

P15-205 PCT seq protocol_ST25.txt
SEQUENCE LISTING

<110> Merck Patent GmbH
<120> Bi-specific antibodies for enhanced tumor selectivity and inhibition and uses thereof
<130> P15/205 PCT - TF
<160> 52
<170> PatentIn version 3.5
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Gly Gln Pro Phe Arg Pro Glu Val His Leu Leu Pro Pro Ser Arg Glu
1 5 10 15

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

Tyr Pro Xaa Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro Glu
35 40 45

Asn Asn Tyr Lys Thr Thr Pro Ser Arg Gln Glu Pro Ser Gln Gly Thr
50 55 60

Thr Thr Phe Ala Val Thr Ser Lys Leu Thr Xaa Asp Lys Ser Arg Trp
65 70 75 80

Gln Gln Gly Asn Val Phe Ser Cys Ser Val Met His Glu Ala Leu His
85 90 95

Asn His Tyr Thr Gln Lys Xaa Ile Ser Leu
100 105

<210> 2
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<220>
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<223> X is V or T

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<222> (103)..(103)
<223> X is T or S

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Gly Gln Pro Phe Arg Pro Glu Val His Leu Leu Pro Pro Ser Arg Glu
1 5 10 15

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

Tyr Pro Xaa Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro Glu
35 40 45

Asn Asn Tyr Lys Thr Thr Pro Ser Arg Gln Glu Pro Ser Gln Gly Thr
50 55 60

Thr Thr Phe Ala Val Thr Ser Lys Leu Thr Xaa Asp Lys Ser Arg Trp
65 70 75 80

Gln Gln Gly Asn Val Phe Ser Cys Ser Val Met His Glu Ala Leu His
85 90 95

Asn His Tyr Thr Gln Lys Xaa Ile Ser Leu
100 105

<210> 3
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 <223> X may be any amino acid

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<400> 3

Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Pro Ser Glu
 1 5 10 15

Glu Leu Ala Leu Asn Glu Xaa Val Thr Leu Thr Cys Leu Val Lys Gly
 20 25 30

Phe Tyr Pro Ser Asp Ile Ala Val Glu Trp Leu Gln Gly Ser Gln Glu
 35 40 45

Leu Pro Arg Glu Lys Tyr Leu Thr Trp Xaa Pro Val Xaa Asp Ser Asp
 50 55 60

Gly Ser Xaa Phe Leu Tyr Ser Ile Leu Arg Val Xaa Ala Xaa Asp Trp
 65 70 75 80

Lys Lys Gly Asp Thr Phe Ser Cys Ser Val Met His Glu Ala Leu His
 85 90 95

Asn His Tyr Thr Gln Lys Ser Leu Asp Arg
 100 105

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<223> X is A or T

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<221> X

<222> (61)..(61)

<223> X is L, V, D or T

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<223> X is F, A, D, E, G, H, K, N, P, Q, R, S or T

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<221> X

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<223> X is A or T

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<222> (78)..(78)

<223> X is E or D

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Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Pro Ser Glu
1 5 10 15

Glu Leu Ala Leu Asn Glu Xaa Val Thr Leu Thr Cys Leu Val Lys Gly
20 25 30

Phe Tyr Pro Ser Asp Ile Ala Val Glu Trp Leu Gln Gly Ser Gln Glu
35 40 45

Leu Pro Arg Glu Lys Tyr Leu Thr Trp Xaa Pro Val Xaa Asp Ser Asp
50 55 60

Gly Ser Xaa Phe Leu Tyr Ser Ile Leu Arg Val Xaa Ala Xaa Asp Trp
65 70 75 80

Lys Lys Gly Asp Thr Phe Ser Cys Ser Val Met His Glu Ala Leu His
85 90 95

Asn His Tyr Thr Gln Lys Ser Leu Asp Arg
100 105

<210> 5

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Gly Gln Pro Phe Arg Pro Glu Val His Leu Leu Pro Pro Ser Arg Glu
1 5 10 15

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

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Tyr Pro Lys Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro Glu
35 40 45

Asn Asn Tyr Lys Thr Thr Pro Ser Arg Gln Glu Pro Ser Gln Gly Thr
50 55 60

Thr Thr Phe Ala Val Thr Ser Lys Leu Thr Val Asp Lys Ser Arg Trp
65 70 75 80

Gln Gln Gly Asn Val Phe Ser Cys Ser Val Met His Glu Ala Leu His
85 90 95

Asn His Tyr Thr Gln Lys Thr Ile Ser Leu
100 105

<210> 6
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<223> GA-SEED

<400> 6

Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Pro Ser Glu
1 5 10 15

Glu Leu Ala Leu Asn Glu Leu Val Thr Leu Thr Cys Leu Val Lys Gly
20 25 30

Phe Tyr Pro Ser Asp Ile Ala Val Glu Trp Leu Gln Gly Ser Gln Glu
35 40 45

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

Gly Ser Phe Phe Leu Tyr Ser Ile Leu Arg Val Ala Ala Glu Asp Trp
65 70 75 80

Lys Lys Gly Asp Thr Phe Ser Cys Ser Val Met His Glu Ala Leu His
85 90 95

Asn His Tyr Thr Gln Lys Ser Leu Asp Arg
100 105

<210> 7
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<223> AG-SEED

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<400> 7

Gly Gln Pro Phe Glu Pro Glu Val His Thr Leu Pro Pro Ser Arg Glu
1 5 10 15

Glu Met Thr Lys Asn Gln Val Ser Leu Thr Cys Leu Val Arg Gly Phe
20 25 30

Tyr Pro Ser Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro Glu
35 40 45

Asn Asn Tyr Lys Thr Thr Pro Ser Arg Leu Glu Pro Ser Gln Gly Thr
50 55 60

Thr Thr Phe Ala Val Thr Ser Lys Leu Thr Val Asp Lys Ser Arg Trp
65 70 75 80

Gln Gln Gly Asn Val Phe Ser Cys Ser Val Met His Glu Ala Leu His
85 90 95

Asn His Tyr Thr Gln Lys Ser Leu Ser Leu
100 105

<210> 8

<211> 106

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<220>

<223> GA-SEED

<400> 8

Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Pro Ser Glu
1 5 10 15

Glu Leu Ala Leu Asn Asn Gln Val Thr Leu Thr Cys Leu Val Lys Gly
20 25 30

Phe Tyr Pro Ser Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro
35 40 45

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

Gly Ser Phe Phe Leu Tyr Ser Ile Leu Arg Val Asp Ala Ser Arg Trp
65 70 75 80

Gln Gln Gly Asn Val Phe Ser Cys Ser Val Met His Glu Ala Leu His
85 90 95

Asn His Tyr Thr Gln Lys Ser Leu Ser Leu
100 105

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<210> 9
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 <223> hu225 VL sequence

<400> 9

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 Gly Thr Asn
 20 25 30

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

Lys Tyr Ala Ser Glu Ser Ile Ser Gly Val Pro Ser Arg Phe Ser Gly
 50 55 60

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

Glu Asp Val Ala Thr Tyr Tyr Cys Gln Gln Asn Asn Asn Trp Pro Thr
 85 90 95

Thr Phe Gly Gln Gly Thr Lys Val Glu Ile Lys
 100 105

<210> 10
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<220>
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 <222> (27)..(27)
 <223> X is Q, M or R

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 <222> (31)..(31)
 <223> X is T or V

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 <223> X is N, R, F or M

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 <222> (49)..(49)
 <223> X is K or M

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 <222> (50)..(50)
 <223> X is Y or W

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 <222> (92)..(92)
 <223> X is N, R, S, Y, or M

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 <222> (93)..(93)
 <223> X is N, or E

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 <222> (96)..(96)
 <223> X is T or N

<400> 10

Xaa 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 Xaa Ser Ile Gly Xaa Xaa
 20 25 30

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

Xaa Xaa Ala Ser Glu Ser Ile Ser Gly Val Pro Ser Arg Phe Ser Gly
 50 55 60

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

Glu Asp Val Ala Thr Tyr Tyr Cys Gln Gln Asn Xaa Xaa Trp Pro Xaa
 85 90 95

Thr Phe Gly Gln Gly Thr Lys Val Glu Ile Lys
 100 105

<210> 11
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<220>
 <223> hu225 VH sequence

<400> 11

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

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Ser Val Lys Val Ser Cys Lys Ala Ser Gly Phe Ser Leu Thr Asn Tyr
20 25 30

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

Gly Val Ile Trp Ser Gly Gly Asn Thr Asp Tyr Asn Thr Pro Phe Thr
50 55 60

Ser Arg Val Thr Ile Thr Ser Asp Lys Ser Thr Ser Thr Ala Tyr Met
65 70 75 80

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

Arg Ala Leu Thr Tyr Tyr Asp Tyr Glu Phe Ala Tyr Trp Gly Gln Gly
100 105 110

Thr Leu Val Thr Val Ser Ser
115

<210> 12
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<223> X is S or W

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<222> (53)..(53)
<223> X is S, A, E, H, Q, or R

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<223> X is N, I, L, M, K, or R

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<222> (58)..(58)
<223> X is D, E, Q, or R

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<222> (74)..(74)
<223> X is S or W

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<222> (100)..(100)
<223> X is T, F, W, or D

<220>

<221> X
 <222> (101)..(101)
 <223> X is Y or W

<220>
 <221> X
 <222> (103)..(103)
 <223> X is D or E

<400> 12

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

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Phe Xaa Leu Thr Asn Tyr
 20 25 30

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

Gly Val Ile Trp Xaa Gly Gly Xaa Thr Xaa Tyr Asn Thr Pro Phe Thr
 50 55 60

Ser Arg Val Thr Ile Thr Ser Asp Lys Xaa Thr Ser Thr Ala Tyr Met
 65 70 75 80

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

Arg Ala Leu Xaa Xaa Tyr Xaa Tyr Glu Phe Ala Tyr Trp Gly Gln Gly
 100 105 110

Thr Leu Val Thr Val Ser Ser
 115

<210> 13
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<220>
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<400> 13

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 Ser Ala Ser Ser Ser Val Thr Tyr Met
 20 25 30

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

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

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

Asp Ile Ala Thr Tyr Tyr Cys Gln Gln Trp Ser Ser His Ile Phe Thr
85 90 95

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

<210> 14
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<212> PRT
<213> Artificial sequence

<220>
<223> hu425 VH sequence

<400> 14

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

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Ser His
20 25 30

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

Gly Glu Phe Asn Pro Ser Asn Gly Arg Thr Asn Tyr Asn Glu Lys Phe
50 55 60

Lys Ser Lys Ala Thr Met Thr Val Asp Thr Ser Thr Asn Thr Ala Tyr
65 70 75 80

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

Ala Ser Arg Asp Tyr Asp Tyr Asp Gly Arg Tyr Phe Asp Tyr Trp Gly
100 105 110

Gln Gly Thr Leu Val Thr Val Ser Ser
115 120

<210> 15
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<400> 15

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

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Thr Ala Arg Ile Thr Cys Gly Gly Asn Asn Ile Ala Arg Lys Ser Val
 20 25 30
 His Trp Tyr Gln Gln Arg Pro Gly Gln Ala Pro Val Leu Val Val Tyr
 35 40 45
 Asp Asp Ser Asp Arg Pro Ser Gly Ile Pro Glu Arg Phe Ser Gly Ser
 50 55 60
 Asn Ser Gly Asn Thr Ala Thr Leu Thr Ile Ser Arg Val Glu Ala Gly
 65 70 75 80
 Asp Glu Ala Asp Tyr Tyr Cys Gln Val Trp Asp Ser Ser Ser Asp Gln
 85 90 95
 Leu Tyr Val Phe Gly Thr Gly Thr Lys Val Thr Val Leu Gly Gln Pro
 100 105 110
 Lys Ala Gly
 115

<210> 16
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 <223> c-MET binder A12 VH sequence
 <400> 16

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

115

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<400> 17

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

Arg Val Thr Ile Ser Cys Phe Gly Ser Ser Ser Asn Val Gly Val Asn
20 25 30

Thr Val Asn Trp Tyr Arg Gln Leu Pro Gly Thr Ala Pro Lys Leu Leu
35 40 45

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

Gly Ser Lys Ser Gly Thr Ser Ala Ser Leu Ala Ile Thr Gly Leu Gln
65 70 75 80

Ala Glu Asp Glu Gly Asp Tyr Tyr Cys Gln Ser Tyr Asp Ser Ser Leu
85 90 95

Ser Asp Val Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu Gly Gln
100 105 110

Pro Lys Ala Gly
115

<210> 18
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<400> 18

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

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

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

Ser Ala Ile Ser Gly Ser Gly Gly Ser Thr Tyr Tyr Ala Asp Ser Val
Seite 13

50

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

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

Ala Lys Asp Arg Arg Ile Thr His Thr Tyr Trp Gly Gln Gly Thr Leu
100 105 110

Val Thr Val Ser Ser
115

<210> 19
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<212> PRT
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<400> 19

Gln Ser Val Leu Thr Gln Pro Pro Ser Val Ser Val Ala Pro Gly Lys
1 5 10 15

Thr Ala Arg Ile Thr Cys Gly Gly Asn Asn Ile Gly Ser Lys Ser Val
20 25 30

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

Asp Asp Ser Asp Arg Pro Ser Gly Ile Pro Glu Arg Phe Ser Gly Ser
50 55 60

Asn Ser Gly Asn Thr Ala Thr Leu Thr Ile Ser Arg Val Glu Ala Gly
65 70 75 80

Asp Glu Ala Asp Tyr Tyr Cys Gln Val Trp Asp Ser Ser Ser Asp Leu
85 90 95

Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu Gly Gln Pro Lys
100 105 110

Ala Gly

<210> 20
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P15-205 PCT seq protocol_ST25.txt

<400> 20

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

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Gly Tyr
20 25 30

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

Gly Trp Ile Asn Pro Asn Ser Gly Gly Thr Asn Tyr Ala Gln Lys Phe
50 55 60

Gln Gly Arg Val Thr Met Thr Arg Asp Thr Ser Ile Ser Thr Ala Tyr
65 70 75 80

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

Ala Ser Leu Asn Phe Pro Asp Ile Ala Val Ala Gly Tyr Gly Asp Tyr
100 105 110

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

<210> 21

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<400> 21

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

Ser Ile Thr Ile Ser Cys Thr Gly Thr Ser Ser Asp Val Gly Gly Tyr
20 25 30

Asp Tyr Val Ser Trp Tyr Gln Gln His Pro Gly Lys Ala Pro Gln Leu
35 40 45

Met Ile Tyr Asp Val Thr Ser Arg Pro Ser Glu Val Ser Asn Arg Phe
50 55 60

Ser Gly Ser Lys Ser Gly Thr Ser Ala Ser Leu Ala Ile Thr Gly Leu
65 70 75 80

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

P15-205 PCT seq protocol_ST25.txt

Ser Thr Leu Val Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu Gly
100 105 110

Gln Pro Lys Ala Gly
115

<210> 22
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<220>
<223> c-MET binder E07 VH sequence
<400> 22

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

Thr Leu Ser Leu Thr Cys Ala Val Ser Gly Gly Ser Ile Ser Ser Ser
20 25 30

Asn Trp Trp Ser Trp Val Arg Gln Pro Pro Gly Lys Gly Leu Glu Trp
35 40 45

Ile Gly Glu Ile Tyr His Ser Gly Ser Thr Asn Tyr Asn Pro Ser Leu
50 55 60

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

Leu Lys Leu Ser Ser Val Thr Ala Ala Asp Thr Ala Val Tyr Tyr Cys
85 90 95

Ala Arg Gly Gly Ser Gly Tyr Asp Phe Asp Tyr Trp Gly Gln Gly Thr
100 105 110

Leu Val Thr Val Ser Ser
115

<210> 23
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<212> PRT
<213> Artificial Sequence

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<223> c-MET binder G02 VL sequence
<400> 23

Gln Ser Ala Leu Thr Gln Pro Pro Ser Ala Ser Gly Ser Pro Gly Gln
1 5 10 15

Ser Val Thr Ile Ser Cys Thr Gly Thr Ser Ser Asp Val Gly Gly Tyr
20 25 30

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

Met Ile Tyr Glu Val Ser Lys Arg Pro Ser Gly Val Pro Asp Arg Phe
50 55 60

Ser Gly Ser Lys Ser Gly Asn Thr Ala Ser Leu Thr Ile Ser Gly Leu
65 70 75 80

Gln Ala Glu Asp Glu Ala Asp Tyr Tyr Cys Trp Ser Tyr Ala Gly Ser
85 90 95

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

Pro Lys Ala Gly
115

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<400> 24

Glu Val Gln Leu Val Glu Thr Gly Gly Gly Val Val Gln Pro Gly Arg
1 5 10 15

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

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

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

Ala Lys Met Gly Tyr Gly Thr Gly Ala Phe Asp Ile Trp Gly Gln Gly
100 105 110

Thr Met Val Thr Val Ser Ser
115

<210> 25

P15-205 PCT seq protocol_ST25.txt

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<400> 25

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

Arg Val Thr Ile Ser Cys Ser Gly Ser Ser Ser Asn Ile Gly Ser Asn
20 25 30

Tyr Val Tyr Trp Tyr Gln His Leu Pro Gly Thr Ala Pro Lys Leu Leu
35 40 45

Ile Tyr Ser Asn Asn Gln Arg Pro Ser Gly Val Pro Asp Arg Phe Ser
50 55 60

Gly Ser Lys Ser Gly Thr Ser Ala Ser Leu Ala Ile Ser Gly Leu Gln
65 70 75 80

Ser Glu Asp Glu Gly Asp Tyr Tyr Cys Ala Ser Trp Asp Asp Asn Leu
85 90 95

Asn Ala His Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Val Ser
100 105 110

Gln Pro Lys Ala Gly
115

<210> 26

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<400> 26

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

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asn Leu
20 25 30

Asp Ile Asn Trp Val Arg Gln Ala Ser Gly Gln Gly Leu Glu Trp Met
35 40 45

Gly Trp Met Asn Pro Asn Ser Gly Asn Thr Gly Tyr Ala Gln Lys Phe
50 55 60

Gln Gly Arg Val Thr Met Thr Arg Ser Thr Ser Val Ser Thr Ala Tyr

1 P15-205 PCT seq protocol_ST25.txt 15
 5 10 15
 Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Ile Tyr
 20 25 30
 Gly Met His Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Leu
 35 40 45
 Ala Phe Ile Arg His Asp Gly Gly Asn Asn Phe Tyr Ala 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
 Met Gln Met Ser Ser Leu Arg Pro Glu Asp Thr Ala Val Tyr Tyr Cys
 85 90 95
 Ala Lys Asp Phe Ala Met Thr Gln Trp Leu Pro Glu Arg Gly Met Asp
 100 105 110
 Val Trp Gly Gln Gly Thr Thr Val Thr Val Ser Ser
 115 120
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 Gln Leu Val Leu Thr Gln Ser Pro Ser Val Ser Val Ala Pro Gly Lys
 1 5 10 15
 Thr Ala Arg Ile Thr Cys Gly Gly Asn Asn Ile Arg Asn Val Gly Val
 20 25 30
 His Trp Tyr Gln Lys Lys Pro Gly Gln Ala Pro Ile Leu Val Val Tyr
 35 40 45
 Asp Asp Asp Asp Arg Pro Ser Gly Val Pro Glu Arg Phe Ser Gly Ser
 50 55 60
 Asn Ser Gly Asn Thr Ala Thr Leu Thr Ile Ser Arg Val Glu Ala Gly
 65 70 75 80
 Asp Glu Ala Asp Tyr Tyr Cys Gln Val Trp Asp Ser Ala Thr Asp Gln
 85 90 95
 Arg Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu Gly Gln Pro Lys
 100 105 110

Ala Gly Cys
115

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

<220>
<223> c-MET binder F06 VH sequence

<400> 30

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

Ser Ala Lys Val Ser Cys Lys Ala Ser Gly Gly Thr Phe Ser Ser Tyr
20 25 30

Ala Ile Ser Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35 40 45

Gly Gly Ile Ile Pro Ile Phe Gly Thr Ala Asn Tyr Ala Gln Lys Phe
50 55 60

Gln Gly Arg Val Thr Ile Thr Ala Asp Glu Ser Thr Ser Thr Ala Tyr
65 70 75 80

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

Ala Arg Asp Gln Arg Gly Tyr Asp Tyr Tyr Tyr Tyr Tyr Gly Met Asp
100 105 110

Val Trp Gly Gln Gly Thr Thr Val Thr Val Ser Ser
115 120

<210> 31
<211> 114
<212> PRT
<213> Artificial Sequence

<220>
<223> c-MET binder B10v5 VL sequence

<400> 31

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

Thr Ala Thr Ile Pro Cys Gly Gly Asp Ser Leu Gly Ser Lys Ile Val
20 25 30

His Trp Tyr Gln Gln Arg Pro Gly Gln Ala Pro Leu Leu Val Val Tyr
35 40 45

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

Lys Ser Gly Thr Thr Ala Thr Leu Thr Ile Ser Ser Val Glu Ala Gly
65 70 75 80

Asp Glu Ala Asp Tyr Phe Cys Gln Val Tyr Asp Tyr His Ser Asp Val
85 90 95

Glu Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu Gly Gln Pro Lys
100 105 110

Ala Ala

<210> 32
<211> 117
<212> PRT
<213> Artificial Sequence

<220>
<223> c-MET binder B10v5 VH sequence

<400> 32

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

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

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

Ser Ala Ile Ser Gly Ser Gly Gly Ser Thr Tyr Tyr Ala 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 Ser Leu Arg Ala Glu Asp Thr Ala Val Tyr Tyr Cys
85 90 95

Ala Lys Asp Arg Arg Ile Thr His Thr Tyr Trp Gly Gln Gly Thr Leu
100 105 110

Val Thr Val Ser Ser
115

<210> 33
<211> 114
<212> PRT
<213> Artificial Sequence

P15-205 PCT seq protocol_ST25.txt

<220>

<223> c-MET binder CS06 VL sequence

<400> 33

Gln Leu Val Leu Thr Gln Ser Pro Ser Val Ser Val Ala Pro Gly Lys
1 5 10 15

Thr Ala Arg Ile Thr Cys Gly Gly Asn Asn Ile Arg Asn Val Gly Val
20 25 30

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

Asp Asp Asp Asp Arg Pro Ser Gly Val Pro Glu Arg Phe Ser Gly Ser
50 55 60

Asn Ser Gly Asn Thr Ala Thr Leu Thr Ile Ser Arg Val Glu Ala Gly
65 70 75 80

Asp Glu Ala Asp Tyr Tyr Cys Gln Val Trp Asp Ser Ala Thr Asp Gln
85 90 95

Arg Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu Gly Gln Pro Lys
100 105 110

Ala Gly

<210> 34

<211> 124

<212> PRT

<213> Artificial sequence

<220>

<223> c-MET binder CS06 VH sequence

<400> 34

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

Ser Ala Lys Val Ser Cys Lys Ala Ser Gly Gly Thr Phe Ser Ser Asn
20 25 30

Ala Ile Ser Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35 40 45

Gly Gly Ile Ile Pro Ile Phe Gly Thr Ala Ile Tyr Ala Gln Lys Phe
50 55 60

Gln Gly Arg Val Thr Ile Thr Ala Asp Glu Ser Thr Ser Thr Ala Tyr
65 70 75 80

Met Glu Leu Ser Ser Leu Arg Ser Glu Asp Thr Ala Val Tyr Tyr Cys

Ala Arg Asp Gln Arg Gly Tyr Tyr Tyr Tyr Tyr Tyr Gly Met Asp
100 105 110

Val Trp Gly Gln Gly Thr Thr Val Thr Val Ser Ser
115 120

<210> 35
<211> 20
<212> PRT
<213> Artificial Sequence

<220>
<223> glycine-serine linker

<400> 35

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

Gly Gly Gly Ser
20

<210> 36
<211> 15
<212> PRT
<213> Artificial Sequence

<220>
<223> hinge 1

<400> 36

Glu Pro Lys Ser Ser Asp Lys Thr His Thr Cys Pro Pro Cys Pro
1 5 10 15

<210> 37
<211> 15
<212> PRT
<213> Artificial Sequence

<220>
<223> hinge 2

<400> 37

Glu Pro Lys Ser Cys Asp Lys Thr His Thr Cys Pro Pro Cys Pro
1 5 10 15

<210> 38
<211> 99
<212> PRT
<213> Artificial Sequence

<220>
<223> CL sequence

<400> 38

Pro Ser Val Thr Leu Phe Pro Pro Ser Ser Glu Glu Leu Gln Ala Asn
Seite 24

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

<210> 39
 <211> 98
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> CH1 sequence

<400> 39

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

<210> 40
 <211> 110
 <212> PRT

<213> Artificial Sequence

<220>

<223> CH2 domain

<400> 40

Ala Pro Glu Leu Leu Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys
1 5 10 15

Pro Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val
20 25 30

Val Val Asp Val Ser His Glu Asp Pro Glu Val Lys Phe Asn Trp Tyr
35 40 45

Val Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu
50 55 60

Gln Tyr Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His
65 70 75 80

Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys
85 90 95

Ala Leu Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys Ala Lys
100 105 110

<210> 41

<211> 110

<212> PRT

<213> Artificial Sequence

<220>

<223> CH3 domain (AG)

<400> 41

Gly Gln Pro Phe Arg Pro Glu Val His Leu Leu Pro Pro Ser Arg Glu
1 5 10 15

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

Tyr Pro Lys Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro Glu
35 40 45

Asn Asn Tyr Lys Thr Thr Pro Ser Arg Gln Glu Pro Ser Gln Gly Thr
50 55 60

Thr Thr Phe Ala Val Thr Ser Lys Leu Thr Val Asp Lys Ser Arg Trp
65 70 75 80

Gln Gln Gly Asn Val Phe Ser Cys Ser Val Met His Glu Ala Leu His
85 90 95

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Asn His Tyr Thr Gln Lys Thr Ile Ser Leu Ser Pro Gly Lys
100 105 110

<210> 42
<211> 110
<212> PRT
<213> Artificial Sequence

<220>
<223> CH3 domain (GA)

<400> 42

Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Pro Ser Glu
1 5 10 15

Glu Leu Ala Leu Asn Glu Leu Val Thr Leu Thr Cys Leu Val Lys Gly
20 25 30

Phe Tyr Pro Ser Asp Ile Ala Val Glu Trp Leu Gln Gly Ser Gln Glu
35 40 45

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

Gly Ser Phe Phe Leu Tyr Ser Ile Leu Arg Val Ala Ala Glu Asp Trp
65 70 75 80

Lys Lys Gly Asp Thr Phe Ser Cys Ser Val Met His Glu Ala Leu His
85 90 95

Asn His Tyr Thr Gln Lys Ser Leu Asp Arg Ser Pro Gly Lys
100 105 110

<210> 43
<211> 119
<212> PRT
<213> Artificial Sequence

<220>
<223> humanized C225 VH kinetic variant S58R (IMGT numbering) hu225-L

<400> 43

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

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Phe Ser Leu Thr Asn Tyr
20 25 30

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

Gly Val Ile Trp Arg Gly Gly Asn Thr Asp Tyr Asn Thr Pro Phe Thr
50 55 60

P15-205 PCT seq protocol_ST25.txt

Ser Arg Val Thr Ile Thr Ser Asp Lys Ser Thr Ser Thr Ala Tyr Met
65 70 75 80

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

Arg Ala Leu Thr Tyr Tyr Asp Tyr Glu Phe Ala Tyr Trp Gly Gln Gly
100 105 110

Thr Leu Val Thr Val Ser Ser
115

<210> 44
<211> 107
<212> PRT
<213> Artificial Sequence

<220>
<223> humanized C225 VL kinetic variant N108Y (IMGT numbering) hu225-M
<400> 44

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 Gly Thr Asn
20 25 30

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

Lys Tyr Ala Ser Glu Ser Ile Ser Gly Val Pro Ser Arg Phe Ser Gly
50 55 60

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

Glu Asp Val Ala Thr Tyr Tyr Cys Gln Gln Asn Tyr Asn Trp Pro Thr
85 90 95

Thr Phe Gly Gln Gly Thr Lys Val Glu Ile Lys
100 105

<210> 45
<211> 119
<212> PRT
<213> Artificial Sequence

<220>
<223> humanized c225 VH T109D kinetic variant (hu225-H)
<400> 45

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

P15-205 PCT seq protocol_ST25.txt

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Phe Ser Leu Thr Asn Tyr
20 25 30

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

Gly Val Ile Trp Ser Gly Gly Asn Thr Asp Tyr Asn Thr Pro Phe Thr
50 55 60

Ser Arg Val Thr Ile Thr Ser Asp Lys Ser Thr Ser Thr Ala Tyr Met
65 70 75 80

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

Arg Ala Leu Asp Tyr Tyr Asp Tyr Glu Phe Ala Tyr Trp Gly Gln Gly
100 105 110

Thr Leu Val Thr Val Ser Ser
115

<210> 46
<211> 107
<212> PRT
<213> Artificial Sequence

<220>
<223> humanized C225 VL N109E, T116N kinetic variant (hu225-H)
<400> 46

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 Gly Thr Asn
20 25 30

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

Lys Tyr Ala Ser Glu Ser Ile Ser Gly Val Pro Ser Arg Phe Ser Gly
50 55 60

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

Glu Asp Val Ala Thr Tyr Tyr Cys Gln Gln Asn Asn Glu Trp Pro Asn
85 90 95

Thr Phe Gly Gln Gly Thr Lys Val Glu Ile Lys
100 105

<210> 47

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<211> 116
<212> PRT
<213> Artificial Sequence

<220>
<223> c-Met binder B10 VL variants comprising single or multiple
      amino acid substitutions selected from V3A, T11V, T14S, R18S,
      R43Q, L45P, E74D, T85N, S86T, A90T, T92S G100A (numbering
      according to IMGT numbering)

<220>
<221> X
<222> (3)..(3)
<223> X is V or A

<220>
<221> X
<222> (10)..(10)
<223> X is T or V

<220>
<221> X
<222> (13)..(13)
<223> X is T or S

<220>
<221> X
<222> (17)..(17)
<223> X is R or S

<220>
<221> X
<222> (38)..(38)
<223> X is R or Q

<220>
<221> X
<222> (40)..(40)
<223> X is L or P

<220>
<221> X
<222> (61)..(61)
<223> X is E or D

<220>
<221> X
<222> (70)..(70)
<223> X is T or N

<220>
<221> X
<222> (71)..(71)
<223> X is S or T

<220>
<221> X
<222> (75)..(75)
<223> X is A or T

<220>
<221> X
<222> (77)..(77)
<223> X is T or S

<220>
<221> X

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P15-205 PCT seq protocol_ST25.txt

<222> (85)..(85)

<223> X is G or A

<400> 47

Gln Ser Xaa Leu Thr Gln Pro Pro Ser Xaa Ser Gly Xaa Pro Gly Gln
1 5 10 15

Xaa Val Thr Ile Ser Cys Phe Gly Ser Ser Ser Asn Val Gly Val Asn
20 25 30

Thr Val Asn Trp Tyr Xaa Gln Xaa Pro Gly Thr Ala Pro Lys Leu Leu
35 40 45

Ile Tyr Asp Asn Asn Leu Arg Pro Ser Gly Val Pro Xaa Arg Phe Ser
50 55 60

Gly Ser Lys Ser Gly Xaa Xaa Ala Ser Leu Xaa Ile Xaa Gly Leu Gln
65 70 75 80

Ala Glu Asp Glu Xaa Asp Tyr Tyr Cys Gln Ser Tyr Asp Ser Ser Leu
85 90 95

Ser Asp Val Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu Gly Gln
100 105 110

Pro Lys Ala Gly
115

<210> 48

<211> 117

<212> PRT

<213> Artificial sequence

<220>

<223> c-MET binder B10 VH kinetic variant Q6E (IMGT numbering)

<220>

<221> X

<222> (6)..(6)

<223> X is Q or E

<400> 48

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

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

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

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

P15-205 PCT seq protocol_ST25.txt

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

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

Ala Lys Asp Arg Arg Ile Thr His Thr Tyr Trp Gly Gln Gly Thr Leu
100 105 110

Val Thr Val Ser Ser
115

<210> 49
<211> 115
<212> PRT
<213> Artificial Sequence

<220>
<223> c-Met binder F06 VL sequence variants comprising single or
multiple amino acid substitutions selected from Q1S, L2Y, S7P,
K44Q, I51V, V71I (numbering acc. to IMGT numbering)

<220>
<221> X
<222> (1)..(1)
<223> X is Q or S

<220>
<221> X
<222> (2)..(2)
<223> X is L or Y

<220>
<221> X
<222> (7)..(7)
<223> X is S or P

<220>
<221> X
<222> (37)..(37)
<223> X is K or Q

<220>
<221> X
<222> (44)..(44)
<223> X is I or V

<220>
<221> X
<222> (57)..(57)
<223> X is V or I

<400> 49

Xaa Xaa Val Leu Thr Gln Xaa Pro Ser Val Ser Val Ala Pro Gly Lys
1 5 10 15

Thr Ala Arg Ile Thr Cys Gly Gly Asn Asn Ile Arg Asn Val Gly Val
20 25 30

P15-205 PCT seq protocol_ST25.txt

His Trp Tyr Gln Xaa Lys Pro Gly Gln Ala Pro Xaa Leu Val Val Tyr
35 40 45

Asp Asp Asp Asp Arg Pro Ser Gly Xaa Pro Glu Arg Phe Ser Gly Ser
50 55 60

Asn Ser Gly Asn Thr Ala Thr Leu Thr Ile Ser Arg Val Glu Ala Gly
65 70 75 80

Asp Glu Ala Asp Tyr Tyr Cys Gln Val Trp Asp Ser Ala Thr Asp Gln
85 90 95

Arg Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu Gly Gln Pro Lys
100 105 110

Ala Gly Cys
115

<210> 50
<211> 124
<212> PRT
<213> Artificial Sequence

<220>
<223> c-Met binder F06 VH variants comprising single or multiple amino acid substitutions selected from Q5V, A19V, M115I, M115L, M115V, M115A, M115F (numbering according to IMGT numbering)

<220>
<221> X
<222> (5)..(5)
<223> X is Q or V

<220>
<221> X
<222> (18)..(18)
<223> X is A or V

<220>
<221> X
<222> (111)..(111)
<223> X is M, I, L, V, A, or F

<400> 50

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

Ser Ala Lys Val Ser Cys Lys Ala Ser Gly Gly Thr Phe Ser Ser Tyr
20 25 30

Ala Ile Ser Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35 40 45

Gly Gly Ile Ile Pro Ile Phe Gly Thr Ala Asn Tyr Ala Gln Lys Phe
50 55 60

P15-205 PCT seq protocol_ST25.txt

Gln Gly Arg Val Thr Ile Thr Ala Asp Glu Ser Thr Ser Thr Ala Tyr
65 70 75 80

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

Ala Arg Asp Gln Arg Gly Tyr Asp Tyr Tyr Tyr Tyr Tyr Gly Met Asp
100 105 110

Val Trp Gly Gln Gly Thr Thr Val Thr Val Ser Ser
115 120

<210> 51
<211> 114
<212> PRT
<213> Artificial Sequence

<220>
<223> c-Met binder B10v5 VL variants comprising single or multiple amino acid substitutions selected from E1S, P2Y, E17Q, T20R, P22T, R45K, L51V, T85N, S93R, F103Y (IMGT numbering)

<220>
<221> X
<222> (1)..(1)
<223> X is E or S

<220>
<221> X
<222> (2)..(2)
<223> X is P or Y

<220>
<221> X
<222> (16)..(16)
<223> X is E or Q

<220>
<221> X
<222> (19)..(19)
<223> X is T or R

<220>
<221> X
<222> (21)..(21)
<223> X is P or T

<220>
<221> X
<222> (38)..(38)
<223> X is R or K

<220>
<221> X
<222> (44)..(44)
<223> X is L or V

<220>
<221> X
<222> (68)..(68)
<223> X is T or N

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<220>
 <221> X
 <222> (76)..(76)
 <223> X is S or R

<220>
 <221> X
 <222> (86)..(86)
 <223> X is F or Y

<400> 51

Xaa Xaa Val Leu Thr Gln Pro Pro Ser Val Ser Val Ala Pro Gly Xaa
 1 5 10 15

Thr Ala Xaa Ile Xaa Cys Gly Gly Asp Ser Leu Gly Ser Lys Ile Val
 20 25 30

His Trp Tyr Gln Gln Xaa Pro Gly Gln Ala Pro Xaa Leu Val Val Tyr
 35 40 45

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

Lys Ser Gly Xaa Thr Ala Thr Leu Thr Ile Ser Xaa Val Glu Ala Gly
 65 70 75 80

Asp Glu Ala Asp Tyr Xaa Cys Gln Val Tyr Asp Tyr His Ser Asp Val
 85 90 95

Glu Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu Gly Gln Pro Lys
 100 105 110

Ala Ala

<210> 52
 <211> 124
 <212> PRT
 <213> Artificial sequence

<220>
 <223> c-Met binder CS06 VH kinetic variants comprising including
 single, double, triple or quadruple combinations of the following
 listed mutations: Q3R, Y37N, N66I, G110S, D111.1Y, Y111.2D, S126Y
 (IMGT numbering)

<220>
 <221> X
 <222> (3)..(3)
 <223> X is Q or R

<220>
 <221> X
 <222> (32)..(32)
 <223> X is N or Y

<220>
 <221> X

<222> (59)..(59)
<223> X is N or I

<220>
<221> X
<222> (102)..(102)
<223> X is S or G

<220>
<221> X
<222> (104)..(104)
<223> X is Y or D

<220>
<221> X
<222> (105)..(105)
<223> X is Y or D

<220>
<221> X
<222> (123)..(123)
<223> X is Y or S

<400> 52

Gln Val Xaa Leu Gln Gln Ser Gly Ala Glu Val Lys Lys Pro Gly Ser
1 5 10 15

Ser Ala Lys Val Ser Cys Lys Ala Ser Gly Gly Thr Phe Ser Ser Xaa
20 25 30

Ala Ile Ser Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met
35 40 45

Gly Gly Ile Ile Pro Ile Phe Gly Thr Ala Xaa Tyr Ala Gln Lys Phe
50 55 60

Gln Gly Arg Val Thr Ile Thr Ala Asp Glu Ser Thr Ser Thr Ala Tyr
65 70 75 80

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

Ala Arg Asp Gln Arg Xaa Tyr Xaa Xaa Tyr Tyr Tyr Tyr Gly Met Asp
100 105 110

Val Trp Gly Gln Gly Thr Thr Val Thr Val Xaa Ser
115 120