

SEQUENCE LISTING

<110> MONOCLONAL ANTIBODIES THERAPEUTICS MATBIOPHARMA
 Institut National de la Santé et de la Recherche Médicale (INSERM)
 DIDIERLAURENT Denis
 CHOSE Olivier
 KADOUCHE Jean

<120> NEW ANTI-CD160 MONOCLONAL ANTIBODIES AND USES THEREOF

<130> D25265

<150> EP09305057.3

<151> 2009-01-21

<160> 29

<170> PatentIn version 3.3

<210> 1

<211> 6

<212> PRT

<213> Artificial

<220>

<223> CDR1 from light chain of anti-CD160 antibody

<400> 1

Gln Ser Ile Ser Asn His

1 5

<210> 2

<211> 9

<212> PRT

<213> Artificial

<220>

<223> CDR3 from light chain of anti-CD160 antibody

<400> 2

Gln Gln Ser Asn Ser Trp Pro Leu Thr

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<210> 3

<211> 8

<212> PRT

<213> Artificial

<220>

<223> CDR1 from heavy chain of anti-CD160 antibody

<400> 3

Gly Tyr Thr Phe Thr Asp Tyr Trp

1 5

<210> 4
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<212> PRT
<213> Artificial

<220>
<223> CDR2 from heavy chain of anti-CD160 antibody

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Ile Tyr Pro Gly Asp Asp Asp Ala
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<210> 5
<211> 14
<212> PRT
<213> Artificial

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<223> CDR3 from heavy chain of anti-CD160 antibody

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Ala Arg Arg Gly Ile Ala Ala Val Val Gly Gly Phe Asp Tyr
1 5 10

<210> 6
<211> 121
<212> PRT
<213> Mus musculus

<400> 6

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1 5 10 15

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

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

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

Arg Gly Lys Ala Thr Leu Thr Ala Asp Lys Ser Ser Ser Thr Ala Tyr
65 70 75 80

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

Ala Arg Arg Gly Ile Ala Ala Val Val Gly Gly Phe Asp Tyr Trp Gly
100 105 110

Gln Gly Thr Thr Leu Thr Val Ser Ser
115 120

<210> 7

<211> 363
<212> DNA
<213> Mus musculus

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cctggacagg gtctggaatg gattgggtct atttatcctg gagatgatga tgctaggtac 180
actcagaagt tcaggggcaa ggccacattg actgcagata aatcctccag cacagcctac 240
atgcagctca gcagcttggc atctgaagac tctgcggtct attactgtgc aagaaggggt 300
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tca 363

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<211> 108
<212> PRT
<213> Mus musculus

<400> 8
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20 25 30
Leu His Trp Tyr Gln Gln Lys Ser His Glu Ser Pro Arg Leu Leu Ile
35 40 45
Lys Tyr Ala Ser Gln Ser Ile Ser Gly Ile Pro Ser Arg Phe Ser Gly
50 55 60
Ser Gly Ser Gly Thr Asp Phe Thr Leu Ser Ile Asn Ser Val Glu Thr
65 70 75 80
Glu Asp Phe Gly Met Tyr Phe Cys Gln Gln Ser Asn Ser Trp Pro Leu
85 90 95
Thr Phe Gly Ala Gly Thr Lys Leu Glu Leu Lys Arg
100 105

<210> 9
<211> 324
<212> DNA
<213> Mus musculus

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catgagtctc caaggcttct catcaagtat gcttcccagt ccattctctgg gatcccctcc 180

aggttcagtg gcagtggatc agggacagat ttcactctca gtatcaacag tgtggagact 240
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<210> 10
<211> 114
<212> PRT
<213> homo sapiens

<400> 10

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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
Lys Val Glu Pro Lys Ser Cys Asp Lys Thr His Thr Cys Pro Pro Cys
100 105 110

Pro Cys

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<211> 345
<212> DNA
<213> homo sapiens

<400> 11

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ggactctact ccctcagcag cgtgggtgacc gtgccctcca gcagcttggg caccagacc 240
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<212> PRT
<213> homo sapiens

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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		
Lys	Val	Glu	Pro	Lys	Ser	Cys	Asp	Lys	Thr	His	Thr	Cys	Pro	Pro	Cys	
			100					105					110			
Pro	Ala	Pro	Glu	Leu	Leu	Gly	Gly	Pro	Ser	Val	Phe	Leu	Phe	Pro	Pro	
		115					120					125				
Lys	Pro	Lys	Asp	Thr	Leu	Met	Ile	Ser	Arg	Thr	Pro	Glu	Val	Thr	Cys	
	130					135					140					
Val	Val	Val	Asp	Val	Ser	His	Glu	Asp	Pro	Glu	Val	Lys	Phe	Asn	Trp	
145					150					155					160	
Tyr	Val	Asp	Gly	Val	Glu	Val	His	Asn	Ala	Lys	Thr	Lys	Pro	Arg	Glu	
			165						170					175		
Glu	Gln	Tyr	Asn	Ser	Thr	Tyr	Arg	Val	Val	Ser	Val	Leu	Thr	Val	Leu	
			180					185					190			
His	Gln	Asp	Trp	Leu	Asn	Gly	Lys	Glu	Tyr	Lys	Cys	Lys	Val	Ser	Asn	
		195					200					205				
Lys	Ala	Leu	Pro	Ala	Pro	Ile	Glu	Lys	Thr	Ile	Ser	Lys	Ala	Lys	Gly	
	210					215					220					
Gln	Pro	Arg	Glu	Pro	Gln	Val	Tyr	Thr	Leu	Pro	Pro	Ser	Arg	Asp	Glu	
225					230					235					240	
Leu	Thr	Lys	Asn	Gln	Val	Ser	Leu	Thr	Cys	Leu	Val	Lys	Gly	Phe	Tyr	
			245						250					255		
Pro	Ser	Asp	Ile	Ala	Val	Glu	Trp	Glu	Ser	Asn	Gly	Gln	Pro	Glu	Asn	
			260					265					270			
Asn	Tyr	Lys	Thr	Thr	Pro	Pro	Val	Leu	Asp	Ser	Asp	Gly	Ser	Phe	Phe	
		275					280					285				
Leu	Tyr	Ser	Lys	Leu	Thr	Val	Asp	Lys	Ser	Arg	Trp	Gln	Gln	Gly	Asn	
	290					295					300					

Val Phe Ser Cys Ser Val Met His Glu Ala Leu His Asn His Tyr Thr
 305 310 315 320

Gln Lys Ser Leu Ser Leu Ser Pro Gly Lys
 325 330

<210> 13
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 <212> DNA
 <213> homo sapiens

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 tggaactcag gcgccctgac cagcggcgctg cacaccttcc cggctgtcct acagtctctca 180
 ggactctact ccctcagcag cgtgggtgacc gtgccctcca gcagcttggg caccagacc 240
 tacatctgca acgtgaatca caagcccagc aacaccaagg tggacaagaa agttgagccc 300
 aaatcttgtg aaaaaactca cacatgccca ccgtgccag cacctgaact cctgggggga 360
 ccgtcagtct tctcttccc cccaaaaccc aaggacaccc tcatgatctc ccggaccct 420
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 agcacgtacc gtgtggtcag cgtcctcacc gtctgcacc aggactgggt gaatggcaag 600
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 aaagccaaag ggcagccccg agaaccacag gtgtacaccc tgcccccatc ccgggatgag 720
 ctgaccaaga accaggtcag cctgacctgc ctgggtcaaag gcttctatcc cagcgacatc 780
 gccgtggagt gggagagcaa tgggcagccg gagaacaact acaagaccac gcctcccgtg 840
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<210> 14
 <211> 327
 <212> PRT
 <213> homo sapiens

<400> 14

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

<210> 15
<211> 984

<212> DNA
<213> homo sapiens

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tggaactcag gcgccctgac cagcggcgctg cacaccttcc cggctgtcct acagtctctca 180
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ttctgttcc ccccaaaacc caaggacact ctcatgatct cccggacccc tgagggtcacg 420
tgcgtgggtg tggacgtgag ccaggaagac cccgaggtcc agttcaactg gtacgtggat 480
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<210> 16
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<212> PRT
<213> homo sapiens

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Gln Leu Lys Ser Gly Thr Ala Ser Val Val Cys Leu Leu Asn Asn Phe
20 25 30
Tyr Pro Arg Glu Ala Lys Val Gln Trp Lys Val Asp Asn Ala Leu Gln
35 40 45
Ser Gly Asn Ser Gln Glu Ser Val Thr Glu Gln Asp Ser Lys Asp Ser
50 55 60
Thr Tyr Ser Leu Ser Ser Thr Leu Thr Leu Ser Lys Ala Asp Tyr Glu
65 70 75 80

Lys His Lys Val Tyr Ala Cys Glu Val Thr His Gln Gly Leu Ser Ser
85 90 95

Pro Val Thr Lys Ser Phe Asn Arg Gly Glu Cys
100 105

<210> 17
<211> 324
<212> DNA
<213> homo sapiens

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tggaaggtgg ataacgccct ccaatcgggt aactcccagg agagtgtcac agagcaggac 180
agcaaggaca gcacctacag cctcagcagc accctgacgc tgagcaaagc agactacgag 240
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agcttcaaca ggggagagtg ttaa 324

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<212> PRT
<213> mus musculus

<400> 18

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20 25 30
Leu Ser Val Thr Pro Gly Asn Ser Val Ser Leu Ser Cys Arg Ala Ser
35 40 45
Gln Ser Ile Ser Asn His Leu His Trp Tyr Gln Gln Lys Ser His Glu
50 55 60
Ser Pro Arg Leu Leu Ile Lys Tyr Ala Ser Gln Ser Ile Ser Gly Ile
65 70 75 80
Pro Ser Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Ser
85 90 95
Ile Asn Ser Val Glu Thr Glu Asp Phe Gly Met Tyr Phe Cys Gln Gln
100 105 110
Ser Asn Ser Trp Pro Leu Thr Phe Gly Ala Gly Thr Lys Leu Glu Leu
115 120 125
Lys Arg
130

<210> 19
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<212> DNA
<213> mus musculus

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gtcagtccttt cctgcagggc cagccaaagt attagcaacc acctacactg gtatcaacaa 180
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ccctccaggt tcagtggcag tggatcaggg acagatttca ctctcagtat caacagtggtg 300
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<210> 20
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<213> mus musculus

<400> 20

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20 25 30
Pro Gly Ala Ser Val Lys Leu Ser Cys Lys Ala Ser Gly Tyr Thr Phe
35 40 45
Thr Asp Tyr Trp Met Gln Trp Ile Lys Gln Arg Pro Gly Gln Gly Leu
50 55 60
Glu Trp Ile Gly Ser Ile Tyr Pro Gly Asp Asp Ala Arg Tyr Thr
65 70 75 80
Gln Lys Phe Arg Gly Lys Ala Thr Leu Thr Ala Asp Lys Ser Ser Ser
85 90 95
Thr Ala Tyr Met Gln Leu Ser Ser Leu Ala Ser Glu Asp Ser Ala Val
100 105 110
Tyr Tyr Cys Ala Arg Arg Gly Ile Ala Ala Val Val Gly Gly Phe Asp
115 120 125
Tyr Trp Gly Gln Gly Thr Thr Leu Thr Val Ser Ser
130 135 140

<210> 21
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<212> DNA
<213> mus musculus

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cagctcagca gcttggcatc tgaagactct gcggtctatt actgtgcaag aaggggtatt 360
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<210> 22
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<212> PRT
<213> homo sapiens

<400> 22
Glu Ile Val Leu Thr Gln Ser Pro Ala Thr Leu Ser Leu Ser Pro Gly
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Glu Arg Ala Thr Leu Ser Cys Arg Ala Ser Gln Ser Ile Asn Asn Tyr
20 25 30
Leu Ala Trp Tyr Gln Gln Lys Asp Gly Gln Ala Pro Arg Leu Leu Ile
35 40 45
Tyr Asp Ala Ser Asn Arg Ala Thr Gly Ile Pro Thr Arg Phe Ser Gly
50 55 60
Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Asn Ser Leu Glu Pro
65 70 75 80
Glu Asp Phe Ala Val Tyr Tyr Cys Gln Gln Arg Asn Asn Trp Pro Leu
85 90 95
Thr Phe Gly Gly Gly Thr Lys Val Glu Ile Lys
100 105

<210> 23
<211> 114
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<213> homo sapiens

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Ser Val Lys Leu Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Ser Tyr
20 25 30
Trp Met His Trp Val Lys Gln Arg Pro Gly Arg Gly Leu Glu Trp Ile
35 40 45

Gly Met Ile Asp Pro Asn Ser Gly Gly Thr Lys Tyr Asn Glu Lys Phe
 50 55 60
 Lys Ser Lys Ala Thr Leu Thr Val Asp Lys Pro Ser Asn Thr Ala Tyr
 65 70 75 80
 Met Gln Leu Ser Ser Leu Thr Ser Glu Asp Ser Ala Val Tyr Tyr Cys
 85 90 95
 Thr Arg Arg Asp Met Asp Tyr Trp Gly Ala Gly Thr Thr Val Thr Val
 100 105 110
 Ser Ser

<210> 24
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<220>
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 <222> (3)..(3)
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<400> 24

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

<210> 25
 <211> 108
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 <213> artificial

<220>
 <223> humanized variable light chain anti-CD160 antibody

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<400> 25

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1 5 10 15

Xaa Xaa Xaa Ser Leu Ser Cys Arg Ala Ser Gln Ser Ile Ser Asn His
20 25 30

Leu His Trp Tyr Gln Gln Lys Ser Xaa Glu Ser Pro Arg Leu Leu Ile
35 40 45

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

Ser Gly Ser Gly Thr Asp Phe Thr Leu Ser Ile Asn Ser Val Glu Xaa
65 70 75 80

Glu Asp Phe Xaa Met Tyr Phe Cys Gln Gln Ser Asn Ser Trp Pro Leu
85 90 95

Thr Phe Gly Xaa Gly Thr Lys Leu Glu Leu Lys Arg
100 105

<210> 26
<211> 121
<212> PRT
<213> artificial

<220>
<223> humanized variable heavy chain anti-CD160 antibody

<220>
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<222> (87)..(87)
<223> X= A or T

<400> 26

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

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

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

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

Arg Gly Lys Ala Thr Leu Thr Ala Asp Lys Ser Ser Ser Thr Ala Tyr
65 70 75 80

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

Ala Arg Arg Gly Ile Ala Ala Val Val Gly Gly Phe Asp Tyr Trp Gly
100 105 110

Gln Gly Thr Thr Leu Thr Val Ser Ser
115 120

<210> 27
<211> 181
<212> PRT
<213> homo sapiens

<400> 27

Met Leu Leu Glu Pro Gly Arg Gly Cys Cys Ala Leu Ala Ile Leu Leu
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 Ala Ile Val Asp Ile Gln Ser Gly Gly Cys Ile Asn Ile Thr Ser Ser
 20 25 30
 Ala Ser Gln Glu Gly Thr Arg Leu Asn Leu Ile Cys Thr Val Trp His
 35 40 45
 Lys Lys Glu Glu Ala Glu Gly Phe Val Val Phe Leu Cys Lys Asp Arg
 50 55 60
 Ser Gly Asp Cys Ser Pro Glu Thr Ser Leu Lys Gln Leu Arg Leu Lys
 65 70 75 80
 Arg Asp Pro Gly Ile Asp Gly Val Gly Glu Ile Ser Ser Gln Leu Met
 85 90 95
 Phe Thr Ile Ser Gln Val Thr Pro Leu His Ser Gly Thr Tyr Gln Cys
 100 105 110
 Cys Ala Arg Ser Gln Lys Ser Gly Ile Arg Leu Gln Gly His Phe Phe
 115 120 125
 Ser Ile Leu Phe Thr Glu Thr Gly Asn Tyr Thr Val Thr Gly Leu Lys
 130 135 140
 Gln Arg Gln His Leu Glu Phe Ser His Asn Glu Gly Thr Leu Ser Ser
 145 150 155 160
 Gly Phe Leu Gln Glu Lys Val Trp Val Met Leu Val Thr Ser Leu Val
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 Ala Leu Gln Ala Leu
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<210> 28
 <211> 129
 <212> PRT
 <213> Mus musculus

<400> 28

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

				85						90					95				
Ile	Asn	Ser	Val	Glu	Thr	Glu	Asp	Phe	Gly	Met	Tyr	Phe	Cys	Gln	Gln				
			100					105					110						
Ser	Asn	Ser	Trp	Pro	Leu	Thr	Phe	Gly	Ala	Gly	Thr	Lys	Leu	Glu	Leu				
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Lys

<210> 29
 <211> 387
 <212> DNA
 <213> Mus musculus

<400> 29	
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gtcagtcttt cctgcagggc cagccaaagt attagcaacc acctacactg gtatcaacaa	180
aatcacatg agtctccaag gcttctcatc aagtatgctt cccagtccat ctctgggatc	240
ccctccaggt tcagtggcag tggatcaggg acagatttca ctctcagtat caacagtgtg	300
gagactgaag attttggcat gtatttctgt caacagagta acagctggcc gctcacgttc	360
ggtgctggga ccaagctgga gctaaaa	387