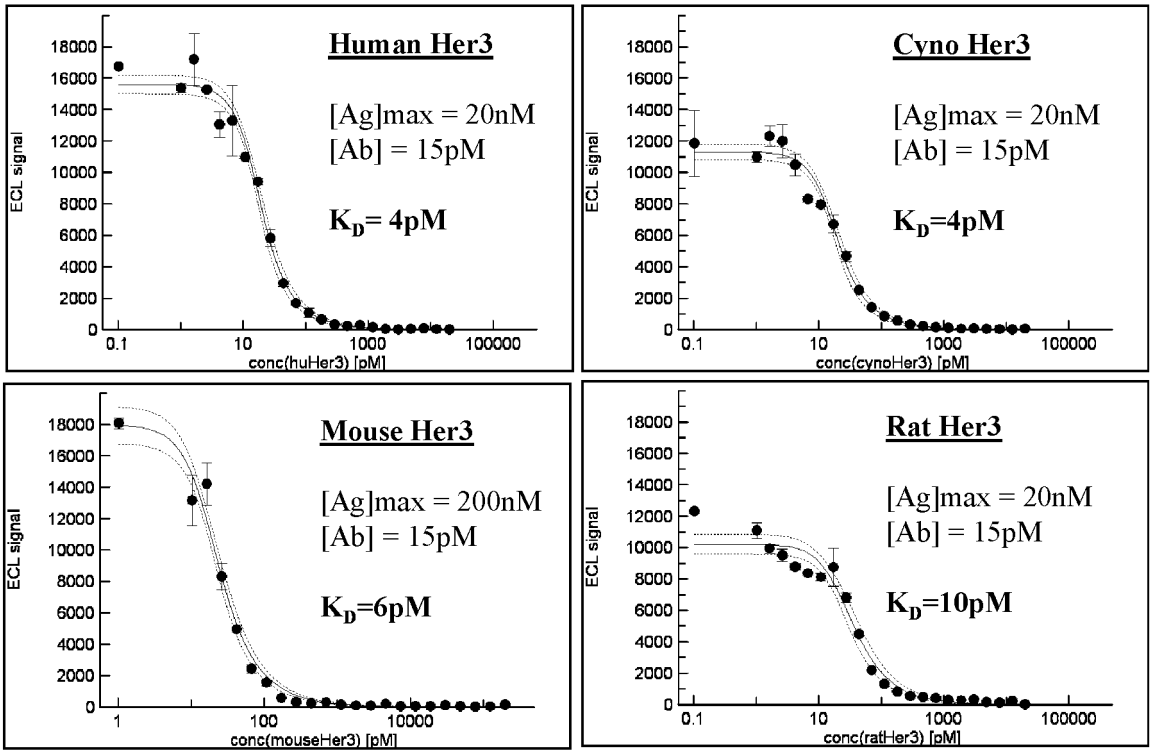

Human Her3
[Ag]max = 20nM
[Ab] = 15pM
K_D= 4pM
ECL signal
conc(huHer3) [pM]
Cyno Her3
[Ag]max = 20nM
[Ab] = 15pM
K_D=4pM
ECL signal
conc(cynoHer3) [pM]
Mouse Her3
[Ag]max = 200nM
[Ab] = 15pM
K_D=6pM
ECL signal
conc(mouseHer3) [pM]
Rat Her3
[Ag]max = 20nM
[Ab] = 15pM
K_D=10pM
ECL signal
conc(ratHer3) [pM]

Fig. 1

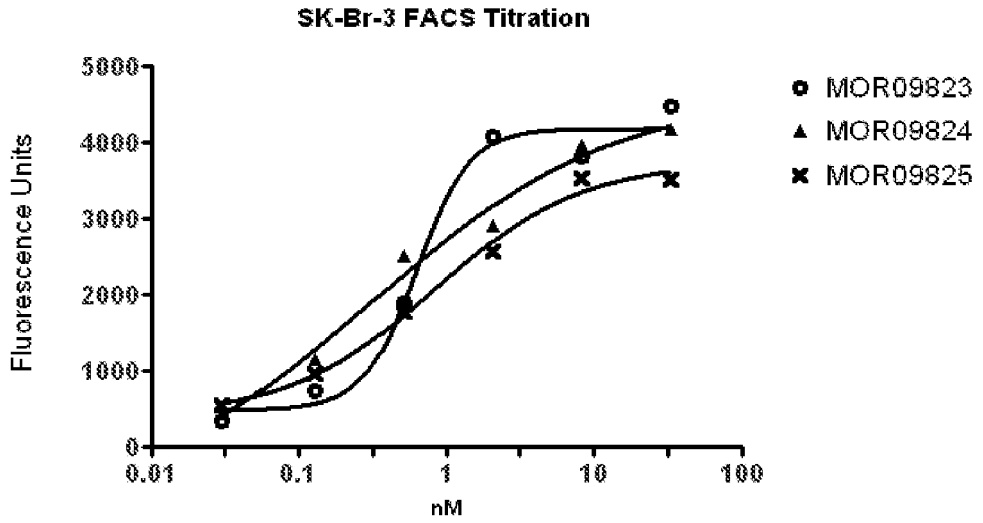

SK-Br-3 FACS Titration
Fluorescence Units
5000
4000
3000
2000
1000
0
0.01
0.1
1
10
100
nM
MOR09823
MOR09824
MOR09825

Fig. 2

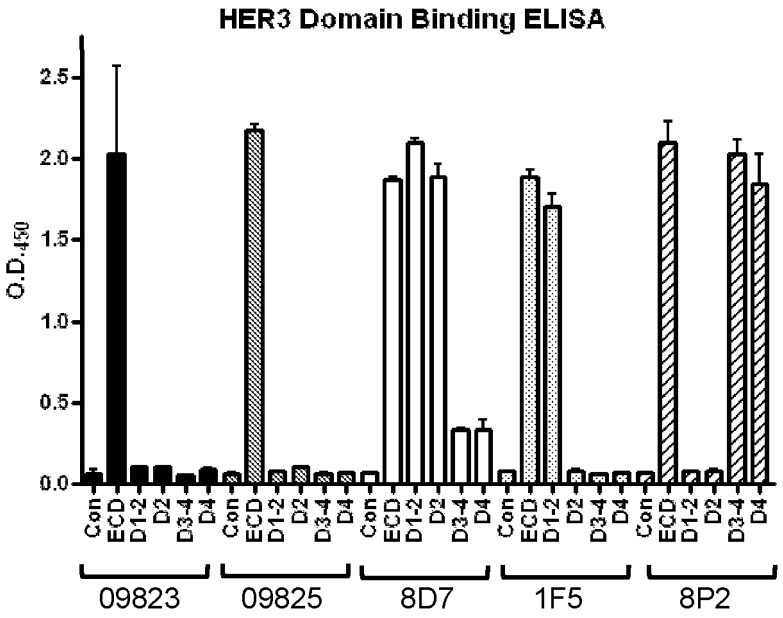

HER3 Domain Binding ELISA
O.D.450
2.5
2.0
1.5
1.0
0.5
0.0
Con
ECD
D1-2
D2
D3-4
D4
09823
09825
8D7
1F5
8P2

Fig. 3


20 SEVGNSQAVC PGTLNGLSVT GDAENQYQTL YKLYERCEVV MGNLEIVLTG HNADLSFLQW IREVTGYVLV AMNEFSTLPL
100 PNLRVVRGTQ VYDGKFAIFV MLNYNTNSSH ALRQLRLTQL TEILSGGVYI EKNDKLCHMD TIDWRDIVRD RDAEIVVKDN
180 GRSCPPCHEV CKGRCWGPGS EDCQTLTKTI CAPQCNGHCF GPNPNQCCHD ECAGGCSGPQ DTDCFACRHF NDSGACVPRC
260 PQPLVYNKLT FQLEPNPHTK YQYGGVCVAS CPHNFVVDQT SCVRACPPDK MEVDKNGLKM CEPCGGLCPK ACEGTGSGSR
340 FQTVDSSNID GFVNCTKILG NLDFLITGLN GDPWHKIPAL DPEKLNVFRT VREITGYLNI QSWPPHMHNF SVFSNLTTIG
420 GRSLYNRGFS LLIMKNLNVT SLGFRSLKEI SAGRIYISAN RQLCYHHSLN WTKVLRGPTE ERLDIKHNRP RRDCVAEGKV
500 CDPLCSSGGC WGPGPGQCLS CRNYSRGGVC VTHCNFLNGE PREFAHEAEC FSCHPECQPM EGTATCNGSG SDTCAQCAHF
580 RDGPHCVSSC PHGVLGAKGP IYKYPDVQNE CRPCHENCTQ GCKGPELQDC LGQTLVLIGK TEFRHDS

Fig. 4A

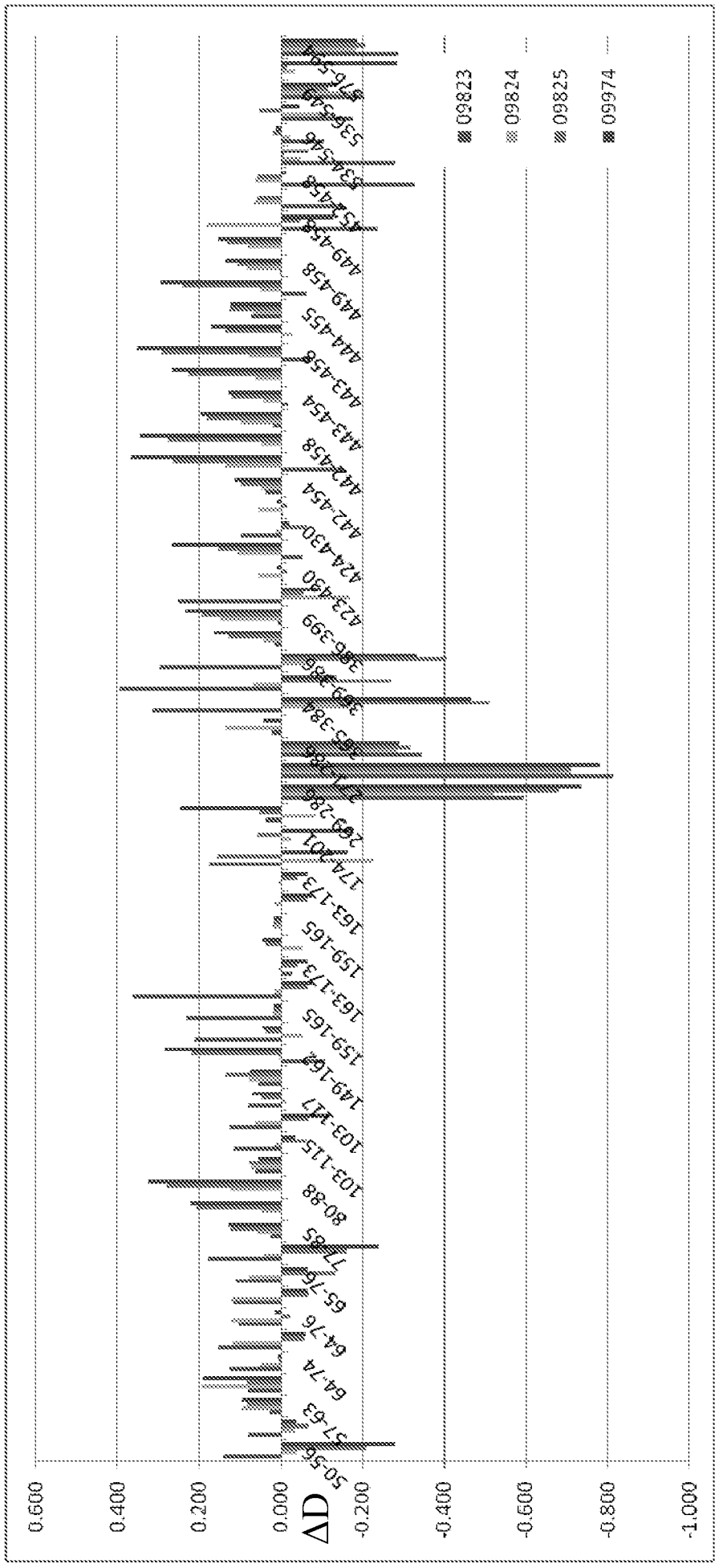

ΔD [HER3: Fab] vs HER3
ΔD
0.600
0.400
0.200
0.000
-0.200
-0.400
-0.600
-0.800
-1.000
09823
09824
09825
09974
50-56
57-63
64-74
64-76
65-76
77-85
80-88
103-115
103-117
149-164
159-165
163-173
159-165
163-173
174-201
269-286
273-285
365-384
379-386
380-399
423-450
424-430
442-454
442-458
443-454
443-458
444-455
449-458
449-455
453-458
534-546
538-546
576-584

Fig. 4B

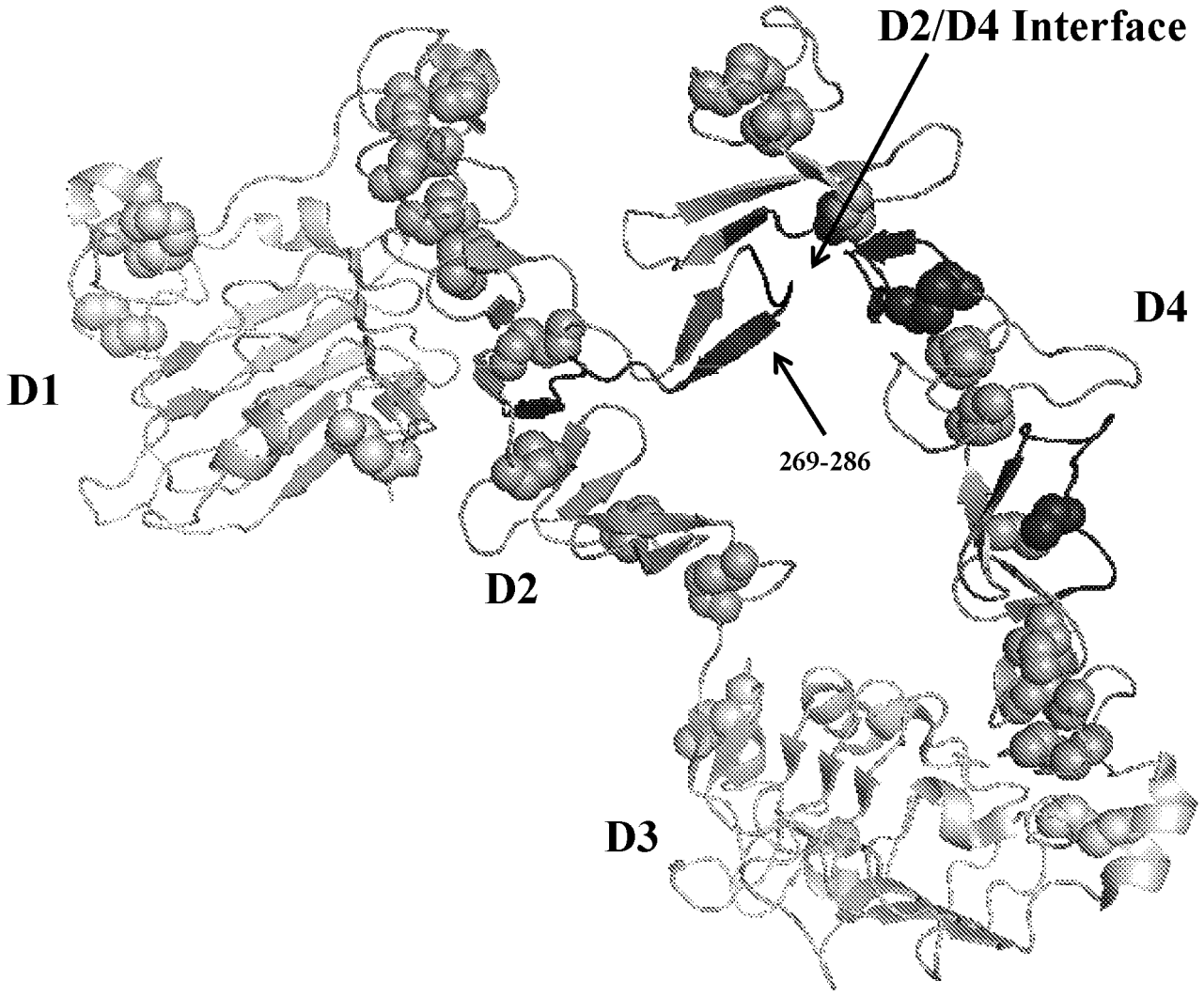


Fig. 4C

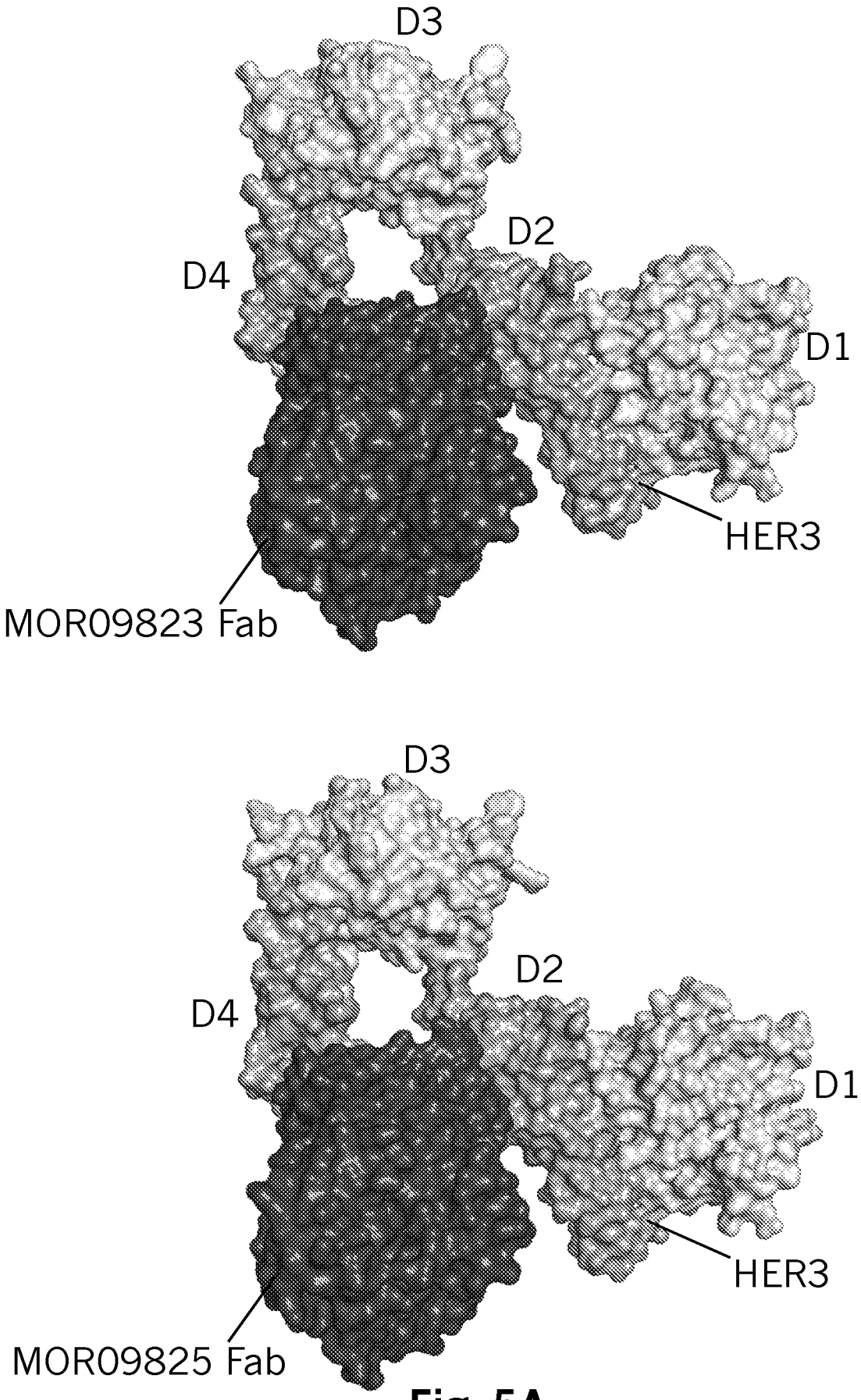


Fig. 5A

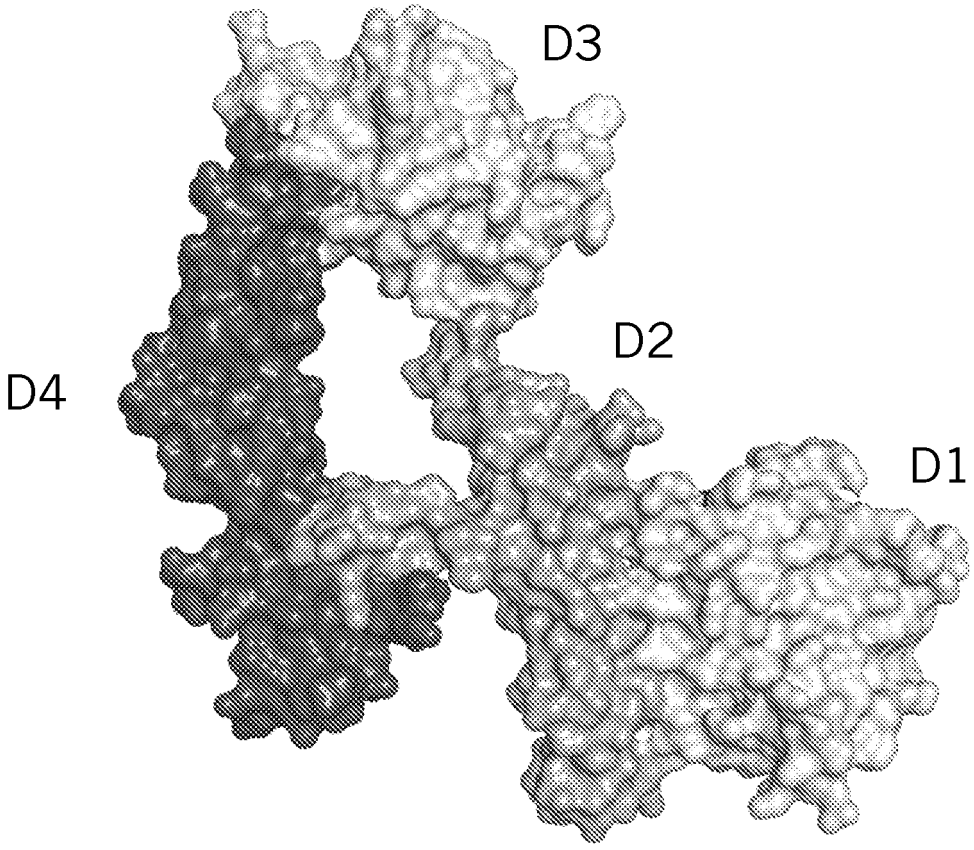


Fig. 5B

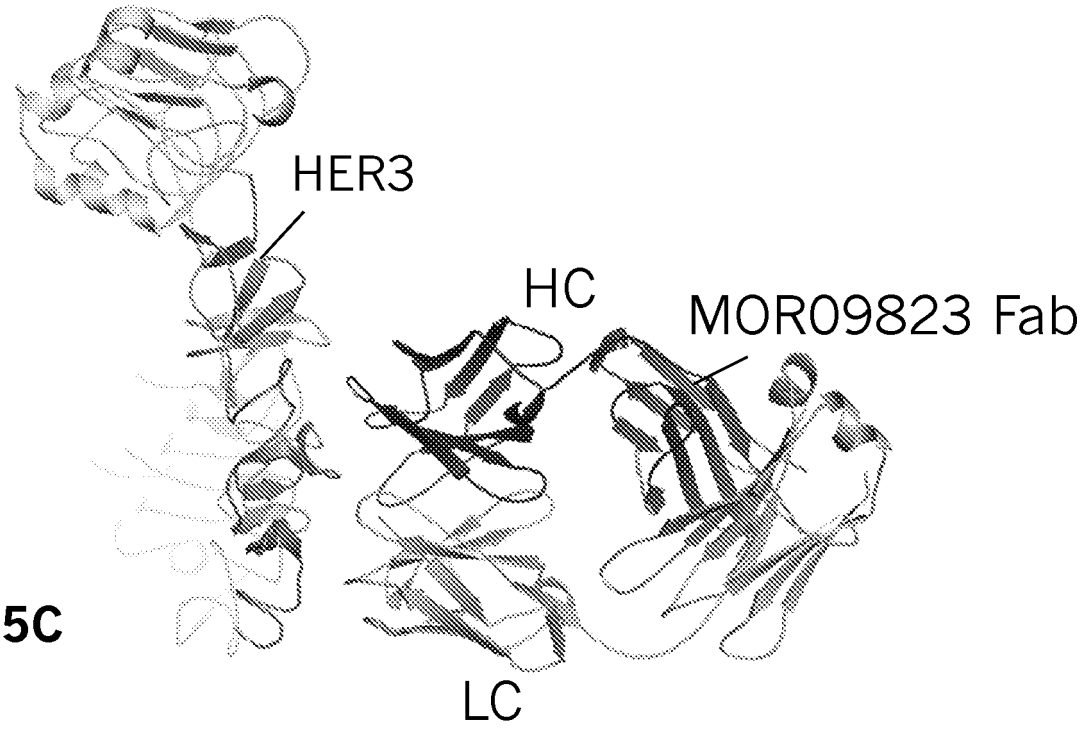

HER3
HC
MOR09823 Fab
LC

Fig. 5C

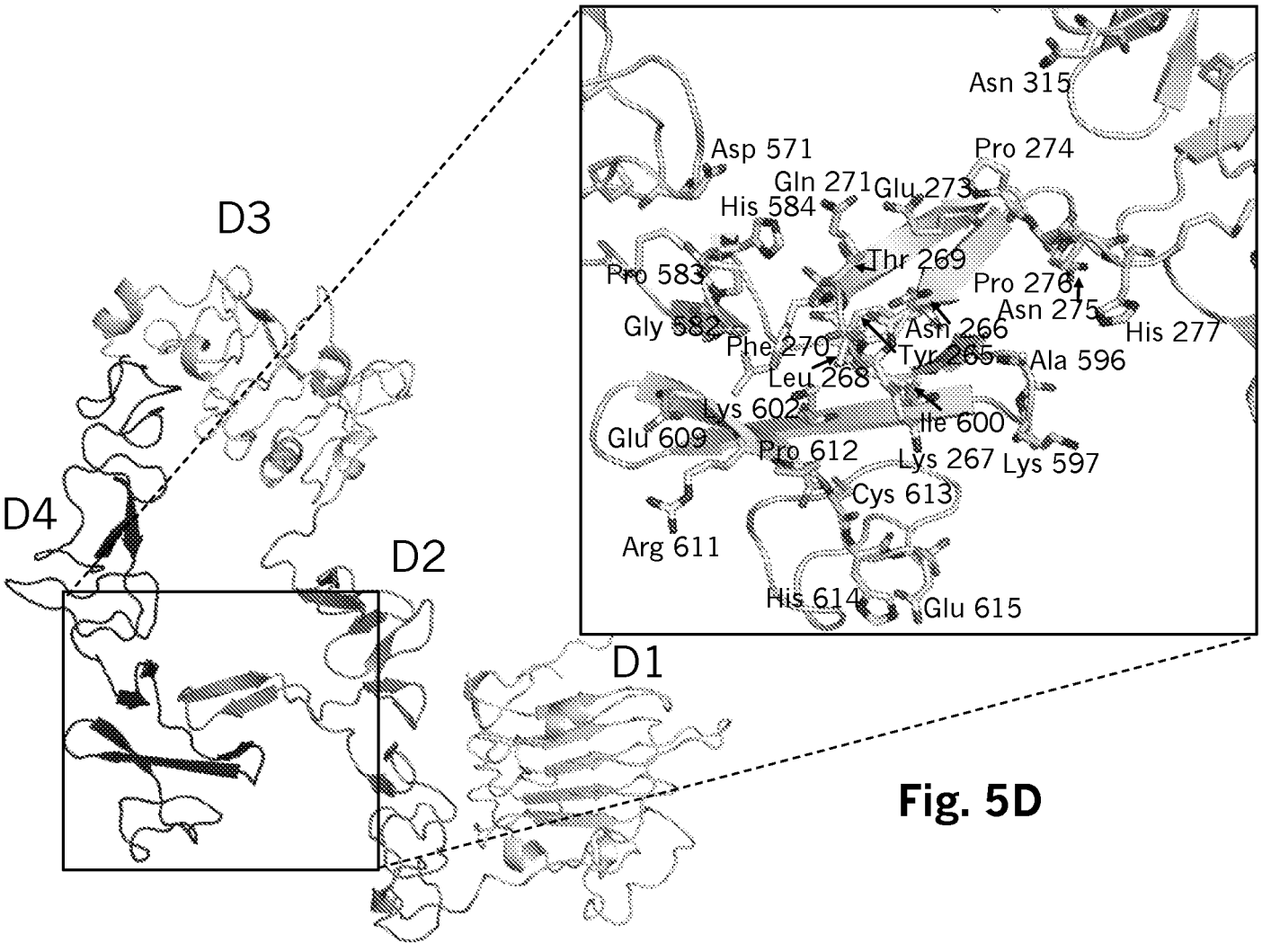

D3
D4
D2
D1
Asn 315
Asp 571
Pro 274
Gln 271
Glu 273
His 584
Thr 269
Pro 583
Pro 276
Asn 275
Gly 582
Asn 266
His 277
Phe 270
Tyr 265
Ala 596
Leu 268
Lys 602
Ile 600
Glu 609
Pro 612
Lys 267
Lys 597
Cys 613
Arg 611
His 614
Glu 615

Fig. 5D

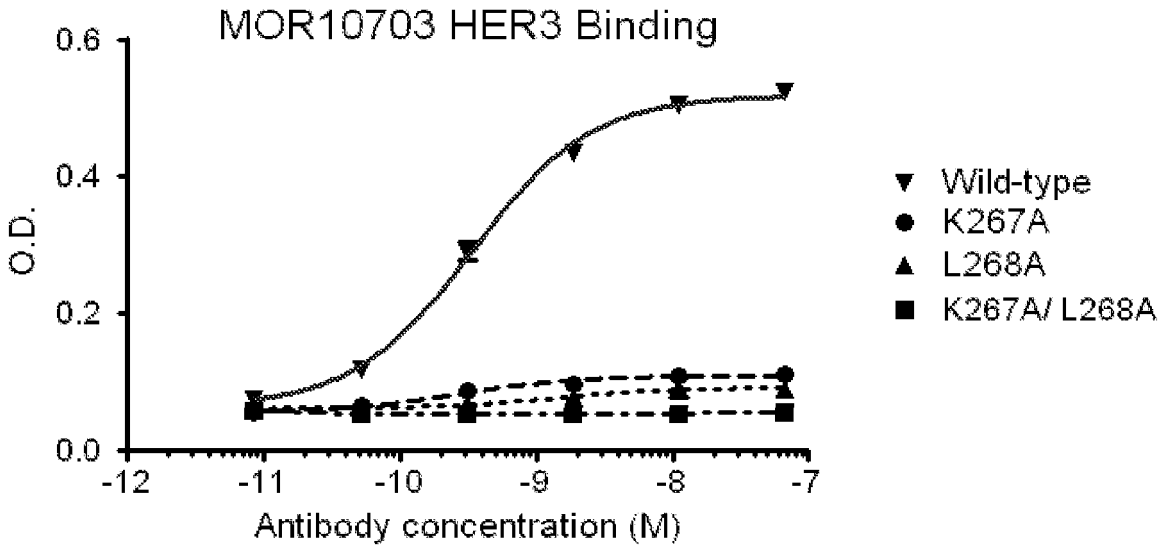

MOR10703 HER3 Binding
0.6
0.4
0.2
0.0
O.D.
-12
-11
-10
-9
-8
-7
Antibody concentration (M)
Wild-type
K267A
L268A
K267A/ L268A

Fig. 5E

Fig. 6A

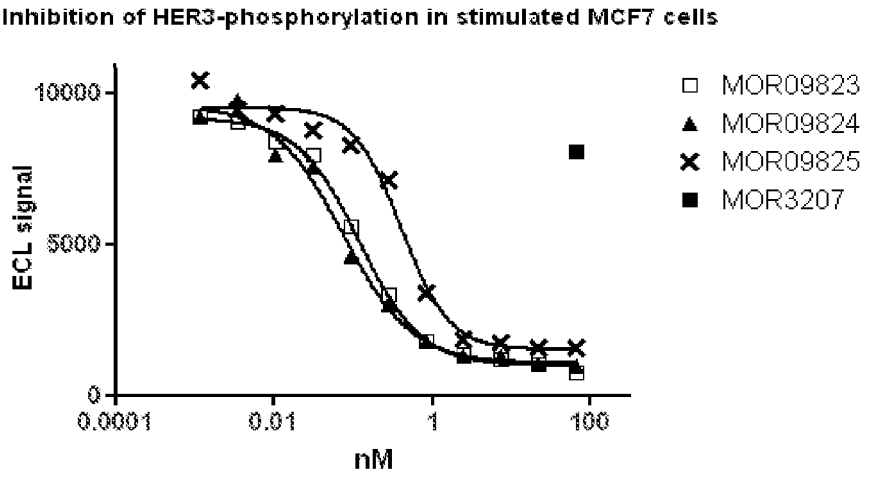

Inhibition of HER3-phosphorylation in stimulated MCF7 cells
ECL signal
10000
5000
0
0.0001
0.01
1
100
nM
MOR09823
MOR09824
MOR09825
MOR3207

Fig. 6B

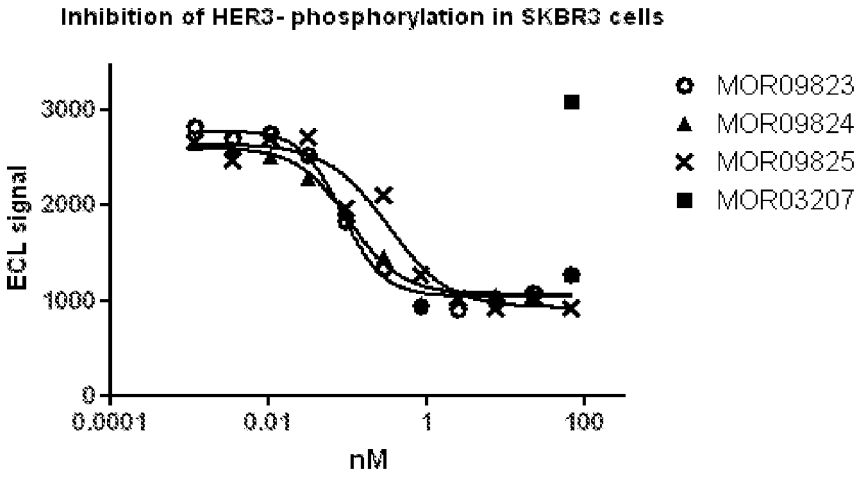

Inhibition of HER3- phosphorylation in SKBR3 cells
ECL signal
3000
2000
1000
0
0.0001
0.01
1
100
nM
MOR09823
MOR09824
MOR09825
MOR03207

Fig. 7A

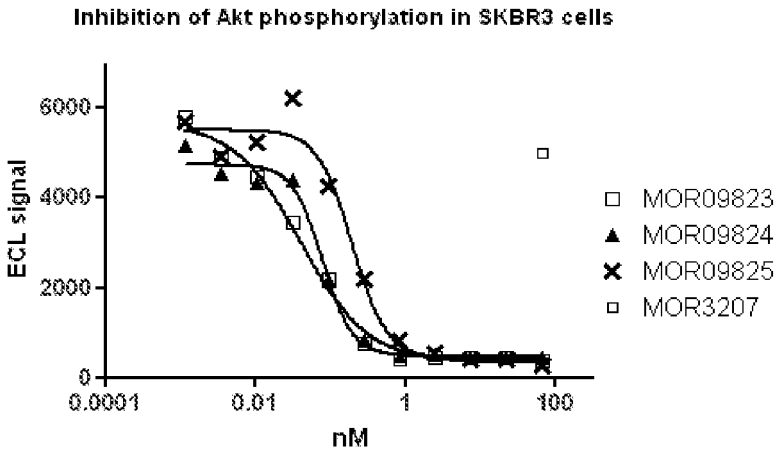

Inhibition of Akt phosphorylation in SKBR3 cells
ECL signal
6000
4000
2000
0
0.0001
0.01
1
100
nM
MOR09823
MOR09824
MOR09825
MOR3207

Fig. 7B

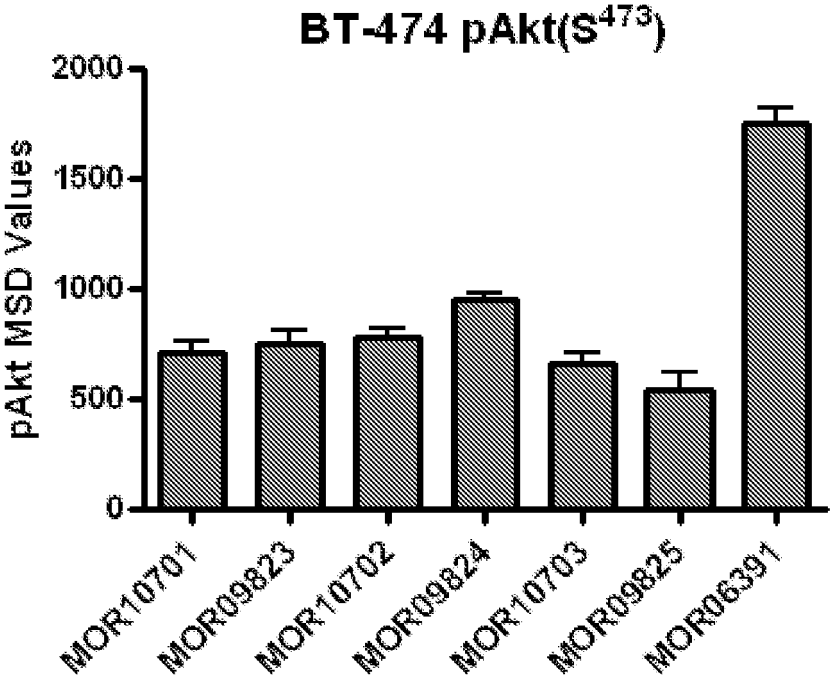

BT-474 pAkt(S473)
pAkt MSD Values
2000
1500
1000
500
0
MOR10701
MOR09823
MOR10702
MOR09824
MOR10703
MOR09825
MOR06391

Fig. 8A

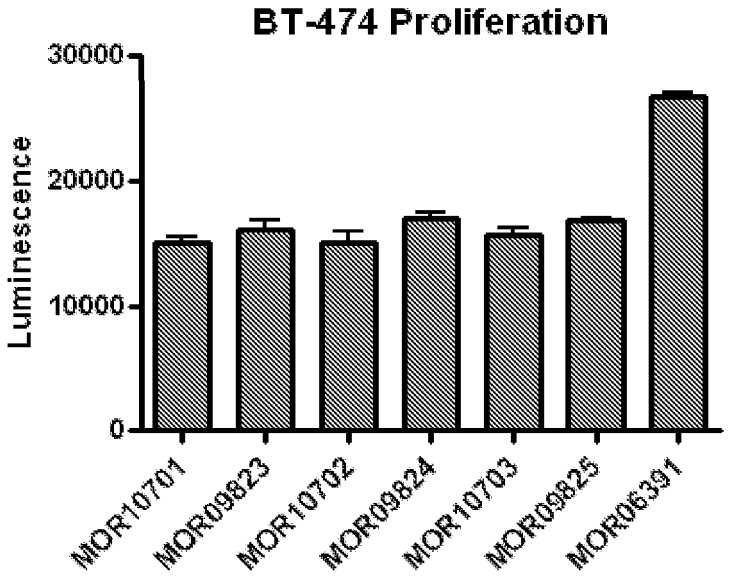
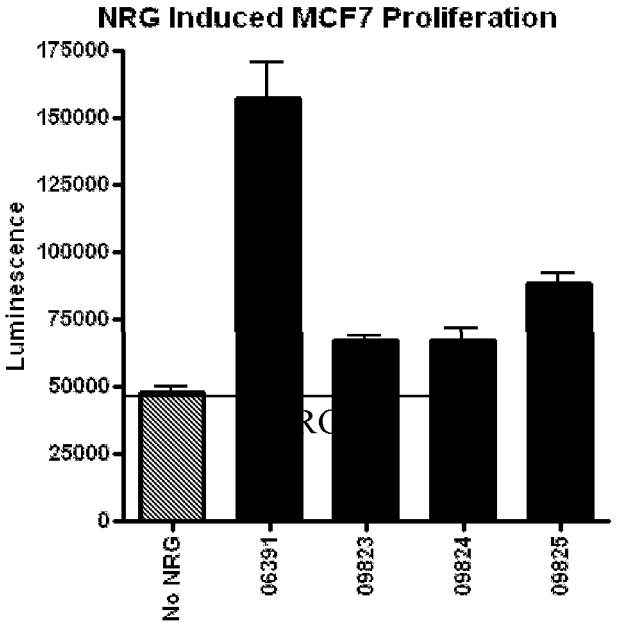

BT-474 Proliferation
Luminescence
30000
20000
10000
0
MOR10701
MOR09823
MOR10702
MOR09824
MOR10703
MOR09825
MOR06391

Fig. 8B


NRG Induced MCF7 Proliferation
Luminescence
175000
150000
125000
100000
75000
50000
25000
0
No NRG
06391
09823
09824
09825

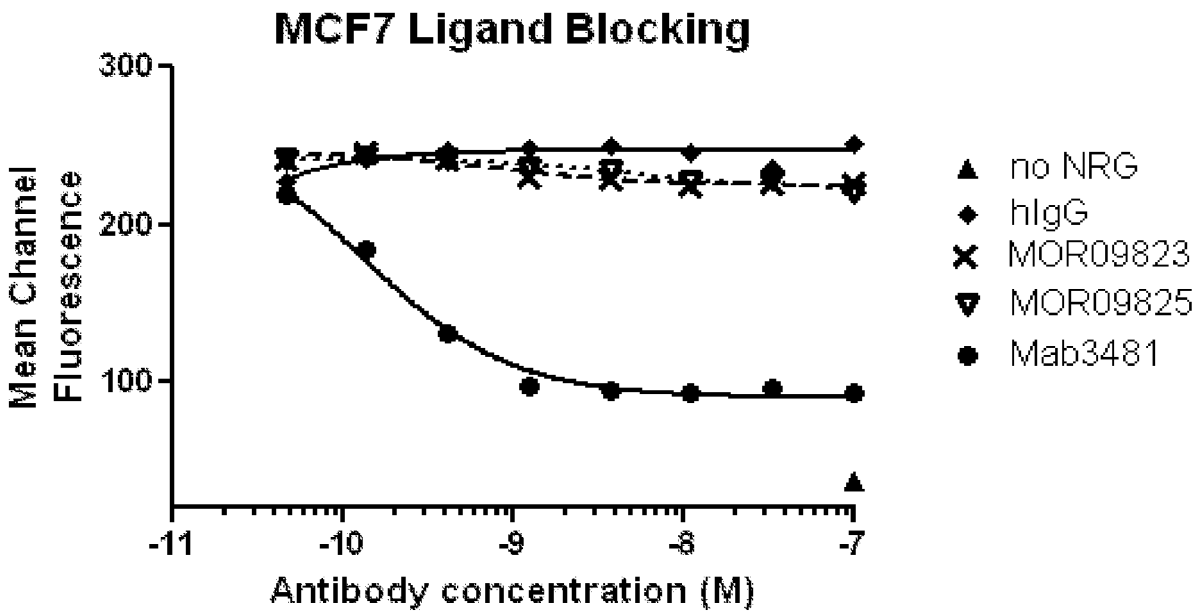

MCF7 Ligand Blocking
Mean Channel Fluorescence
300
200
100
-11
-10
-9
-8
-7
Antibody concentration (M)
no NRG
hIgG
MOR09823
MOR09825
Mab3481

Fig. 9

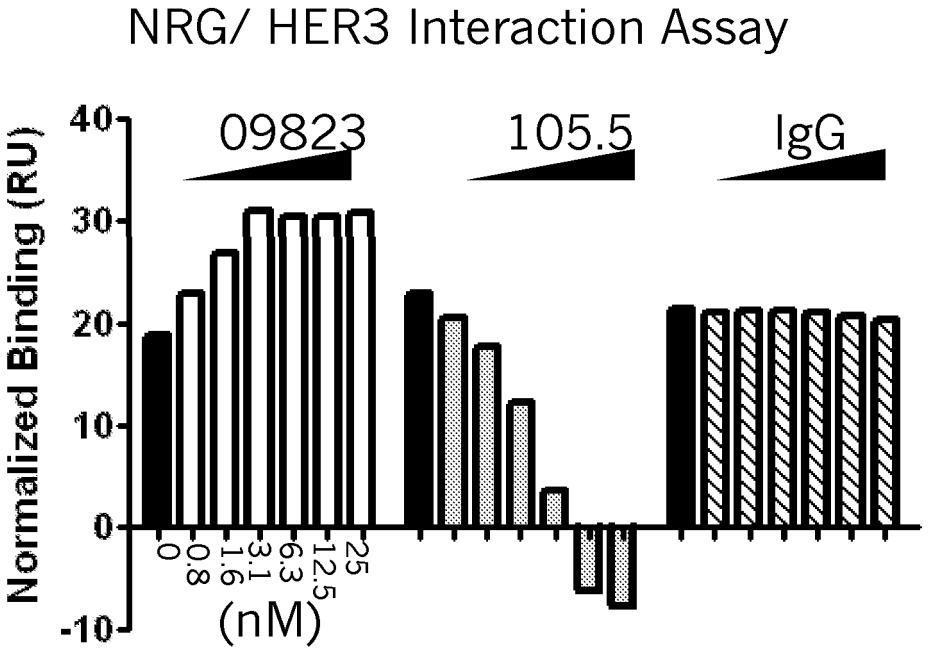

NRG/ HER3 Interaction Assay
09823
105.5
IgG
Normalized Binding (RU)
40
30
20
10
0
-10
0
0.8
1.6
3.1
6.3
12.5
25
(nM)

Fig. 10

Fig. 11A

MOR0923 treated BT474 xenografts

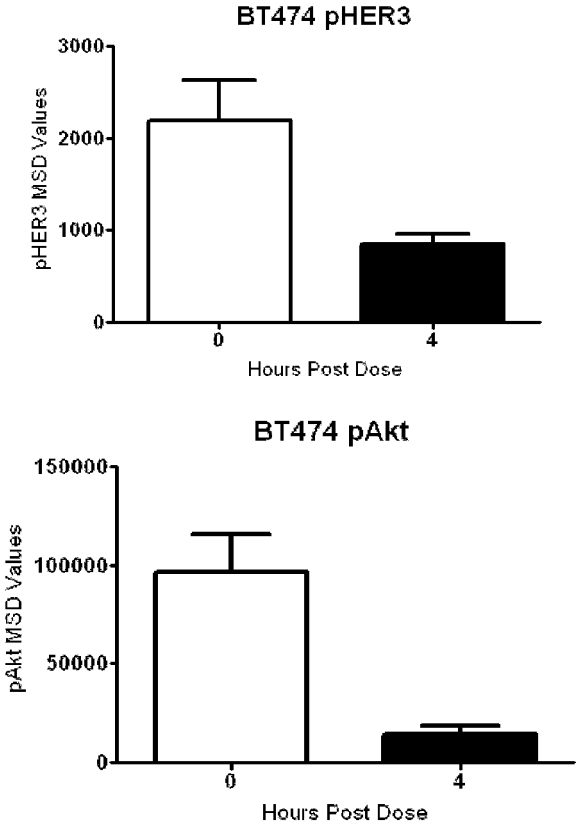


Fig. 11B

MOR09823 treated BxPC3 xenografts

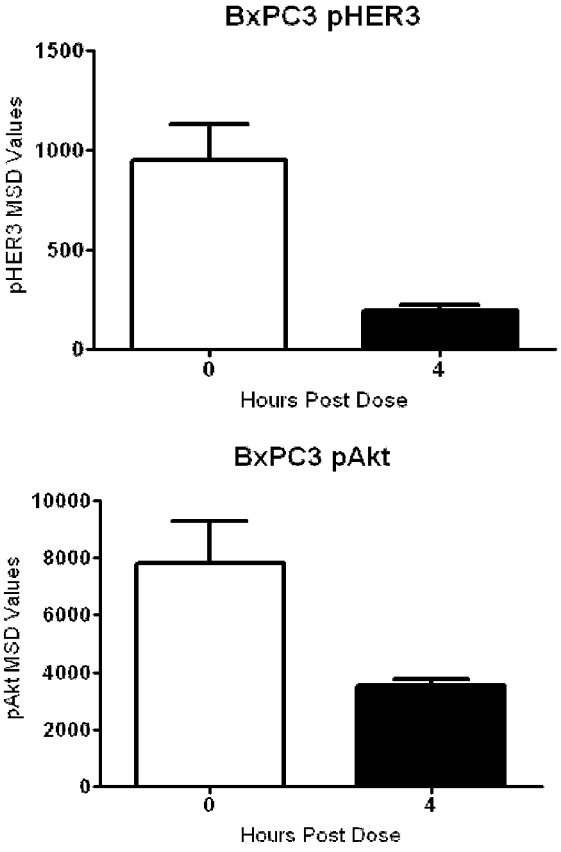


Fig. 12A

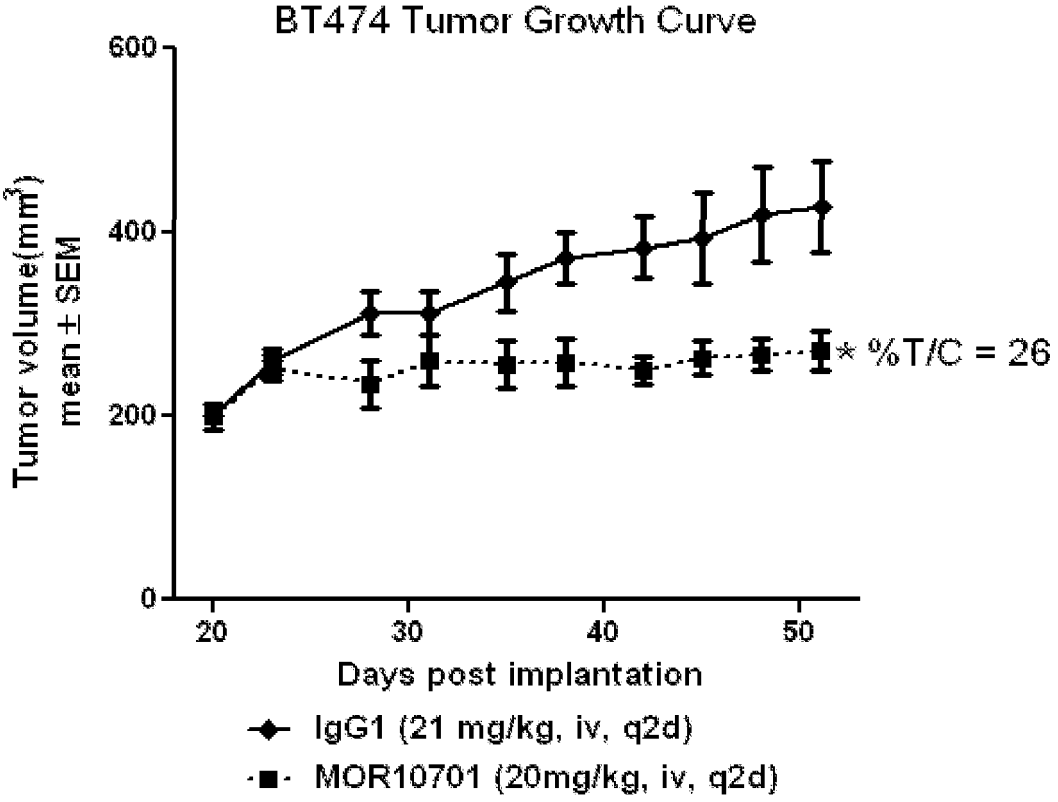
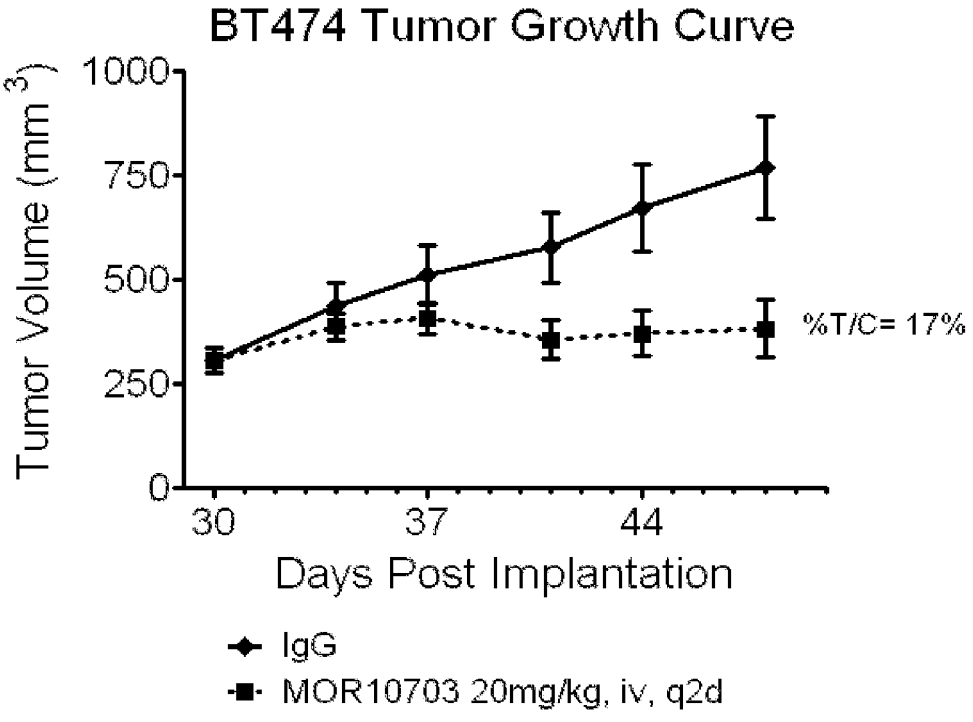


Fig. 12B



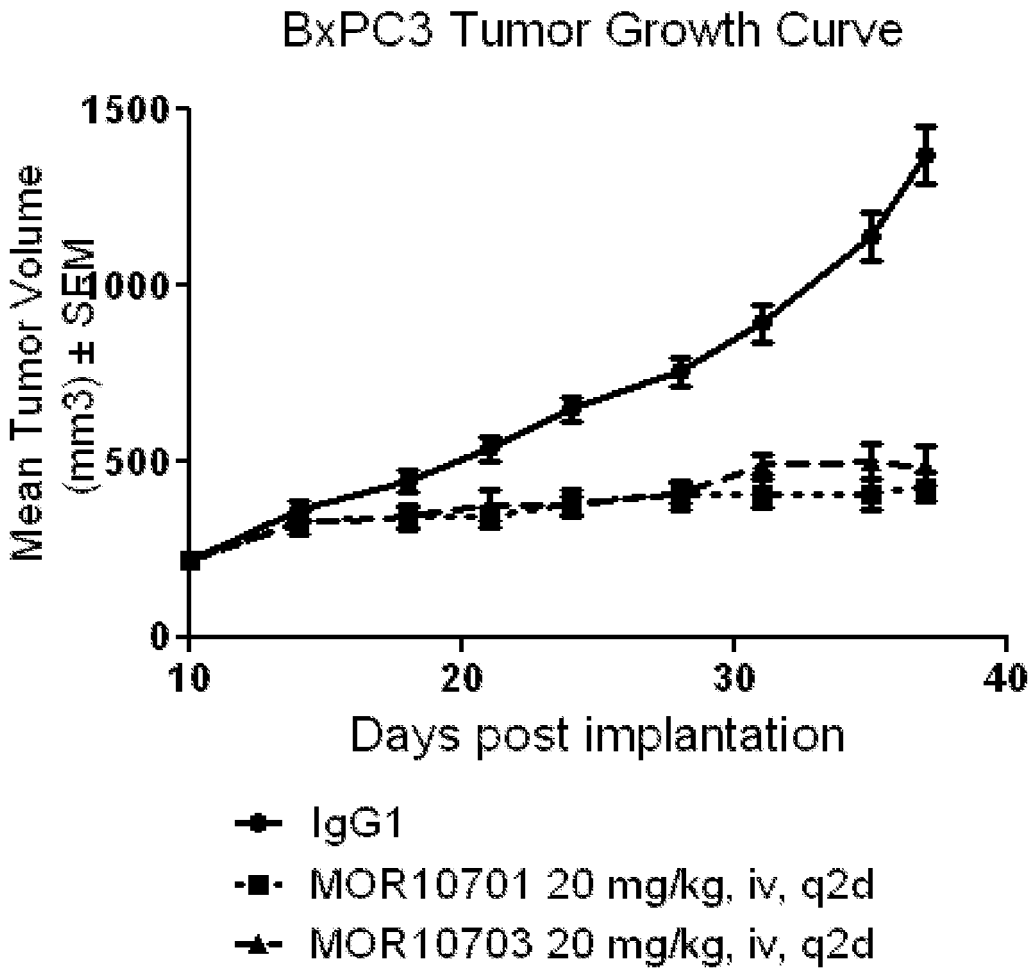

BxPC3 Tumor Growth Curve
Mean Tumor Volume (mm3) ± SEM
1500
1000
500
0
10
20
30
40
Days post implantation
IgG1
MOR10701 20 mg/kg, iv, q2d
MOR10703 20 mg/kg, iv, q2d

Fig. 13

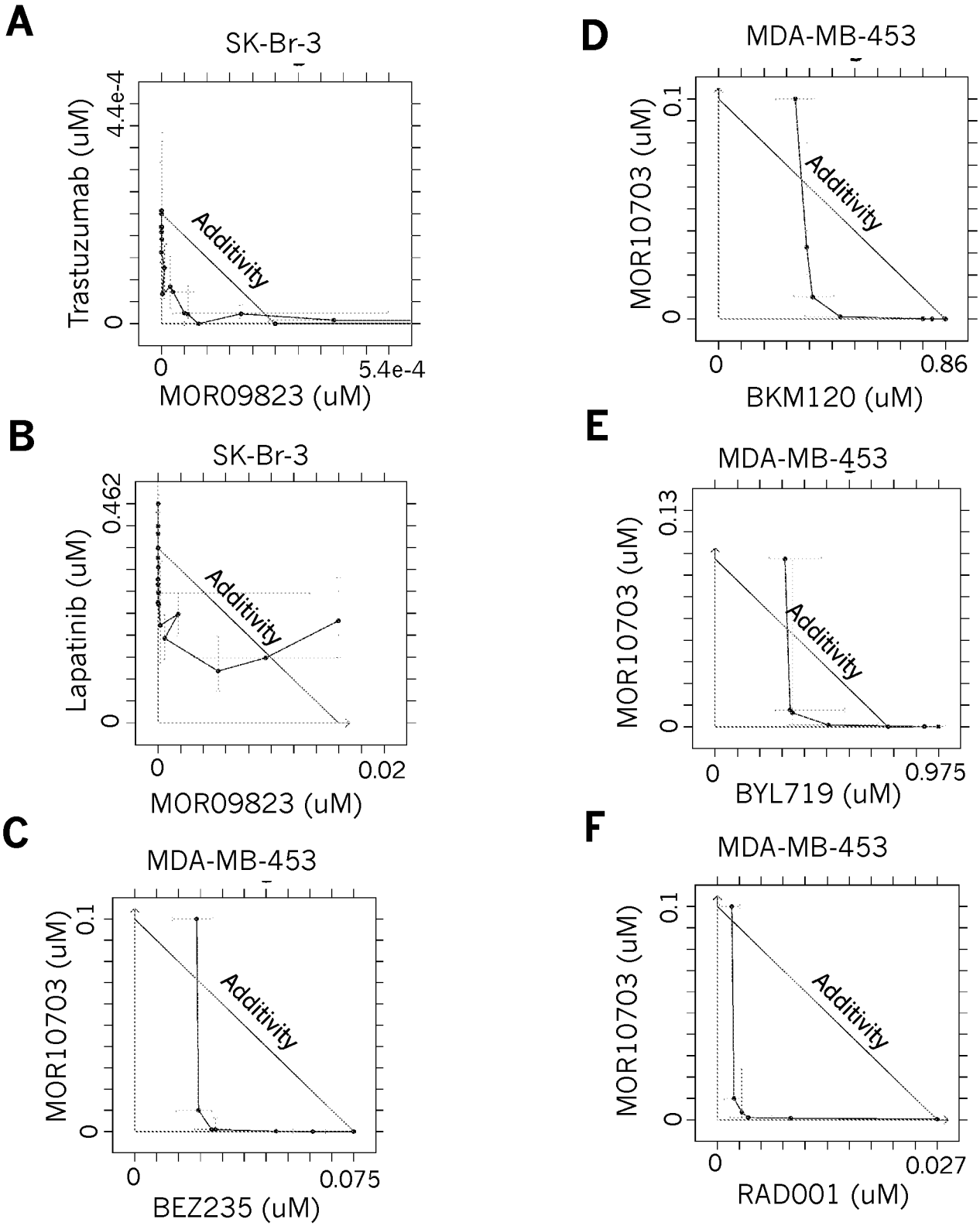
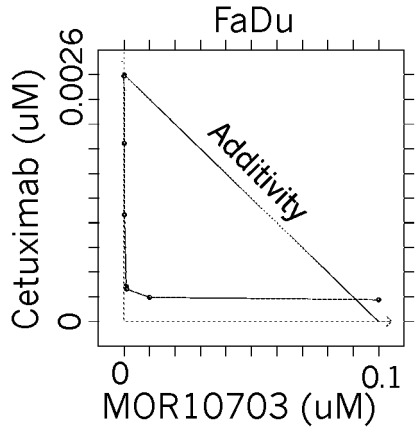

A
SK-Br-3
Trastuzumab (uM)
4.4e-4
0
0
5.4e-4
MOR09823 (uM)
Additivity
B
SK-Br-3
Lapatinib (uM)
0.462
0
0
0.02
MOR09823 (uM)
Additivity
C
MDA-MB-453
MOR10703 (uM)
0.1
0
0
0.075
BEZ235 (uM)
Additivity
D
MDA-MB-453
MOR10703 (uM)
0.1
0
0
0.86
BKM120 (uM)
Additivity
E
MDA-MB-453
MOR10703 (uM)
0.13
0
0
0.975
BYL719 (uM)
Additivity
F
MDA-MB-453
MOR10703 (uM)
0.1
0
0
0.027
RAD001 (uM)
Additivity

Fig. 14

G



H

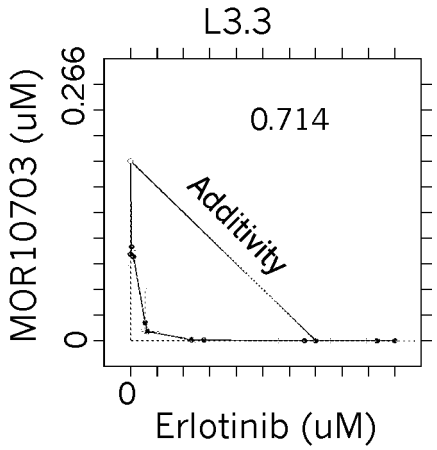
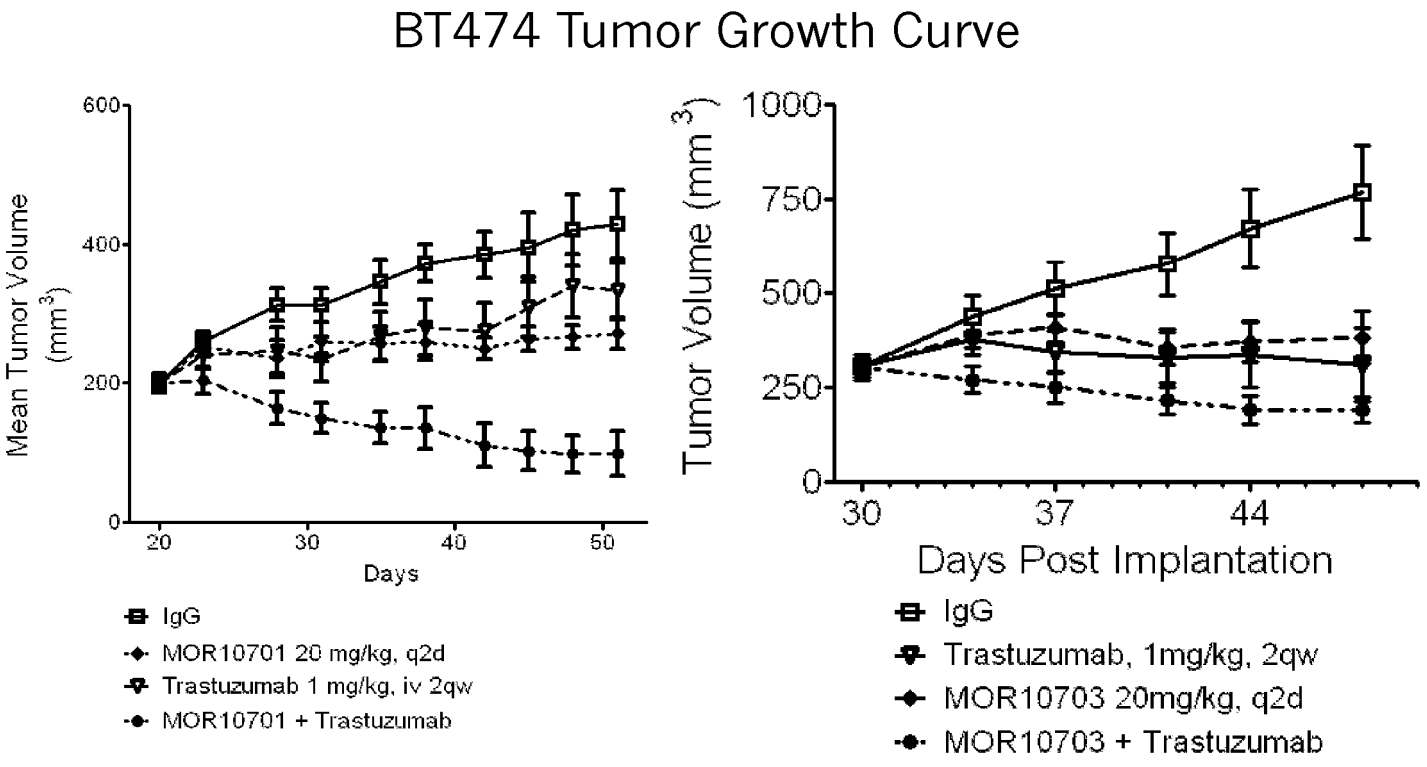


Fig. 14 (cont.)

Fig. 15A



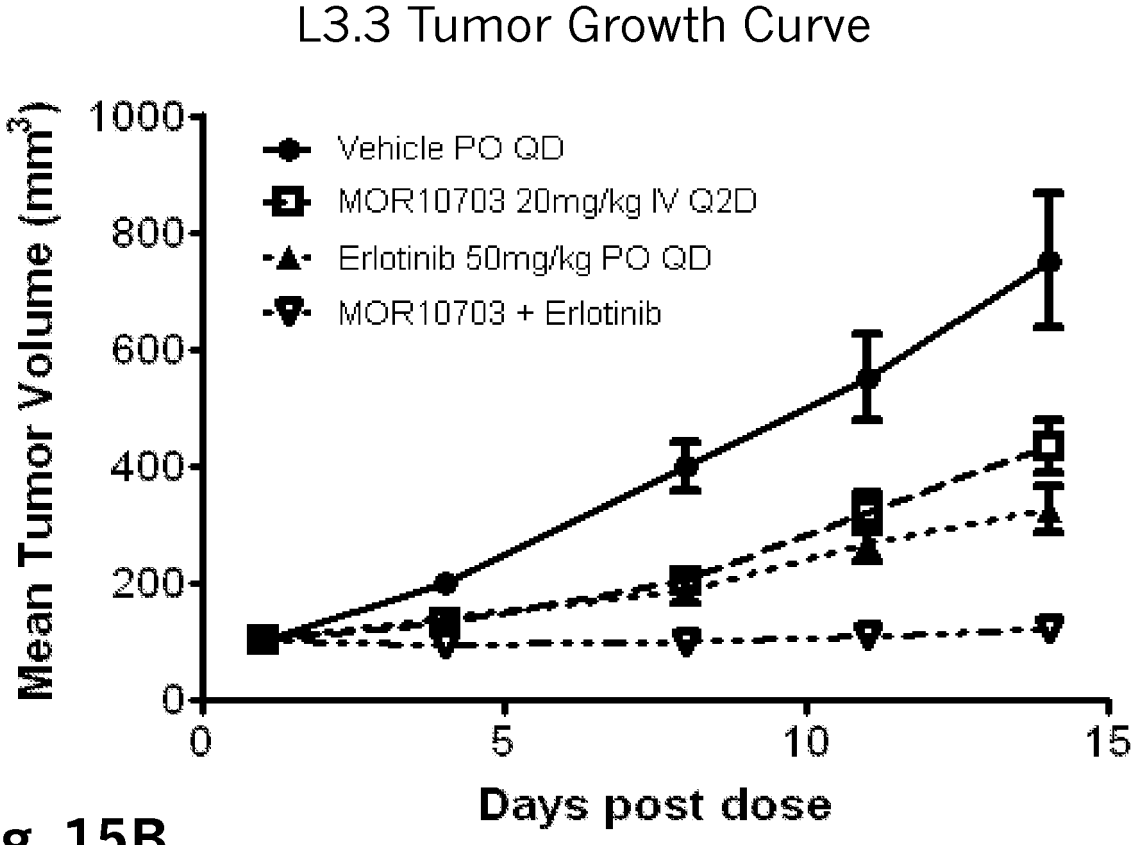


Fig. 15B