

LISTADO DE SECUENCIAS

<110> JSC "BIOCAD"

<120> Anticuerpos monoclonales que se unen específicamente a la región beta de la familia TRBV-9

del receptor de células T humano, y los métodos para su uso.

<130> MA042.docx

<160> 22

<170> PatentIn version 3.5

<210> 1

<211> 5

<212> PRT

<213> Artificial

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<223> La secuencia HCDR1 del anticuerpo original TRBV9-2

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Asp Tyr Leu Val His

1 5

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<223> La secuencia HCDR2 del anticuerpo original TRBV9-2

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Gly

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<223> La secuencia HCDR3 del anticuerpo original TRBV9-2

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Ser Trp Arg Arg Gly Leu Arg Gly Ile Gly Phe Asp Tyr

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<223> La secuencia LCDR1 del anticuerpo original TRBV9-2

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Lys Ala Ser Lys Ser Ile Asn Lys Tyr Leu Ala

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Gln Gln His Asn Glu Tyr Pro Pro Thr

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<223> La secuencia de nucleótidos del dominio variable
de la cadena pesada del anticuerpo original TRBV9-2

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cccggtaagg gattgaaatg gatgggatgg atcaatactt ataccggcac acctacatat 180

gcagacgatt tcgaggggcg atttggttc agtttgagg cctctgccag cacggcgaac 240

ctgcagatat cgaatctcaa gaatgaggac accgccacgt atttctgcgc tagatcttgg 300

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tcatcc 366

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<223> La secuencia de aminoácidos del dominio variable
de la cadena pesada del anticuerpo original TRBV9-2

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Gln Ile Gln Leu Val Gln Ser Gly Pro Glu Leu Arg Glu Pro Gly Glu

1 5 10 15

Ser Val Lys Leu Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asp Tyr

20 25 30

Leu Val His Trp Val Lys Gln Ala Pro Gly Lys Gly Leu Lys Trp Met

35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Thr Pro Thr Tyr Ala Asp Asp Phe

50 55 60

Glu Gly Arg Phe Val Phe Ser Leu Glu Ala Ser Ala Ser Thr Ala Asn

65 70 75 80

Leu Gln Ile Ser Asn Leu Lys Asn Glu Asp Thr Ala Thr Tyr Phe Cys

85 90 95

Ala Arg Ser Trp Arg Arg Gly Leu Arg Gly Ile Gly Phe Asp Tyr Trp

100 105 110

Gly Gln Gly Val Phe Val Thr Val Ser Ser

115

120

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<223> La secuencia de nucleótidos del dominio variable de la cadena
liviana del anticuerpo original TRBV9-2

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ggaaagccca acaagctcct gatctatgac ggctctacac tgcaatctgg cataccttcg 180

cggttttctg gctcgggggc cgggactgac ttactctta caatacgagg acttgaaccc 240

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<223> La secuencia de nucleótidos del dominio variable de la cadena
liviana del anticuerpo original TRBV9-2

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Asp Val Gln Met Thr Gln Ser Pro Tyr Asn Leu Ala Ala Ser Pro Gly

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Glu Ser Val Ser Ile Asn Cys Lys Ala Ser Lys Ser Ile Asn Lys Tyr

20 25 30

Leu Ala Trp Tyr Gln Gln Lys Pro Gly Lys Pro Asn Lys Leu Leu Ile

35 40 45

Tyr Asp Gly Ser Thr Leu Gln Ser Gly Ile Pro Ser Arg Phe Ser Gly

50 55 60

Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Arg Gly Leu Glu Pro

65 70 75 80

Glu Asp Phe Gly Leu Tyr Tyr Cys Gln Gln His Asn Glu Tyr Pro Pro

85 90 95

Thr Phe Gly Ala Gly Thr Lys Leu Glu Leu Lys

100 105

<210> 11

<211> 639

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<223> La secuencia de nucleótidos de la cadena liviana
del anticuerpo original TRBV9-2

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ggcaagccca acaagctgct gatctacgac ggctccaccc tgcagtccgg catcccctcc 180

aggttctccg gctccggctc cggcaccgac ttcaccctga ccatcagggg cctggagccc 240

gaggacttcg gcctgtacta ctgccagcag cacaacgagt accccccac cttcggcgcc 300

ggcaccaagc tggagctgaa gcgaactgtg gctgcacat ctgtcttcat cttcccgcca 360

tctgatgagc agttgaaatc tggaactgcc tctgtcgtgt gcctgctgaa taacttctat 420

cccagagagg ccaaagtaca gtggaagggt gataacgccc tccaatcggg taactcccag 480

gagagtgtca cagagcagga cagcaaggac agcacctaca gcctcagcag caccctgacg 540

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<223> La secuencia de aminoácidos de la cadena liviana
del anticuerpo original TRBV9-2

<400> 12

Asp Val Gln Met Thr Gln Ser Pro Tyr Asn Leu Ala Ala Ser Pro Gly

1 5 10 15

Glu Ser Val Ser Ile Asn Cys Lys Ala Ser Lys Ser Ile Asn Lys Tyr

20 25 30

Leu Ala Trp Tyr Gln Gln Lys Pro Gly Lys Pro Asn Lys Leu Leu Ile

35 40 45

Tyr Asp Gly Ser Thr Leu Gln Ser Gly Ile Pro Ser Arg Phe Ser Gly

50 55 60

Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Arg Gly Leu Glu Pro

65 70 75 80

Glu Asp Phe Gly Leu Tyr Tyr Cys Gln Gln His Asn Glu Tyr Pro Pro

85 90 95

Thr Phe Gly Ala Gly Thr Lys Leu Glu Leu Lys Arg Thr Val Ala Ala

100 105 110

Pro Ser Val Phe Ile Phe Pro Pro Ser Asp Glu Gln Leu Lys Ser Gly

115 120 125

Thr Ala Ser Val Val Cys Leu Leu Asn Asn Phe Tyr Pro Arg Glu Ala

130 135 140

Lys Val Gln Trp Lys Val Asp Asn Ala Leu Gln Ser Gly Asn Ser Gln

145 150 155 160

Glu Ser Val Thr Glu Gln Asp Ser Lys Asp Ser Thr Tyr Ser Leu Ser

165 170 175

Ser Thr Leu Thr Leu Ser Lys Ala Asp Tyr Glu Lys His Lys Val Tyr

180 185 190

Ala Cys Glu Val Thr His Gln Gly Leu Ser Ser Pro Val Thr Lys Ser

195 200 205

Phe Asn Arg Gly Glu

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<210> 13

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<223> La secuencia de nucleótidos de la cadena pesada
del anticuerpo original TRBV9-2

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gccgacgact tcgagggcag gttcgtgttc tccctggagg cctccgcctc caccgccaac 240

ctgcagatct ccaacctgaa gaacgaggac accgccacct acttctgcgc caggtcctgg 300

aggaggggcc tgaggggcat cggcttcgac tactggggcc agggcgtgtt cgtgaccgtg 360

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 tctgggggca cagcggccct gggctgcctg gtcaaggact acttccccga accggtgacg 480

 gtgtcgtgga actcaggcgc cctgaccagc ggcgtgcaca ccttcccggc tgcctacag 540

 tcctcaggac tctactccct cagcagcgtg gtgactgtgc cctctagcag cttgggcacc 600

 cagacctaca tctgcaacgt gaatcacaag cccagcaaca ccaagggtgga caagaaagtt 660

 gagccccga aatcttgtga caaaactcac acatgccac cgtagccagc acctgaactc 720

 ctggggggac cgtcagtctt cctcttcccc ccaaaacca aggacaccct catgatctcc 780

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 aatggcaagg agtacaagtg caaggtctcc aacaaagccc tccagcccc catcgagaaa 1020

 accatctcca aagccaaagg gcagccccga gaaccacagg tgtacaccct gccccatcc 1080

 cgggatgagc tgaccaagaa ccaggtcagc ctgacctgcc tggtaaagg cttctatccc 1140

agcgacatcg ccgtggagtg ggagagcaat gggcagccgg agaacaacta caagaccacg 1200

cctcccgtgc tggactccga cggctccttc ttctctaca gcaagctcac cgtggacaag 1260

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del anticuerpo original TRBV9-2

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Ser Val Lys Leu Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asp Tyr

20 25 30

Leu Val His Trp Val Lys Gln Ala Pro Gly Lys Gly Leu Lys Trp Met

35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Thr Pro Thr Tyr Ala Asp Asp Phe

50 55 60

Glu Gly Arg Phe Val Phe Ser Leu Glu Ala Ser Ala Ser Thr Ala Asn

65 70 75 80

Leu Gln Ile Ser Asn Leu Lys Asn Glu Asp Thr Ala Thr Tyr Phe Cys

85 90 95

Ala Arg Ser Trp Arg Arg Gly Leu Arg Gly Ile Gly Phe Asp Tyr Trp

100 105 110

Gly Gln Gly Val Phe Val Thr Val Ser Ser Ala Ser Thr Lys Gly Pro

115 120 125

Ser Val Phe Pro Leu Ala Pro Ser Ser Lys Ser Thr Ser Gly Gly Thr

130 135 140

Ala Ala Leu Gly Cys Leu Val Lys Asp Tyr Phe Pro Glu Pro Val Thr

145 150 155 160

Val Ser Trp Asn Ser Gly Ala Leu Thr Ser Gly Val His Thr Phe Pro

165 170 175

Ala Val Leu Gln Ser Ser Gly Leu Tyr Ser Leu Ser Ser Val Val Thr

180 185 190

Val Pro Ser Ser Ser Leu Gly Thr Gln Thr Tyr Ile Cys Asn Val Asn

195 200 205

His Lys Pro Ser Asn Thr Lys Val Asp Lys Lys Val Glu Pro Pro Lys

210 215 220

Ser Cys Asp Lys Thr His Thr Cys Pro Pro Cys Pro Ala Pro Glu Leu

225 230 235 240

Leu Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro Lys Asp Thr

245 250 255

Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val Val Asp Val

260 265 270

Ser His Glu Asp Pro Glu Val Lys Phe Asn Trp Tyr Val Asp Gly Val

275 280 285

Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln Tyr Asn Ser

290 295 300

Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His Gln Asp Trp Leu

305 310 315 320

Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Ala Leu Pro Ala

325 330 335

Pro Ile Glu Lys Thr Ile Ser Lys Ala Lys Gly Gln Pro Arg Glu Pro

340 345 350

Gln Val Tyr Thr Leu Pro Pro Ser Arg Asp Glu Leu Thr Lys Asn Gln

355 360 365

Val Ser Leu Thr Cys Leu Val Lys Gly Phe Tyr Pro Ser Asp Ile Ala

370 375 380

Val Glu Trp Glu Ser Asn Gly Gln Pro Glu Asn Asn Tyr Lys Thr Thr

385 390 395 400

Pro Pro Val Leu Asp Ser Asp Gly Ser Phe Phe Leu Tyr Ser Lys Leu

405 410 415

Thr Val Asp Lys Ser Arg Trp Gln Gln Gly Asn Val Phe Ser Cys Ser

420 425 430

Val Met His Glu Ala Leu His Asn His Tyr Thr Gln Lys Ser Leu Ser

435 440 445

Leu Ser Pro Gly Lys

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del anticuerpo MA-042

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ccagggtcaag gactggaatg gatgggatgg atcaatacct atacagggac acccacatat 180

gccgatgact ttgagggacg gtttgtcttc tcttgata ccagtgttc cactgctaac 240

ctccagataa gcagcctgaa ggcagaggac accgccgttt atttctgcgc ccgatcatgg 300

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<211> 122

<212> PRT

<213> Artificial

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<223> La secuencia de aminoácidos del dominio variable
de la cadena pesada del anticuerpo original MA-042

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Gln Val Gln Leu Val Gln Ser Gly Ser Glu Leu Lys Lys Pro Gly Glu

1 5 10 15

Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asp Tyr

20 25 30

Leu Val His Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met

35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Thr Pro Thr Tyr Ala Asp Asp Phe

Glu Gly Arg Phe Val Phe Ser Leu Asp Thr Ser Val Ser Thr Ala Asn

Leu Gln Ile Ser Ser Leu Lys Ala Glu Asp Thr Ala Val Tyr Phe Cys

Ala Arg Ser Trp Arg Arg Gly Leu Arg Gly Ile Gly Phe Asp Tyr Trp

Gly Gln Gly Thr Leu Val Thr Val Ser Ser

<210> 17

<211> 321

<212> ADN

<213> Artificial

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<223> La secuencia de nucleótidos de la cadena liviana variable

del anticuerpo MA-042

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ggaaagccta acaagctatt aatatacgat gggctaccc tccaatccgg ggtcccttca 180

cgattttctg gaagcggctc aggaaccgat ttcacgctga ccatcagtag cttggagcct 240

gaggactttg ccacttatta ttgccagcag cacaacgagt atcctccac cttcggacag 300

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<212> PRT

<213> Artificial

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<223> La secuencia de aminoácidos de la cadena liviana variable
del anticuerpo MA-042

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Asp Arg Val Thr Ile Thr Cys Lys Ala Ser Lys Ser Ile Asn Lys Tyr

20 25 30

Leu Ala Trp Phe Gln Gln Lys Pro Gly Lys Pro Asn Lys Leu Leu Ile

35 40 45

Tyr Asp Gly Ser Thr Leu Gln 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 Glu Pro

65 70 75 80

Glu Asp Phe Ala Thr Tyr Tyr Cys Gln Gln His Asn Glu Tyr Pro Pro

85 90 95

Thr Phe Gly Gln Gly Thr Lys Leu Glu Ile Lys

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ccaggtaag gactggaatg gatgggatgg atcaatacct atacaggga acccacatat 180

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tctagcgcta gcaccaaggg cccatcggtc tccccctgg caccctctc caagagcacc 420

tctgggggca cagcgccct gggctgcctg gtcaaggact acttccccga accggtgacg 480

gtgtcgtgga actcaggcgc cctgaccagc ggcgtgcaca ccttcccggc tgtcctacag 540

tcctcaggac tctactccct cagcagcgtg gtgaccgtgc cctccagcag cttgggcacc 600

cagacctaca tctgcaacgt gaatcacaag cccagcaaca ccaagggtgga caagagagtt 660

gagcccaaatt cttgtgacaa aactcacaca tgcccaccgt gcccagcacc tgaactcctg 720

gggggaccgt cagtcttctt cttccccca aaaccaagg acaccctcat gatctcccgg 780

acccctgagg tcacatgcgt ggtggtggac gtgagccacg aagaccctga ggtcaagttc 840

aactggtacg tggacggcgt ggaggtgcat aatgccaaga caaagccgcg ggaggagcag 900

tacaacagca cgtaccgtgt ggtcagcgtc ctcaccgtcc tgcaccagga ctggctgaat 960

ggcaaggagt acaagtgcaa ggtctccaac aaagccctcc cagccccat cgagaaaacc 1020

atctccaaag ccaaaggga gccccgagaa ccacaggtgt acaccctgcc cccatcccgg 1080

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cccgtgctgg actccgacgg ctcttcttc ctctatagca agctcaccgt ggacaagagc 1260

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<212> PRT

<213> Artificial

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<223> La secuencia de aminoácidos de la cadena pesada del anticuerpo MA-042

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Gln Val Gln Leu Val Gln Ser Gly Ser Glu Leu Lys Lys Pro Gly Glu

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Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Asp Tyr

20 25 30

Leu Val His Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met

35 40 45

Gly Trp Ile Asn Thr Tyr Thr Gly Thr Pro Thr Tyr Ala Asp Asp Phe

50

55

60

Glu Gly Arg Phe Val Phe Ser Leu Asp Thr Ser Val Ser Thr Ala Asn

65

70

75

80

Leu Gln Ile Ser Ser Leu Lys Ala Glu Asp Thr Ala Val Tyr Phe Cys

85

90

95

Ala Arg Ser Trp Arg Arg Gly Leu Arg Gly Ile Gly Phe Asp Tyr Trp

100

105

110

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

115

120

125

Ser Val Phe Pro Leu Ala Pro Ser Ser Lys Ser Thr Ser Gly Gly Thr

130

135

140

Ala Ala Leu Gly Cys Leu Val Lys Asp Tyr Phe Pro Glu Pro Val Thr

145 150 155 160

Val Ser Trp Asn Ser Gly Ala Leu Thr Ser Gly Val His Thr Phe Pro

165 170 175

Ala Val Leu Gln Ser Ser Gly Leu Tyr Ser Leu Ser Ser Val Val Thr

180 185 190

Val Pro Ser Ser Ser Leu Gly Thr Gln Thr Tyr Ile Cys Asn Val Asn

195 200 205

His Lys Pro Ser Asn Thr Lys Val Asp Lys Arg Val Glu Pro Lys Ser

210 215 220

Cys Asp Lys Thr His Thr Cys Pro Pro Cys Pro Ala Pro Glu Leu Leu

225 230 235 240

Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro Lys Asp Thr Leu

245 250 255

Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val Val Asp Val Ser

260 265 270

His Glu Asp Pro Glu Val Lys Phe Asn Trp Tyr Val Asp Gly Val Glu

275 280 285

Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln Tyr Asn Ser Thr

290 295 300

Tyr Arg Val Val Ser Val Leu Thr Val Leu His Gln Asp Trp Leu Asn

305 310 315 320

Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Ala Leu Pro Ala Pro

325 330 335

Ile Glu Lys Thr Ile Ser Lys Ala Lys Gly Gln Pro Arg Glu Pro Gln

340 345 350

Val Tyr Thr Leu Pro Pro Ser Arg Glu Glu Met Thr Lys Asn Gln Val

355 360 365

Ser Leu Thr Cys Leu Val Lys Gly Phe Tyr Pro Ser Asp Ile Ala Val

370 375 380

Glu Trp Glu Ser Asn Gly Gln Pro Glu Asn Asn Tyr Lys Thr Thr Pro

385 390 395 400

Pro Val Leu Asp Ser Asp Gly Ser Phe Phe Leu Tyr Ser Lys Leu Thr

405 410 415

Val Asp Lys Ser Arg Trp Gln Gln Gly Asn Val Phe Ser Cys Ser Val

420 425 430

Met His Glu Ala Leu His Asn His Tyr Thr Gln Lys Ser Leu Ser Leu

435 440 445

Ser Pro Gly Lys

450

<210> 21

<211> 642

<212> ADN

<213> Artificial

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<223> La secuencia de nucleótidos de la cadena liviana del anticuerpo MA-042

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ggaaagccta acaagctatt aatatacgat gggctctacc tccaatccgg ggtcccttca 180

cgattttctg gaagcggctc aggaaccgat ttcacgctga ccatcagtag cttggagcct 240

gaggactttg ccacttatta ttgccagcag cacaacgagt atcctccac cttcggacag 300

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<223> La secuencia de aminoácidos de una cadena liviana del anticuerpo MA-042

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Asp Ile Gln Met Thr Gln Ser Pro Tyr Ser Leu Ser Ala Ser Val Gly

1 5 10 15

Asp Arg Val Thr Ile Thr Cys Lys Ala Ser Lys Ser Ile Asn Lys Tyr

20 25 30

Leu Ala Trp Phe Gln Gln Lys Pro Gly Lys Pro Asn Lys Leu Leu Ile

35 40 45

Tyr Asp Gly Ser Thr Leu Gln 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 Glu Pro

65 70 75 80

Glu Asp Phe Ala Thr Tyr Tyr Cys Gln Gln His Asn Glu Tyr Pro Pro

85 90 95

Thr Phe Gly Gln Gly Thr Lys Leu Glu Ile Lys Arg Thr Val Ala Ala

100 105 110

Pro Ser Val Phe Ile Phe Pro Pro Ser Asp Glu Gln Leu Lys Ser Gly

115 120 125

Thr Ala Ser Val Val Cys Leu Leu Asn Asn Phe Tyr Pro Arg Glu Ala

130 135 140

Lys Val Gln Trp Lys Val Asp Asn Ala Leu Gln Ser Gly Asn Ser Gln

145 150 155 160

Glu Ser Val Thr Glu Gln Asp Ser Lys Asp Ser Thr Tyr Ser Leu Ser

165 170 175

Ser Thr Leu Thr Leu Ser Lys Ala Asp Tyr Glu Lys His Lys Val Tyr

180 185 190

Ala Cys Glu Val Thr His Gln Gly Leu Ser Ser Pro Val Thr Lys Ser

195 200 205

Phe Asn Arg Gly Glu Cys

210