	H-CDR1	H-CDR2	H-CDR3
MOR9464	SYAIS (SEQ ID NO:3)	NIIP-MTGQTYYAQKFQG (SEQ ID NO:9)	AAYHPLVFDN (SEQ ID NO: 5)
MOR9464N30K	SYAIS (SEQ ID NO:3)	NIIP-MTGQTYYAQKFQG (SEQ ID NO:9)	AAYHPLVFDN (SEQ ID NO: 5)
MOR10222N30SM54I	SYAIS (SEQ ID NO:3)	NIIP-ITGQTYYAQKFQG (SEQ ID NO:13)	AAYHPLVFDN (SEQ ID NO: 5)
MOR8775	SYAIS (SEQ ID NO:3)	GIIP-IYGTANYAQKFQG (SEQ ID NO:4)	AAYHPLVFDN (SEQ ID NO: 5)
MOR9465	SYAIS (SEQ ID NO:3)	NIIP-HYGFAYYAQKFQG (SEQ ID NO:11)	AAYHPLVFDN (SEQ ID NO: 5)
MOR9466	SYAIS (SEQ ID NO:3)	NIIP-YSGFAYYAQKFQG (SEQ ID NO:12)	AAYHPLVFDN (SEQ ID NO: 5)
MOR10222	SYAIS (SEQ ID NO:3)	NIIP-MTGQTYYAQKFQG (SEQ ID NO:9)	AAYHPLVFDN (SEQ ID NO: 5)
MOR9441	SYAIS (SEQ ID NO:3)	WINPFYIGETFYAQKFQG (SEQ ID NO:10)	AAYHPLVFDN (SEQ ID NO: 5)
MOR10579	SYAIS (SEQ ID NO:3)	WINPFYIGETFYAQKFQG (SEQ ID NO:10)	AAYHPLVFDN (SEQ ID NO: 5)
MOR13363	SYAIH (SEQ ID NO:106)	VISGEGSNTYYADSVKG (SEQ ID NO:107)	VMIGYGFDY (SEQ ID NO:108)
MOR13361	SYAIH (SEQ ID NO:106)	TIQSSGENKFYADSVKG (SEQ ID NO:122)	VMIGYGFDY (SEQ ID NO:108)
MOR13341	TFSIS (SEQ ID NO:120)	GIIP-IFGTANYAQKFQG (SEQ ID NO:121)	GGYGGYYYFDY (SEQ ID NO:123)
MOR13342	TFSIS (SEQ ID NO:120)	GIIP-IFGTANYAQKFQG (SEQ ID NO:121)	GGYGGYYYFDY (SEQ ID NO:123)
MOR13347	TFSIS (SEQ ID NO:120)	GIIP-IFGTANYAQKFQG (SEQ ID NO:121)	GGYGGYYYFDY (SEQ ID NO:123)
MOR8776	TGSYYWN (SEQ ID NO:74)	EINHMGITYYNPSLKG (SEQ ID NO:75)	TTRYWMSHILAYGMDY (SEQ ID NO: 79)
MOR10497	TGSYYWN (SEQ ID NO:74)	EIQSPGYTFYNPSLKS (SEQ ID NO:78)	TTRYWMSHILAYGMDY (SEQ ID NO: 79)
MOR10501	TGSYYWN (SEQ ID NO:74)	EIWHSGPTFYNPSLKS (SEQ ID NO:76)	TTRYWMSHILAYGMDY (SEQ ID NO: 79)
MOR10502	TGSYYWN (SEQ ID NO:74)	EIHGHGFTFYNPSLKS (SEQ ID NO:77)	TTRYWMSHILAYGMDY (SEQ ID NO: 79)

Figure 1

	L-CDR1	L-CDR2	L-CDR3
MOR9464	SGSSSNIGNHYVN (SEQ ID NO:6)	RNNHRPS (SEQ ID NO:7)	QSWDYSGFSTV (SEQ ID NO:8)
MOR9464N30K	SGSSSNIGNHYVN (SEQ ID NO:6)	RNNHRPS (SEQ ID NO:7)	QSWDYSGFSTV (SEQ ID NO:8)
MOR10222N30SM54I	SGSSSNIGNHYVN (SEQ ID NO:6)	RNNHRPS (SEQ ID NO:7)	QSWDYSGFSTV (SEQ ID NO:8)
MOR8775	SGSSSNIGNHYVN (SEQ ID NO:6)	RNNHRPS (SEQ ID NO:7)	QSWDYSGFSTV (SEQ ID NO:8)
MOR9465	SGSSSNIGNHYVN (SEQ ID NO:6)	RNNHRPS (SEQ ID NO:7)	QSWDYSGFSTV (SEQ ID NO:8)
MOR9466	SGSSSNIGNHYVN (SEQ ID NO:6)	RNNHRPS (SEQ ID NO:7)	QSWDYSGFSTV (SEQ ID NO:8)
MOR10579	SGSSSNIGNHYVN (SEQ ID NO:6)	RNNHRPS (SEQ ID NO:7)	QSWDYSGFSTV (SEQ ID NO:8)
MOR10222	SGSSSNIGNHYVN (SEQ ID NO:6)	RNNHRPS (SEQ ID NO:7)	QSWDYSGFSTV (SEQ ID NO:8)
MOR9441	SGSSSNIGNHYVN (SEQ ID NO:6)	RNNHRPS (SEQ ID NO:7)	QSWDYSGFSTV (SEQ ID NO:8)
MOR13363	RASQSIFNYLN (SEQ ID NO:109)	DSSTLQS (SEQ ID NO:110)	LQYSGFLFT (SEQ ID NO:111)
MOR13361	RASQSIFNYLN (SEQ ID NO:109)	DSSTLQS (SEQ ID NO:110)	HQYSGLLFT (SEQ ID NO:126)
MOR13341	RASQSISNRLN (SEQ ID NO:124)	KGSTLQS (SEQ ID NO:125)	QQHKVWLTT (SEQ ID NO:127)
MOR13342	RASQSISNRLN (SEQ ID NO:124)	KGSTLQS (SEQ ID NO:125)	QQHYVWSTT (SEQ ID NO:128)
MOR13347	RASQSISNRLN (SEQ ID NO:124)	KGSTLQS (SEQ ID NO:125)	QQHYQWLTT (SEQ ID NO:129)
MOR8776	SGSSSNIGNHYVS (SEQ ID NO:80)	ANTKRPS (SEQ ID NO:81)	SSYDGSQSIV (SEQ ID NO:82)
MOR10497	SGSSSNIGNHYVS (SEQ ID NO:80)	ANTKRPS (SEQ ID NO:81)	SSYDGSQSIV (SEQ ID NO:82)
MOR10501	SGSSSNIGNHYVS (SEQ ID NO:80)	ANTKRPS (SEQ ID NO:81)	SSYDGSQSIV (SEQ ID NO:82)
MOR10502	SGSSSNIGNHYVS (SEQ ID NO:80)	ANTKRPS (SEQ ID NO:81)	SSYDGSQSIV (SEQ ID NO:82)

Figure 2

Figure 3(A)

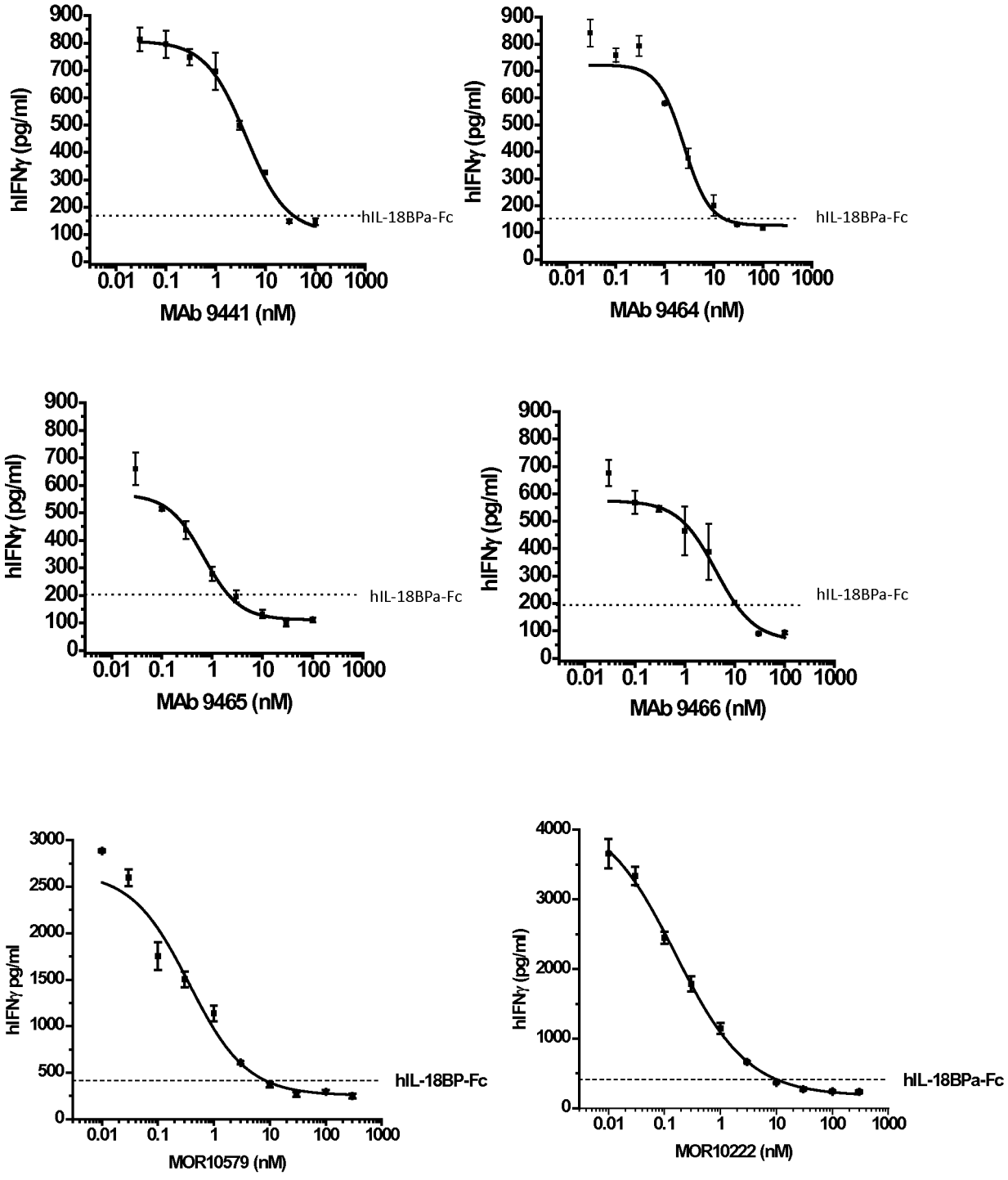

hIFNγ (pg/ml)
MAb 9441 (nM)
hIL-18BPa-Fc
MAb 9464 (nM)
MAb 9465 (nM)
MAb 9466 (nM)
hIFNγ pg/ml
hIL-18BP-Fc
MOR10579 (nM)
MOR10222 (nM)

Figure 3(B)

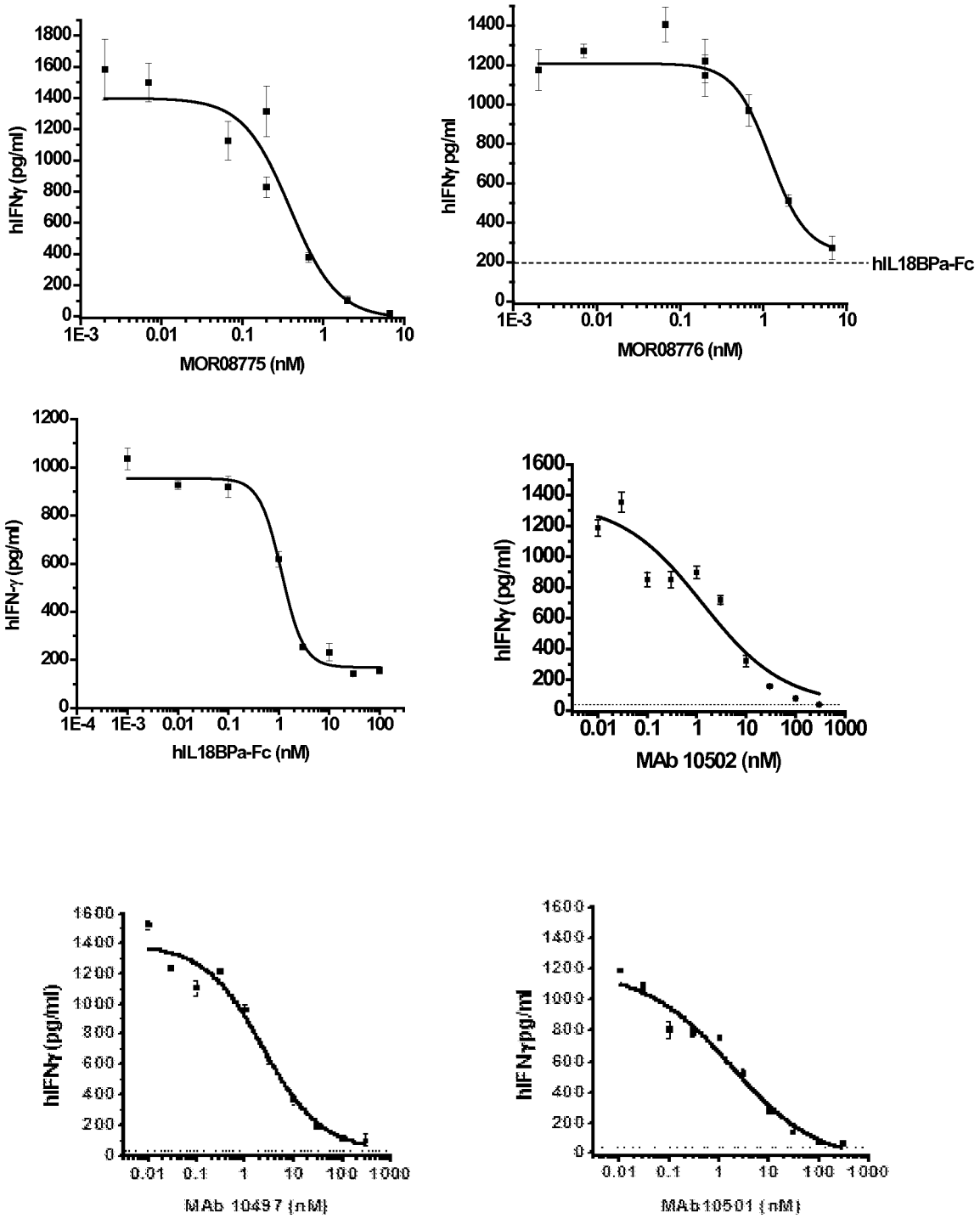

hIFNγ (pg/ml)
MOR08775 (nM)
hIFNγ pg/ml
MOR08776 (nM)
hIL18BPa-Fc
hIFN-γ (pg/ml)
hIL18BPa-Fc (nM)
hIFNγ (pg/ml)
MAb 10502 (nM)
hIFNγ (pg/ml)
MAb 10497 (nM)
hIFNγpg/ml
MAb10501 (nM)

Figure 3(C)

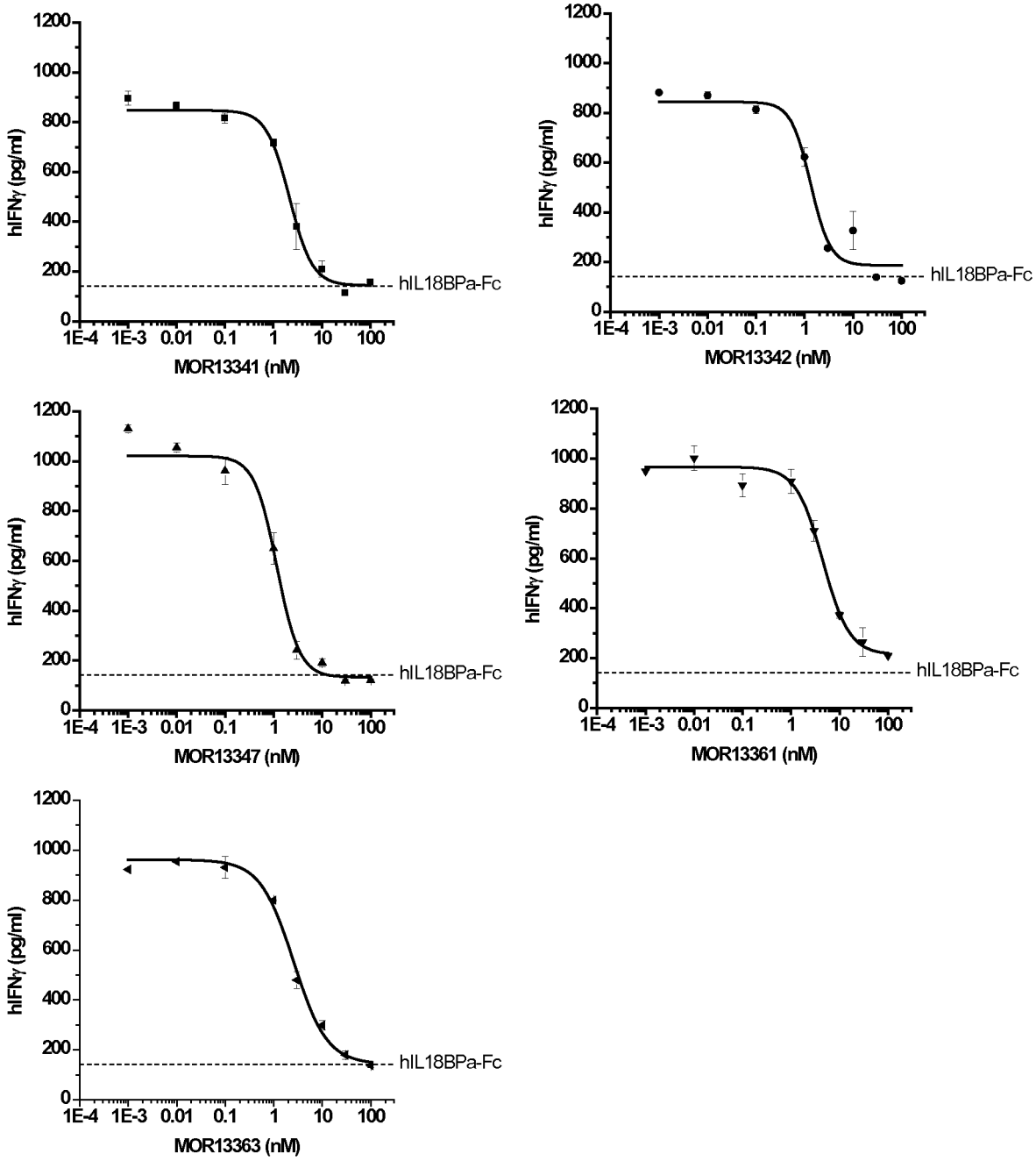

hIFNγ (pg/ml)
1200
1000
800
600
400
200
0
1E-4 1E-3 0.01 0.1 1 10 100
hIL18BPa-Fc
MOR13341 (nM)
MOR13342 (nM)
MOR13347 (nM)
MOR13361 (nM)
MOR13363 (nM)

Figure 3(D)

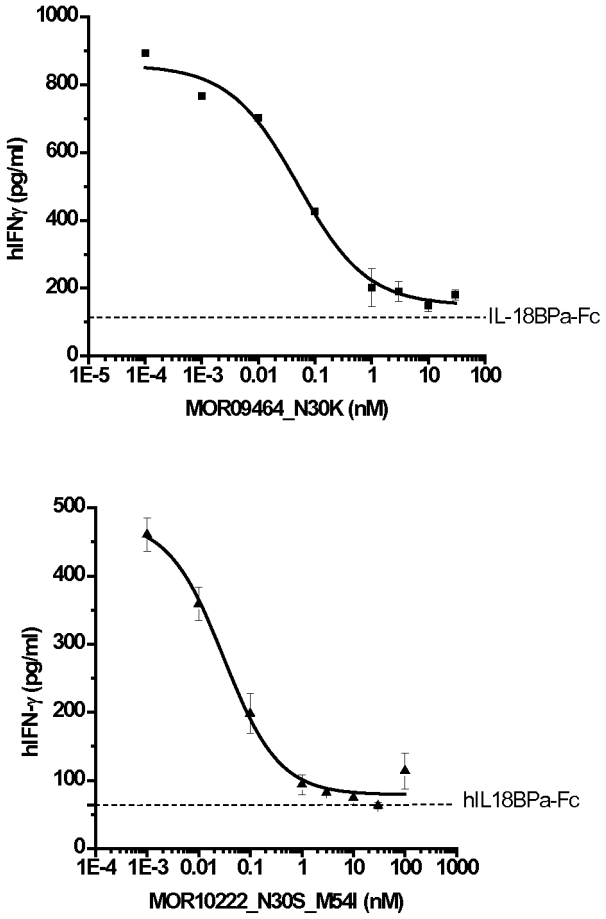
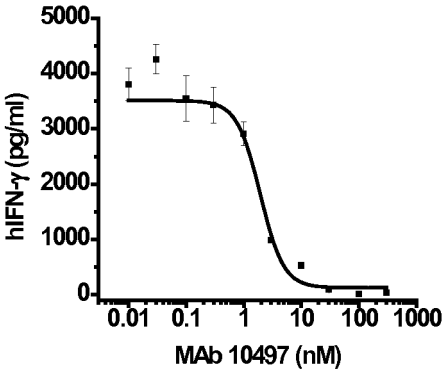
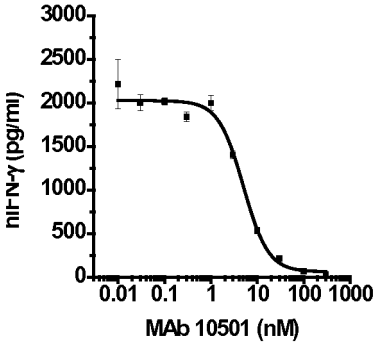
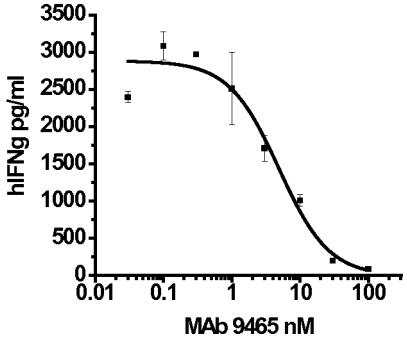
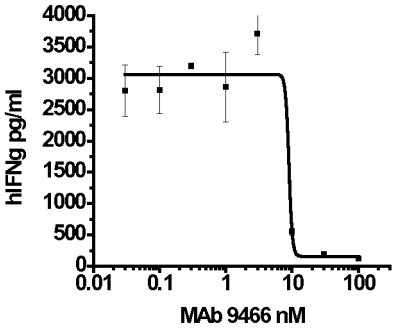

1000
800
600
400
200
0
hIFNγ (pg/ml)
IL-18BPa-Fc
1E-5 1E-4 1E-3 0.01 0.1 1 10 100
MOR09464_N30K (nM)
500
400
300
200
100
0
hIFN-γ (pg/ml)
hIL18BPa-Fc
1E-4 1E-3 0.01 0.1 1 10 100 1000
MOR10222_N30S_M54I (nM)

Figure 4(A)


hIFN-γ (pg/ml)
5000
4000
3000
2000
1000
0
0.01 0.1 1 10 100 1000
MAb 10497 (nM)


hIFN-γ (pg/ml)
3000
2500
2000
1500
1000
500
0
0.01 0.1 1 10 100 1000
MAb 10501 (nM)


hIFNg pg/ml
3500
3000
2500
2000
1500
1000
500
0
0.01 0.1 1 10 100
MAb 9465 nM


hIFNg pg/ml
4000
3500
3000
2500
2000
1500
1000
500
0
0.01 0.1 1 10 100
MAb 9466 nM

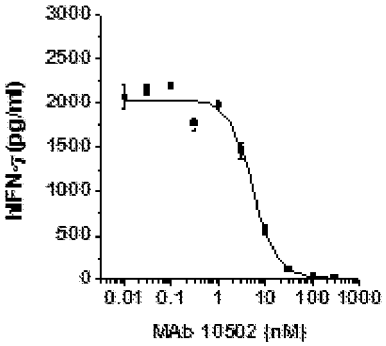
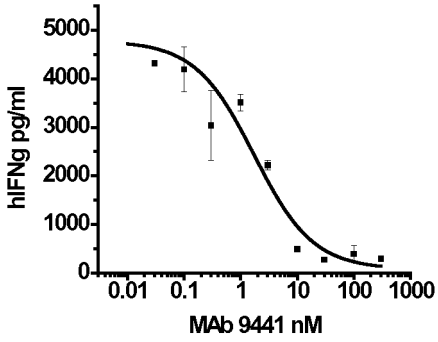
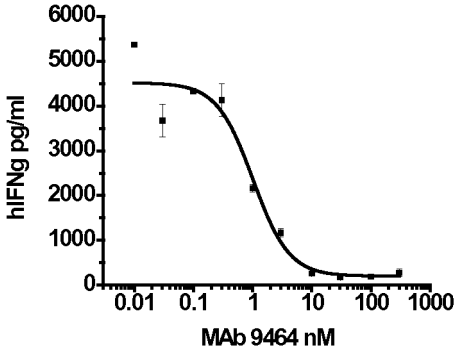
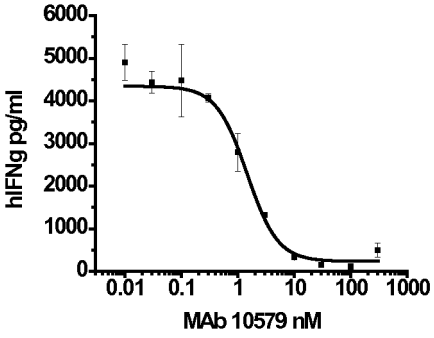
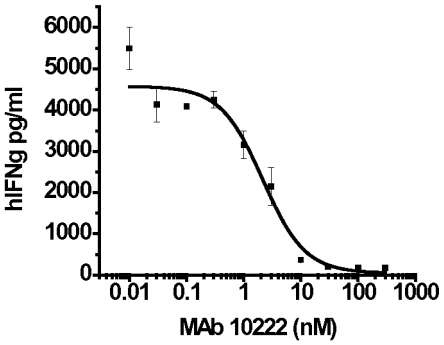

hIFN-γ (pg/ml)
3000
2500
2000
1500
1000
500
0
0.01 0.1 1 10 100 1000
MAb 10502 (nM)

Figure 4 (B)


hIFNg pg/ml
5000
4000
3000
2000
1000
0
0.01 0.1 1 10 100 1000
MAb 9441 nM


hIFNg pg/ml
6000
5000
4000
3000
2000
1000
0
0.01 0.1 1 10 100 1000
MAb 9464 nM


hIFNg pg/ml
6000
5000
4000
3000
2000
1000
0
0.01 0.1 1 10 100 1000
MAb 10579 nM


hIFNg pg/ml
6000
5000
4000
3000
2000
1000
0
0.01 0.1 1 10 100 1000
MAb 10222 (nM)

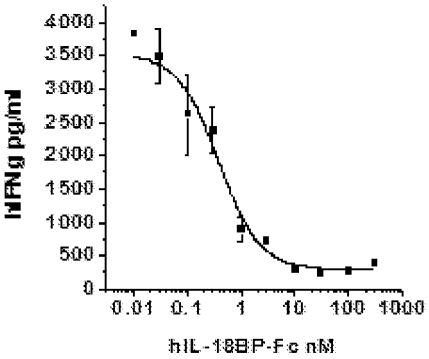

hIFNg pg/ml
4000
3500
3000
2500
2000
1500
1000
500
0
0.01 0.1 1 10 100 1000
hIL-18BP-Fc nM

Figure 5(A)

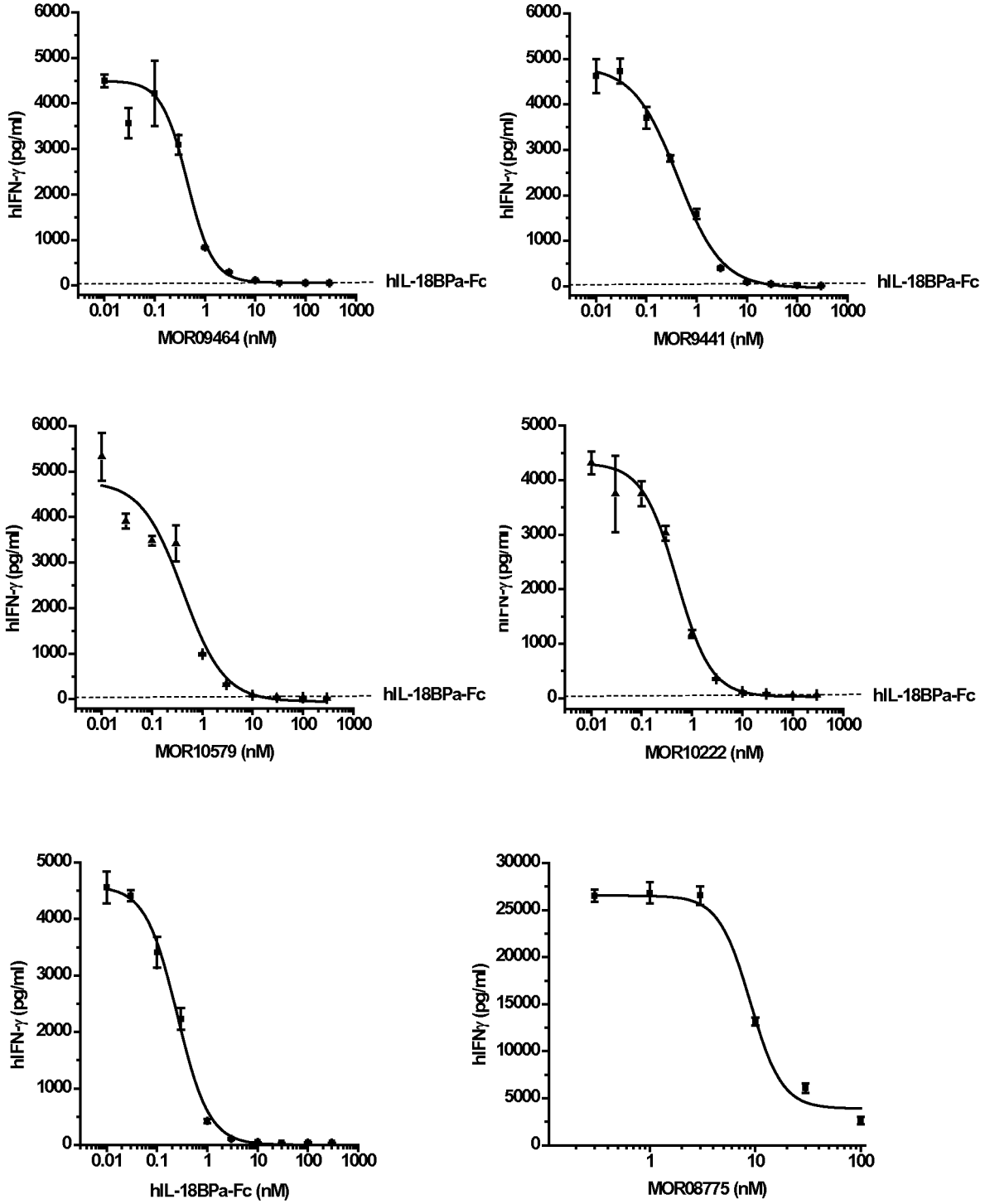
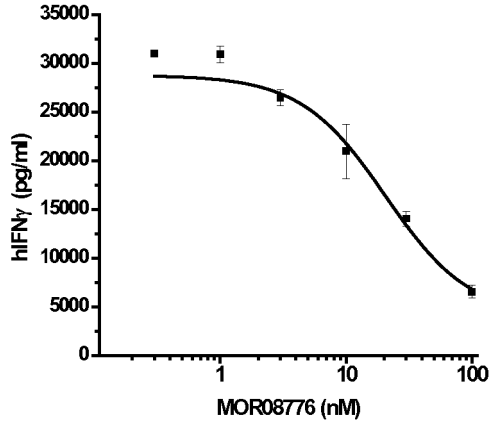
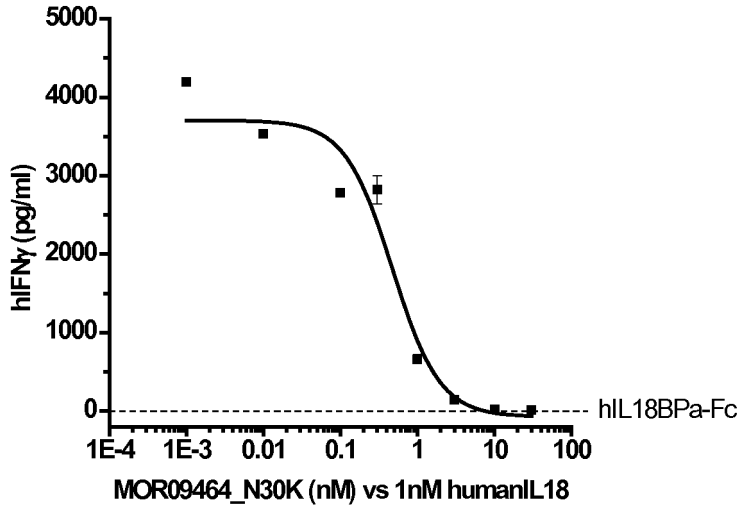

hIFN-γ (pg/ml)
MOR09464 (nM)
hIL-18BPa-Fc
MOR9441 (nM)
MOR10579 (nM)
MOR10222 (nM)
hIL-18BPa-Fc (nM)
MOR08775 (nM)

Figure 5(B)


35000
30000
25000
20000
15000
10000
5000
0
hIFNγ (pg/ml)
1
10
100
MOR08776 (nM)


5000
4000
3000
2000
1000
0
hIFNγ (pg/ml)
hIL18BPa-Fc
1E-4
1E-3
0.01
0.1
1
10
100
MOR09464_N30K (nM) vs 1nM humanIL18

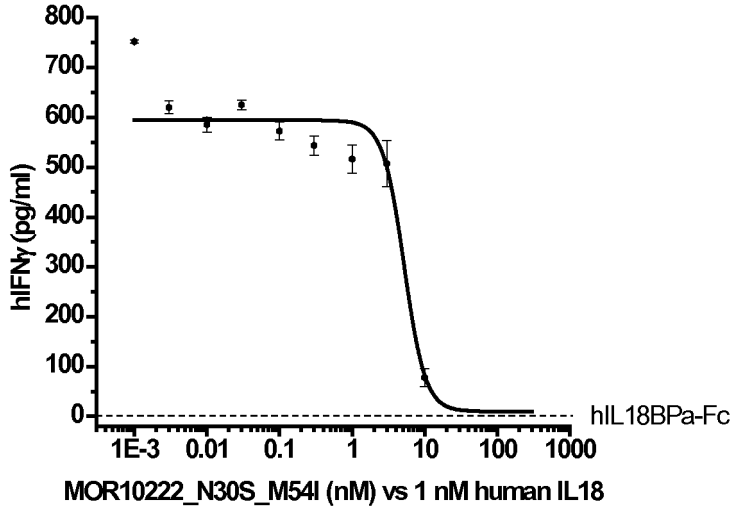
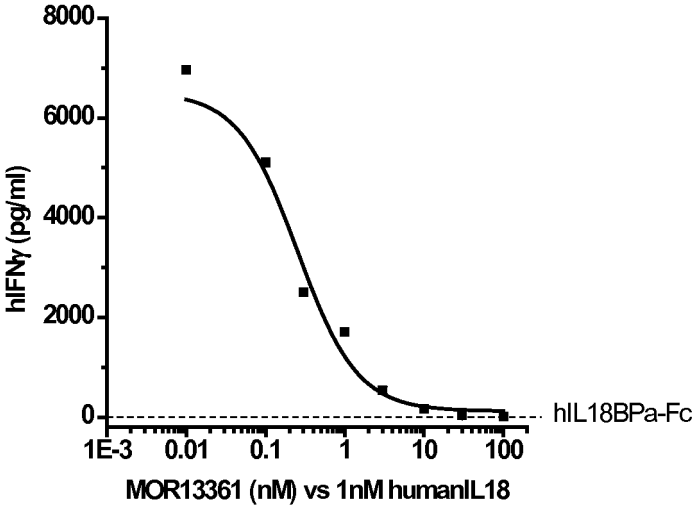

800
700
600
500
400
300
200
100
0
hIFNγ (pg/ml)
hIL18BPa-Fc
1E-3
0.01
0.1
1
10
100
1000
MOR10222_N30S_M54I (nM) vs 1 nM human IL18

Figure 5(C)


8000
6000
4000
2000
0
hIFNγ (pg/ml)
hIL18BPa-Fc
1E-3
0.01
0.1
1
10
100
MOR13361 (nM) vs 1nM humanIL18

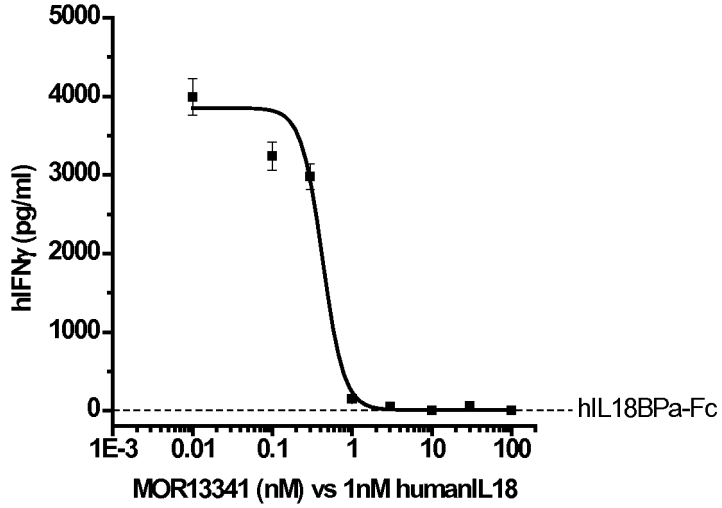

5000
4000
3000
2000
1000
0
hIFNγ (pg/ml)
hIL18BPa-Fc
1E-3
0.01
0.1
1
10
100
MOR13341 (nM) vs 1nM humanIL18

Figure 6(A)

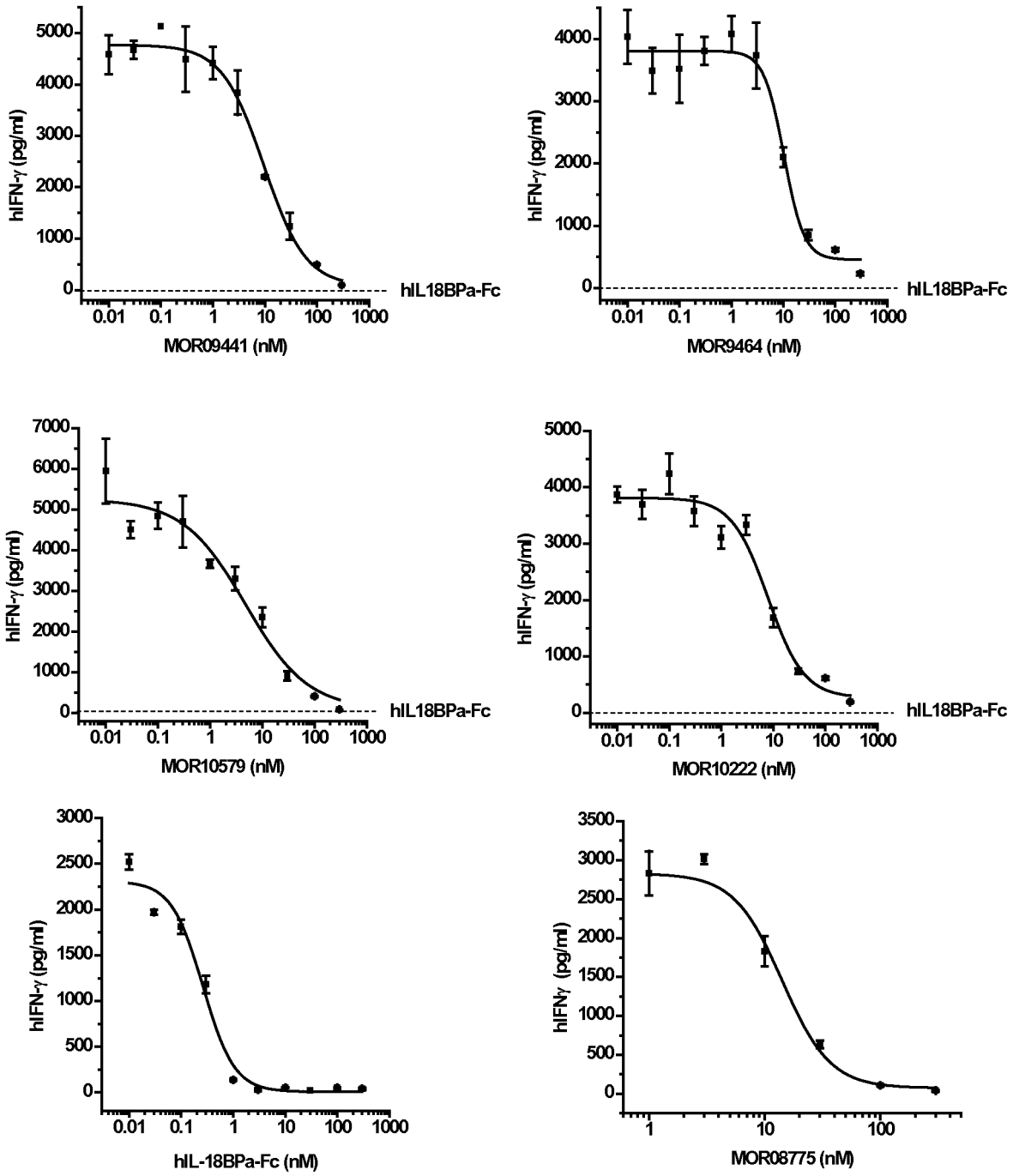
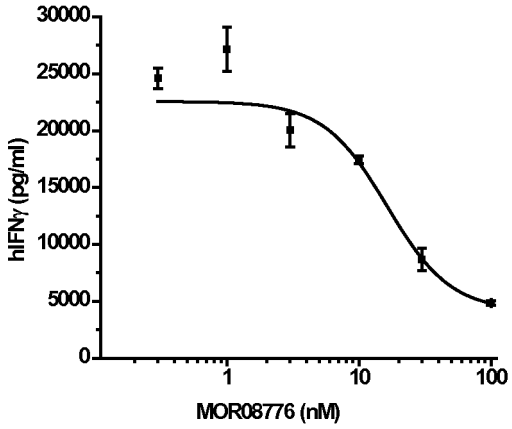
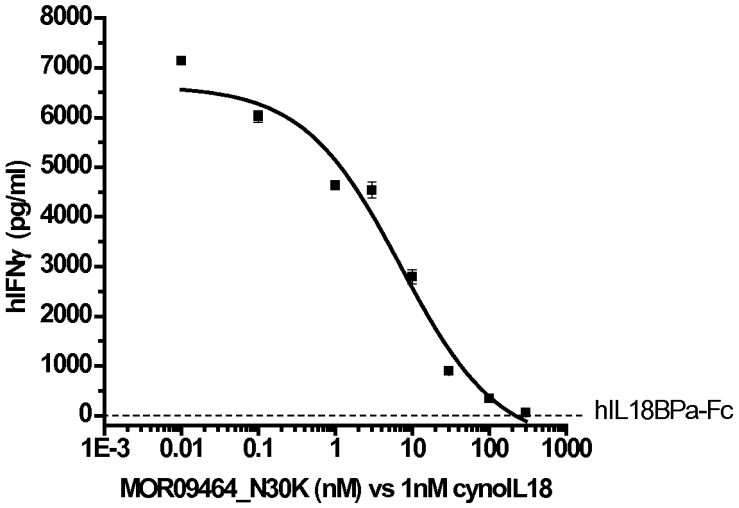

hIFN-γ (pg/ml)
MOR09441 (nM)
hIL18BPa-Fc
MOR9464 (nM)
MOR10579 (nM)
MOR10222 (nM)
hIL-18BPa-Fc (nM)
hIFNγ (pg/ml)
MOR08775 (nM)

Figure 6(B)


30000
25000
20000
15000
10000
5000
0
hIFNγ (pg/ml)
1
10
100
MOR08776 (nM)


8000
7000
6000
5000
4000
3000
2000
1000
0
hIFNγ (pg/ml)
hIL18BPa-Fc
1E-3
0.01
0.1
1
10
100
1000
MOR09464_N30K (nM) vs 1nM cynoIL18

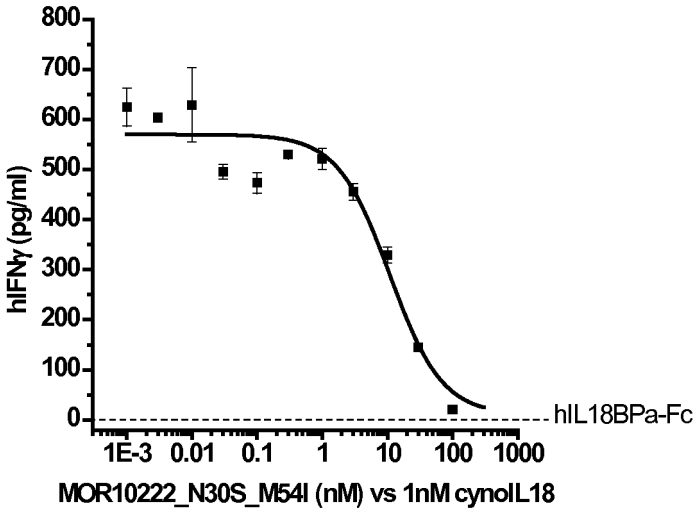

800
700
600
500
400
300
200
100
0
hIFNγ (pg/ml)
hIL18BPa-Fc
1E-3
0.01
0.1
1
10
100
1000
MOR10222_N30S_M54I (nM) vs 1nM cynoIL18

Figure 6(C)

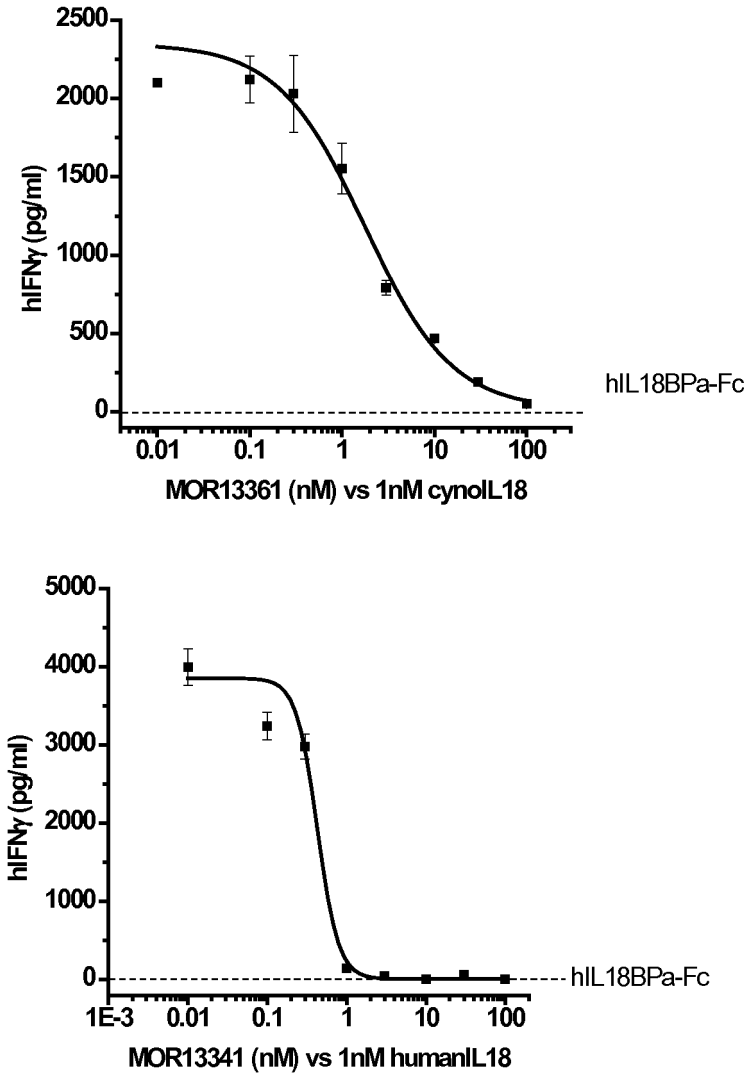

2500
2000
1500
1000
500
0
hIFNγ (pg/ml)
0.01
0.1
1
10
100
MOR13361 (nM) vs 1nM cynoIL18
hIL18BPa-Fc
5000
4000
3000
2000
1000
0
hIFNγ (pg/ml)
1E-3
0.01
0.1
1
10
100
MOR13341 (nM) vs 1nM humanIL18
hIL18BPa-Fc

Figure 7(A)

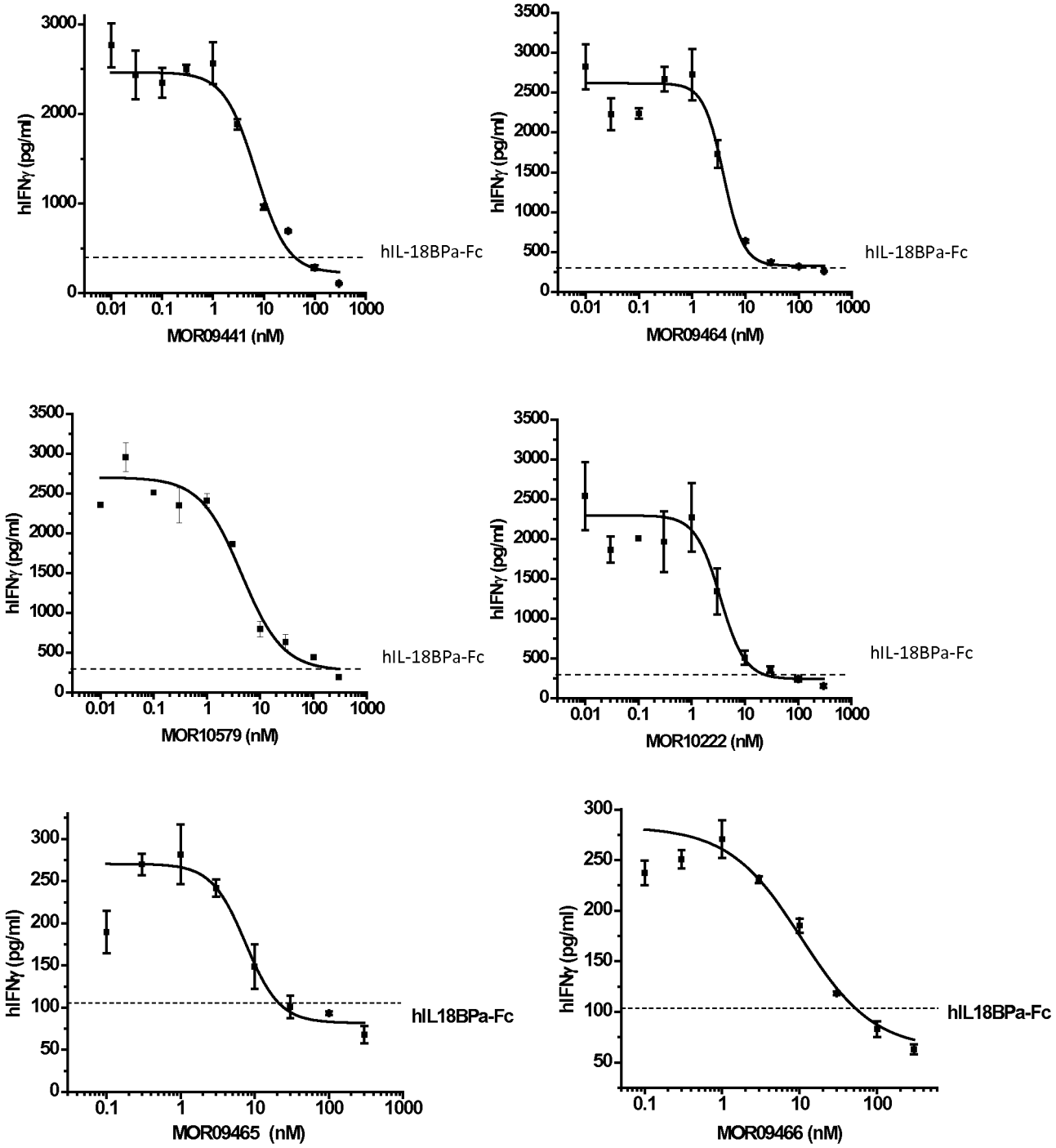

hIFNγ (pg/ml)
MOR09441 (nM)
hIL-18BPa-Fc
MOR09464 (nM)
MOR10579 (nM)
MOR10222 (nM)
MOR09465 (nM)
hIL18BPa-Fc
MOR09466 (nM)

Figure 7(B)

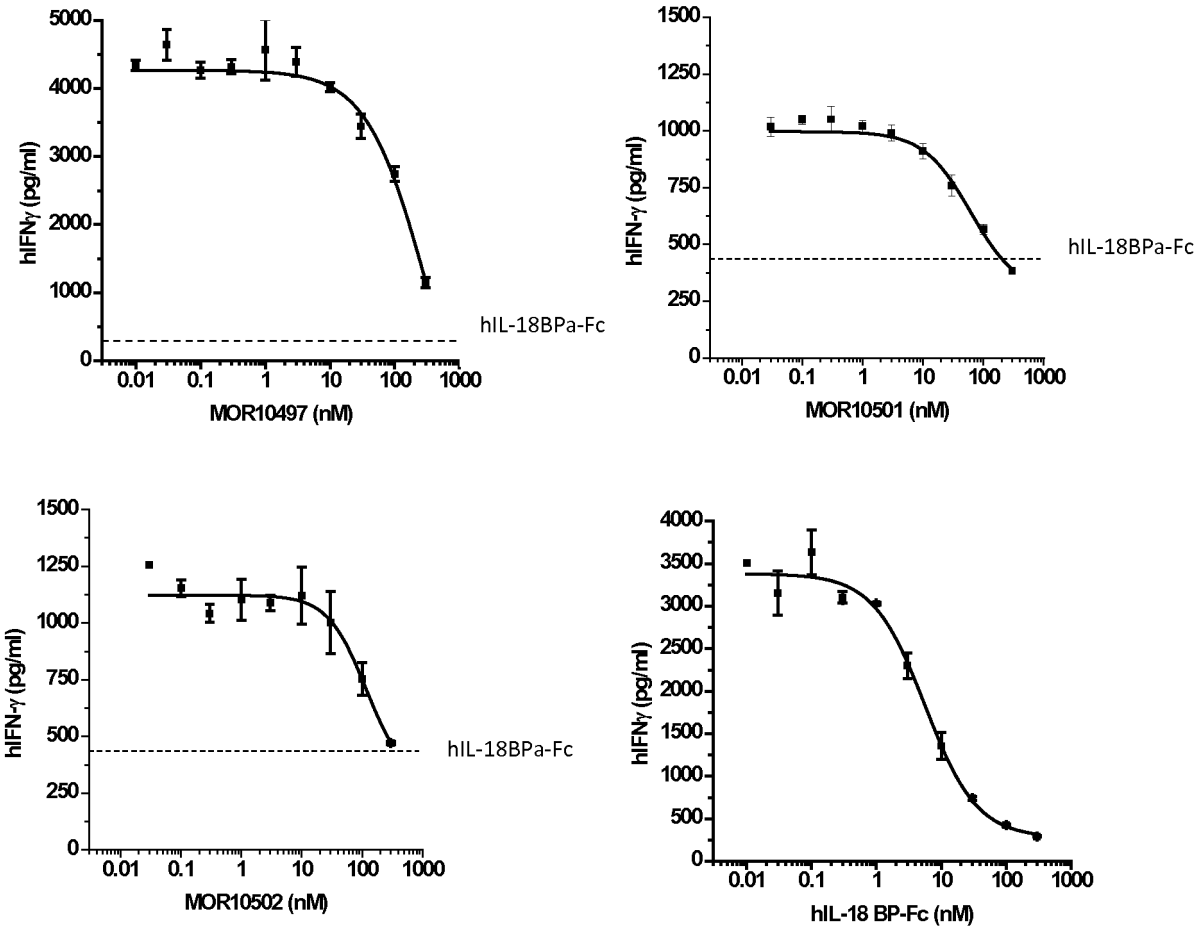

hIFNγ (pg/ml)
MOR10497 (nM)
hIL-18BPa-Fc
hIFN-γ (pg/ml)
MOR10501 (nM)
hIL-18BPa-Fc
hIFN-γ (pg/ml)
MOR10502 (nM)
hIL-18BPa-Fc
hIFNγ (pg/ml)
hIL-18 BP-Fc (nM)

Figure 7(C)

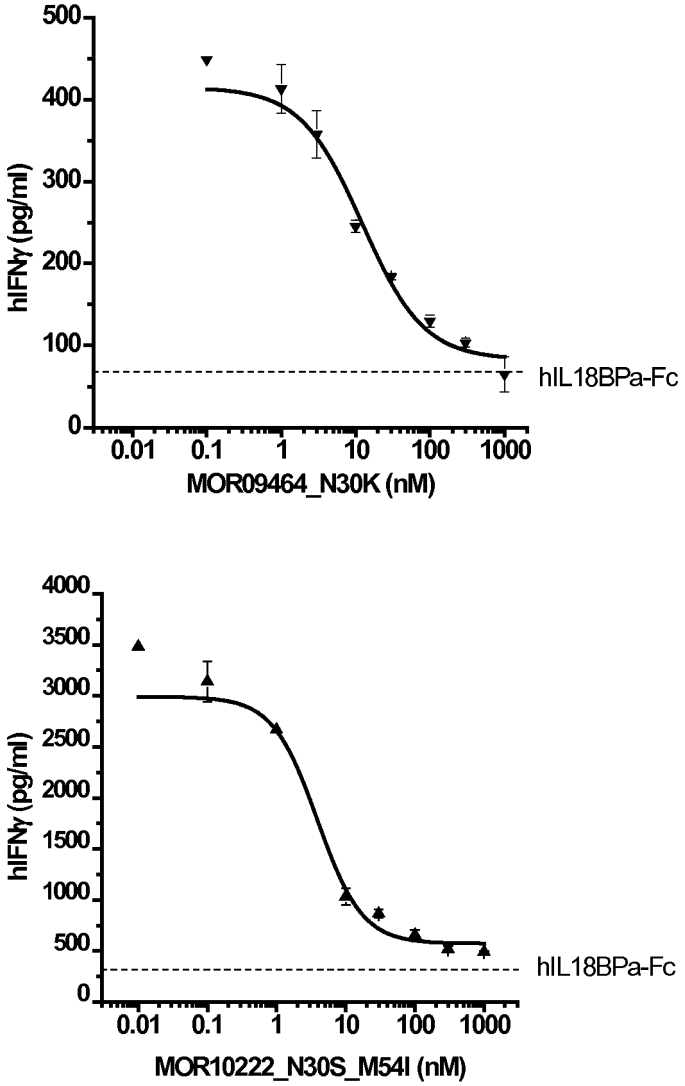

hIFNγ (pg/ml)
500
400
300
200
100
0
0.01
0.1
1
10
100
1000
MOR09464_N30K (nM)
hIL18BPa-Fc
hIFNγ (pg/ml)
4000
3500
3000
2500
2000
1500
1000
500
0
0.01
0.1
1
10
100
1000
MOR10222_N30S_M54I (nM)
hIL18BPa-Fc

Figure 7(D)

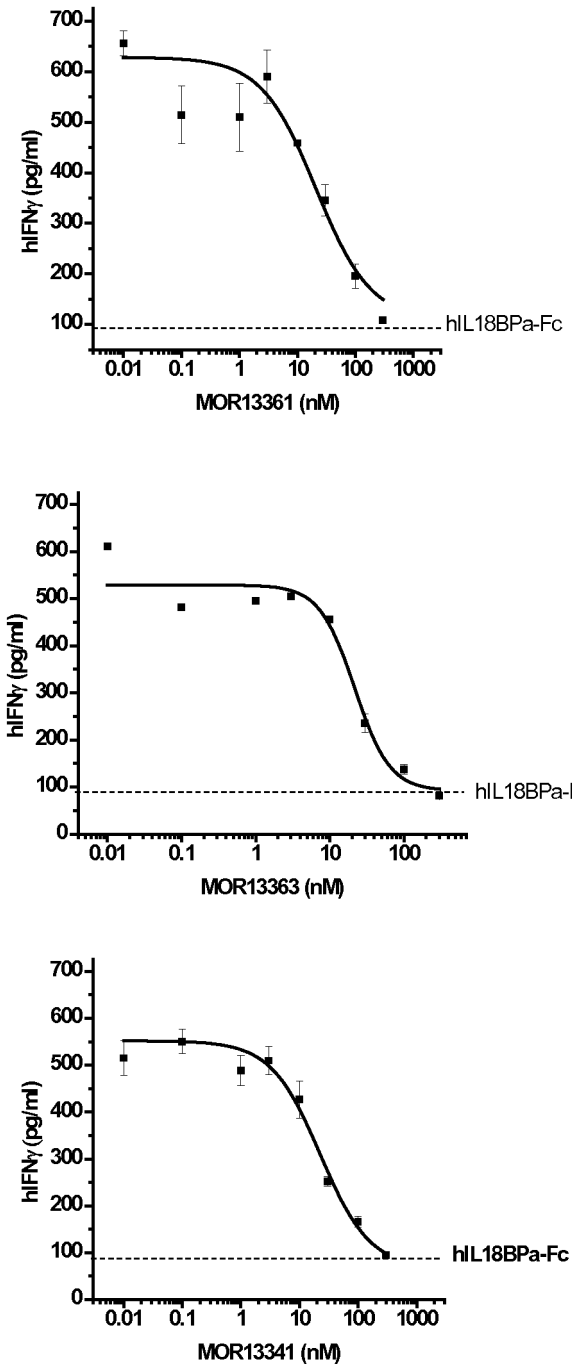

hIFNγ (pg/ml)
700
600
500
400
300
200
100
0
0.01
0.1
1
10
100
1000
hIL18BPa-Fc
MOR13361 (nM)
MOR13363 (nM)
MOR13341 (nM)

Figure 7(E)

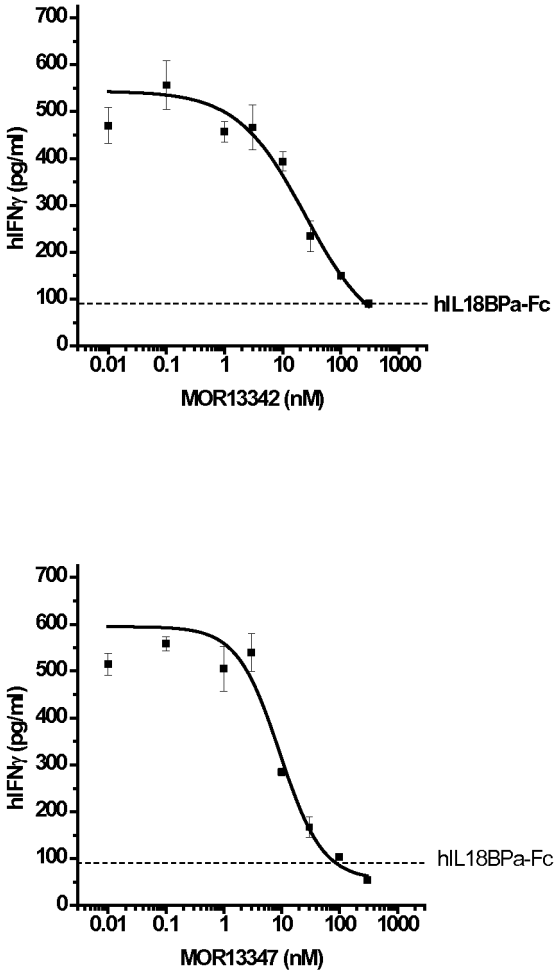

hIFNγ (pg/ml)
700
600
500
400
300
200
100
0
0.01
0.1
1
10
100
1000
MOR13342 (nM)
hIL18BPa-Fc
hIFNγ (pg/ml)
700
600
500
400
300
200
100
0
0.01
0.1
1
10
100
1000
MOR13347 (nM)
hIL18BPa-Fc

Figure 8(A)

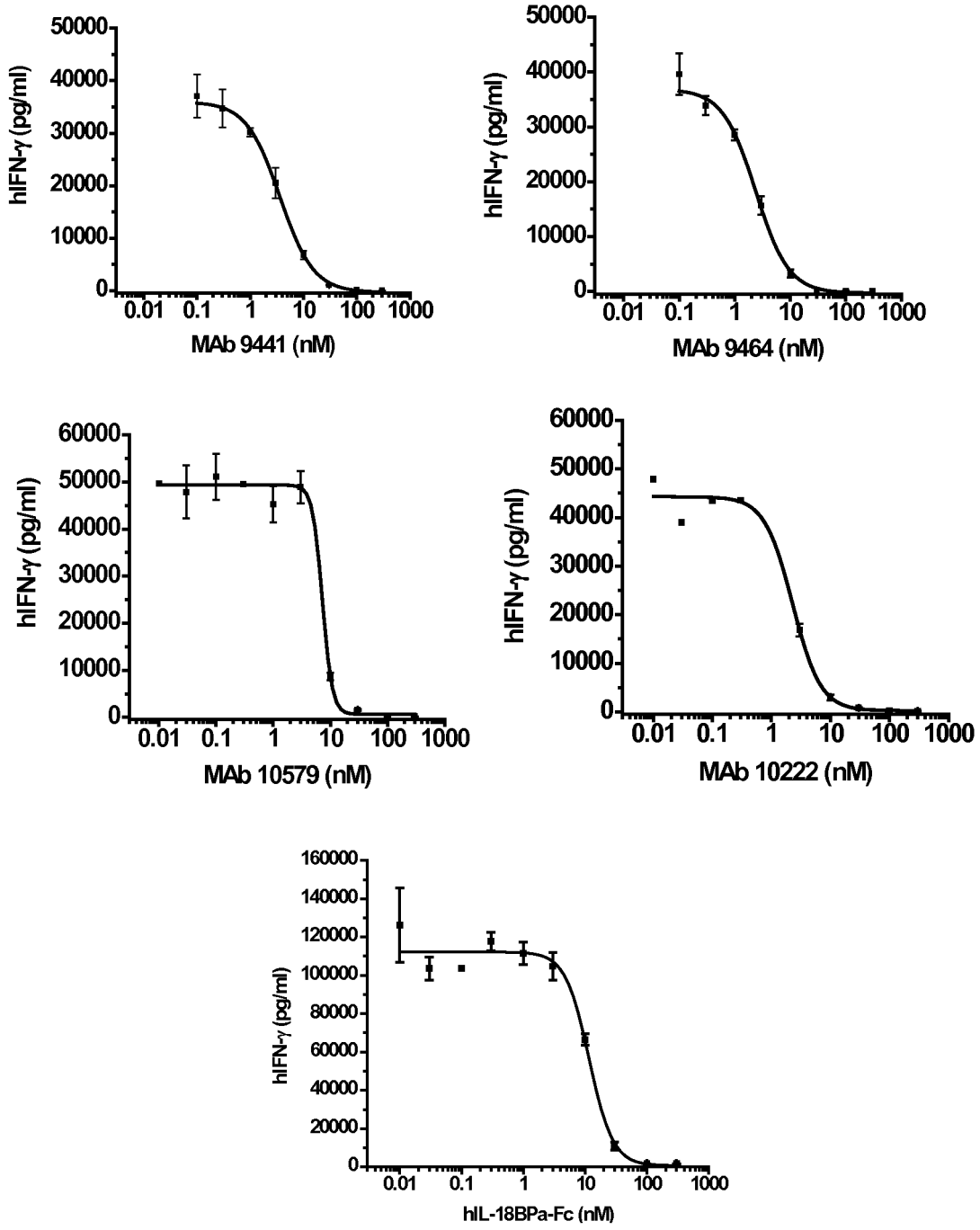

hIFN-γ (pg/ml)
MAb 9441 (nM)
MAb 9464 (nM)
MAb 10579 (nM)
MAb 10222 (nM)
hIL-18BPa-Fc (nM)

Figure 8(B)

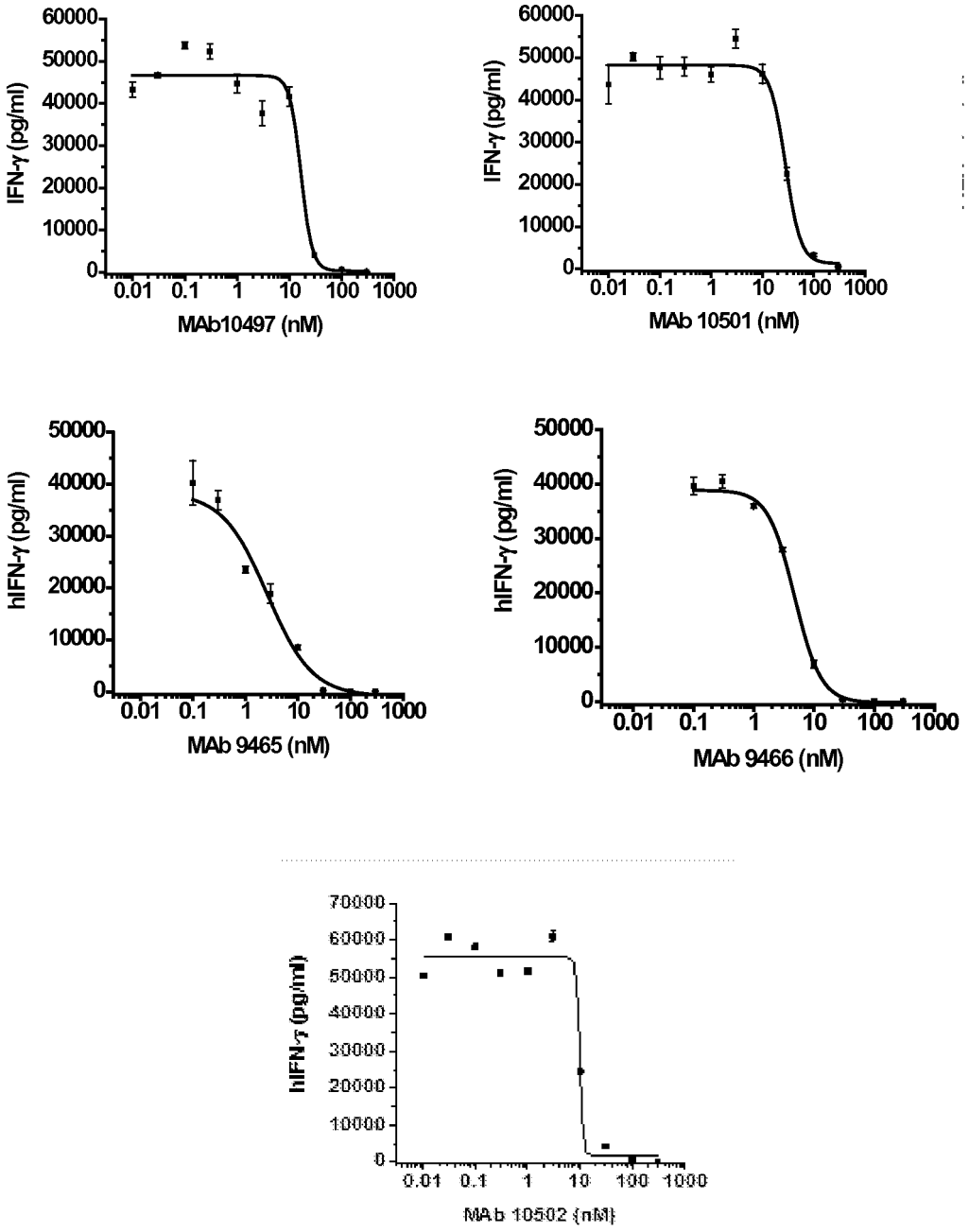
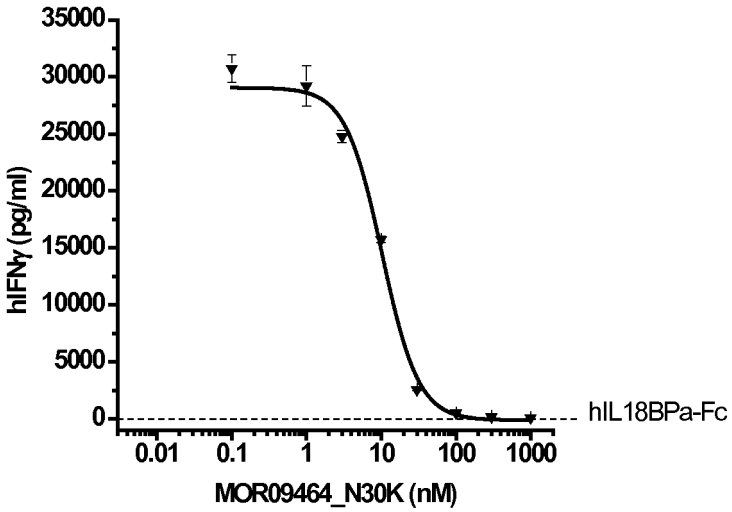

IFN-γ (pg/ml)
60000
50000
40000
30000
20000
10000
0
0.01 0.1 1 10 100 1000
MAb10497 (nM)
IFN-γ (pg/ml)
MAb 10501 (nM)
hIFN-γ (pg/ml)
50000
40000
30000
20000
10000
0
MAb 9465 (nM)
hIFN-γ (pg/ml)
MAb 9466 (nM)
hIFN-γ (pg/ml)
70000
60000
50000
40000
30000
20000
10000
0
MAb 10502 (nM)

Figure 8(C)


35000
30000
25000
20000
15000
10000
5000
0
hIFNγ (pg/ml)
hIL18BPa-Fc
0.01
0.1
1
10
100
1000
MOR09464_N30K (nM)

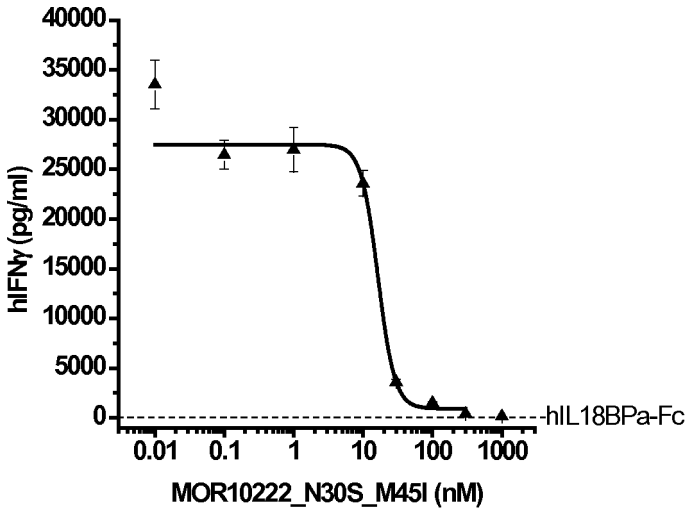
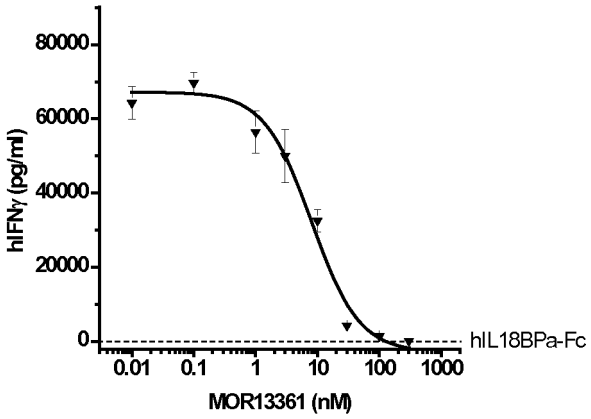
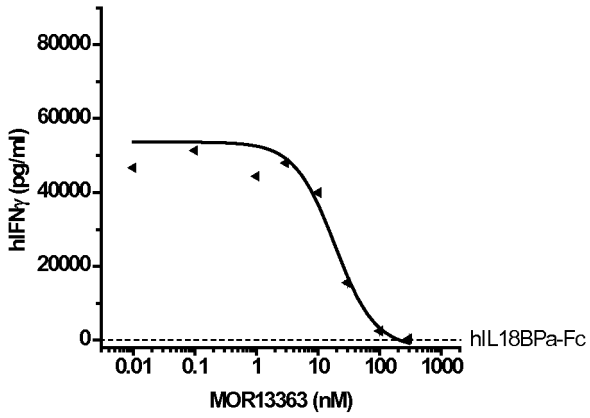

40000
35000
30000
25000
20000
15000
10000
5000
0
hIFNγ (pg/ml)
hIL18BPa-Fc
0.01
0.1
1
10
100
1000
MOR10222_N30S_M45I (nM)

Figure 8(D)


80000
60000
40000
20000
0
hIFNγ (pg/ml)
hIL18BPa-Fc
0.01
0.1
1
10
100
1000
MOR13361 (nM)


80000
60000
40000
20000
0
hIFNγ (pg/ml)
hIL18BPa-Fc
0.01
0.1
1
10
100
1000
MOR13363 (nM)

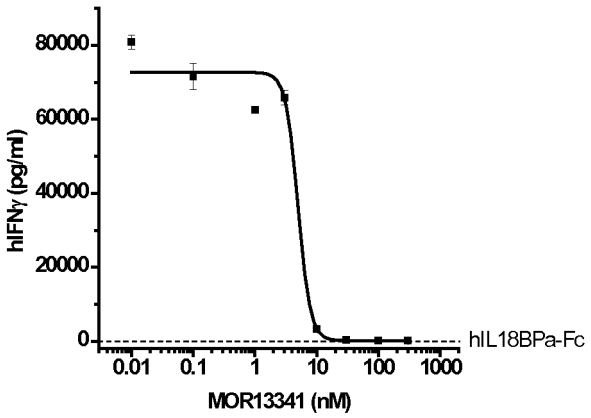

80000
60000
40000
20000
0
hIFNγ (pg/ml)
hIL18BPa-Fc
0.01
0.1
1
10
100
1000
MOR13341 (nM)

Figure 8(E)

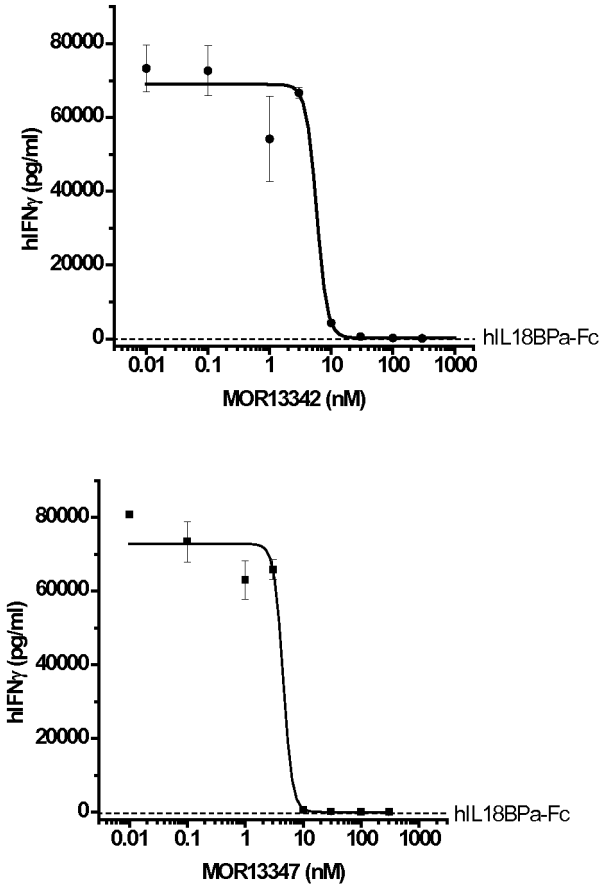

80000
60000
40000
20000
0
hIFNγ (pg/ml)
0.01
0.1
1
10
100
1000
MOR13342 (nM)
hIL18BPa-Fc
80000
60000
40000
20000
0
hIFNγ (pg/ml)
0.01
0.1
1
10
100
1000
MOR13347 (nM)
hIL18BPa-Fc

Figure 9 (A)

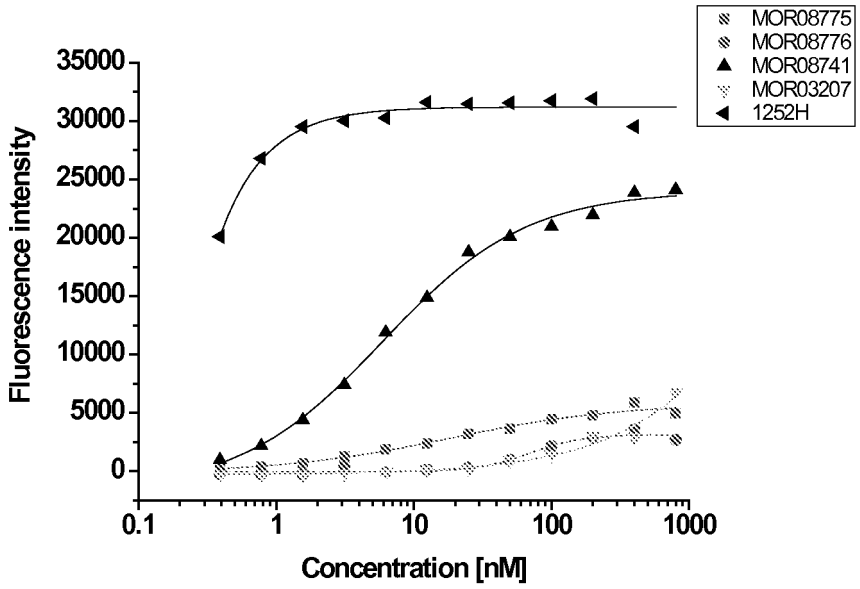
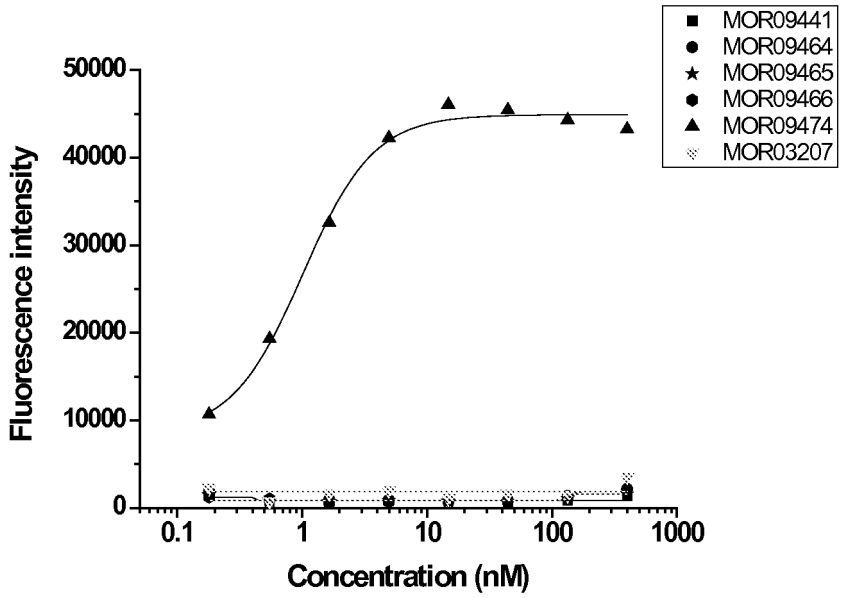

MOR08775
MOR08776
MOR08741
MOR03207
1252H
Fluorescence intensity
35000
30000
25000
20000
15000
10000
5000
0
0.1
1
10
100
1000
Concentration [nM]

Figure 9 (B)


MOR09441
MOR09464
MOR09465
MOR09466
MOR09474
MOR03207
Fluorescence intensity
50000
40000
30000
20000
10000
0
0.1
1
10
100
1000
Concentration (nM)

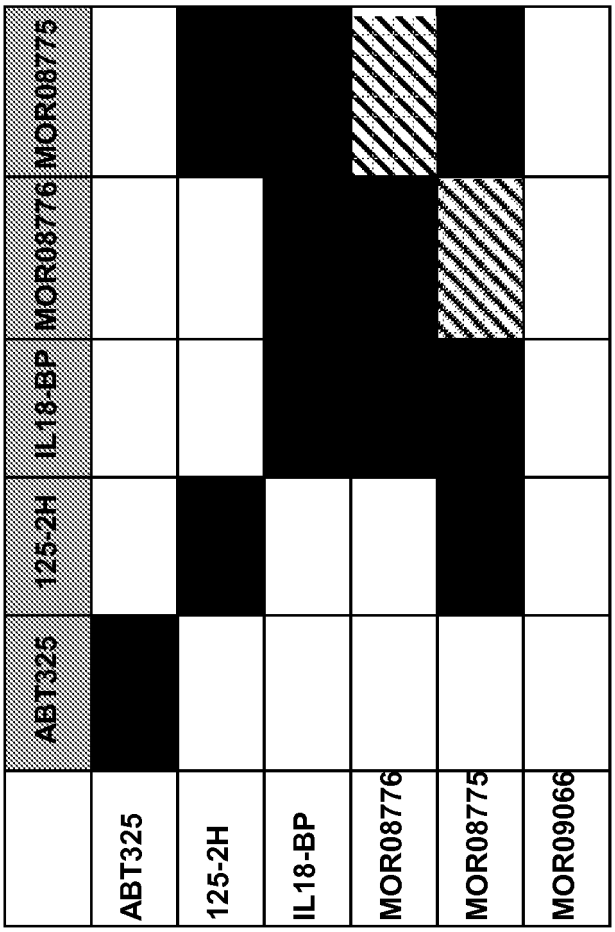


Figure 10(A)

Figure 10(B)

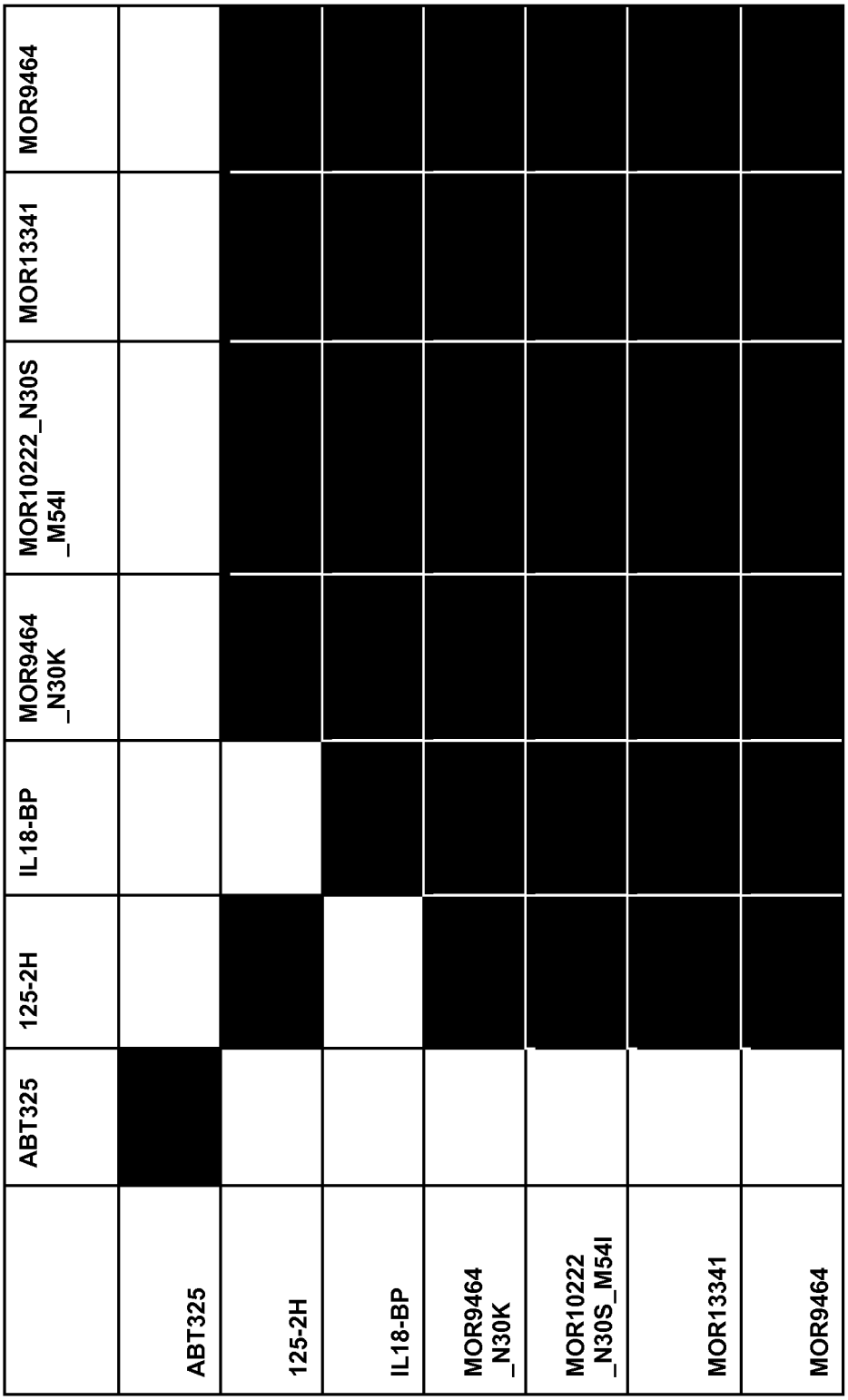

ABT325
125-2H
IL18-BP
MOR9464_N30K
MOR10222_N30S_M54I
MOR13341
MOR9464

Figure 11(A)

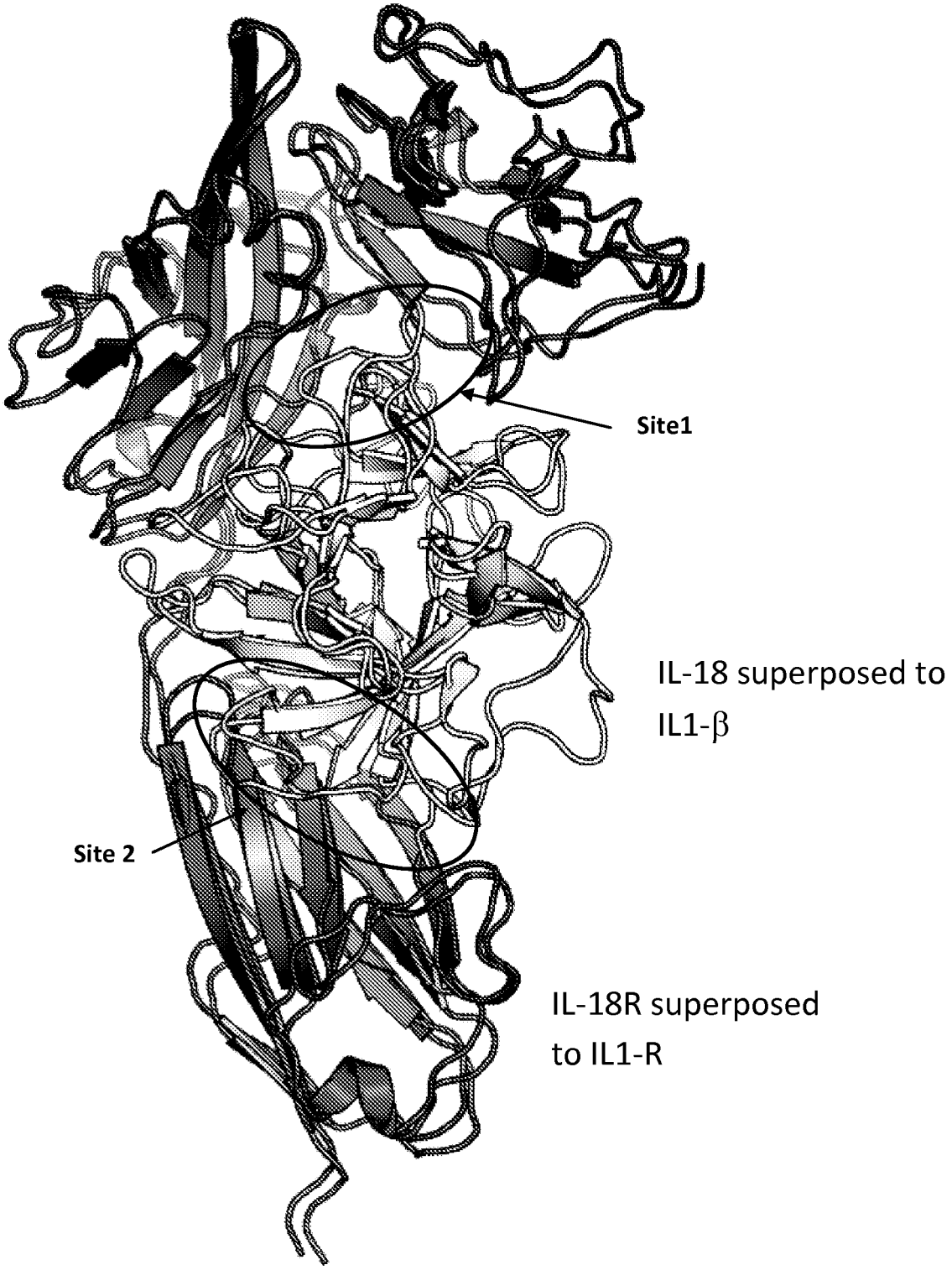
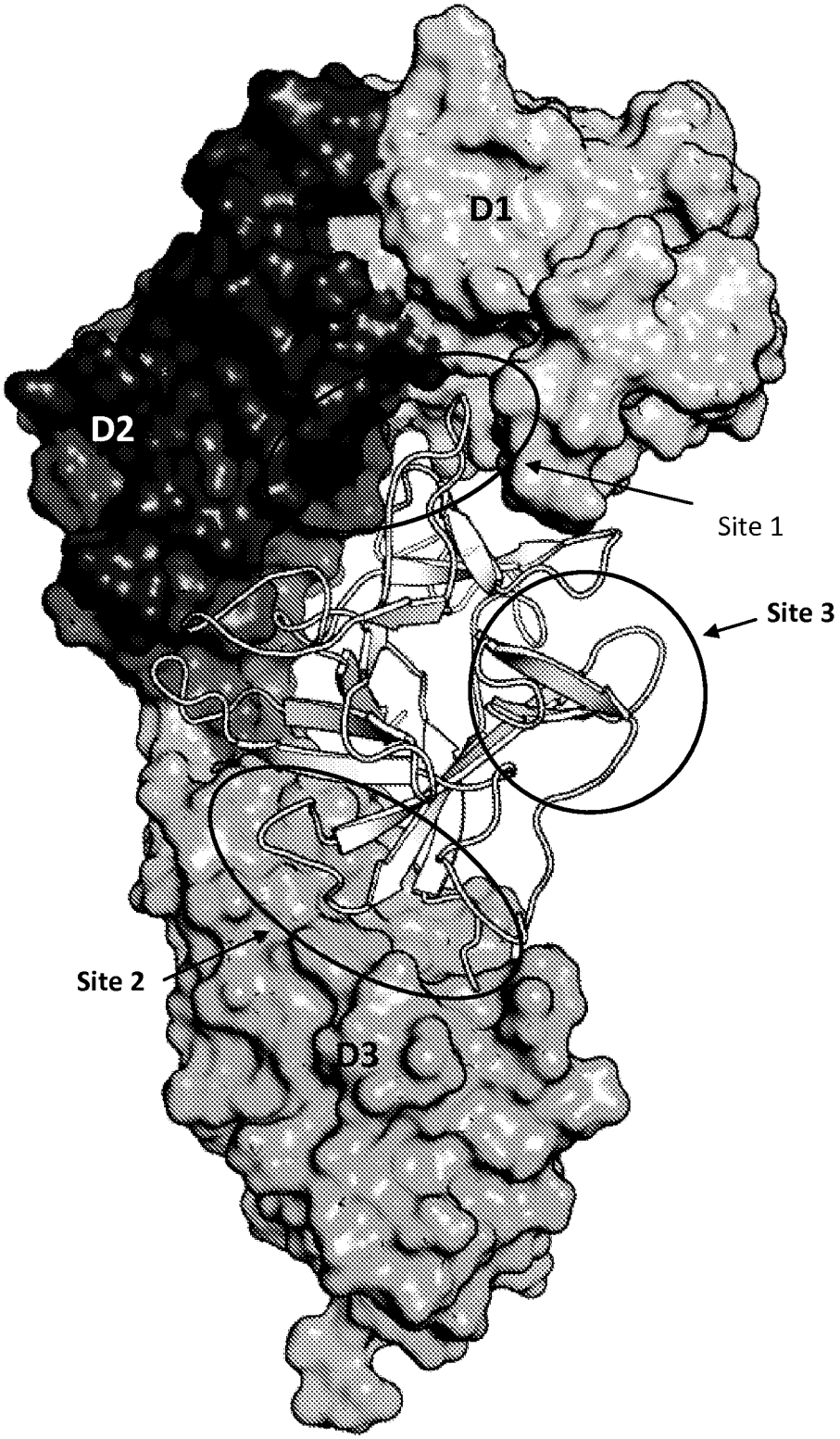

Site1
IL-18 superposed to IL1-β
Site 2
IL-18R superposed to IL1-R

Figure 11(B)



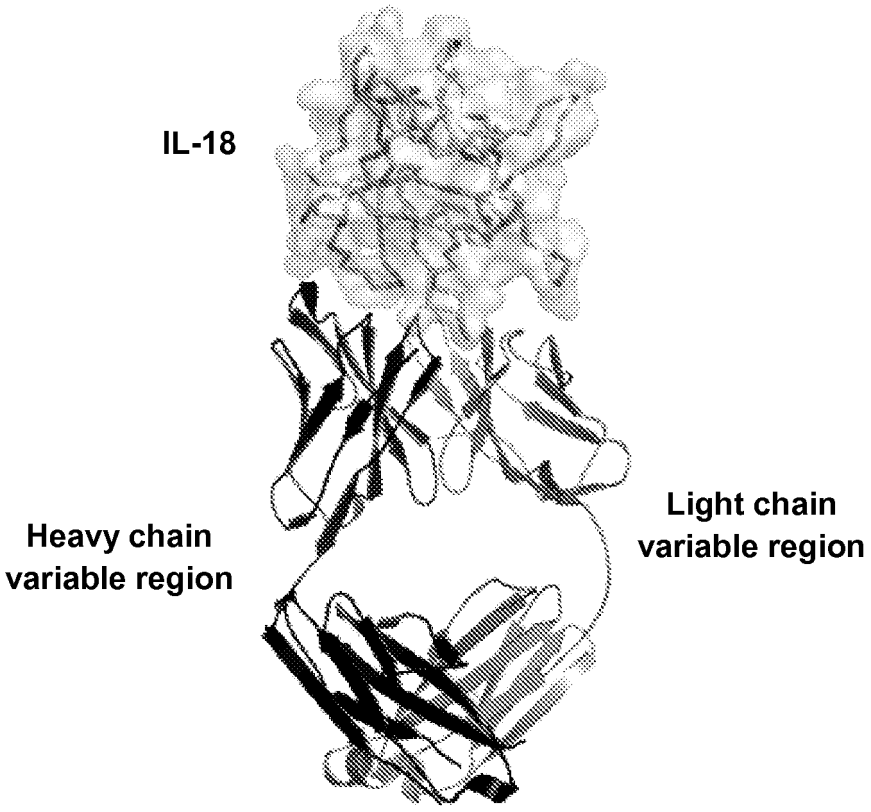
1) Human IL-18	YFGK^{40}LESKLSVIRN^{50}LNDQVLFIDQ^{60}GNRPLFEDMT^{70}DSDCRDNAPR^{80}TIFIISMYKD^{90}SQPRGMAVTI^{100}SVKCENKI
2) Kim et all	E I M K M
3) Krumm et all	YFG L K I MYKD^{90}S PRGMAV
4) IL-18Rα	K^{40}L K R D M D R M
5) H/DxMS	MYKD^{90}SQPRGMAVT

1) Human IL-18	STL^{110}SCENKIISFK^{120}EMNPPDNIKD^{130}TKSDIIFFQR^{140}SVPGHDNKMQ^{150}FESSSYEGYF^{160}LACEKERDLF^{170}KLILK
2) Kim et all	N QR^{140}S DNKM
3) Krumm et all	N QR^{140}S DN M
4) IL-18Rα	R D
5) H/DxMS	FK^{120}EMNPPDNIKD^{130}TKSDIIFFQR^{140}SVPGHDNKMQ^{150}FESSSYEGYF

1) Human IL-18	KEDEL^{180}GDRSIMFTVQ^{190}NED
2) Kim et all	V N
3) Krumm et all	V
4) IL-18Rα	
5) H/DxMS	

Figure 12

Figure 13


IL-18
Heavy chain variable region
Light chain variable region

	47	86	99	115
Human IL-18	YFGKLESKLS**V**IRNLNDQVLFIDQGNRPLFEDMTDSDCRDNAPRTIFII**S**MYKDSQPRGMAV**T**ISVKCEN**K**ISTLSCE			
Cyno. IL-18	YFGKLESKLS**I**IRNLNDQVLFIDQGNRPLFEDMTDSDCRDNAPRTIFII**N**MYKDSQPRGMAV**A**ISVKCEN**R**ISTLSCE			
	*	*	*	*

	170	177
Human IL-18	NKIISFKEMNPPDNIKDTKSDIIFFQRSVPGHDNKMQFESSSYEGYFLACEKERDL**F**KLILKK**E**DELGDRSIMFTVQNED	
Cyno. IL-18	NKIISFKEMNPPDNIKDTKSDIIFFQRSVPGHDNKMQFESSSYEGYFLACEKERDL**Y**KLILKK**K**DELGDRSIMFTVQNED	
	*	*

Figure 14(A)

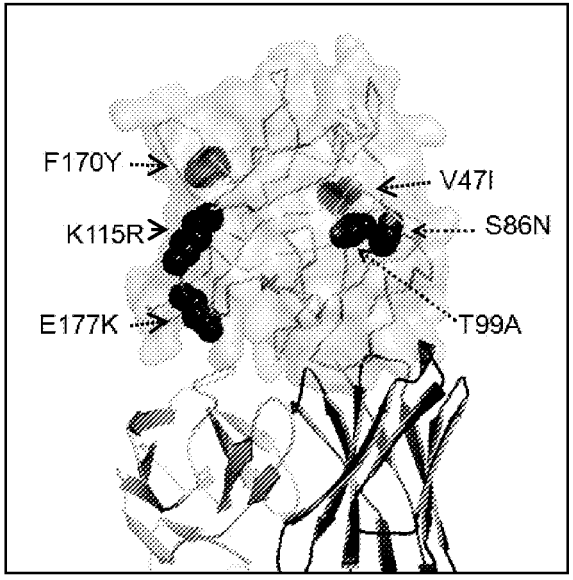
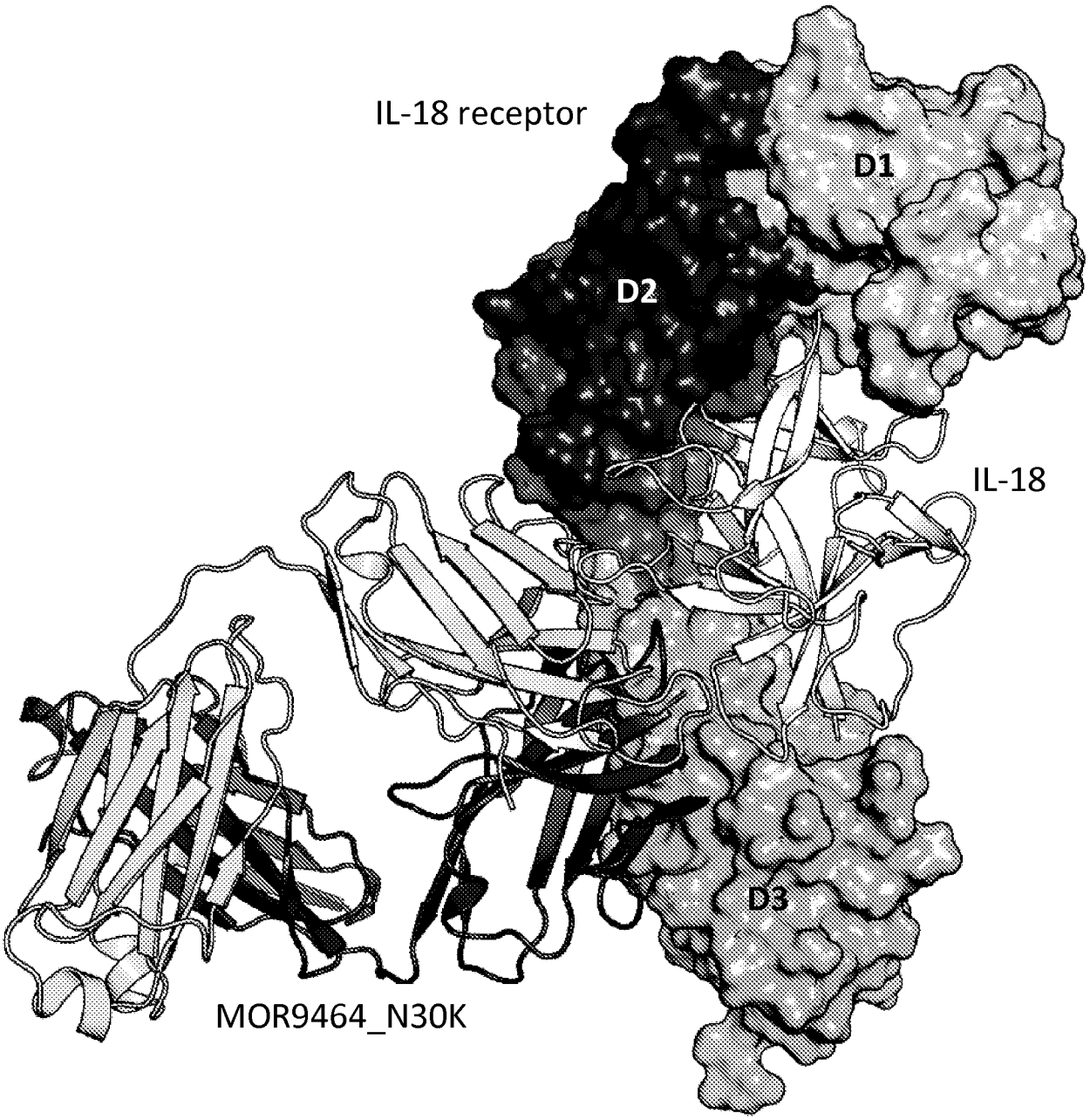


Figure 14(B)

Figure 15



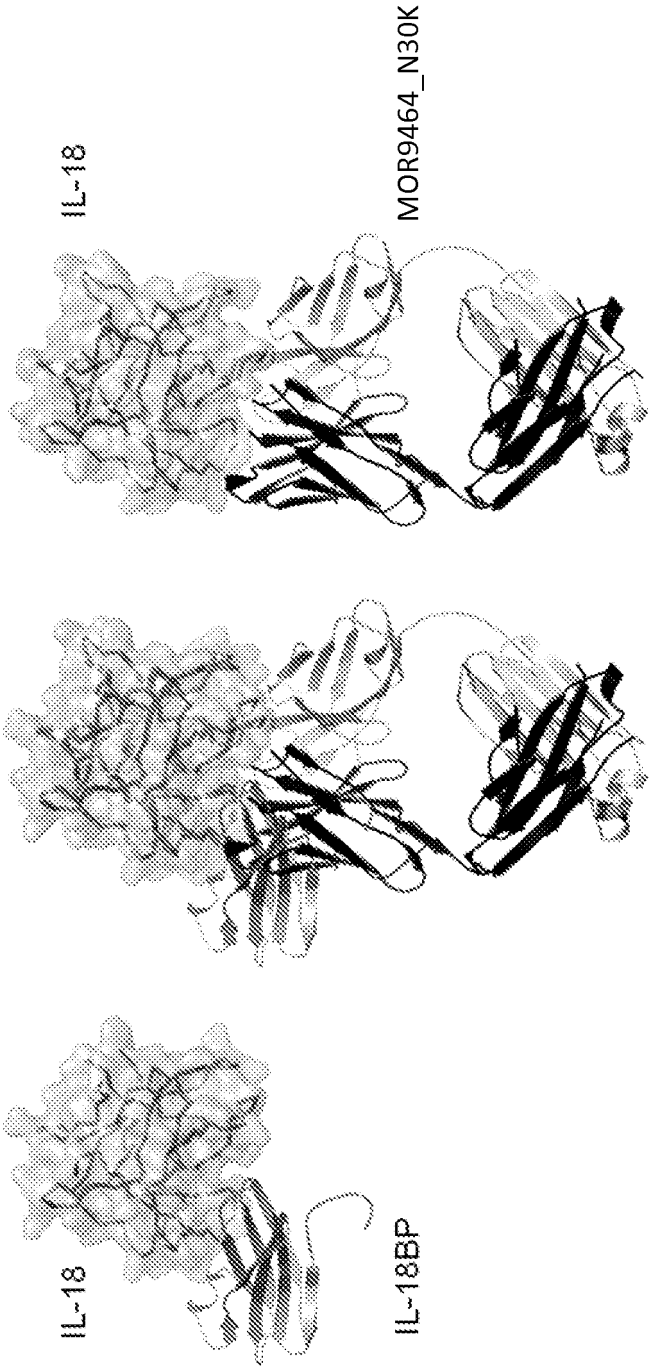


Figure 16

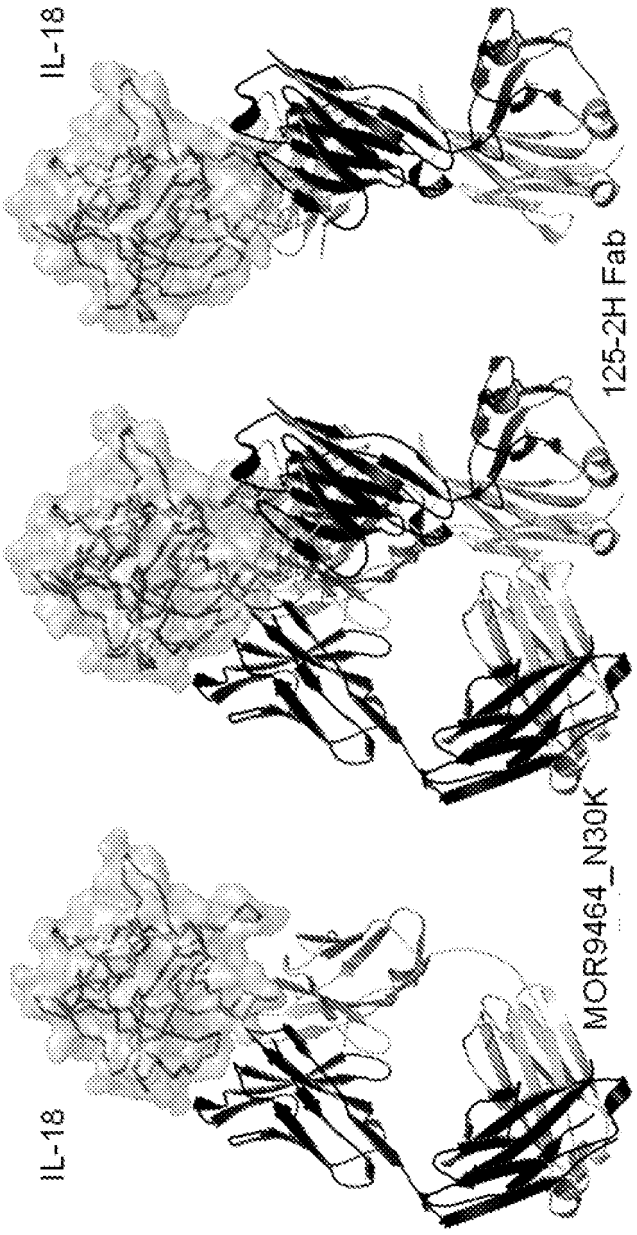


Figure 17

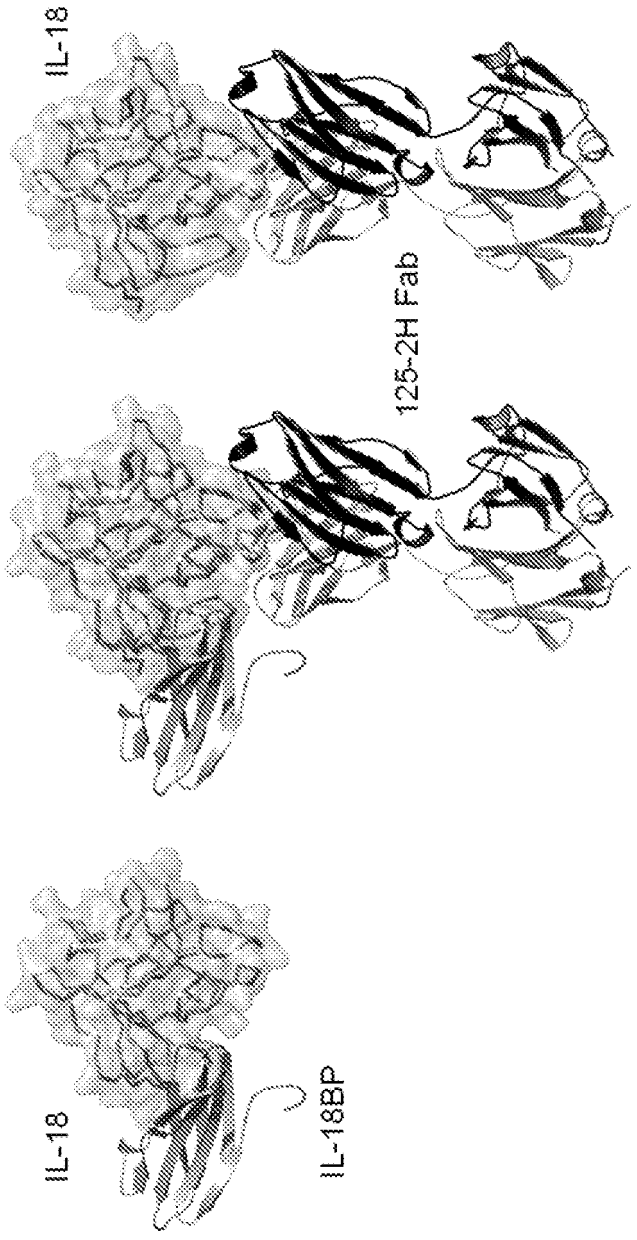


Figure 18

Figure 19

