

SEQUENCE LISTING

<110> Immunwork Inc.

<120> MOLECULAR CONSTRUCTS FOR INHIBITING THE FORMATION OF BLOOD CLOT AND/OR TREATING THROMBOSIS

<130> P2955-PCT

<150> US 62/308,349

<151> 2016-03-15

<150> PCT/CN2016/071184

<151> 2016-01-18

<150> US 62/213,012

<151> 2015-09-01

<160> 35

<170> BiSSAP 1.3

<210> 1

<211> 4

<212> PRT

<213> Artificial Sequence

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<223> filler sequence-1

<400> 1

Gly Gly Gly Ser

1

<210> 2

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Gly Ser Gly Ser

1

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<223> filler sequence-3

<400> 3

Gly Gly Ser Gly

1

<210> 4

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<400> 4

Gly Ser Gly Gly Ser

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Ser Gly Gly Ser Gly

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Gly Gly Gly Gly Ser

1

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<210> 7

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<223> filler sequence-7

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Gly Gly Ser Gly Gly Ser
1 5

<210> 8

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Gly Gly Ser Gly Gly Ser Gly
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<210> 9

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Ser Gly Ser Gly Gly Ser Gly Ser
1 5

<210> 10

<211> 9

<212> PRT

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<223> filler sequence-10

<400> 10

Gly Ser Gly Gly Ser Gly Ser Gly Ser
1 5

<210> 11

<211> 10

<212> PRT

<213> Artificial Sequence

<220>

<223> filler sequence-11

<400> 11

Ser Gly Gly Ser Gly Gly Ser Gly Ser Gly
1 5 10

<210> 12

<211> 11

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<223> filler sequence-12

<400> 12

Gly Gly Ser Gly Gly Ser Gly Gly Ser Gly Ser
1 5 10

<210> 13

<211> 12

<212> PRT

<213> Artificial Sequence

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<223> filler sequence-13

<400> 13

Ser Gly Gly Ser Gly Gly Ser Gly Ser Gly Gly Ser
1 5 10

<210> 14

<211> 13

<212> PRT

<213> Artificial Sequence

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<223> filler sequence-14

<400> 14

Gly Gly Gly Gly Ser Gly Gly Ser Gly Gly Gly Gly Ser
1 5 10

<210> 15

<211> 14

<212> PRT

<213> Artificial Sequence

<220>

<223> filler sequence-15

<400> 15

Gly Gly Gly Ser Gly Ser Gly Ser Gly Ser Gly Gly Gly Ser
1 5 10

<210> 16

<211> 15

<212> PRT

<213> Artificial Sequence

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<223> filler sequence-16

<400> 16

Ser Gly Ser Gly Gly Gly Gly Gly Ser Gly Gly Ser Gly Ser Gly
1 5 10 15

<210> 17

<211> 16

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<220>

<221> MOD_RES

<222> 1

<223> ACETYLATION

<220>

<223> polypeptide core

<400> 17

Cys Gly Gly Ser Gly Gly Ser Gly Gly Ser Lys Gly Ser Gly Ser Lys
1 5 10 15

<210> 18

<211> 19

<212> PRT

<213> Artificial Sequence

<220>

<221> MOD_RES

<222> 1

<223> ACETYLATION

<220>

<223> polypeptitde core-1

<400> 18

Cys Gly Gly Ser Gly Gly Ser Gly Gly Ser Lys Gly Ser Gly Ser Lys
1 5 10 15
Gly Ser Lys

<210> 19

<211> 31

<212> PRT

<213> Artificial Sequence

<220>

<221> MOD_RES

<222> 1

<223> ACETYLATION

<220>

<223> polypeptide core

<400> 19

Cys Gly Ser Lys Gly Ser Lys Gly Ser Lys Gly Ser Lys Gly Ser Lys
1 5 10 15
Gly Ser Lys Gly Ser Lys Gly Ser Lys Gly Ser Lys
20 25 30

<210> 20

<211> 16

<212> PRT

<213> Artificial Sequence

<220>

<221> MOD_RES

<222> 1

<223> Xaa is homopropargylglycine

<220>

<221> MOD_RES

<222> 1

<223> ACETYLATION

<220>

<223> polypeptide core

<400> 20

Xaa Gly Gly Ser Gly Gly Ser Gly Gly Ser Lys Gly Ser Gly Ser Lys
1 5 10 15

<210> 21
<211> 19
<212> PRT
<213> Artificial Sequence

<220>
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<222> 1
<223> Xaa is homopropargylglycine

<220>
<221> MOD_RES
<222> 1
<223> ACETYLATION

<220>
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<400> 21
Xaa Gly Gly Ser Gly Gly Ser Gly Gly Ser Lys Gly Ser Gly Ser Lys
1 5 10 15
Gly Ser Lys

<210> 22
<211> 19
<212> PRT
<213> Artificial Sequence

<220>
<221> MOD_RES
<222> 1
<223> Xaa is L-azidohomoalanine

<220>
<221> MOD_RES
<222> 1
<223> ACETYLATION

<220>
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<400> 22

Xaa Gly Gly Ser Gly Gly Ser Gly Gly Ser Lys Gly Ser Gly Ser Lys
 1 5 10 15
 Gly Ser Lys

<210> 23
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 <223> Xaa is homopropargylglycine

<220>
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 <222> 1
 <223> ACETYLATION

<220>
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<400> 23
 Xaa Gly Gly Ser Gly Gly Ser Gly Gly Ser Lys Gly Ser Gly Ser Lys
 1 5 10 15
 Gly Ser Gly Ser Cys
 20

<210> 24
 <211> 7
 <212> PRT
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<220>
 <221> MOD_RES
 <222> 1
 <223> ACETYLATION

<220>
 <223> polypeptide core

<220>
 <221> MOD_RES
 <222> 2,4,6
 <223> Xaa is PEGylated amino acid with two EG units

<400> 24

Cys Xaa Lys Xaa Lys Xaa Lys
1 5

<210> 25
<211> 11
<212> PRT
<213> Artificial Sequence

<220>
<221> MOD_RES
<222> 1
<223> ACETYLTATION

<220>
<223> polypeptide core

<220>
<221> MOD_RES
<222> 2,4,6,8,10
<223> Xaa is PEGylated amino acid with six EG units

<400> 25
Cys Xaa Lys Xaa Lys Xaa Lys Xaa Lys Xaa Lys
1 5 10

<210> 26
<211> 16
<212> PRT
<213> Artificial Sequence

<220>
<223> polypeptide core-2

<400> 26
Cys Gly Ser Lys Gly Ser Lys Gly Ser Lys Gly Ser Lys Gly Ser Lys
1 5 10 15

<210> 27
<211> 250
<212> PRT
<213> Artificial Sequence

<220>
<223> anti-fibrin scFv 102-10

<400> 27

Ala	Gln	Ala	Leu	Gly	Leu	Gly	Lys	His	Asn	Tyr	Cys	Arg	Asn	Pro	Asp
50						55					60				
Gly	Asp	Ala	Lys	Pro	Trp	Cys	His	Val	Leu	Lys	Asn	Arg	Arg	Leu	Thr
65					70					75					80
Trp	Glu	Tyr	Cys	Asp	Val	Pro	Ser	Cys	Ser	Thr	Cys	Gly	Leu	Arg	Gln
			85						90					95	
Tyr	Ser	Gln	Pro	Gln	Phe	Arg	Ile	Lys	Gly	Gly	Leu	Phe	Ala	Asp	Ile
			100					105						110	
Ala	Ser	His	Pro	Trp	Gln	Ala	Ala	Ile	Phe	Ala	Lys	His	Arg	Arg	Ser
		115						120				125			
Pro	Gly	Glu	Arg	Phe	Leu	Cys	Gly	Gly	Ile	Leu	Ile	Ser	Ser	Cys	Trp
						135					140				
Ile	Leu	Ser	Ala	Ala	His	Cys	Phe	Gln	Glu	Arg	Phe	Pro	Pro	His	His
145					150					155					160
Leu	Thr	Val	Ile	Leu	Gly	Arg	Thr	Tyr	Arg	Val	Val	Pro	Gly	Glu	Glu
				165					170					175	
Glu	Gln	Lys	Phe	Glu	Val	Glu	Lys	Tyr	Ile	Val	His	Lys	Glu	Phe	Asp
			180					185					190		
Asp	Asp	Thr	Tyr	Asp	Asn	Asp	Ile	Ala	Leu	Leu	Gln	Leu	Lys	Ser	Asp
		195					200					205			
Ser	Ser	Arg	Cys	Ala	Gln	Glu	Ser	Ser	Val	Val	Arg	Thr	Val	Cys	Leu
		210				215					220				
Pro	Pro	Ala	Asp	Leu	Gln	Leu	Pro	Asp	Trp	Thr	Glu	Cys	Glu	Leu	Ser
225					230					235					240
Gly	Tyr	Gly	Lys	His	Glu	Ala	Leu	Ser	Pro	Phe	Tyr	Ser	Glu	Arg	Leu
				245					250					255	
Lys	Glu	Ala	His	Val	Arg	Leu	Tyr	Pro	Ser	Ser	Arg	Cys	Thr	Ser	Gln
			260					265					270		
His	Leu	Leu	Asn	Arg	Thr	Val	Thr	Asp	Asn	Met	Leu	Cys	Ala	Gly	Asp
		275					280					285			
Thr	Arg	Ser	Gly	Gly	Pro	Gln	Ala	Asn	Leu	His	Asp	Ala	Cys	Gln	Gly
						295					300				
Asp	Ser	Gly	Gly	Pro	Leu	Val	Cys	Leu	Asn	Asp	Gly	Arg	Met	Thr	Leu
305					310					315					320
Val	Gly	Ile	Ile	Ser	Trp	Gly	Leu	Gly	Cys	Gly	Gln	Lys	Asp	Val	Pro
				325					330					335	
Gly	Val	Tyr	Thr	Lys	Val	Thr	Asn	Tyr	Leu	Asp	Trp	Ile	Arg	Asp	Asn
			340					345					350		
Met	Arg	Pro	Gly	Gly	Gly	Gly	Ser	Gly	Gly	Gly	Gly	Ser	Cys		
		355					360					365			

<210> 29

<211> 244

<212> PRT

<213> Artificial Sequence

<220>

<223> Phage-displayed anti-fibrin scFv D10

<400> 29

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

<210> 30

<211> 599

<212> PRT

<213> Artificial Sequence

<220>

<223> 2-chain IgG4.Fc fusion protein containing reteplase

<400> 30

Ser Tyr Gln Gly Asn Ser Asp Cys Tyr Phe Gly Asn Gly Ser Ala Tyr
1 5 10 15
Arg Gly Thr His Ser Leu Thr Glu Ser Gly Ala Ser Cys Leu Pro Trp
20 25 30

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

Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln
 420 425 430
 Phe Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His Gln
 435 440 445
 Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Gly
 450 455 460
 Leu Pro Ser Ser Ile Glu Lys Thr Ile Ser Lys Ala Lys Gly Gln Pro
 465 470 475 480
 Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Ser Gln Glu Glu Met Thr
 485 490 495
 Lys Asn Gln Val Ser Leu Thr Cys Leu Val Lys Gly Phe Tyr Pro Ser
 500 505 510
 Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro Glu Asn Asn Tyr
 515 520 525
 Lys Thr Thr Pro Pro Val Leu Asp Ser Asp Gly Ser Phe Phe Leu Tyr
 530 535 540
 Ser Arg Leu Thr Val Asp Lys Ser Arg Trp Gln Glu Gly Asn Val Phe
 545 550 555 560
 Ser Cys Ser Val Met His Glu Ala Leu His Asn His Tyr Thr Gln Lys
 565 570 575
 Ser Leu Ser Leu Ser Leu Gly Lys Gly Gly Gly Ser Gly Gly Gly
 580 585 590
 Gly Ser Gly Gly Gly Ser
 595

<210> 31

<211> 838

<212> PRT

<213> Artificial Sequence

<220>

<223> (reteplase)-hIgG4.Fc-(scFv α fibrin 102-10)

<400> 31

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

Ala 115	Ser 120	His 125	Pro 130	Trp 135	Gln 140	Ala 145	Ile 150	Phe 155	Ala 160	Lys 165	His 170	Arg 175	Arg 180	Ser 185
Pro 130	Gly 135	Glu 140	Arg 145	Phe 150	Leu 155	Cys 160	Gly 165	Gly 170	Ile 175	Leu 180	Ile 185	Ser 190	Ser 195	Cys 200
Ile 145	Leu 150	Ser 155	Ala 160	Ala 165	His 170	Cys 175	Phe 180	Gln 185	Glu 190	Arg 195	Phe 200	Pro 205	Pro 210	His 215
Leu 165	Thr 170	Val 175	Ile 180	Leu 185	Gly 190	Arg 195	Thr 200	Tyr 205	Arg 210	Val 215	Val 220	Pro 225	Gly 230	Glu 235
Glu 180	Gln 185	Lys 190	Phe 195	Glu 200	Val 205	Glu 210	Lys 215	Tyr 220	Ile 225	Val 230	His 235	Lys 240	Glu 245	Phe 250
Asp 195	Asp 200	Thr 205	Tyr 210	Asp 215	Asn 220	Asp 225	Ile 230	Ala 235	Leu 240	Leu 245	Gln 250	Leu 255	Lys 260	Ser 265
Ser 210	Ser 215	Arg 220	Cys 225	Ala 230	Gln 235	Glu 240	Ser 245	Ser 250	Val 255	Val 260	Arg 265	Thr 270	Val 275	Cys 280
Pro 225	Pro 230	Ala 235	Asp 240	Leu 245	Gln 250	Leu 255	Pro 260	Asp 265	Trp 270	Thr 275	Glu 280	Cys 285	Glu 290	Leu 295
Gly 240	Tyr 245	Gly 250	Lys 255	His 260	Glu 265	Ala 270	Leu 275	Ser 280	Pro 285	Phe 290	Tyr 295	Ser 300	Glu 305	Arg 310
Lys 255	Glu 260	Ala 265	His 270	Val 275	Arg 280	Leu 285	Tyr 290	Pro 295	Ser 300	Ser 305	Arg 310	Cys 315	Thr 320	Ser 325
His 270	Leu 275	Leu 280	Asn 285	Arg 290	Thr 295	Val 300	Thr 305	Asp 310	Asn 315	Met 320	Leu 325	Cys 330	Ala 335	Gly 340
Thr 290	Arg 295	Ser 300	Gly 305	Gly 310	Pro 315	Gln 320	Ala 325	Asn 330	Leu 335	His 340	Asp 345	Ala 350	Cys 355	Gln 360
Asp 305	Ser 310	Gly 315	Gly 320	Pro 325	Leu 330	Val 335	Cys 340	Leu 345	Asn 350	Asp 355	Gly 360	Arg 365	Met 370	Thr 375
Val 320	Gly 325	Ile 330	Ile 335	Ser 340	Trp 345	Gly 350	Leu 355	Gly 360	Cys 365	Gly 370	Gln 375	Lys 380	Asp 385	Val 390
Gly 335	Val 340	Tyr 345	Thr 350	Lys 355	Val 360	Thr 365	Asn 370	Tyr 375	Leu 380	Asp 385	Trp 390	Ile 395	Arg 400	Asp 405
Met 350	Arg 355	Pro 360	Ala 365	Ser 370	Gly 375	Gly 380	Ser 385	Pro 390	Pro 395	Cys 400	Pro 405	Ser 410	Cys 415	Pro 420
Pro 365	Glu 370	Phe 375	Leu 380	Gly 385	Gly 390	Pro 395	Ser 400	Val 405	Phe 410	Leu 415	Phe 420	Pro 425	Pro 430	Lys 435
Lys 385	Asp 390	Thr 395	Leu 400	Met 405	Ile 410	Ser 415	Arg 420	Thr 425	Pro 430	Glu 435	Val 440	Thr 445	Cys 450	Val 455
Val 400	Asp 405	Val 410	Ser 415	Gln 420	Glu 425	Asp 430	Pro 435	Glu 440	Val 445	Gln 450	Phe 455	Asn 460	Trp 465	Tyr 470
Asp 415	Gly 420	Val 425	Glu 430	Val 435	His 440	Asn 445	Ala 450	Lys 455	Thr 460	Lys 465	Pro 470	Arg 475	Glu 480	Gln 485
Phe 430	Asn 435	Ser 440	Thr 445	Tyr 450	Arg 455	Val 460	Val 465	Ser 470	Val 475	Leu 480	Thr 485	Val 490	Leu 495	His 500
Asp 445	Trp 450	Leu 455	Asn 460	Gly 465	Lys 470	Glu 475	Tyr 480	Lys 485	Cys 490	Lys 495	Val 500	Ser 505	Asn 510	Lys 515
Leu 465	Pro 470	Ser 475	Ser 480	Ile 485	Glu 490	Lys 495	Thr 500	Ile 505	Ser 510	Lys 515	Ala 520	Lys 525	Gly 530	Gln 535
Arg 480	Glu 485	Pro 490	Gln 495	Val 500	Tyr 505	Thr 510	Leu 515	Pro 520	Pro 525	Ser 530	Gln 535	Glu 540	Glu 545	Met 550

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

<210> 32
 <211> 611
 <212> PRT

<213> Artificial Sequence

<220>

<223> (reteplase)-(scFv α fibrin 102-10)

<400> 32

Ser	Tyr	Gln	Gly	Asn	Ser	Asp	Cys	Tyr	Phe	Gly	Asn	Gly	Ser	Ala	Tyr
1				5					10					15	
Arg	Gly	Thr	His	Ser	Leu	Thr	Glu	Ser	Gly	Ala	Ser	Cys	Leu	Pro	Trp
		20						25					30		
Asn	Ser	Met	Ile	Leu	Ile	Gly	Lys	Val	Tyr	Thr	Ala	Gln	Asn	Pro	Ser
		35					40					45			
Ala	Gln	Ala	Leu	Gly	Leu	Gly	Lys	His	Asn	Tyr	Cys	Arg	Asn	Pro	Asp
	50					55					60				
Gly	Asp	Ala	Lys	Pro	Trp	Cys	His	Val	Leu	Lys	Asn	Arg	Arg	Leu	Thr
65					70					75					80
Trp	Glu	Tyr	Cys	Asp	Val	Pro	Ser	Cys	Ser	Thr	Cys	Gly	Leu	Arg	Gln
				85					90					95	
Tyr	Ser	Gln	Pro	Gln	Phe	Arg	Ile	Lys	Gly	Gly	Leu	Phe	Ala	Asp	Ile
		100						105					110		
Ala	Ser	His	Pro	Trp	Gln	Ala	Ala	Ile	Phe	Ala	Lys	His	Arg	Arg	Ser
	115					120						125			
Pro	Gly	Glu	Arg	Phe	Leu	Cys	Gly	Gly	Ile	Leu	Ile	Ser	Ser	Cys	Trp
	130					135					140				
Ile	Leu	Ser	Ala	Ala	His	Cys	Phe	Gln	Glu	Arg	Phe	Pro	Pro	His	His
145					150					155					160
Leu	Thr	Val	Ile	Leu	Gly	Arg	Thr	Tyr	Arg	Val	Val	Pro	Gly	Glu	Glu
				165					170					175	
Glu	Gln	Lys	Phe	Glu	Val	Glu	Lys	Tyr	Ile	Val	His	Lys	Glu	Phe	Asp
			180					185					190		
Asp	Asp	Thr	Tyr	Asp	Asn	Asp	Ile	Ala	Leu	Leu	Gln	Leu	Lys	Ser	Asp
		195					200					205			
Ser	Ser	Arg	Cys	Ala	Gln	Glu	Ser	Ser	Val	Val	Arg	Thr	Val	Cys	Leu
	210					215					220				
Pro	Pro	Ala	Asp	Leu	Gln	Leu	Pro	Asp	Trp	Thr	Glu	Cys	Glu	Leu	Ser
225					230					235					240
Gly	Tyr	Gly	Lys	His	Glu	Ala	Leu	Ser	Pro	Phe	Tyr	Ser	Glu	Arg	Leu
			245						250					255	
Lys	Glu	Ala	His	Val	Arg	Leu	Tyr	Pro	Ser	Ser	Arg	Cys	Thr	Ser	Gln
			260					265					270		
His	Leu	Leu	Asn	Arg	Thr	Val	Thr	Asp	Asn	Met	Leu	Cys	Ala	Gly	Asp
	275						280					285			
Thr	Arg	Ser	Gly	Gly	Pro	Gln	Ala	Asn	Leu	His	Asp	Ala	Cys	Gln	Gly
	290					295					300				
Asp	Ser	Gly	Gly	Pro	Leu	Val	Cys	Leu	Asn	Asp	Gly	Arg	Met	Thr	Leu
305					310					315					320
Val	Gly	Ile	Ile	Ser	Trp	Gly	Leu	Gly	Cys	Gly	Gln	Lys	Asp	Val	Pro
				325					330					335	

Gly Val Tyr Thr Lys Val Thr Asn Tyr Leu Asp Trp Ile Arg Asp Asn
 340 345 350
 Met Arg Pro Ala Ser Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gly
 355 360 365
 Gly Gly Gly Ser Asp Ile Gln Met Thr Gln Ser Pro Ser Ser Leu Ser
 370 375 380
 Ala Ser Leu Gly Gly Lys Val Thr Ile Thr Cys Lys Ala Ser Gln Asp
 385 390 395 400
 Ile Asn Lys Tyr Ile Ala Trp Phe Gln His Lys Pro Gly Lys Gly Pro
 405 410 415
 Arg Leu Leu Ile His Tyr Thr Ser Thr Leu Gln Pro Gly Ile Pro Ser
 420 425 430
 Arg Phe Ser Gly Ser Gly Ser Gly Arg Asp Tyr Ser Phe Ser Ile Ser
 435 440 445
 Asn Leu Glu Pro Glu Asp Leu Ala Thr Tyr Tyr Cys Leu Gln Tyr Asp
 450 455 460
 Asn Leu Thr Trp Thr Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys Arg
 465 470 475 480
 Gly Ser Thr Ser Gly Ser Gly Lys Pro Gly Ser Gly Glu Gly Ser Thr
 485 490 495
 Lys Gly Gln Ile Gln Leu Val Gln Ser Gly Pro Glu Leu Lys Lys Pro
 500 505 510
 Gly Glu Thr Val Lys Ile Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr
 515 520 525
 Asn Tyr Gly Met Asn Trp Val Lys Gln Ala Pro Gly Lys Gly Leu Lys
 530 535 540
 Trp Met Gly Trp Ile Asn Thr Tyr Thr Gly Glu Ala Thr Tyr Ala Asp
 545 550 555 560
 Asp Phe Lys Gly Arg Phe Ala Phe Ser Leu Glu Thr Ser Ala Asn Thr
 565 570 575
 Ala Tyr Val Gln Ile Asn Asn Leu Lys Asn Glu Asp Thr Ala Thr Tyr
 580 585 590
 Phe Cys Ala Arg Leu Met Asp Tyr Trp Gly Gln Gly Thr Ser Val Thr
 595 600 605
 Val Ser Ser
 610

<210> 33

<211> 771

<212> PRT

<213> Artificial Sequence

<220>

<223> 2-chain IgG4.Fc fusion protein containing TNK-tPA

<400> 33

Ser Tyr Gln Val Ile Cys Arg Asp Glu Lys Thr Gln Met Ile Tyr Gln
 1 5 10 15

Gln	His	Gln	Ser	Trp	Leu	Arg	Pro	Val	Leu	Arg	Ser	Asn	Arg	Val	Glu	
			20					25					30			
Tyr	Cys	Trp	Cys	Asn	Ser	Gly	Arg	Ala	Gln	Cys	His	Ser	Val	Pro	Val	
		35					40					45				
Lys	Ser	Cys	Ser	Glu	Pro	Arg	Cys	Phe	Asn	Gly	Gly	Thr	Cys	Gln	Gln	
	50					55					60					
Ala	Leu	Tyr	Phe	Ser	Asp	Phe	Val	Cys	Gln	Cys	Pro	Glu	Gly	Phe	Ala	
65					70					75					80	
Gly	Lys	Cys	Cys	Glu	Ile	Asp	Thr	Arg	Ala	Thr	Cys	Tyr	Glu	Asp	Gln	
				85					90					95		
Gly	Ile	Ser	Tyr	Arg	Gly	Asn	Trp	Ser	Thr	Ala	Glu	Ser	Gly	Ala	Glu	
			100					105					110			
Cys	Thr	Gln	Trp	Asn	Ser	Ser	Ala	Leu	Ala	Gln	Lys	Pro	Tyr	Ser	Gly	
		115					120					125				
Arg	Arg	Pro	Asp	Ala	Ile	Arg	Leu	Gly	Leu	Gly	Asn	His	Asn	Tyr	Cys	
	130					135					140					
Arg	Asn	Pro	Asp	Arg	Asp	Ser	Lys	Pro	Trp	Cys	Tyr	Val	Phe	Lys	Ala	
145					150					155					160	
Gly	Lys	Tyr	Ser	Ser	Glu	Phe	Cys	Ser	Thr	Pro	Ala	Cys	Ser	Glu	Gly	
				165					170					175		
Asn	Ser	Asp	Cys	Tyr	Phe	Gly	Asn	Gly	Ser	Ala	Tyr	Arg	Gly	Thr	His	
			180					185					190			
Ser	Leu	Thr	Glu	Ser	Gly	Ala	Ser	Cys	Leu	Pro	Trp	Asn	Ser	Met	Ile	
		195					200					205				
Leu	Ile	Gly	Lys	Val	Tyr	Thr	Ala	Gln	Asn	Pro	Ser	Ala	Gln	Ala	Leu	
	210					215					220					
Gly	Leu	Gly	Lys	His	Asn	Tyr	Cys	Arg	Asn	Pro	Asp	Gly	Asp	Ala	Lys	
225					230					235					240	
Pro	Trp	Cys	His	Val	Leu	Lys	Asn	Arg	Arg	Leu	Thr	Trp	Glu	Tyr	Cys	
			245						250					255		
Asp	Val	Pro	Ser	Cys	Ser	Thr	Cys	Gly	Leu	Arg	Gln	Tyr	Ser	Gln	Pro	
			260					265					270			
Gln	Phe	Arg	Ile	Lys	Gly	Gly	Leu	Phe	Ala	Asp	Ile	Ala	Ser	His	Pro	
		275					280					285				
Trp	Gln	Ala	Ala	Ala	Ala	Ala	Lys	His	Arg	Arg	Ser	Pro	Gly	Glu	Arg	
	290					295					300					
Phe	Leu	Cys	Gly	Gly	Ile	Leu	Ile	Ser	Ser	Cys	Trp	Ile	Leu	Ser	Ala	
305					310					315					320	
Ala	His	Cys	Phe	Gln	Glu	Arg	Phe	Pro	Pro	His	His	Leu	Thr	Val	Ile	
				325						330				335		
Leu	Gly	Arg	Thr	Tyr	Arg	Val	Val	Pro	Gly	Glu	Glu	Glu	Gln	Lys	Phe	
			340					345					350			
Glu	Val	Glu	Lys	Tyr	Ile	Val	His	Lys	Glu	Phe	Asp	Asp	Asp	Thr	Tyr	
		355					360					365				
Asp	Asn	Asp	Ile	Ala	Leu	Leu	Gln	Leu	Lys	Ser	Asp	Ser	Ser	Arg	Cys	
	370					375					380					
Ala	Gln	Glu	Ser	Ser	Val	Val	Arg	Thr	Val	Cys	Leu	Pro	Pro	Ala	Asp	
385					390					395					400	

Leu Gln Leu Pro Asp Trp Thr Glu Cys Glu Leu Ser Gly Tyr Gly Lys
 405 410 415
 His Glu Ala Leu Ser Pro Phe Tyr Ser Glu Arg Leu Lys Glu Ala His
 420 425 430
 Val Arg Leu Tyr Pro Ser Ser Arg Cys Thr Ser Gln His Leu Leu Asn
 435 440 445
 Arg Thr Val Thr Asp Asn Met Leu Cys Ala Gly Asp Thr Arg Ser Gly
 450 455 460
 Gly Pro Gln Ala Asn Leu His Asp Ala Cys Gln Gly Asp Ser Gly Gly
 465 470 475 480
 Pro Leu Val Cys Leu Asn Asp Gly Arg Met Thr Leu Val Gly Ile Ile
 485 490 495
 Ser Trp Gly Leu Gly Cys Gly Gln Lys Asp Val Pro Gly Val Tyr Thr
 500 505 510
 Lys Val Thr Asn Tyr Leu Asp Trp Ile Arg Asp Asn Met Arg Pro Ala
 515 520 525
 Ser Gly Gly Ser Pro Pro Cys Pro Ser Cys Pro Ala Pro Glu Phe Leu
 530 535 540
 Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro Lys Asp Thr Leu
 545 550 555 560
 Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val Val Asp Val Ser
 565 570 575
 Gln Glu Asp Pro Glu Val Gln Phe Asn Trp Tyr Val Asp Gly Val Glu
 580 585 590
 Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln Phe Asn Ser Thr
 595 600 605
 Tyr Arg Val Val Ser Val Leu Thr Val Leu His Gln Asp Trp Leu Asn
 610 615 620
 Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Gly Leu Pro Ser Ser
 625 630 635 640
 Ile Glu Lys Thr Ile Ser Lys Ala Lys Gly Gln Pro Arg Glu Pro Gln
 645 650 655
 Val Tyr Thr Leu Pro Pro Ser Gln Glu Glu Met Thr Lys Asn Gln Val
 660 665 670
 Ser Leu Thr Cys Leu Val Lys Gly Phe Tyr Pro Ser Asp Ile Ala Val
 675 680 685
 Glu Trp Glu Ser Asn Gly Gln Pro Glu Asn Asn Tyr Lys Thr Thr Pro
 690 695 700
 Pro Val Leu Asp Ser Asp Gly Ser Phe Phe Leu Tyr Ser Arg Leu Thr
 705 710 715 720
 Val Asp Lys Ser Arg Trp Gln Glu Gly Asn Val Phe Ser Cys Ser Val
 725 730 735
 Met His Glu Ala Leu His Asn His Tyr Thr Gln Lys Ser Leu Ser Leu
 740 745 750
 Ser Leu Gly Lys Gly Gly Gly Gly Ser Gly Gly Gly Gly Ser Gly Gly
 755 760 765
 Gly Gly Ser
 770

<210> 34
 <211> 1015
 <212> PRT
 <213> Artificial Sequence

 <220>
 <223> (TNK-tPA)-hIgG4.Fc-(scFv α fibrin D10)

<400> 34
 Ser Tyr Gln Val Ile Cys Arg Asp Glu Lys Thr Gln Met Ile Tyr Gln
 1 5 10 15
 Gln His Gln Ser Trp Leu Arg Pro Val Leu Arg Ser Asn Arg Val Glu
 20 25 30
 Tyr Cys Trp Cys Asn Ser Gly Arg Ala Gln Cys His Ser Val Pro Val
 35 40 45
 Lys Ser Cys Ser Glu Pro Arg Cys Phe Asn Gly Gly Thr Cys Gln Gln
 50 55 60
 Ala Leu Tyr Phe Ser Asp Phe Val Cys Gln Cys Pro Glu Gly Phe Ala
 65 70 75 80
 Gly Lys Cys Cys Glu Ile Asp Thr Arg Ala Thr Cys Tyr Glu Asp Gln
 85 90 95
 Gly Ile Ser Tyr Arg Gly Asn Trp Ser Thr Ala Glu Ser Gly Ala Glu
 100 105 110
 Cys Thr Gln Trp Asn Ser Ser Ala Leu Ala Gln Lys Pro Tyr Ser Gly
 115 120 125
 Arg Arg Pro Asp Ala Ile Arg Leu Gly Leu Gly Asn His Asn Tyr Cys
 130 135 140
 Arg Asn Pro Asp Arg Asp Ser Lys Pro Trp Cys Tyr Val Phe Lys Ala
 145 150 155 160
 Gly Lys Tyr Ser Ser Glu Phe Cys Ser Thr Pro Ala Cys Ser Glu Gly
 165 170 175
 Asn Ser Asp Cys Tyr Phe Gly Asn Gly Ser Ala Tyr Arg Gly Thr His
 180 185 190
 Ser Leu Thr Glu Ser Gly Ala Ser Cys Leu Pro Trp Asn Ser Met Ile
 195 200 205
 Leu Ile Gly Lys Val Tyr Thr Ala Gln Asn Pro Ser Ala Gln Ala Leu
 210 215 220
 Gly Leu Gly Lys His Asn Tyr Cys Arg Asn Pro Asp Gly Asp Ala Lys
 225 230 235 240
 Pro Trp Cys His Val Leu Lys Asn Arg Arg Leu Thr Trp Glu Tyr Cys
 245 250 255
 Asp Val Pro Ser Cys Ser Thr Cys Gly Leu Arg Gln Tyr Ser Gln Pro
 260 265 270
 Gln Phe Arg Ile Lys Gly Gly Leu Phe Ala Asp Ile Ala Ser His Pro
 275 280 285
 Trp Gln Ala Ala Ala Ala Lys His Arg Arg Ser Pro Gly Glu Arg
 290 295 300

Phe Leu Cys Gly Gly Ile Leu Ile Ser Ser Cys Trp Ile Leu Ser Ala
305 310 315 320
Ala His Cys Phe Gln Glu Arg Phe Pro Pro His His Leu Thr Val Ile
325 330 335
Leu Gly Arg Thr Tyr Arg Val Val Pro Gly Glu Glu Glu Gln Lys Phe
340 345 350
Glu Val Glu Lys Tyr Ile Val His Lys Glu Phe Asp Asp Asp Thr Tyr
355 360 365
Asp Asn Asp Ile Ala Leu Leu Gln Leu Lys Ser Asp Ser Ser Arg Cys
370 375 380
Ala Gln Glu Ser Ser Val Val Arg Thr Val Cys Leu Pro Pro Ala Asp
385 390 395 400
Leu Gln Leu Pro Asp Trp Thr Glu Cys Glu Leu Ser Gly Tyr Gly Lys
405 410 415
His Glu Ala Leu Ser Pro Phe Tyr Ser Glu Arg Leu Lys Glu Ala His
420 425 430
Val Arg Leu Tyr Pro Ser Ser Arg Cys Thr Ser Gln His Leu Leu Asn
435 440 445
Arg Thr Val Thr Asp Asn Met Leu Cys Ala Gly Asp Thr Arg Ser Gly
450 455 460
Gly Pro Gln Ala Asn Leu His Asp Ala Cys Gln Gly Asp Ser Gly Gly
465 470 475 480
Pro Leu Val Cys Leu Asn Asp Gly Arg Met Thr Leu Val Gly Ile Ile
485 490 495
Ser Trp Gly Leu Gly Cys Gly Gln Lys Asp Val Pro Gly Val Tyr Thr
500 505 510
Lys Val Thr Asn Tyr Leu Asp Trp Ile Arg Asp Asn Met Arg Pro Ala
515 520 525
Ser Gly Gly Ser Pro Pro Cys Pro Ser Cys Pro Ala Pro Glu Phe Leu
530 535 540
Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro Lys Asp Thr Leu
545 550 555 560
Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val Val Asp Val Ser
565 570 575
Gln Glu Asp Pro Glu Val Gln Phe Asn Trp Tyr Val Asp Gly Val Glu
580 585 590
Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln Phe Asn Ser Thr
595 600 605
Tyr Arg Val Val Ser Val Leu Thr Val Leu His Gln Asp Trp Leu Asn
610 615 620
Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Gly Leu Pro Ser Ser
625 630 635 640
Ile Glu Lys Thr Ile Ser Lys Ala Lys Gly Gln Pro Arg Glu Pro Gln
645 650 655
Val Tyr Thr Leu Pro Pro Ser Gln Glu Glu Met Thr Lys Asn Gln Val
660 665 670
Ser Leu Thr Cys Leu Val Lys Gly Phe Tyr Pro Ser Asp Ile Ala Val
675 680 685

Glu	Trp	Glu	Ser	Asn	Gly	Gln	Pro	Glu	Asn	Asn	Tyr	Lys	Thr	Thr	Pro
690						695					700				
Pro	Val	Leu	Asp	Ser	Asp	Gly	Ser	Phe	Phe	Leu	Tyr	Ser	Arg	Leu	Thr
705					710					715					720
Val	Asp	Lys	Ser	Arg	Trp	Gln	Glu	Gly	Asn	Val	Phe	Ser	Cys	Ser	Val
				725					730					735	
Met	His	Glu	Ala	Leu	His	Asn	His	Tyr	Thr	Gln	Lys	Ser	Leu	Ser	Leu
			740					745					750		
Ser	Leu	Gly	Lys	Gly	Gly	Gly	Gly	Ser	Gly	Gly	Gly	Gly	Ser	Gly	Gly
		755				760						765			
Gly	Gly	Ser	Asp	Ile	Gln	Met	Thr	Gln	Ser	Pro	Ser	Ser	Leu	Ser	Ala
		770				775					780				
Ser	Val	Gly	Asp	Arg	Val	Thr	Ile	Thr	Cys	Arg	Ala	Ser	Gln	Asp	Val
785					790					795					800
Gly	Phe	Tyr	Val	Ala	Trp	Tyr	Gln	Gln	Lys	Pro	Gly	Lys	Ala	Pro	Lys
				805					810					815	
Leu	Leu	Ile	Ser	Tyr	Pro	Ser	Gly	Leu	Tyr	Ser	Gly	Val	Pro	Ser	Arg
			820					825					830		
Phe	Ser	Gly	Ser	Gly	Ser	Gly	Thr	Asp	Phe	Thr	Leu	Thr	Ile	Ser	Ser
		835					840					845			
Leu	Gln	Pro	Glu	Asp	Phe	Ala	Thr	Tyr	Tyr	Cys	Gln	Gln	Tyr	Tyr	Asp
		850				855					860				
Tyr	Pro	Ile	Thr	Phe	Gly	Gln	Gly	Thr	Lys	Val	Glu	Ile	Lys	Arg	Gly
865					870					875					880
Ser	Thr	Ser	Gly	Ser	Gly	Lys	Pro	Gly	Ser	Gly	Glu	Gly	Ser	Thr	Lys
				885					890					895	
Gly	Glu	Val	Gln	Leu	Val	Glu	Ser	Gly	Gly	Gly	Leu	Val	Gln	Pro	Gly
			900					905					910		
Gly	Ser	Leu	Arg	Leu	Ser	Cys	Ala	Ala	Ser	Gly	Phe	Thr	Ile	Asn	Asp
		915					920					925			
Phe	Ser	Ile	His	Trp	Val	Arg	Gln	Ala	Pro	Gly	Lys	Gly	Leu	Glu	Trp
		930				935					940				
Val	Ala	Gly	Ile	Trp	Pro	Tyr	Gly	Gly	Tyr	Thr	Phe	Tyr	Ala	Asp	Ser
945					950					955					960
Val	Lys	Gly	Arg	Phe	Thr	Ile	Ser	Ala	Asp	Thr	Ser	Lys	Asn	Thr	Ala
				965					970					975	
Tyr	Leu	Arg	Met	Asn	Ser	Leu	Arg	Ala	Glu	Asp	Thr	Ala	Val	Tyr	Tyr
			980					985					990		
Cys	Ala	Arg	Phe	Gly	Tyr	Tyr	Ser	Phe	Gly	Asp	Tyr	Trp	Gly	Gln	Gly
		995					1000					1005			
Thr	Leu	Val	Thr	Val	Ser	Ser									
	1010					1015									

<210> 35

<211> 788

<212> PRT

<213> Artificial Sequence

<220>

<223> (TNK-tPA)-(scFv α fibrin)

<400> 35

Ser	Tyr	Gln	Val	Ile	Cys	Arg	Asp	Glu	Lys	Thr	Gln	Met	Ile	Tyr	Gln
1				5					10					15	
Gln	His	Gln	Ser	Trp	Leu	Arg	Pro	Val	Leu	Arg	Ser	Asn	Arg	Val	Glu
			20					25					30		
Tyr	Cys	Trp	Cys	Asn	Ser	Gly	Arg	Ala	Gln	Cys	His	Ser	Val	Pro	Val
		35				40						45			
Lys	Ser	Cys	Ser	Glu	Pro	Arg	Cys	Phe	Asn	Gly	Gly	Thr	Cys	Gln	Gln
	50					55				60					
Ala	Leu	Tyr	Phe	Ser	Asp	Phe	Val	Cys	Gln	Cys	Pro	Glu	Gly	Phe	Ala
65					70					75					80
Gly	Lys	Cys	Cys	Glu	Ile	Asp	Thr	Arg	Ala	Thr	Cys	Tyr	Glu	Asp	Gln
				85					90					95	
Gly	Ile	Ser	Tyr	Arg	Gly	Asn	Trp	Ser	Thr	Ala	Glu	Ser	Gly	Ala	Glu
			100					105					110		
Cys	Thr	Gln	Trp	Asn	Ser	Ser	Ala	Leu	Ala	Gln	Lys	Pro	Tyr	Ser	Gly
		115					120					125			
Arg	Arg	Pro	Asp	Ala	Ile	Arg	Leu	Gly	Leu	Gly	Asn	His	Asn	Tyr	Cys
	130					135					140				
Arg	Asn	Pro	Asp	Arg	Asp	Ser	Lys	Pro	Trp	Cys	Tyr	Val	Phe	Lys	Ala
145					150					155					160
Gly	Lys	Tyr	Ser	Ser	Glu	Phe	Cys	Ser	Thr	Pro	Ala	Cys	Ser	Glu	Gly
				165					170					175	
Asn	Ser	Asp	Cys	Tyr	Phe	Gly	Asn	Gly	Ser	Ala	Tyr	Arg	Gly	Thr	His
			180					185					190		
Ser	Leu	Thr	Glu	Ser	Gly	Ala	Ser	Cys	Leu	Pro	Trp	Asn	Ser	Met	Ile
	195						200						205		
Leu	Ile	Gly	Lys	Val	Tyr	Thr	Ala	Gln	Asn	Pro	Ser	Ala	Gln	Ala	Leu
	210					215						220			
Gly	Leu	Gly	Lys	His	Asn	Tyr	Cys	Arg	Asn	Pro	Asp	Gly	Asp	Ala	Lys
225					230					235					240
Pro	Trp	Cys	His	Val	Leu	Lys	Asn	Arg	Arg	Leu	Thr	Trp	Glu	Tyr	Cys
			245						250					255	
Asp	Val	Pro	Ser	Cys	Ser	Thr	Cys	Gly	Leu	Arg	Gln	Tyr	Ser	Gln	Pro
		260						265					270		
Gln	Phe	Arg	Ile	Lys	Gly	Gly	Leu	Phe	Ala	Asp	Ile	Ala	Ser	His	Pro
		275					280					285			
Trp	Gln	Ala	Ala	Ala	Ala	Ala	Lys	His	Arg	Arg	Ser	Pro	Gly	Glu	Arg
	290					295					300				
Phe	Leu	Cys	Gly	Gly	Ile	Leu	Ile	Ser	Ser	Cys	Trp	Ile	Leu	Ser	Ala
305					310					315					320
Ala	His	Cys	Phe	Gln	Glu	Arg	Phe	Pro	Pro	His	His	Leu	Thr	Val	Ile
				325					330					335	
Leu	Gly	Arg	Thr	Tyr	Arg	Val	Val	Pro	Gly	Glu	Glu	Glu	Gln	Lys	Phe
			340					345					350		

Glu	Val	Glu	Lys	Tyr	Ile	Val	His	Lys	Glu	Phe	Asp	Asp	Asp	Thr	Tyr
		355					360					365			
Asp	Asn	Asp	Ile	Ala	Leu	Leu	Gln	Leu	Lys	Ser	Asp	Ser	Ser	Arg	Cys
	370					375					380				
Ala	Gln	Glu	Ser	Ser	Val	Val	Arg	Thr	Val	Cys	Leu	Pro	Pro	Ala	Asp
385					390					395					400
Leu	Gln	Leu	Pro	Asp	Trp	Thr	Glu	Cys	Glu	Leu	Ser	Gly	Tyr	Gly	Lys
				405					410					415	
His	Glu	Ala	Leu	Ser	Pro	Phe	Tyr	Ser	Glu	Arg	Leu	Lys	Glu	Ala	His
			420					425					430		
Val	Arg	Leu	Tyr	Pro	Ser	Ser	Arg	Cys	Thr	Ser	Gln	His	Leu	Leu	Asn
		435					440					445			
Arg	Thr	Val	Thr	Asp	Asn	Met	Leu	Cys	Ala	Gly	Asp	Thr	Arg	Ser	Gly
	450					455					460				
Gly	Pro	Gln	Ala	Asn	Leu	His	Asp	Ala	Cys	Gln	Gly	Asp	Ser	Gly	Gly
465					470					475					480
Pro	Leu	Val	Cys	Leu	Asn	Asp	Gly	Arg	Met	Thr	Leu	Val	Gly	Ile	Ile
				485					490					495	
Ser	Trp	Gly	Leu	Gly	Cys	Gly	Gln	Lys	Asp	Val	Pro	Gly	Val	Tyr	Thr
			500					505					510		
Lys	Val	Thr	Asn	Tyr	Leu	Asp	Trp	Ile	Arg	Asp	Asn	Met	Arg	Pro	Ala
		515					520					525			
Ser	Gly	Gly	Gly	Gly	Ser	Gly	Gly	Gly	Gly	Ser	Gly	Gly	Gly	Gly	Ser
	530					535					540				
Asp	Ile	Gln	Met	Thr	Gln	Ser	Pro	Ser	Ser	Leu	Ser	Ala	Ser	Val	Gly
545					550					555					560
Asp	Arg	Val	Thr	Ile	Thr	Cys	Arg	Ala	Ser	Gln	Asp	Val	Gly	Phe	Tyr
				565				570						575	
Val	Ala	Trp	Tyr	Gln	Gln	Lys	Pro	Gly	Lys	Ala	Pro	Lys	Leu	Leu	Ile
			580					585					590		
Ser	Tyr	Pro	Ser	Gly	Leu	Tyr	Ser	Gly	Val	Pro	Ser	Arg	Phe	Ser	Gly
		595					600					605			
Ser	Gly	Ser	Gly	Thr	Asp	Phe	Thr	Leu	Thr	Ile	Ser	Ser	Leu	Gln	Pro
	610					615					620				
Glu	Asp	Phe	Ala	Thr	Tyr	Tyr	Cys	Gln	Gln	Tyr	Tyr	Asp	Tyr	Pro	Ile
625					630					635					640
Thr	Phe	Gly	Gln	Gly	Thr	Lys	Val	Glu	Ile	Lys	Arg	Gly	Ser	Thr	Ser
				645				650						655	
Gly	Ser	Gly	Lys	Pro	Gly	Ser	Gly	Glu	Gly	Ser	Thr	Lys	Gly	Glu	Val
			660					665					670		
Gln	Leu	Val	Glu	Ser	Gly	Gly	Gly	Leu	Val	Gln	Pro	Gly	Gly	Ser	Leu
		675					680					685			
Arg	Leu	Ser	Cys	Ala	Ala	Ser	Gly	Phe	Thr	Ile	Asn	Asp	Phe	Ser	Ile
	690					695					700				
His	Trp	Val	Arg	Gln	Ala	Pro	Gly	Lys	Gly	Leu	Glu	Trp	Val	Ala	Gly
705					710					715					720
Ile	Trp	Pro	Tyr	Gly	Gly	Tyr	Thr	Phe	Tyr	Ala	Asp	Ser	Val	Lys	Gly
				725					730					735	

Arg	Phe	Thr	Ile	Ser	Ala	Asp	Thr	Ser	Lys	Asn	Thr	Ala	Tyr	Leu	Arg
			740					745					750		
Met	Asn	Ser	Leu	Arg	Ala	Glu	Asp	Thr	Ala	Val	Tyr	Tyr	Cys	Ala	Arg
		755					760						765		
Phe	Gly	Tyr	Tyr	Ser	Phe	Gly	Asp	Tyr	Trp	Gly	Gln	Gly	Thr	Leu	Val
	770					775					780				
Thr	Val	Ser	Ser												
785															