	Lys	Leu	Gly	Pro	Leu	Pro	Arg	Leu	Leu	Asn	Asp
705					710					715					720
Ile	Ser	Thr	Ala	Leu	Arg	Asn	Pro	Asn	Ile	Gln	Arg	Gln	Pro	Ser	Arg
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Gln	Ser	Glu	Arg	Pro	Arg	Pro	Gln	Pro	Val	Val	Leu	Arg	Gly	Pro	Ser
			740					745					750		
Ala	Glu	Met	Gln	Gly	Tyr	Met	Met	Arg	Asp	Leu	Asn	Ser	Ser	Met	Asp
		755					760					765			
Met	Ala	Arg	Leu	Pro	Ser	Pro	Thr	Lys	Glu	Lys	Pro	Pro	Pro	Pro	Pro
	770					775					780				
Pro	Gly	Gly	Gly	Lys	Asp	Leu	Phe	Tyr	Val	Ser	Arg	Pro	Pro	Leu	Ala
785					790					795					800
Arg	Ser	Ser	Pro	Ala	Tyr	Cys	Thr	Ser	Ser	Ser	Asp	Ile	Thr	Glu	Pro
				805					810					815	
Glu	Gln	Lys	Met	Leu	Ser	Val	Asn	Lys	Ser	Val	Ser	Met	Leu	Asp	Leu
			820					825					830		

Gln Gly Asp Gly Pro Gly Gly Arg Leu Asn Ser Ser Ser Val Ser Asn
835 840 845

Leu Ala Ala Val Gly Asp Leu Leu His Ser Ser Gln Ala Ser Leu Thr
850 855 860

Ala Ala Leu Gly Leu Arg Pro Ala Pro Ala Gly Arg Leu Ser Gln Gly
865 870 875 880

Ser Gly Ser Ser Ile Thr Ala Ala Gly Met Arg Leu Ser Gln Met Gly
885 890 895

Val Thr Thr Asp Gly Val Pro Ala Gln Gln Leu Arg Ile Pro Leu Ser
900 905 910

Phe Gln Asn Pro Leu Phe His Met Ala Ala Asp Gly Pro Gly Pro Pro
915 920 925

Gly Gly His Gly Gly Gly Gly Gly His Gly Pro Pro Ser Ser His His
930 935 940

His His His His His His His His Arg Gly Gly Glu Pro Pro Gly Asp
945 950 955 960

Thr Phe Ala Pro Phe His Gly Tyr Ser Lys Ser Glu Asp Leu Ser Ser
965 970 975

Gly Val Pro Lys Pro Pro Ala Ala Ser Ile Leu His Ser His Ser Tyr
980 985 990

Ser Asp Glu Phe Gly Pro Ser Gly Thr Asp Phe Thr Arg Arg Gln Leu
995 1000 1005

Ser Leu Gln Asp Asn Leu Gln His Met Leu Ser Pro Pro Gln Ile
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Thr Ile Gly Pro Gln Arg Pro Ala Pro Ser Gly Pro Gly Gly Gly

1025

1030

1035

Ser Gly Gly Gly Ser Gly Gly Gly Gly Gly Gly Gln Pro Pro Pro
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Leu Gln Arg Gly Lys Ser Gln Gln Leu Thr Val Ser Ala Ala Gln
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Lys Pro Arg Pro Ser Ser Gly Asn Leu Leu Gln Ser Pro Glu Pro
 1070 1075 1080

Ser Tyr Gly Pro Ala Arg Pro Arg Gln Gln Ser Leu Ser Lys Glu
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Gly Ser Ile Gly Gly Ser Gly Gly Ser Gly Gly Gly Gly Gly Gly
 1100 1105 1110

Gly Leu Lys Pro Ser Ile Thr Lys Gln His Ser Gln Thr Pro Ser
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Thr Leu Asn Pro Thr Met Pro Ala Ser Glu Arg Thr Val Ala Trp
 1130 1135 1140

Val Ser Asn Met Pro His Leu Ser Ala Asp Ile Glu Ser Ala His
 1145 1150 1155

Ile Glu Arg Glu Glu Tyr Lys Leu Lys Glu Tyr Ser Lys Ser Met
 1160 1165 1170

Asp Glu Ser Arg Leu Asp Arg Glu Tyr Glu Glu Glu Ile His Ser
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Leu Lys Glu Arg Leu His Met Ser Asn Arg Lys Leu Glu Glu Tyr
 1190 1195 1200

Glu Arg Arg Leu Leu Ser Gln Glu Glu Gln Thr Ser Lys Ile Leu
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Met Gln Tyr Gln Ala Arg Leu Glu Gln Ser Glu Lys Arg Leu Arg
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Gln Gln Gln Ala Glu Lys Asp Ser Gln Ile Lys Ser Ile Ile Gly
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Arg Leu Met Leu Val Glu Glu Glu Leu Arg Arg Asp His Pro Ala
 1250 1255 1260

Met Ala Glu Pro Leu Pro Glu Pro Lys Lys Arg Leu Leu Asp Ala
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