

Fig.1

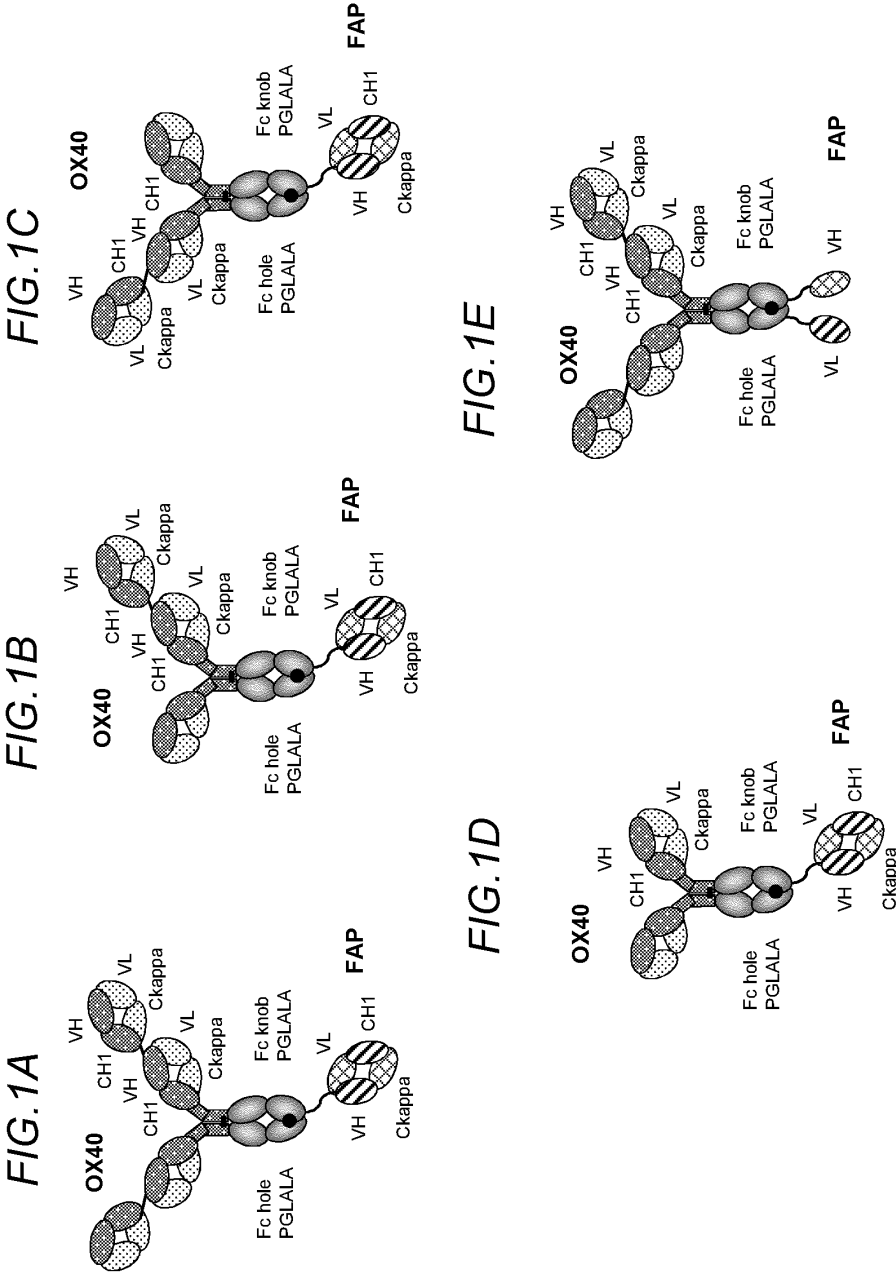


Fig.2

FIG.2A

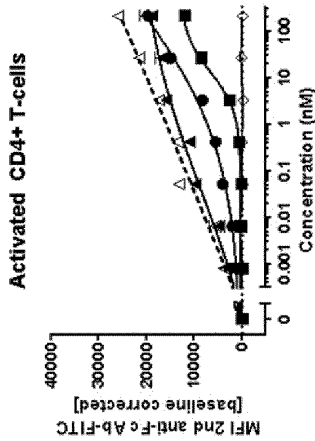


FIG.2B

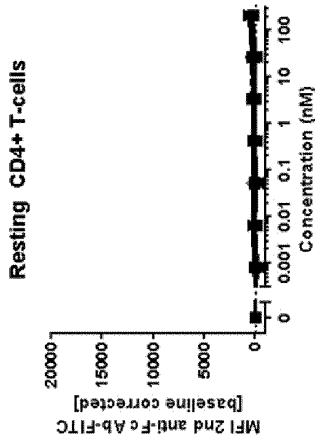


FIG.2C

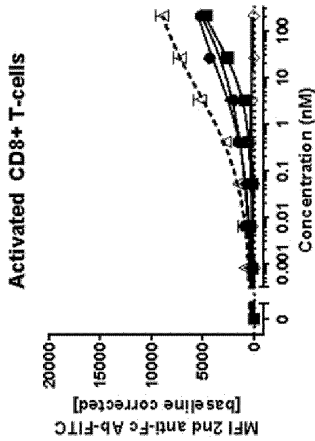
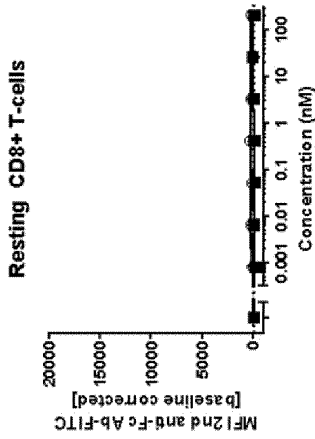


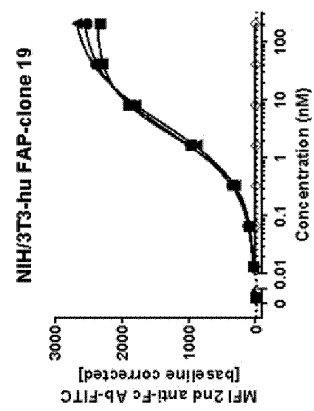
FIG.2D



- ◇ Isotype Control
- ▲ 4+1 49B4/H212 xFab (P1AE6838)
- 3+1 49B4/H212 (P1AE8786)
- 2+1 49B4/H212 (P1AE6840)

Fig.2

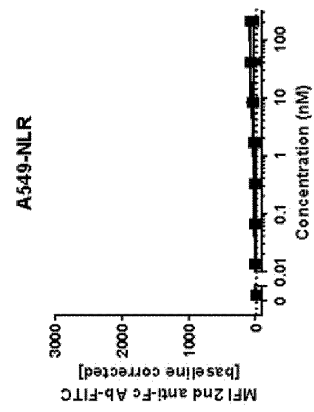
FIG.2E



Isotype Control

4+1 49B4/4B9 VhVI (P1AD4524)

FIG.2F



4+1 49B4/H212 xFab (P1AE6838)

3+1 49B4/H212 (P1AE8786)

2+1 49B4/H212 (P1AE6840)

Fig.3

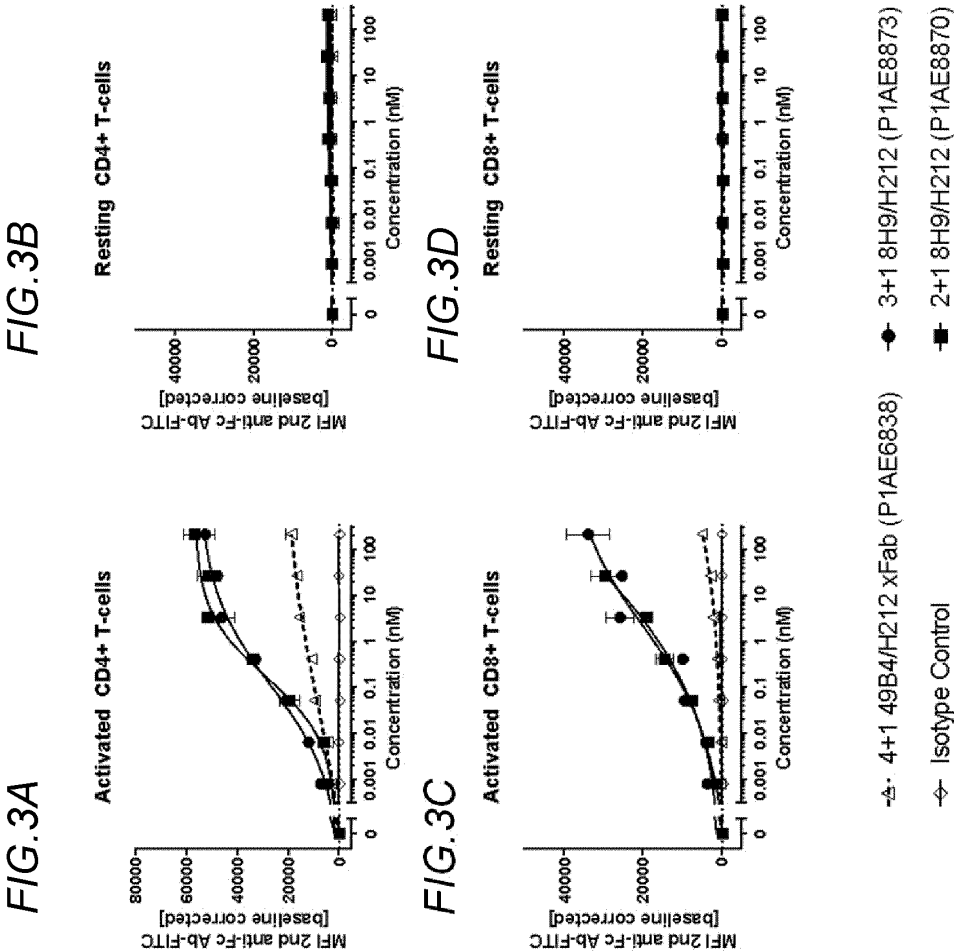
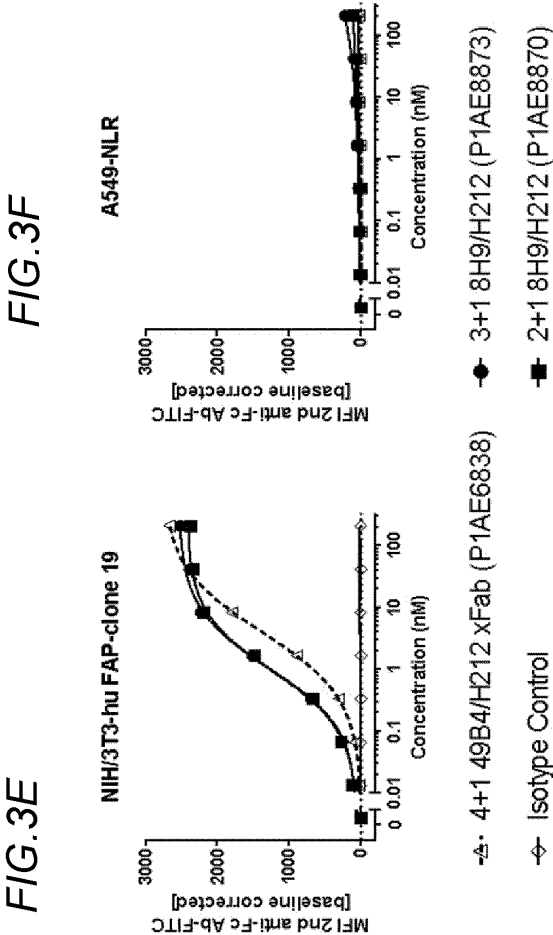


Fig.3



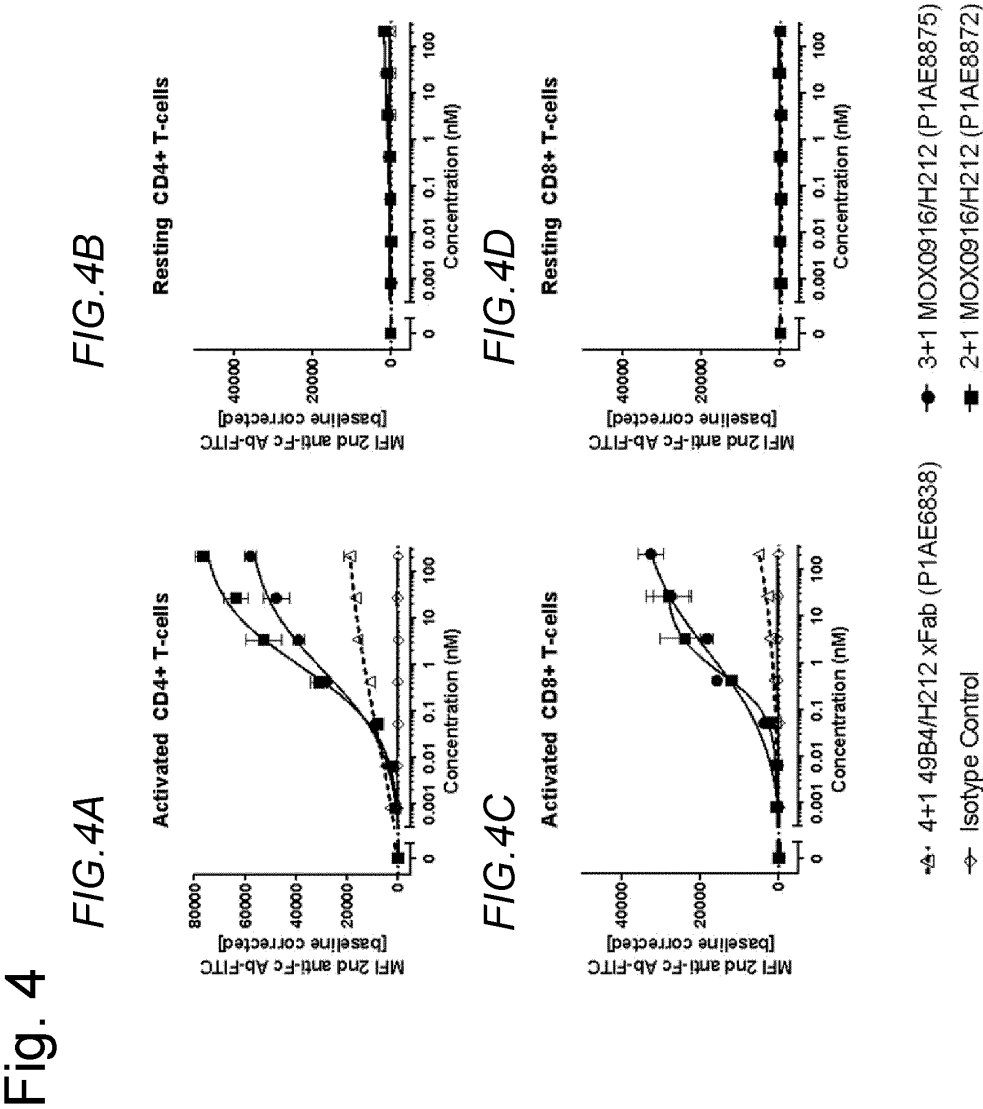


Fig. 4

FIG.4E

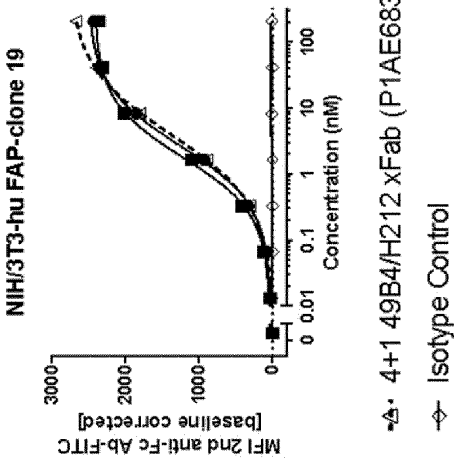


FIG.4F

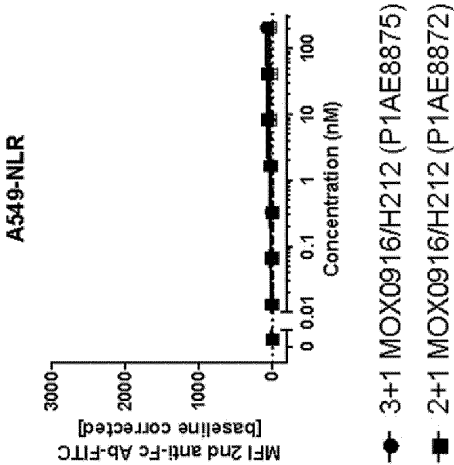


Fig.5

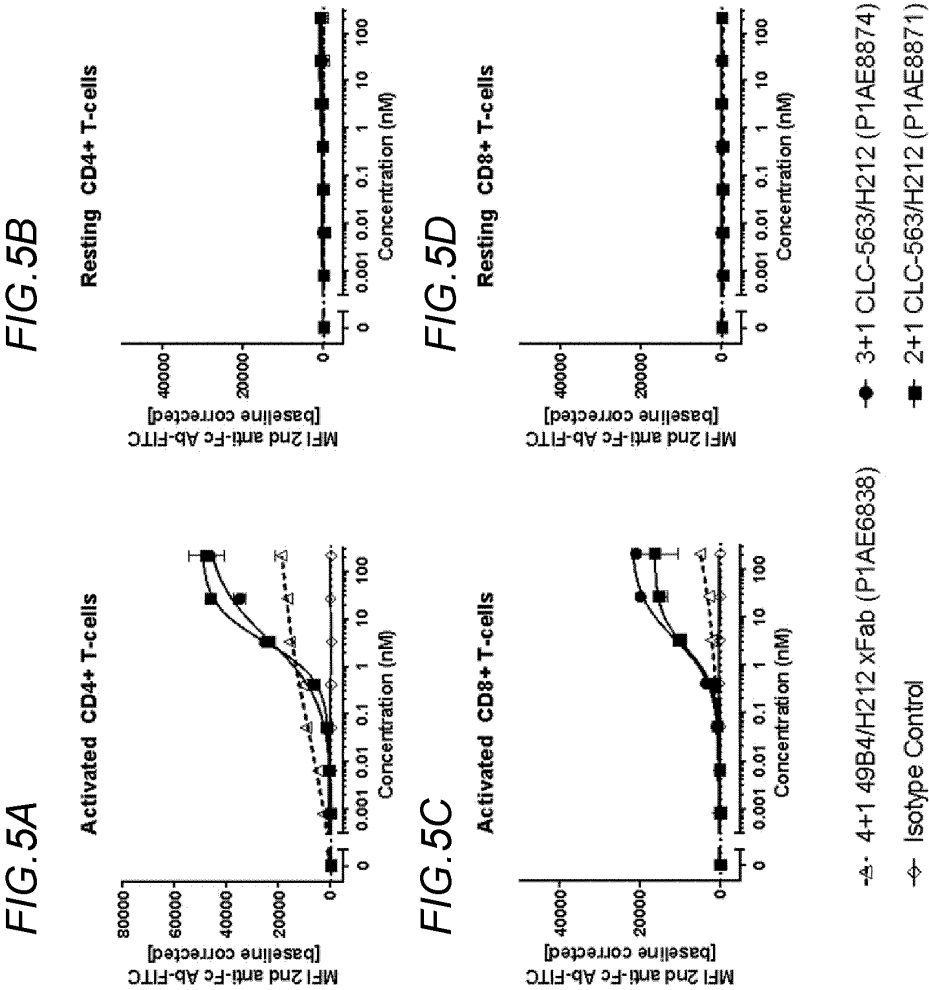


Fig.5

FIG.5E

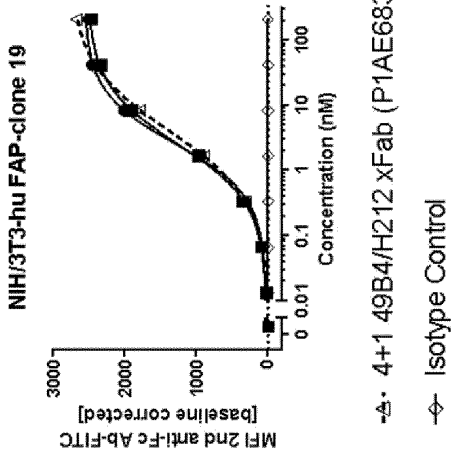


FIG.5F

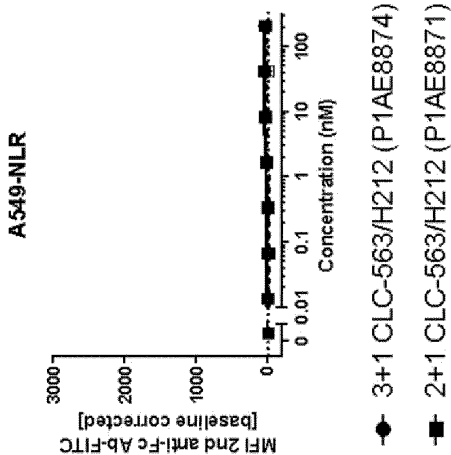


Fig.6

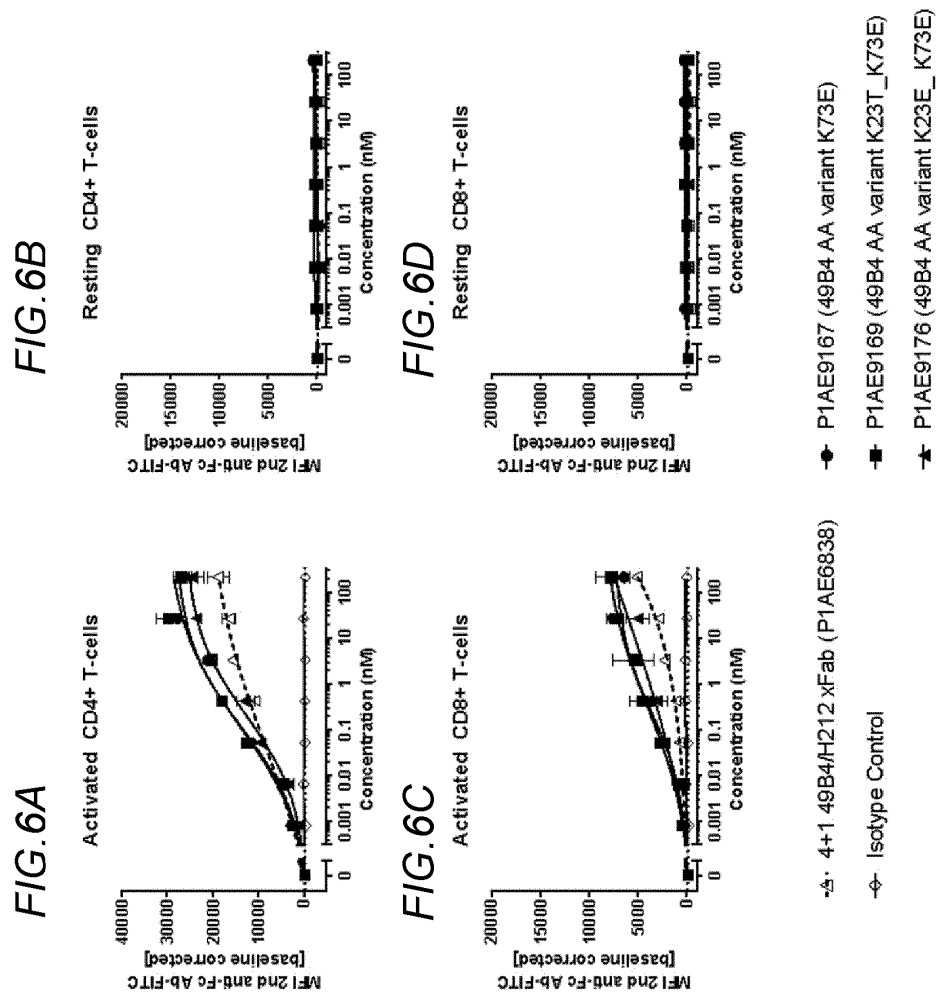


Fig.6

FIG.6E

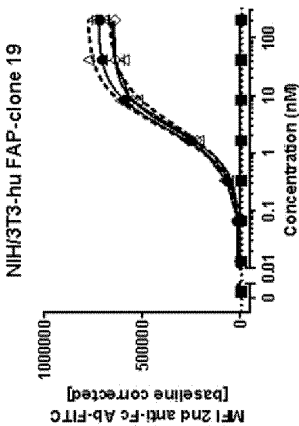
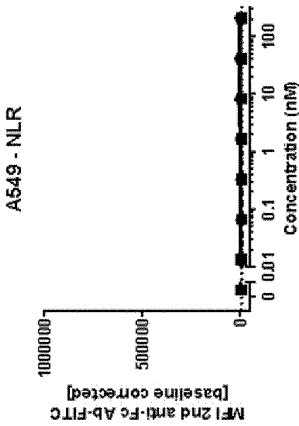


FIG.6F



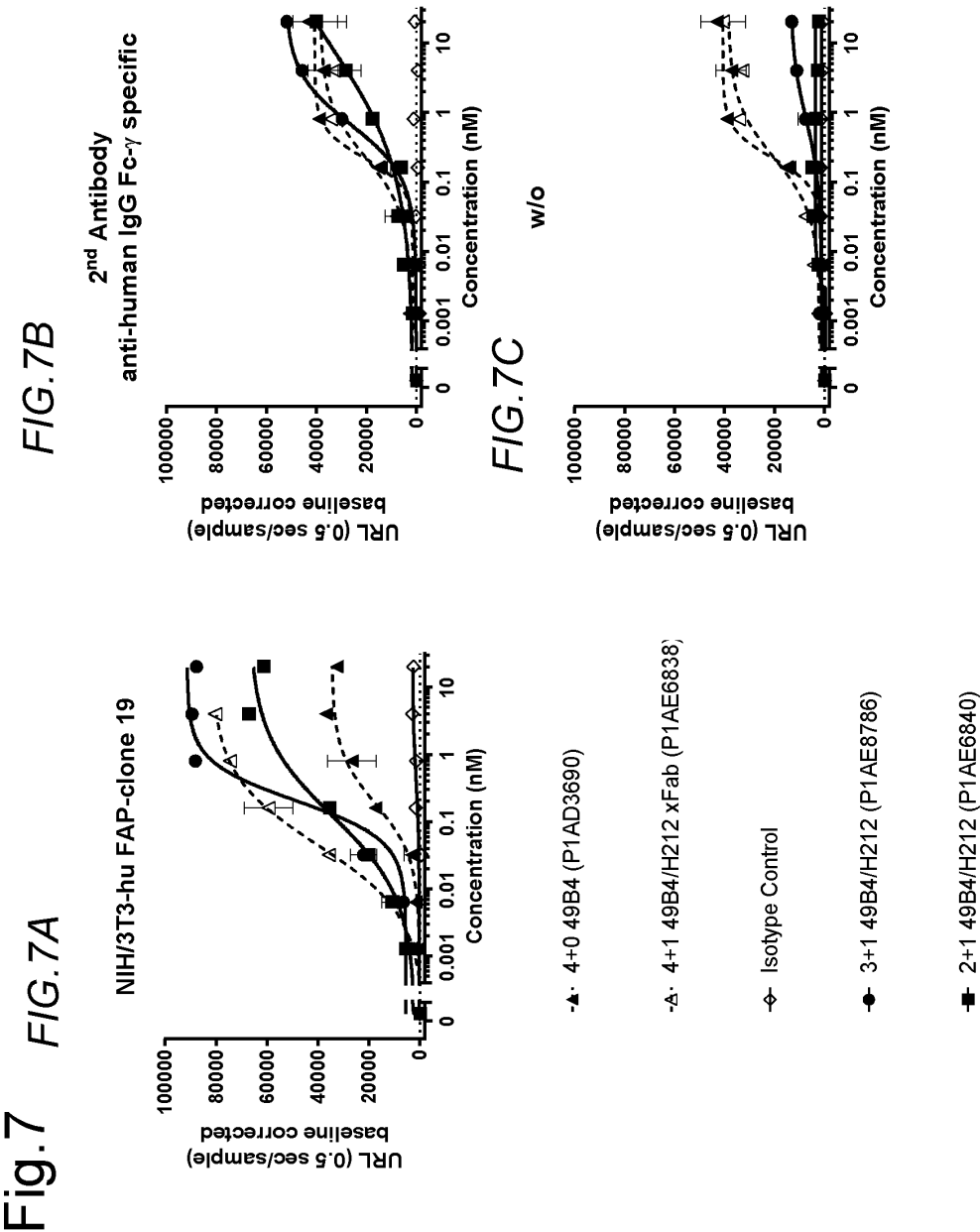
△ 4+1 49B4/H212 xFab (P1AE6838)

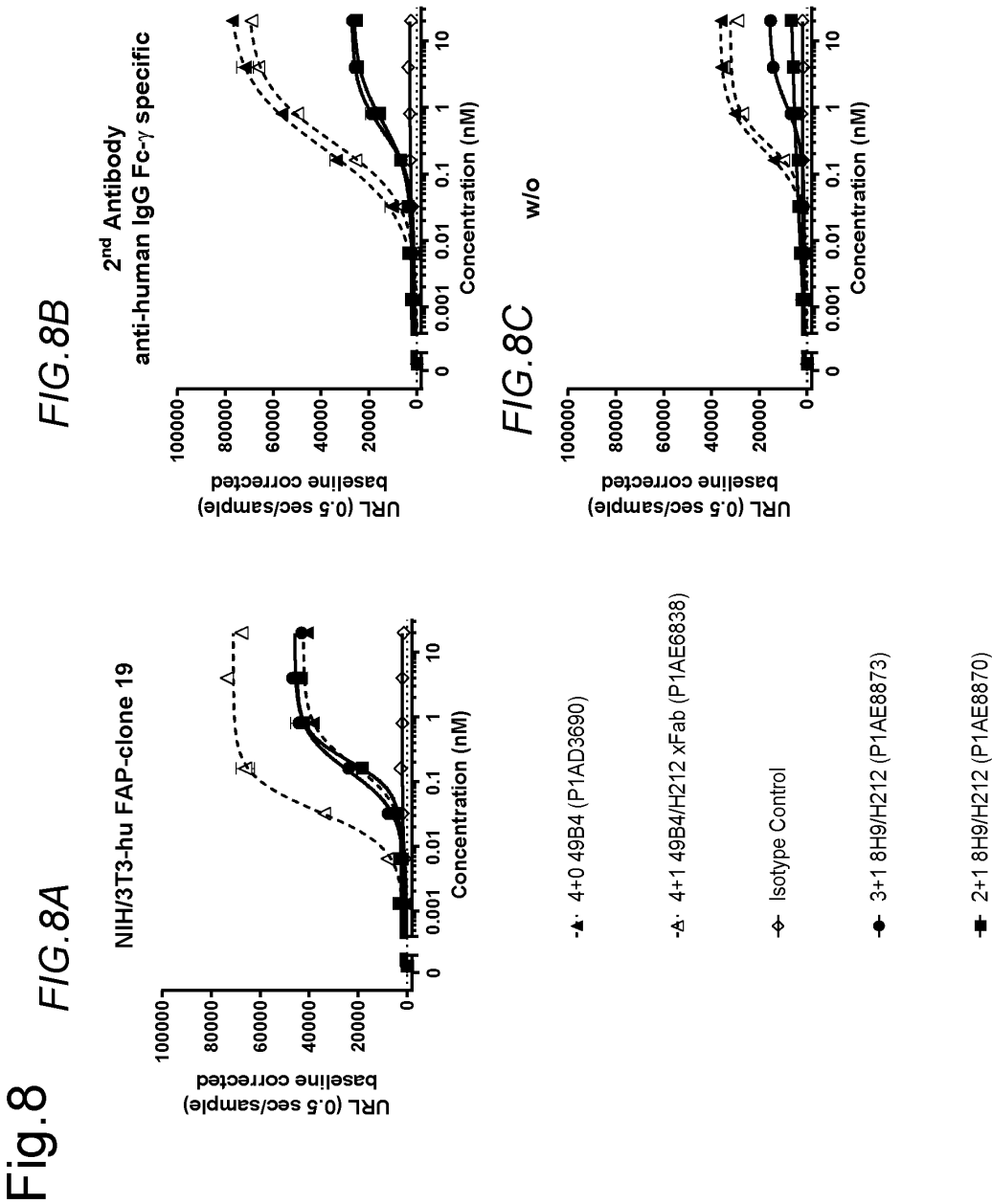
○ Isotype Control

● P1AE9167 (49B4 AA variant K73E)

■ P1AE9169 (49B4 AA variant K23T_K73E)

▲ P1AE9176 (49B4 AA variant K23E_K73E)





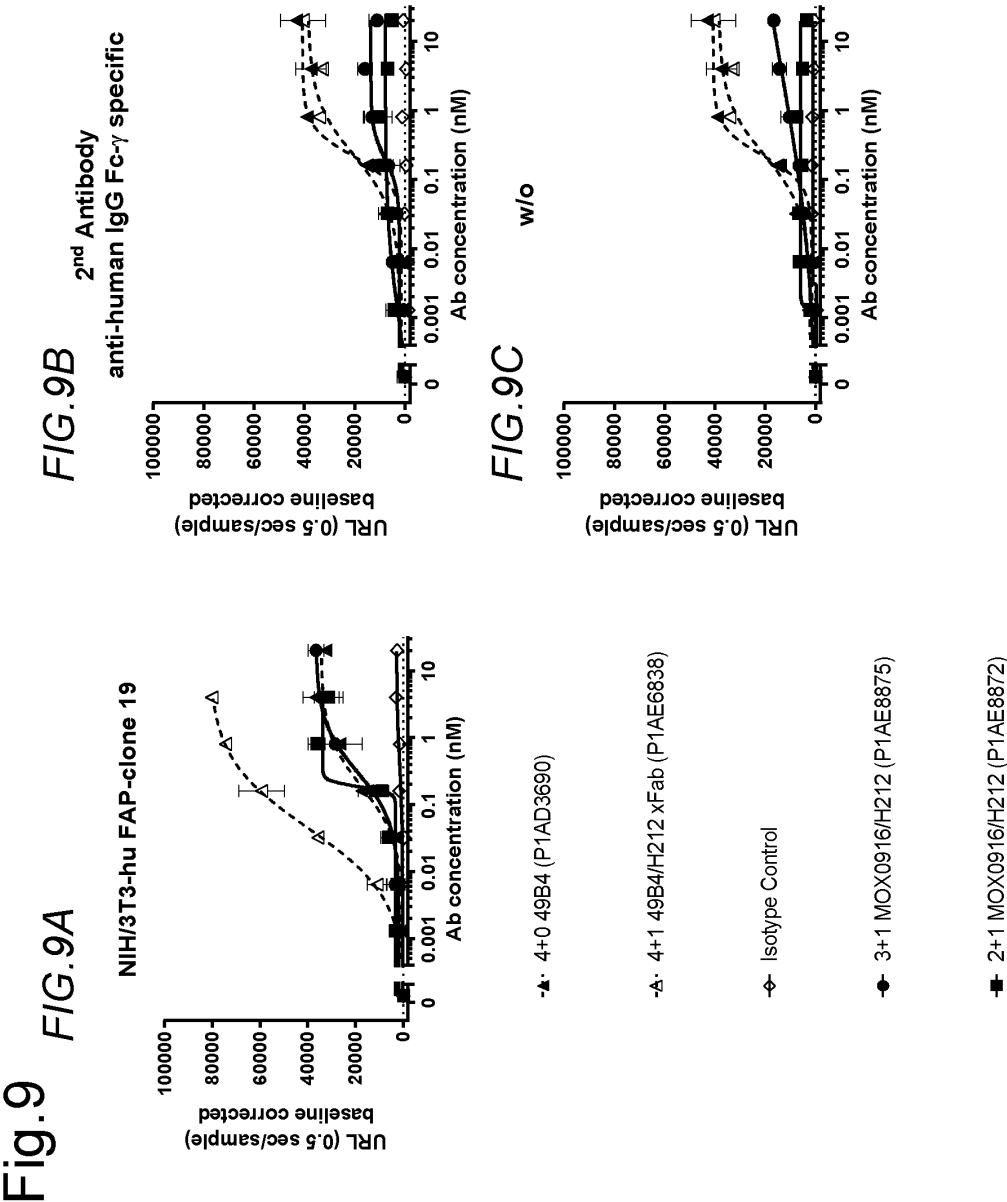
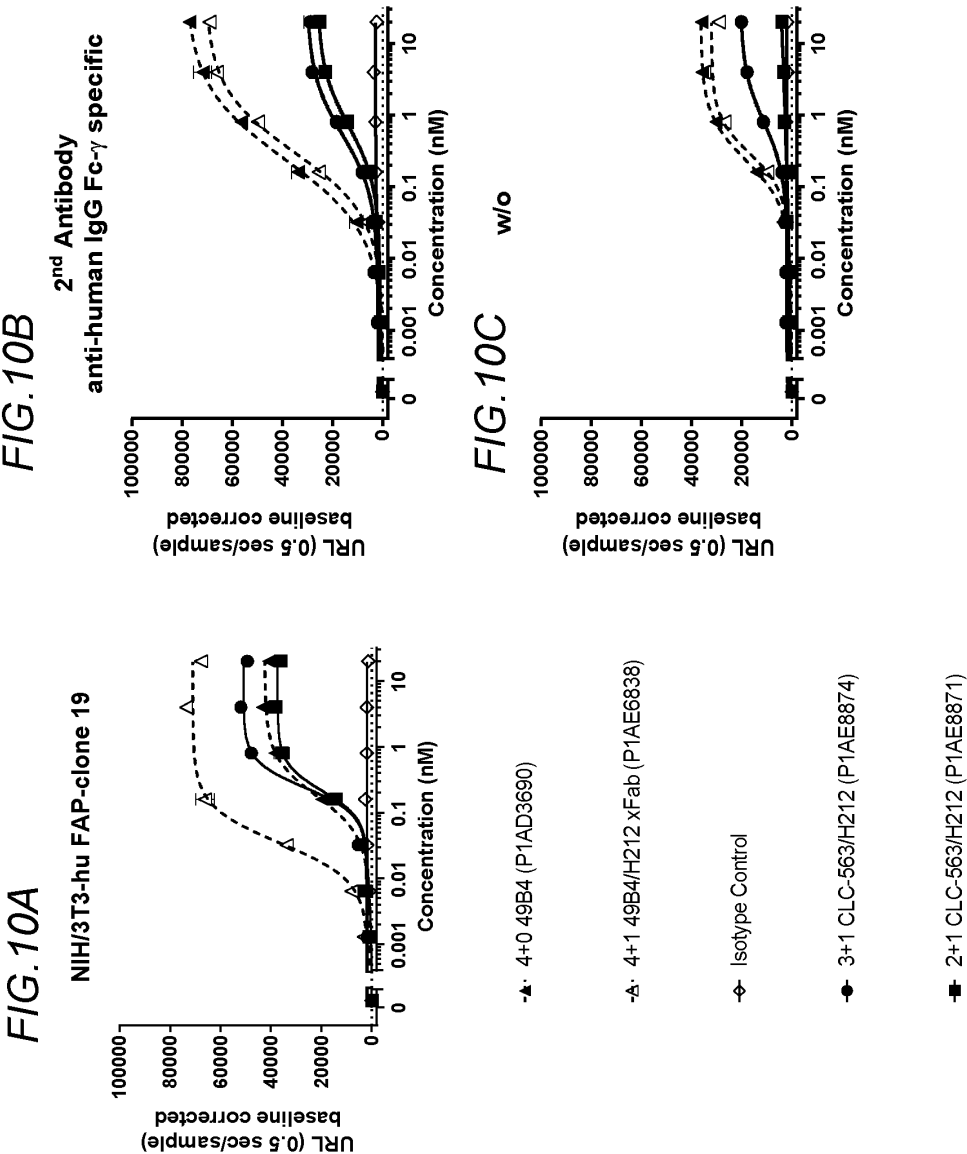


Fig.10



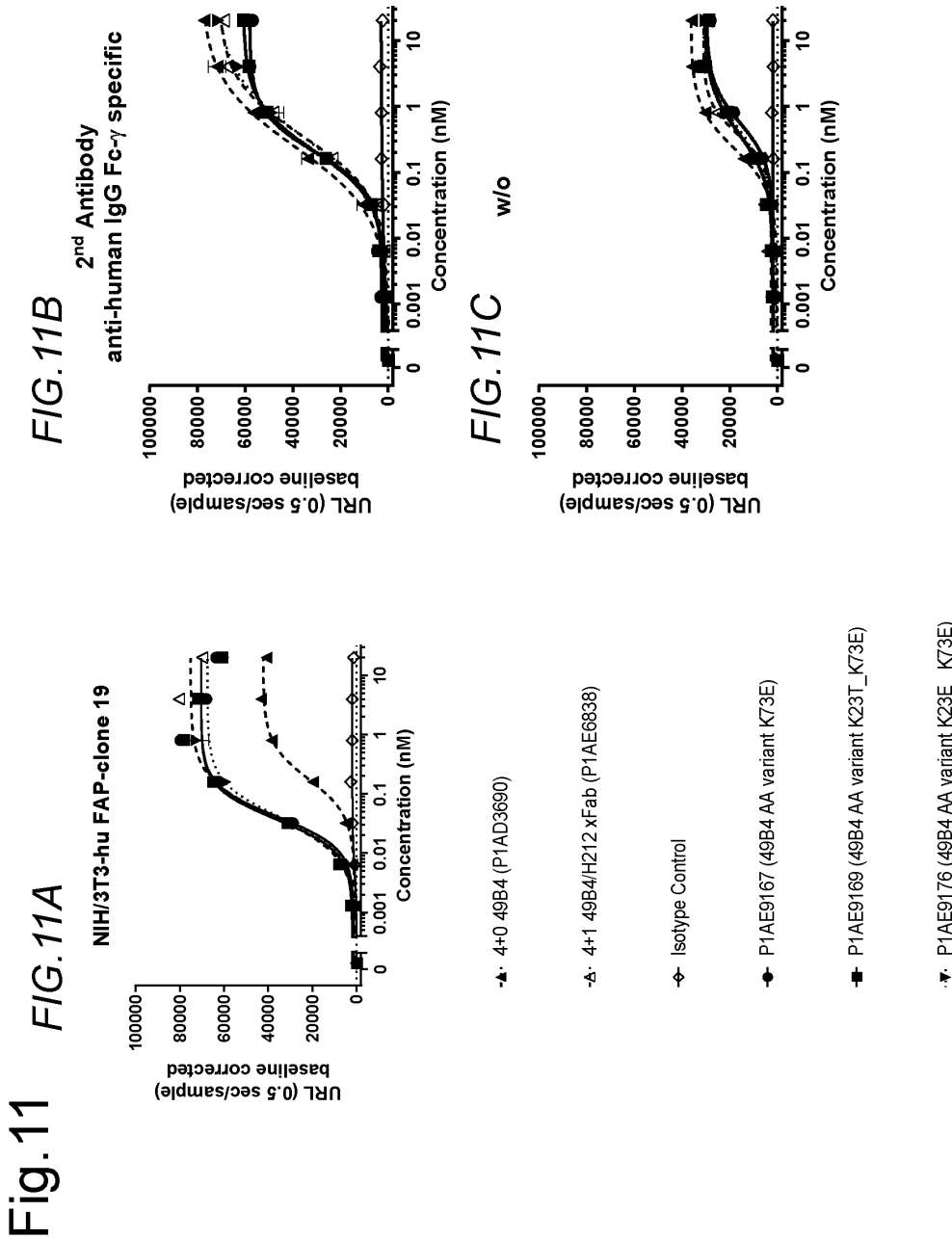


Fig.12

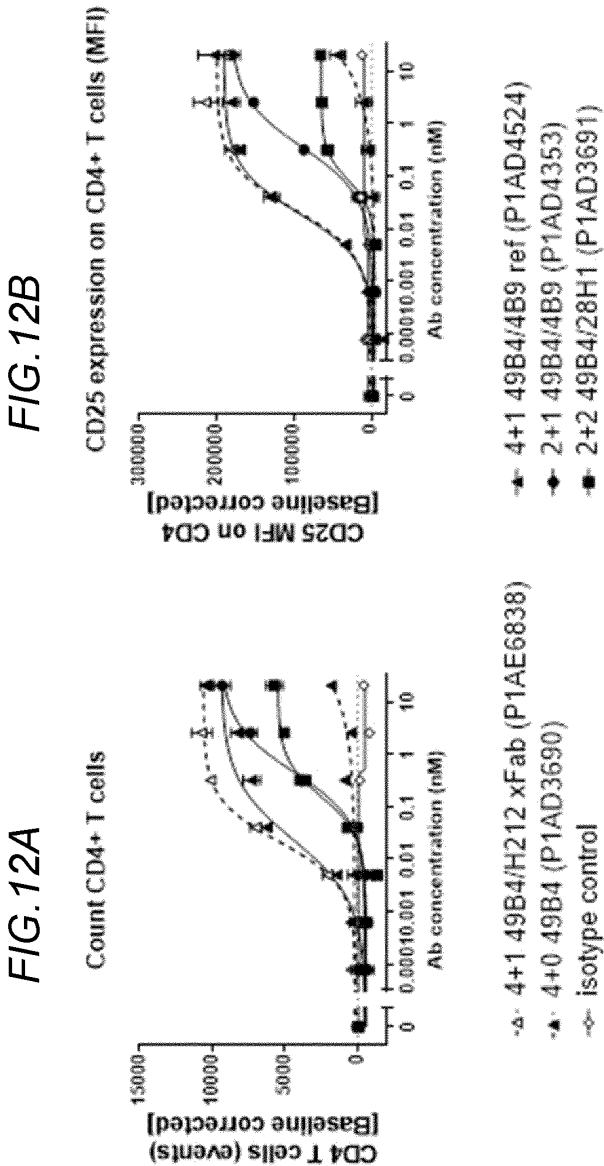


Fig.13

FIG.13A

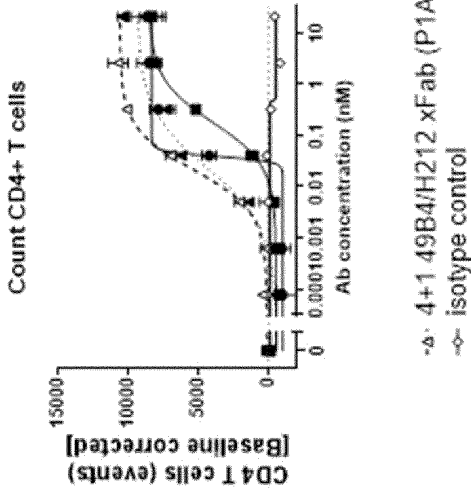


FIG.13B

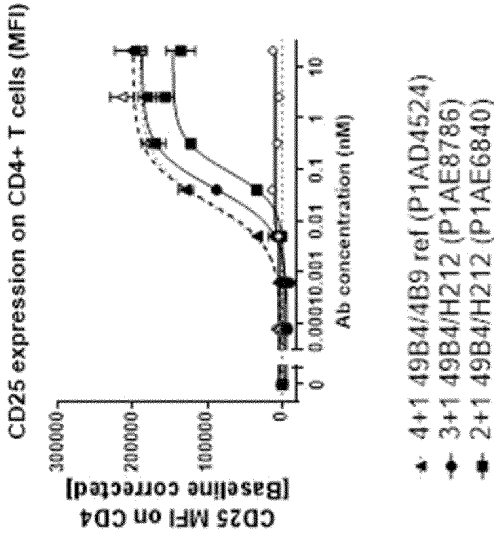


Fig.14

FIG.14A

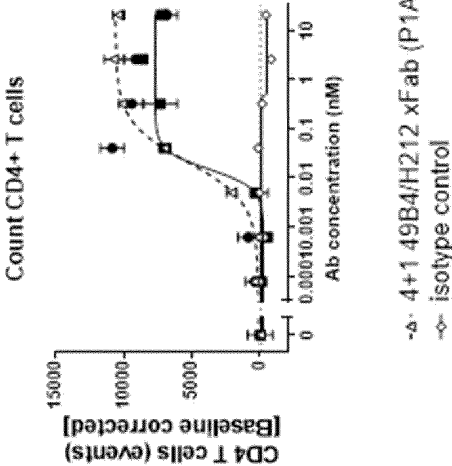


FIG.14B

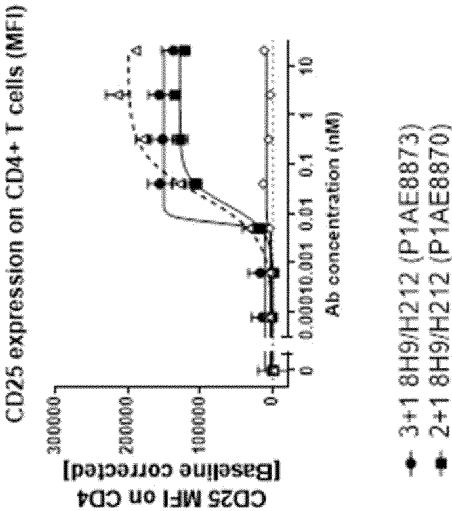


Fig.15

FIG.15A

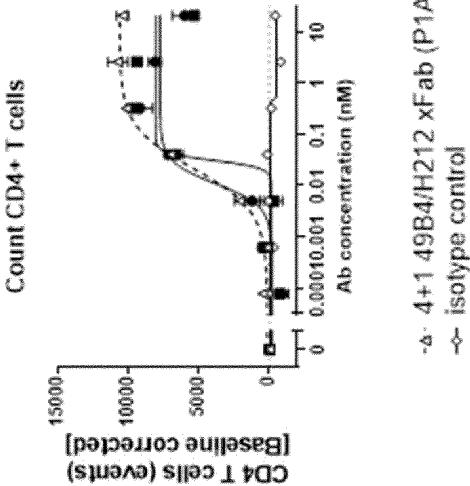


FIG.15B

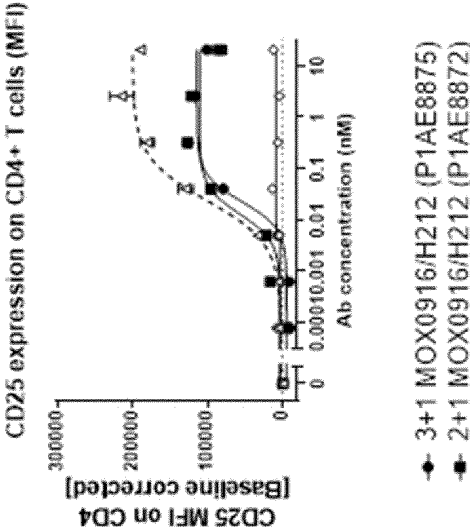


Fig. 16

FIG. 16A

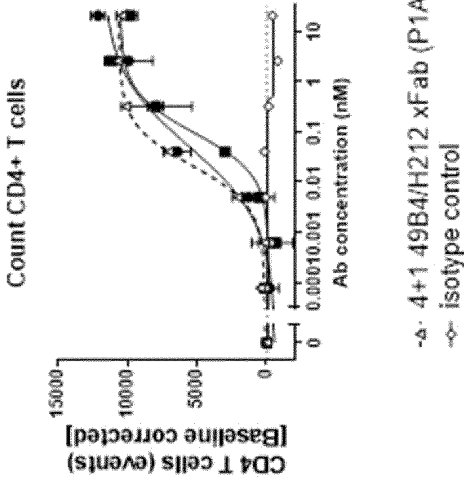


FIG. 16B

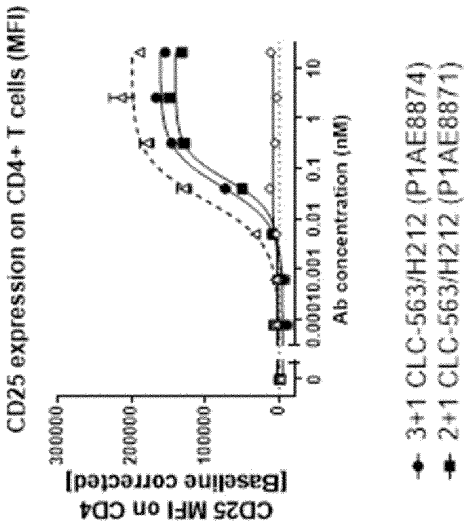
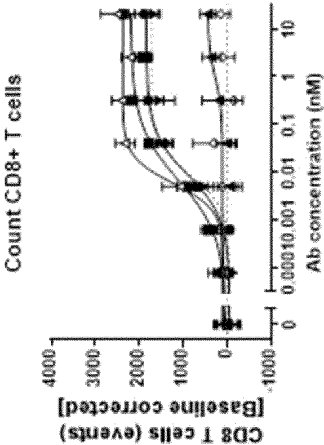


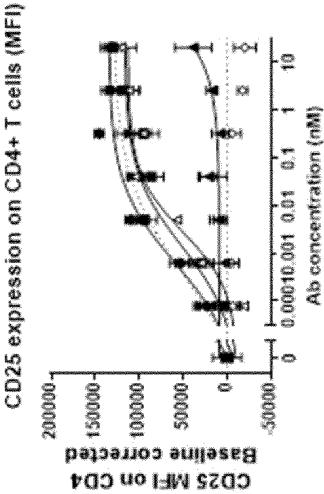
Fig.17

FIG.17A



- △ 4+1 49B4/H212 xFab (P1AE6838)
- ▲ 4+0 49B4 (P1AD3690)
- isotype control

FIG.17B



- P1AE9167 (49B4 AA variant K73E)
- P1AE9169 (49B4 AA variant K23T_K73E)
- ▼ P1AE9176 (49B4 AA variant K23E_K73E)

Fig.18

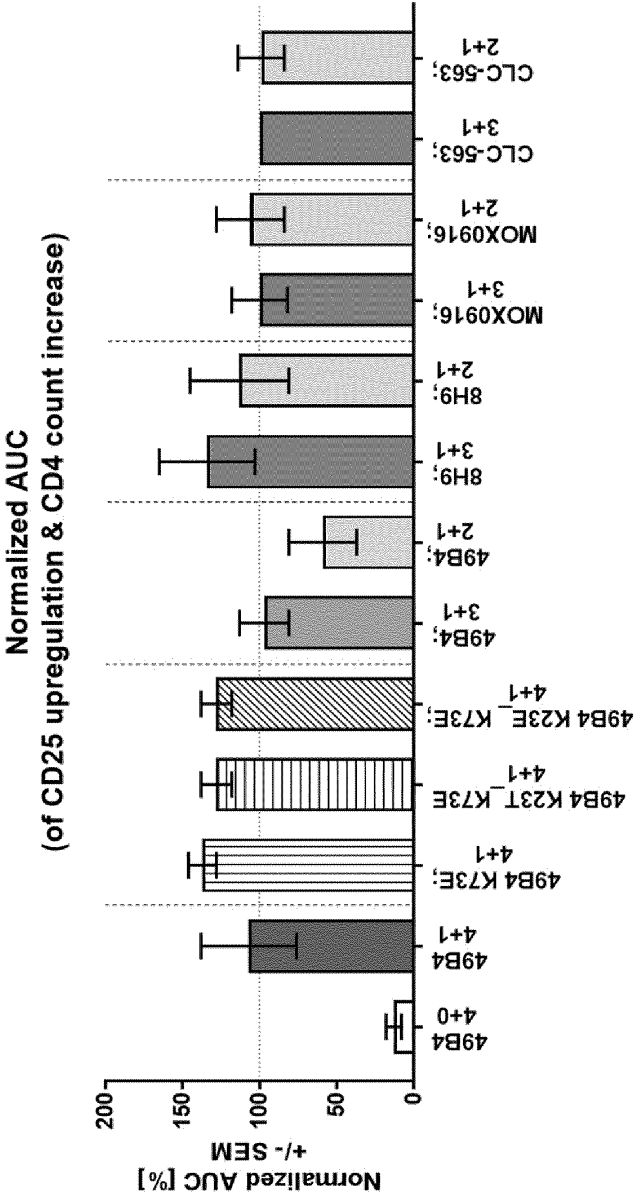


Fig.19

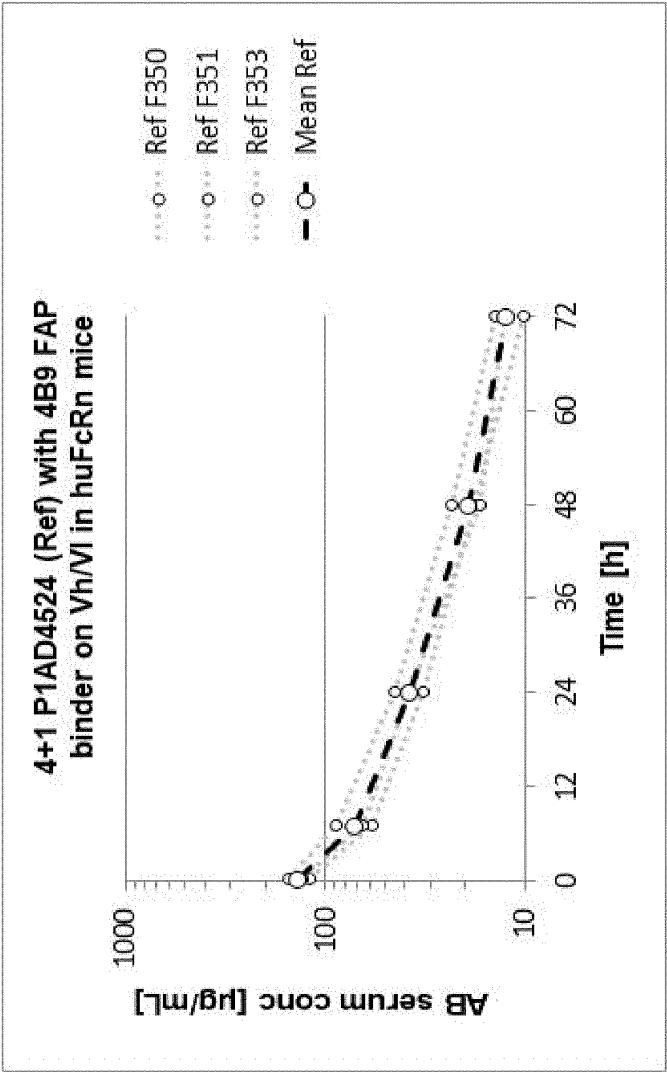


Fig. 20

FIG.20A

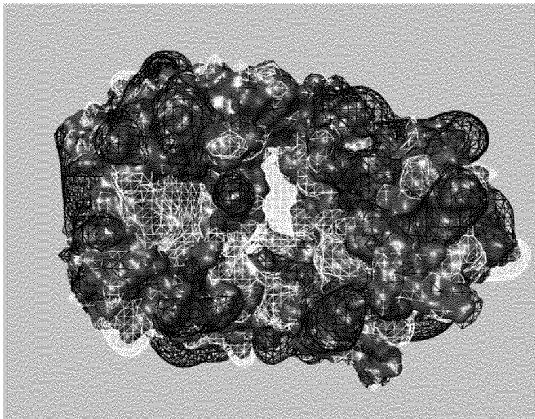


FIG.20B

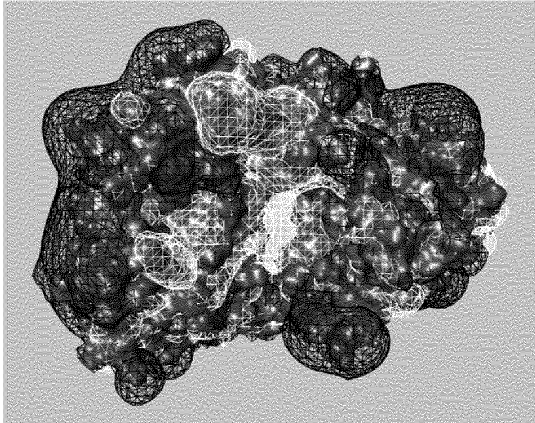
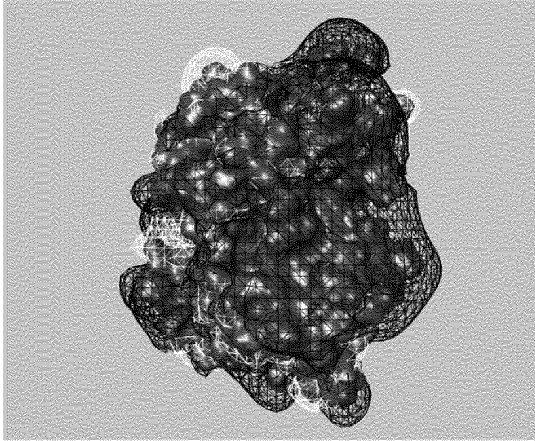


FIG.20C



4B9 FAP clone

Fig. 20

FIG.20D

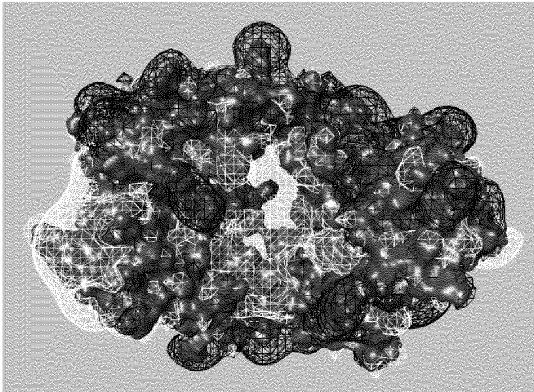


FIG.20E

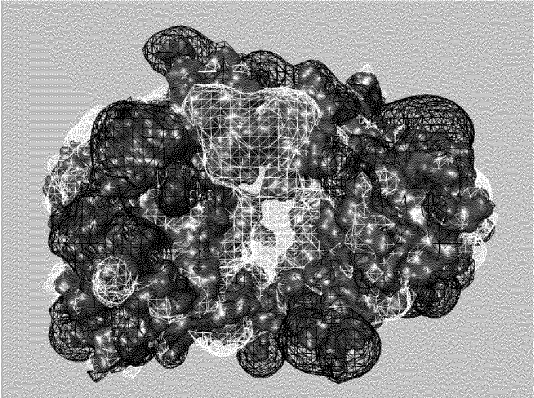
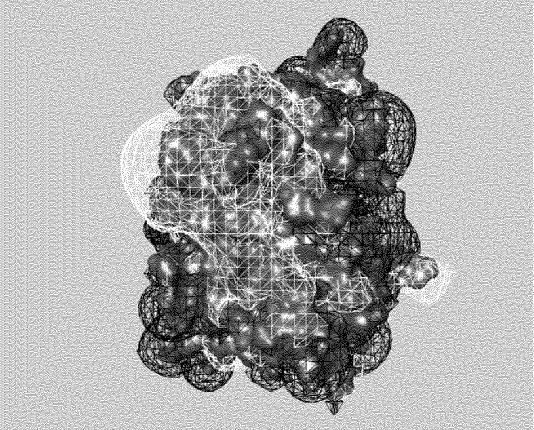


FIG.20F



1G1a (h212) FAP clone

Fig. 21

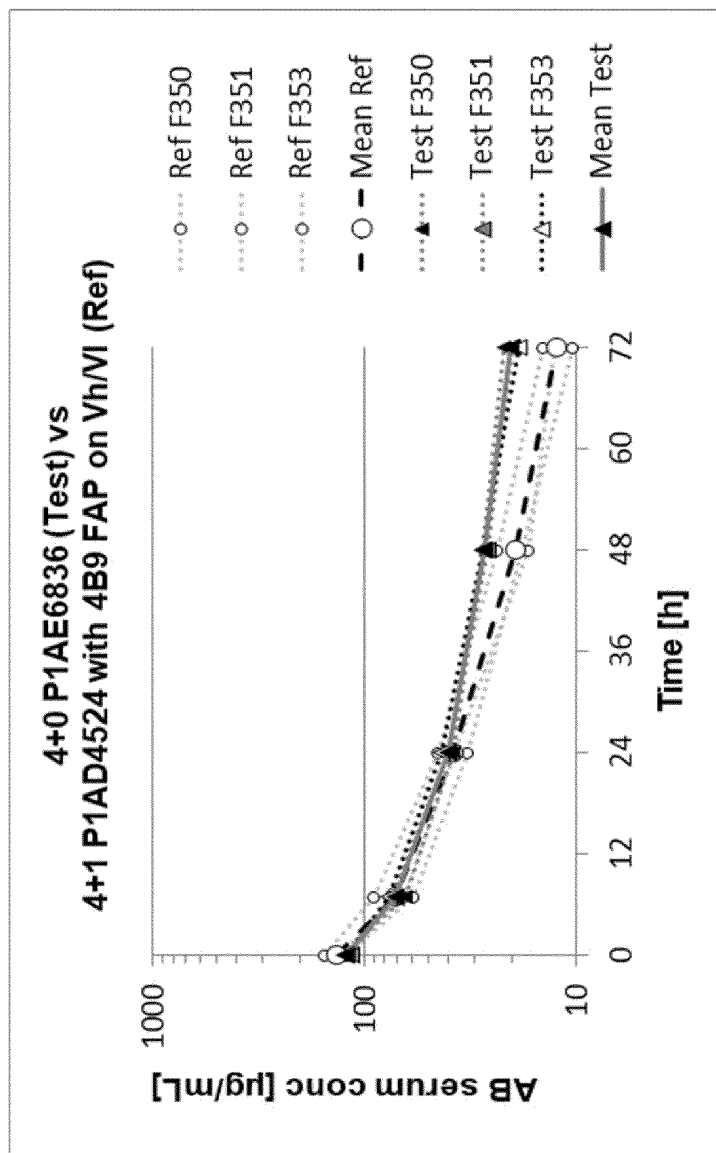


Fig. 22

FIG.22A

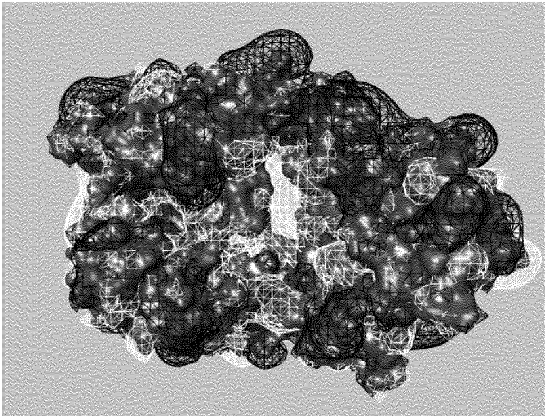


FIG.22B

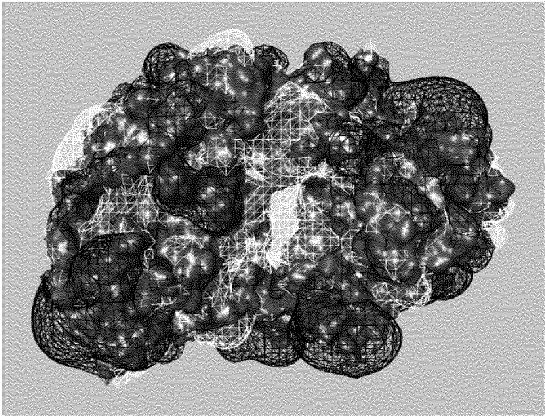
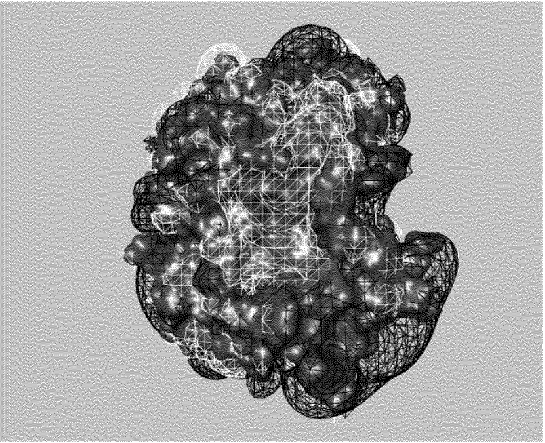


FIG.22C



49B4 OX40 clone

Fig. 23

FIG.23A

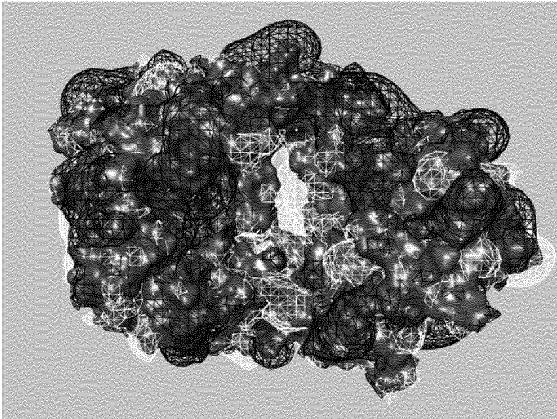


FIG.23B

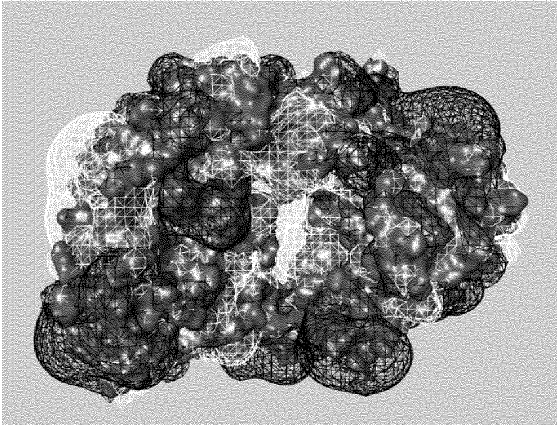
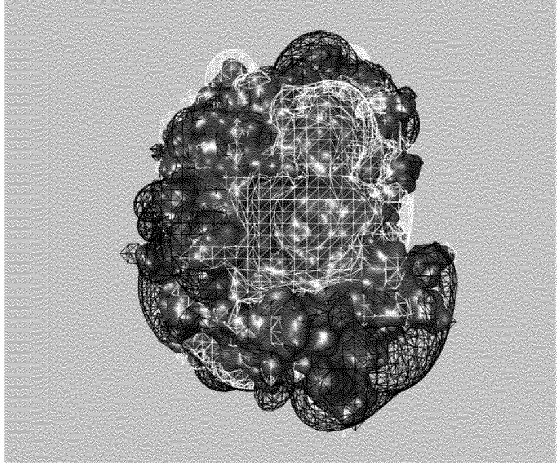


FIG.23C



8H9 OX40 clone

Fig. 24

FIG.24A

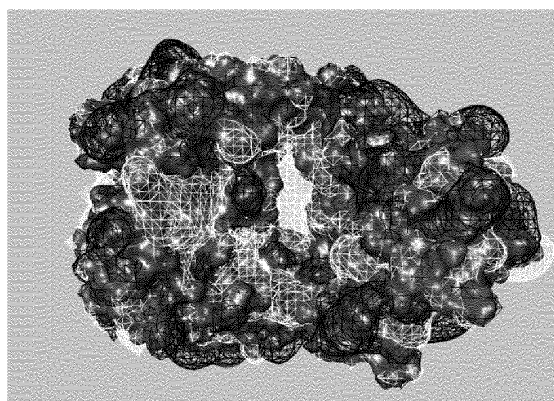


FIG.24B

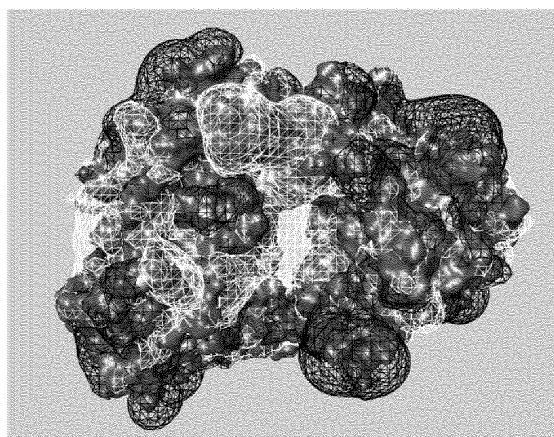
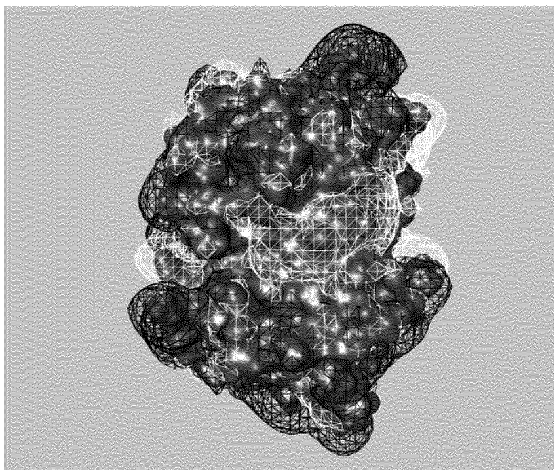


FIG.24C



CLC563 OX40 clone

Fig. 25

FIG.25A

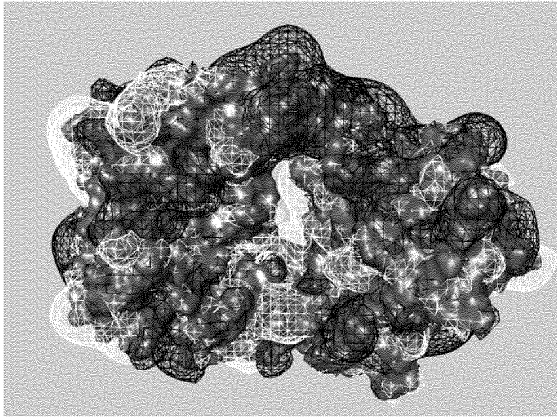


FIG.25B

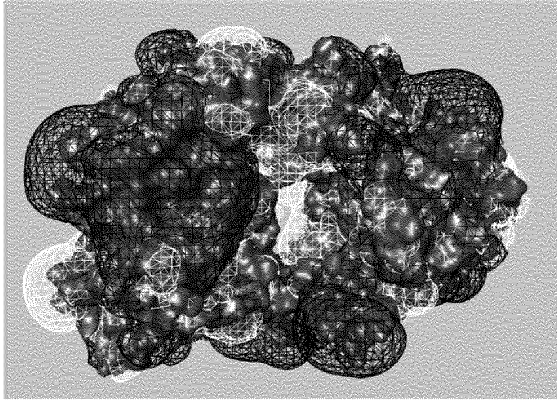
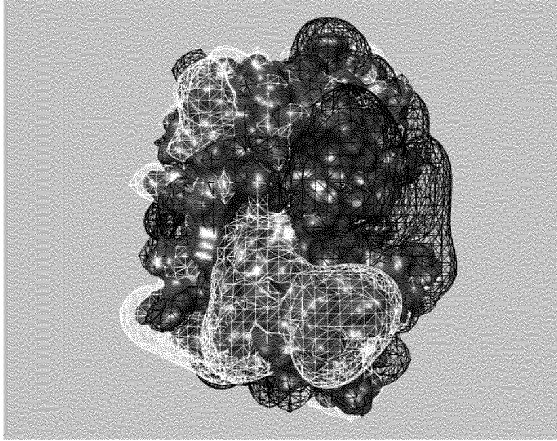


FIG.25C



MOXR0916 OX40 clone

Fig. 26

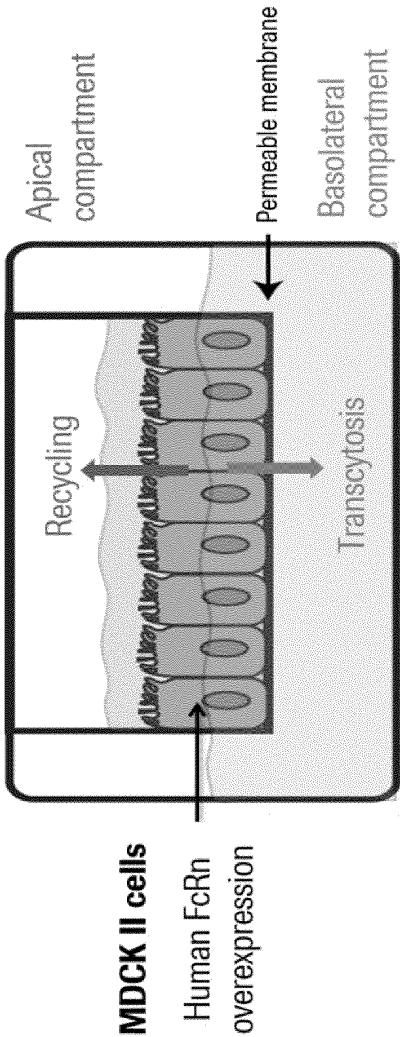


Fig. 27

FIG.27A

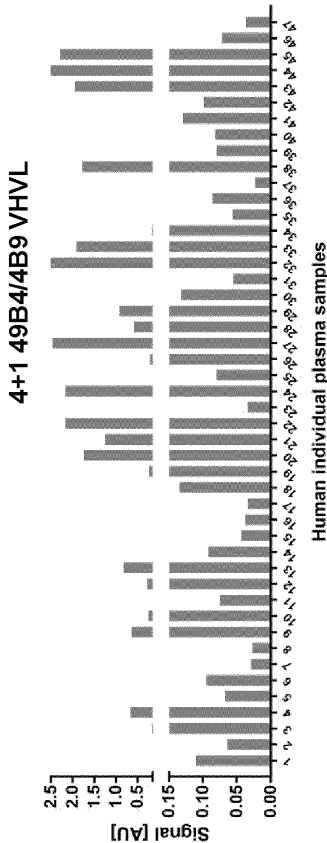


FIG.27B

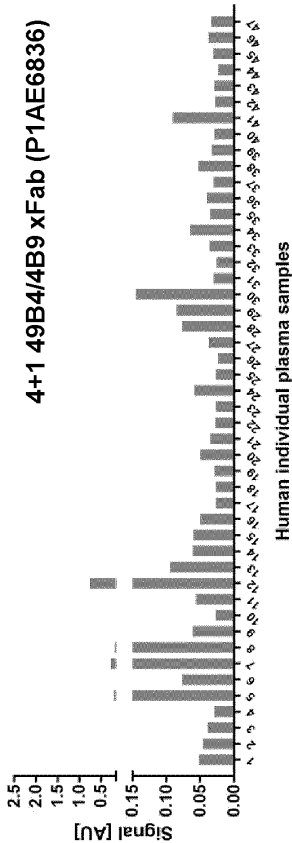
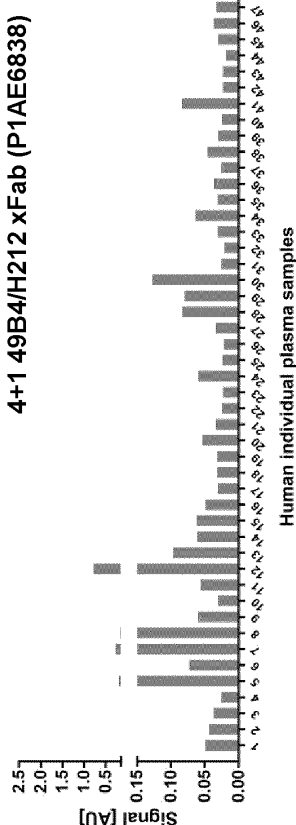


FIG.27C



FAPOX40 4+1 constructs

Fig. 28

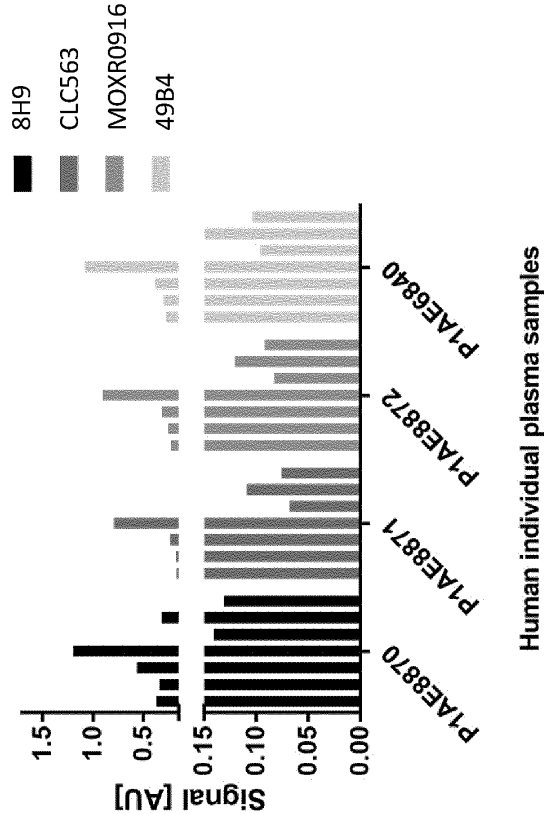


Fig. 29

FIG.29A

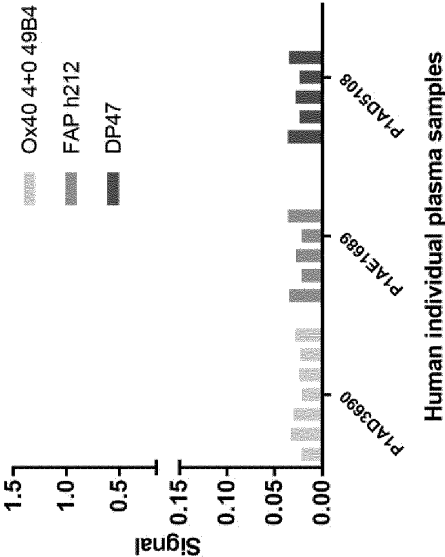


FIG.29B

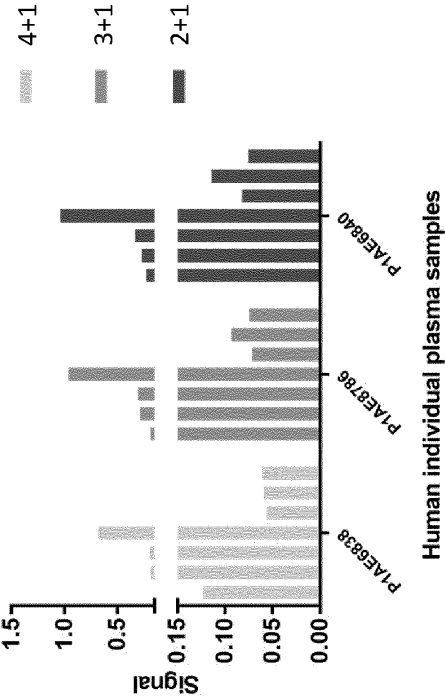


Fig. 30

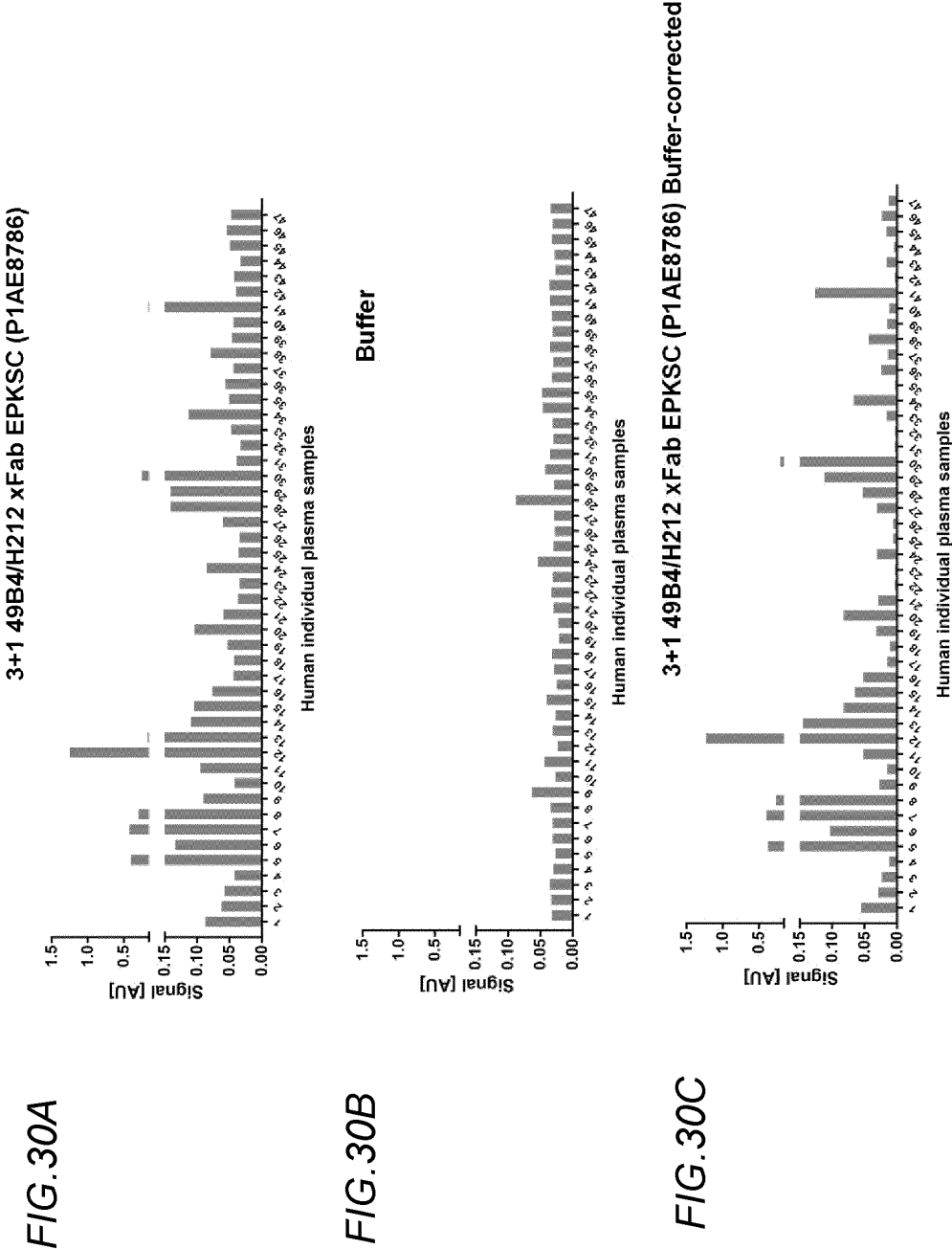


Fig. 31

FIG.31A

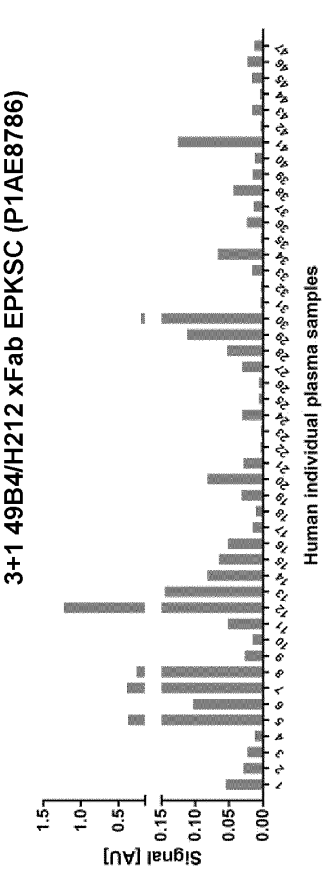


FIG.31B

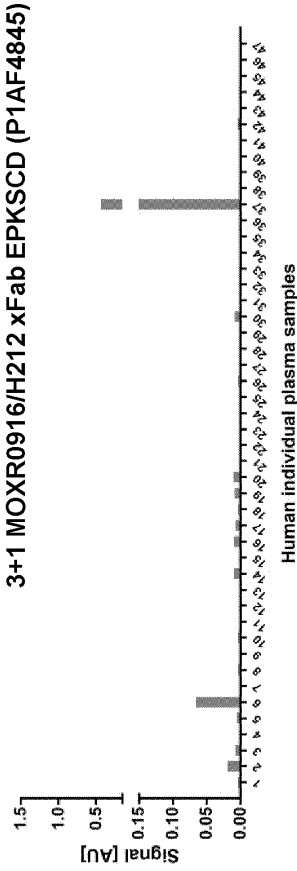


FIG.31C

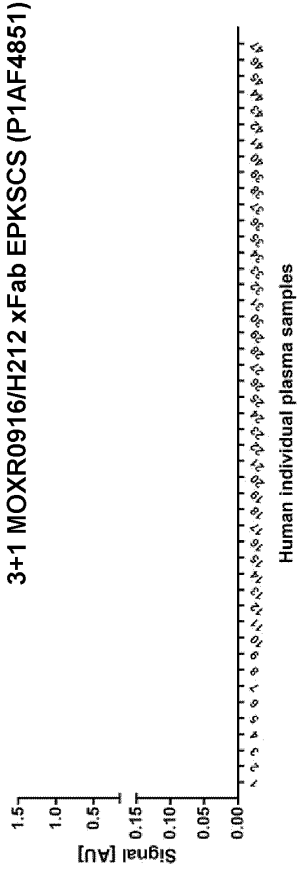


Fig. 32
FIG.32A

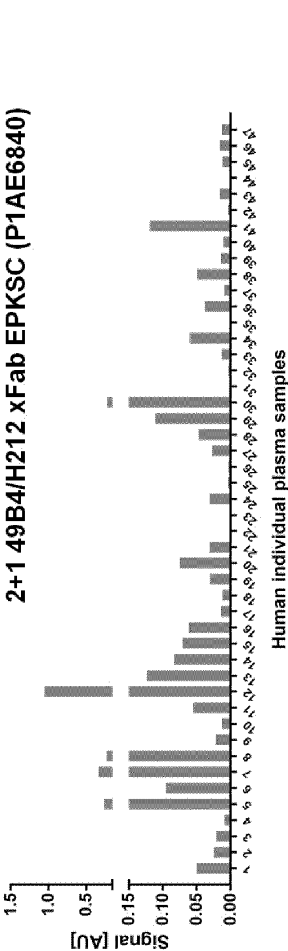


FIG.32B

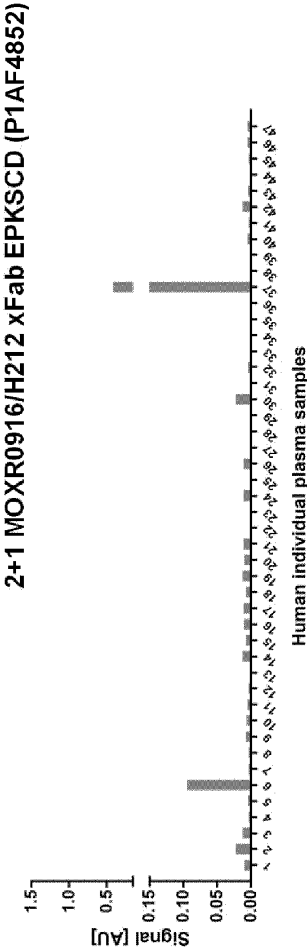


FIG.32C

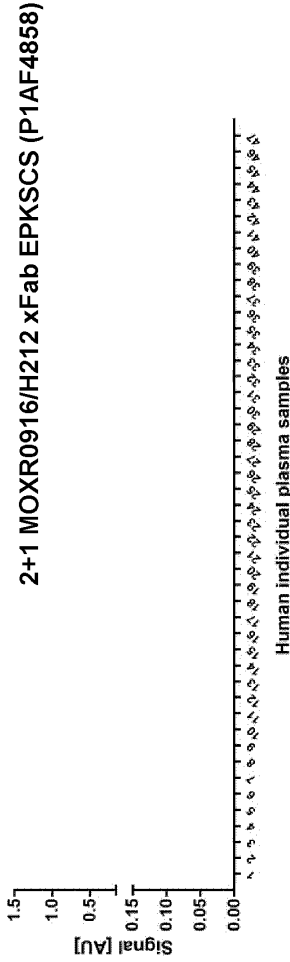


Fig. 33

FIG. 33A

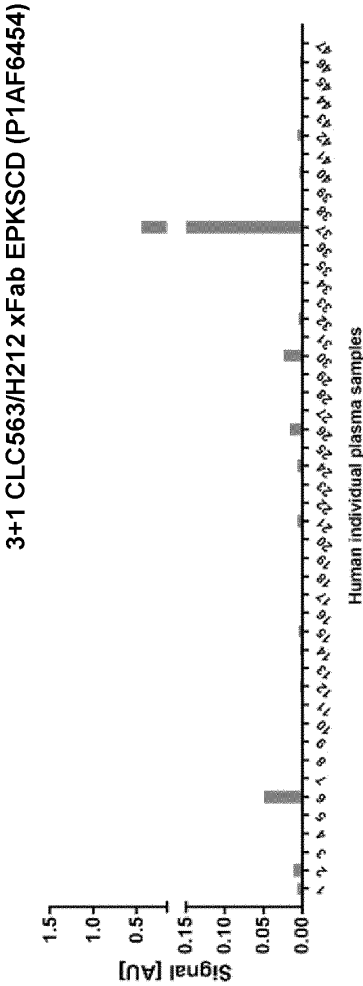


FIG. 33B

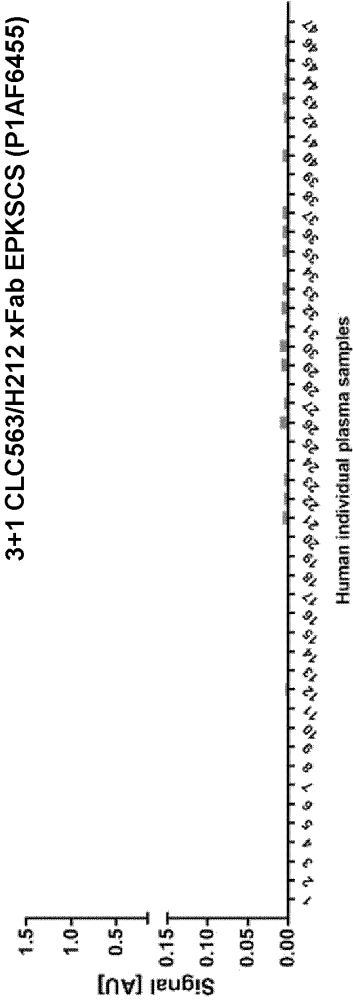


Fig. 33

FIG. 33C

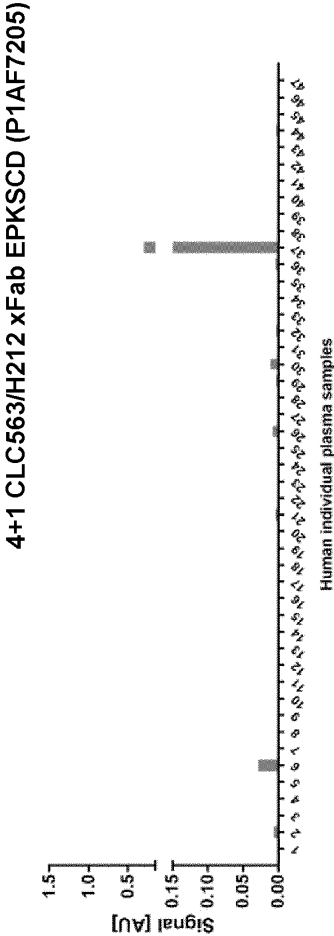


FIG. 33D

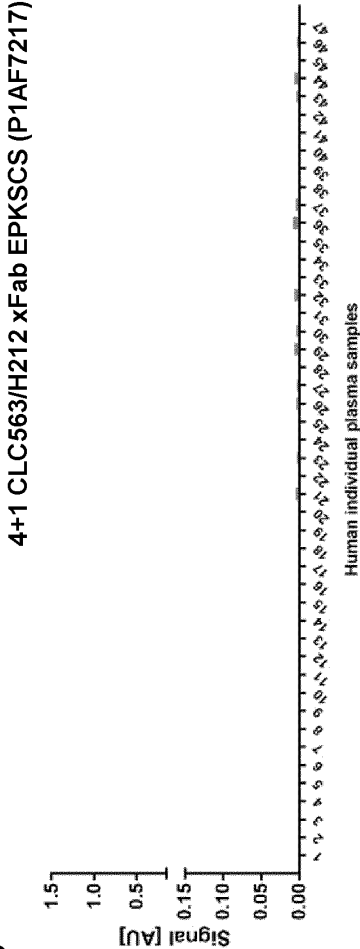


Fig. 33

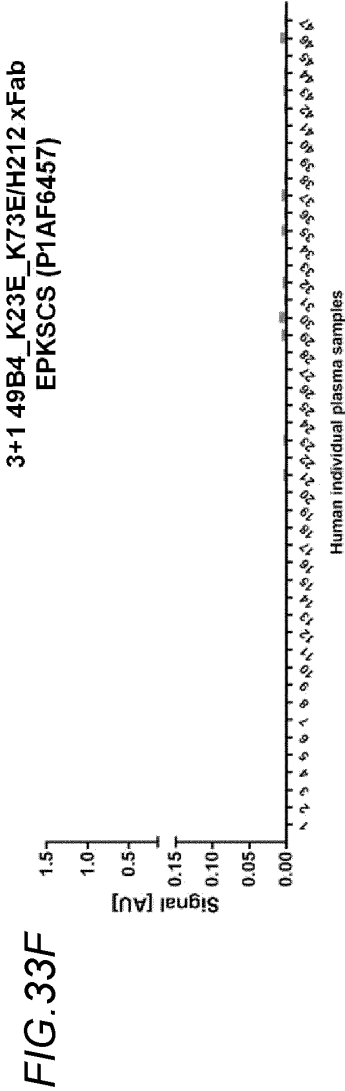
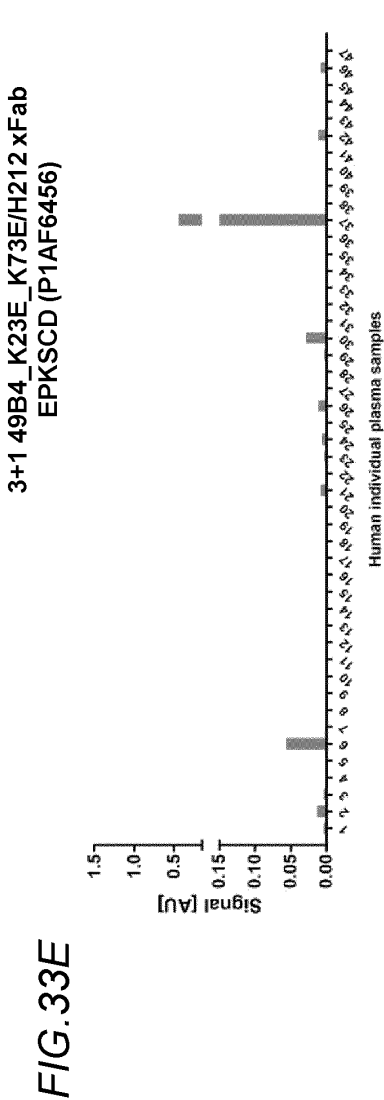


Fig. 34

FIG.34A

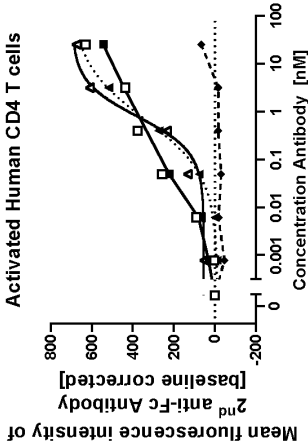
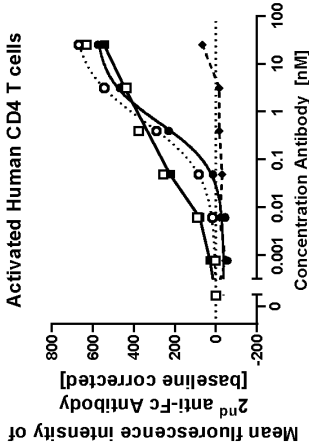


FIG.34C



- 4 49B4 (P1AD3690)
- 4+1 49B4/4B9 VhVI (P1AD4524)
- ◆ Isotype Control

FIG.34B

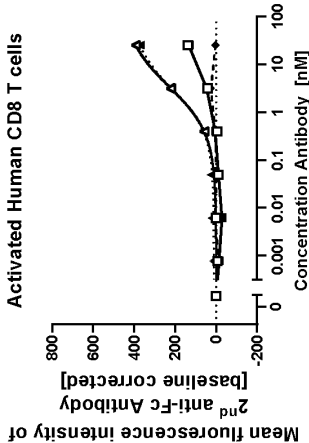
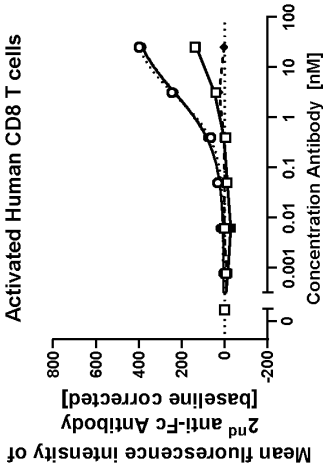


FIG.34D



- ▲ 3+1 CLC563/H212-D (P1AF6454)
- ▲ 3+1 CLC563/H212-S (P1AF6455)
- 4+1 CLC563/H212-D (P1AF7205)
- 4+1 CLC563/H212-S (P1AF7217)

Fig. 34

FIG. 34E

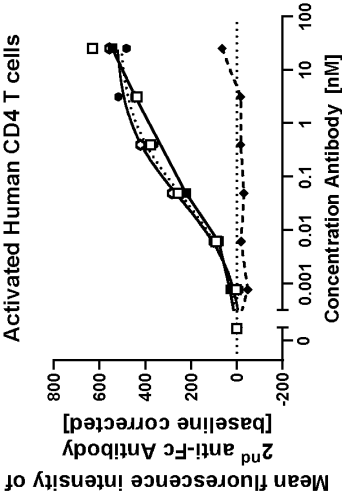


FIG. 34F

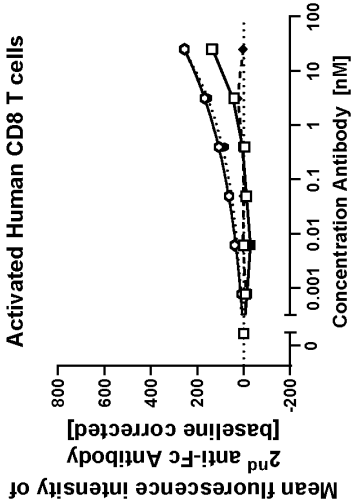


Fig. 35

FIG.35A

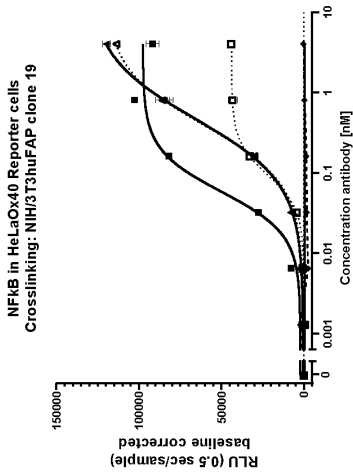
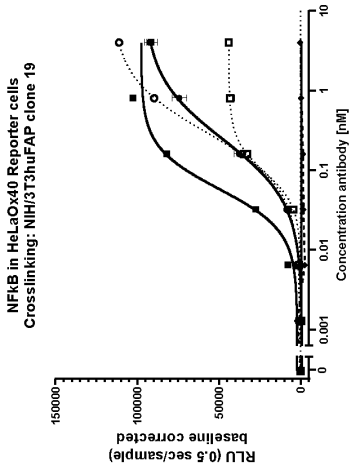


FIG.35C



- 4 49B4 (P1AD3690)
- 4+1 49B4/4B9 VhVI (P1AD4524)
- ◆ Isotype Control

FIG.35B

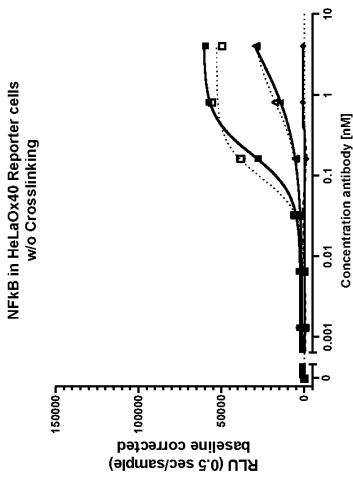
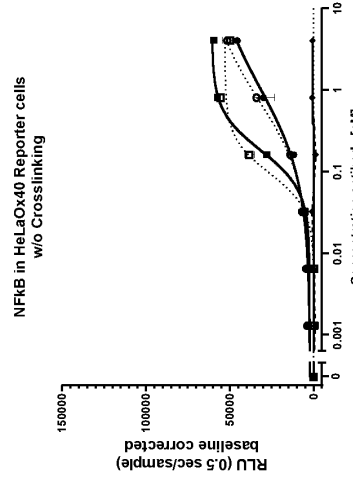


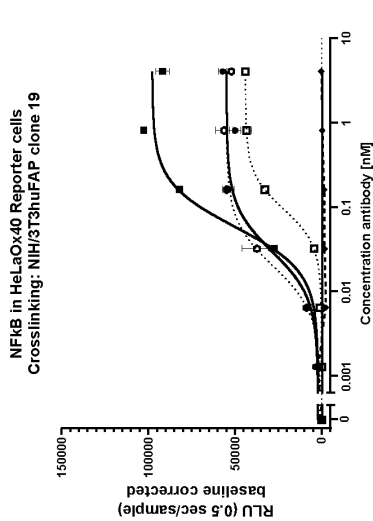
FIG.35D



- 3+1 CLC563/H212-D (P1AF6454)
- ▲ 3+1 CLC563/H212-S (P1AF6455)
- 4+1 CLC563/H212-D (P1AF7205)
- 4+1 CLC563/H212-S (P1AF7217)

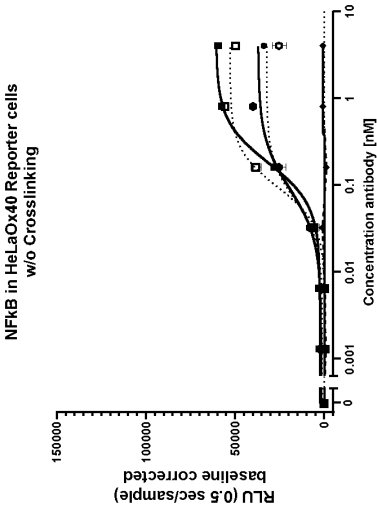
Fig. 35

FIG. 35E



- 4 49B4 (P1AD3690)
- 4+1 49B4/4B9 VhVl (P1AD4524)
- ◆ Isotype Control
- 4+1 49B4 K26E_K73E/H212 - S (P1AF6457)

FIG. 35F



- 4+1 49B4 K26E_K73E/H212 - D (P1AF6456)
- 4+1 49B4 K26E_K73E/H212 - S (P1AF6457)
- ◆ Isotype Control
- 4 49B4 (P1AD3690)

Fig. 36

FIG.36A

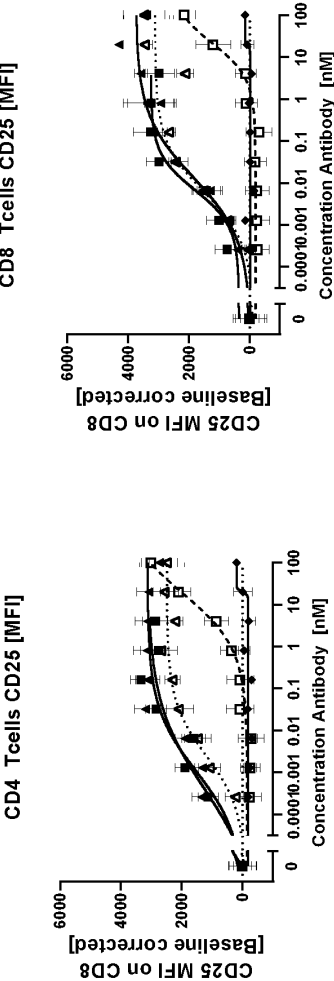
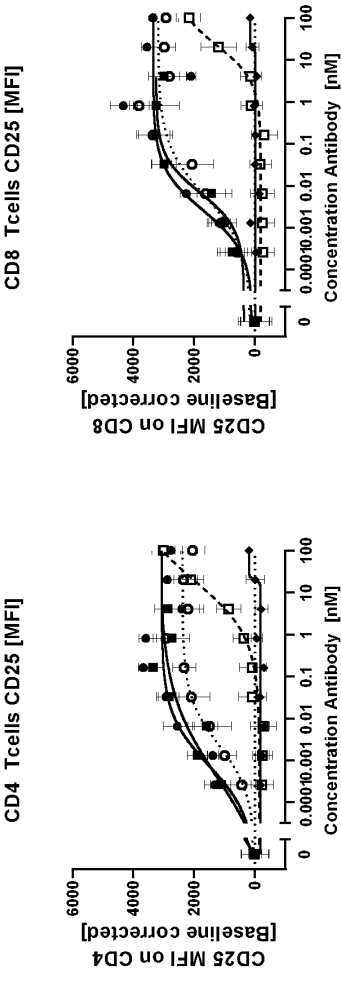


FIG.36C



- 4 49B4 (P1AD3690)
- 4+1 49B4/4B9 VhVI (P1AD4524)
- ◆ Isotype Control

FIG.36B

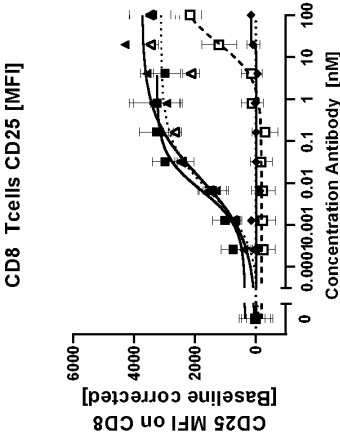
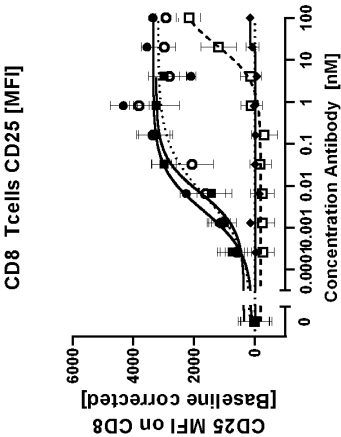


FIG.36D



- ▲ 3+1 CLC563/H212-D (P1AF6454)
- △ 3+1 CLC563/H212-S (P1AF6455)
- 4+1 CLC563/H212-D (P1AF7205)
- 4+1 CLC563/H212-S (P1AF7217)

Fig. 36

FIG. 36E

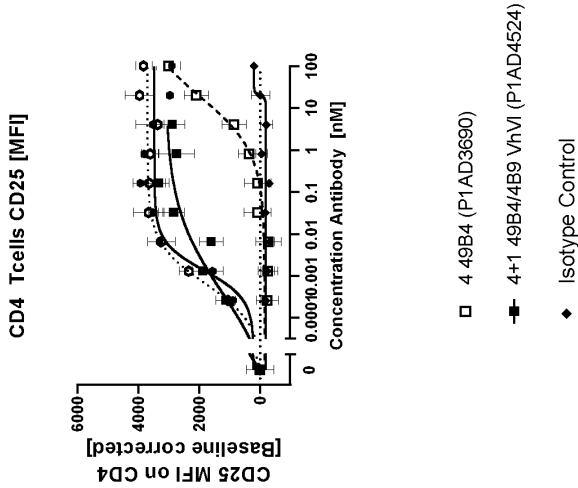


FIG. 36F

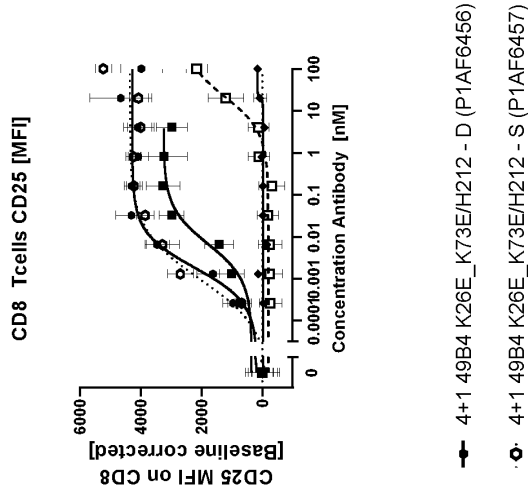


Fig. 37

FIG.37A

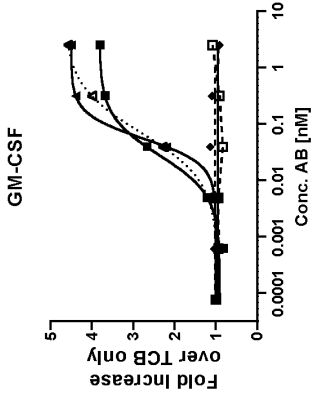


FIG.37B

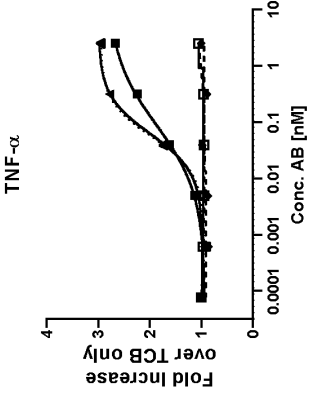


FIG.37C

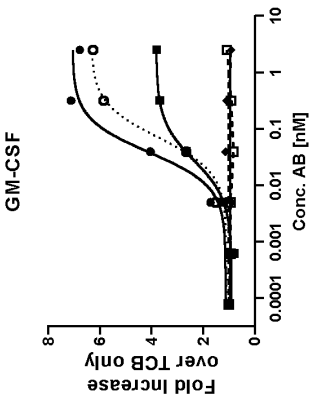
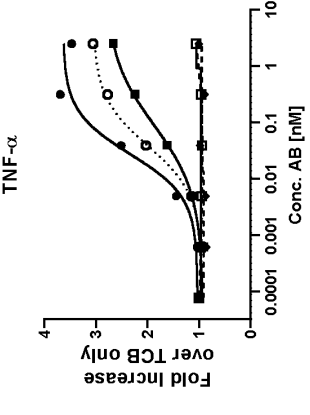


FIG.37D



- 4 49B4 (P1AD3690)
- 4+1 49B4/4B9 VhVI (P1AD4524)
- ◆ Isotype Control

- 3+1 CLC563/H212-D (P1AF6454)
- Δ· 3+1 CLC563/H212-S (P1AF6455)
- 4+1 CLC563/H212-D (P1AF7205)
- 4+1 CLC563/H212-S (P1AF7217)

Fig. 37

FIG.37E

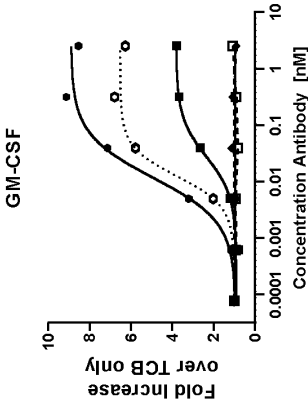
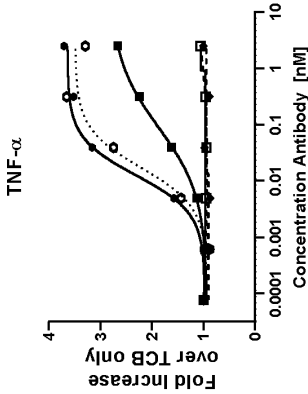


FIG.37F

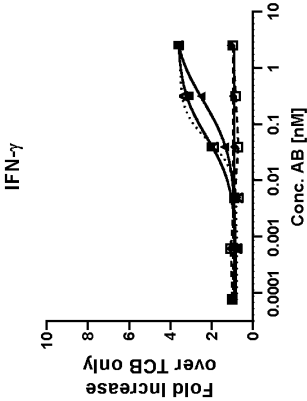


- 4 49B4 (P1AD3690)
- 4+1 49B4/4B9 VhVl (P1AD4524)
- ◆ Isotype Control

- 4+1 49B4 K26E_K73E/H212 - D (P1AF6456)
- 4+1 49B4 K26E_K73E/H212 - S (P1AF6457)

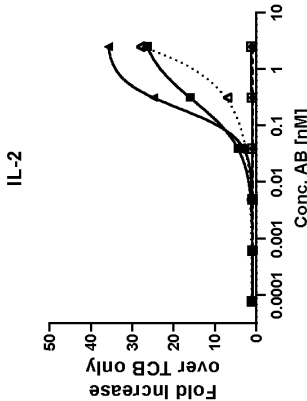
Fig. 38

FIG.38A



- 4 49B4 (P1AD3690)
- 4+1 49B4/4B9 VhVI (P1AD4524)
- 4+1 CLC563/H212-D (P1AF7205)
- 4+1 CLC563/H212-S (P1AF7217)

FIG.38B



- 3+1 CLC563/H212-D (P1AF6454)
- △ 3+1 CLC563/H212-S (P1AF6455)
- 4+1 CLC563/H212-D (P1AF7205)
- 4+1 CLC563/H212-S (P1AF7217)

FIG.38C

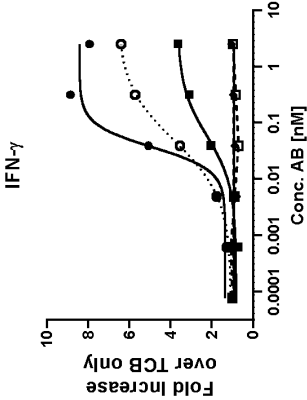


FIG.38D

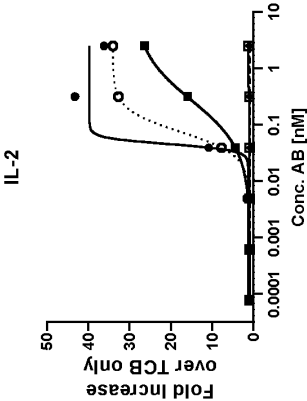


Fig. 38

FIG. 38E

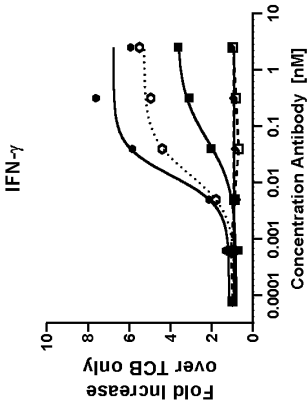
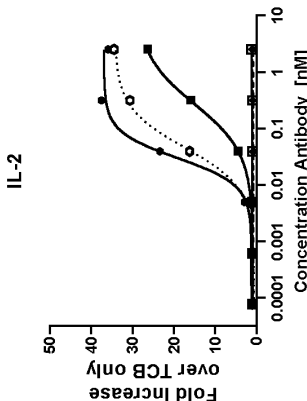


FIG. 38F



- 4 49B4 (P1AD3690)
- 4+1 49B4/4B9 VhVl (P1AD4524)
- ◆ Isotype Control

- 4+1 49B4 K26E_K73E/H212 - D (P1AF6456)
- ◐ 4+1 49B4 K26E_K73E/H212 - S (P1AF6457)

Fig. 39

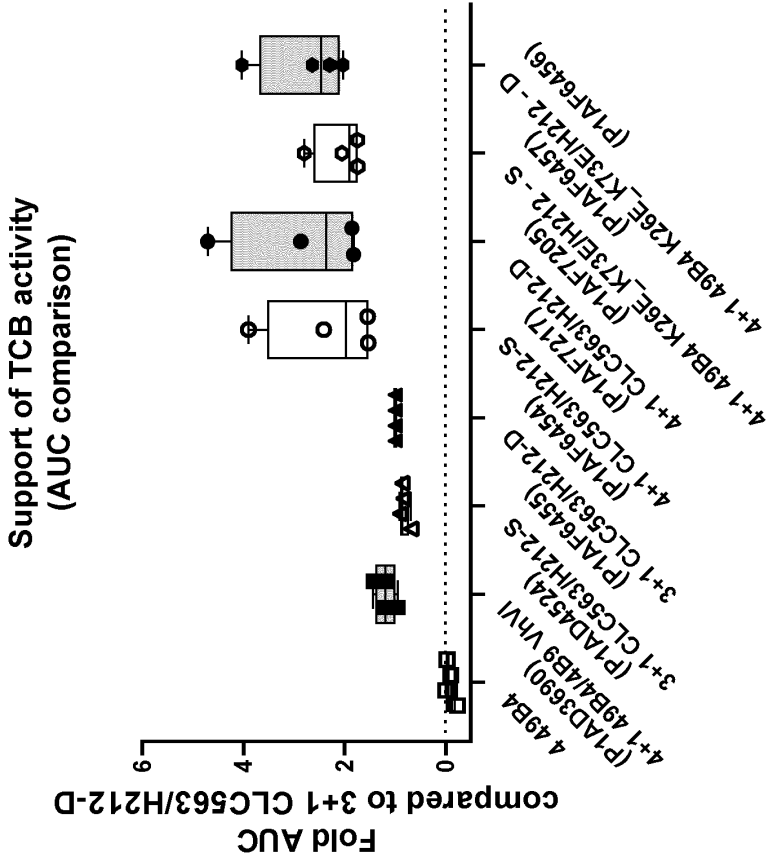


Fig. 40

FIG.40A

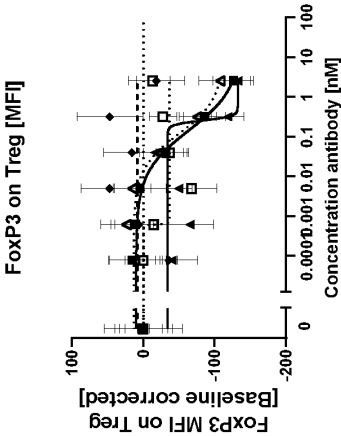


FIG.40B

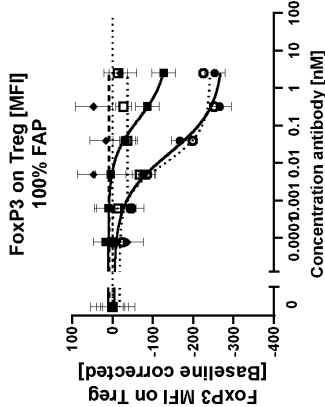
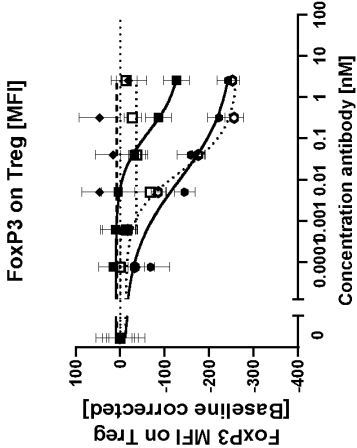


FIG.40C



- 4 49B4 (P1AD3690)
- 4+1 49B4/4B9 VhVl (P1AD4524)
- ▲ 3+1 CLC563/H212-D (P1AF6454)
- ▲ 3+1 CLC563/H212-S (P1AF6455)
- 4+1 CLC563/H212-D (P1AF7205)
- 4+1 CLC563/H212-S (P1AF7217)
- 4+1 49B4 K26E_K73E/H212 - D (P1AF6456)
- 4+1 49B4 K26E_K73E/H212 - S (P1AF6457)
- ◆ Isotype Control

Fig. 41

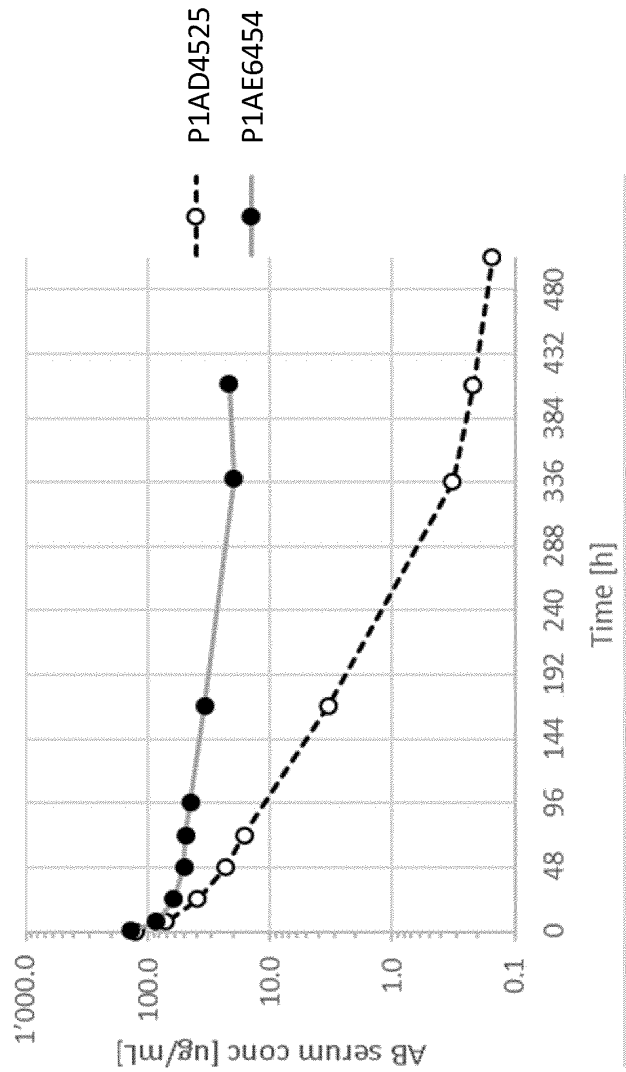


Fig. 42A

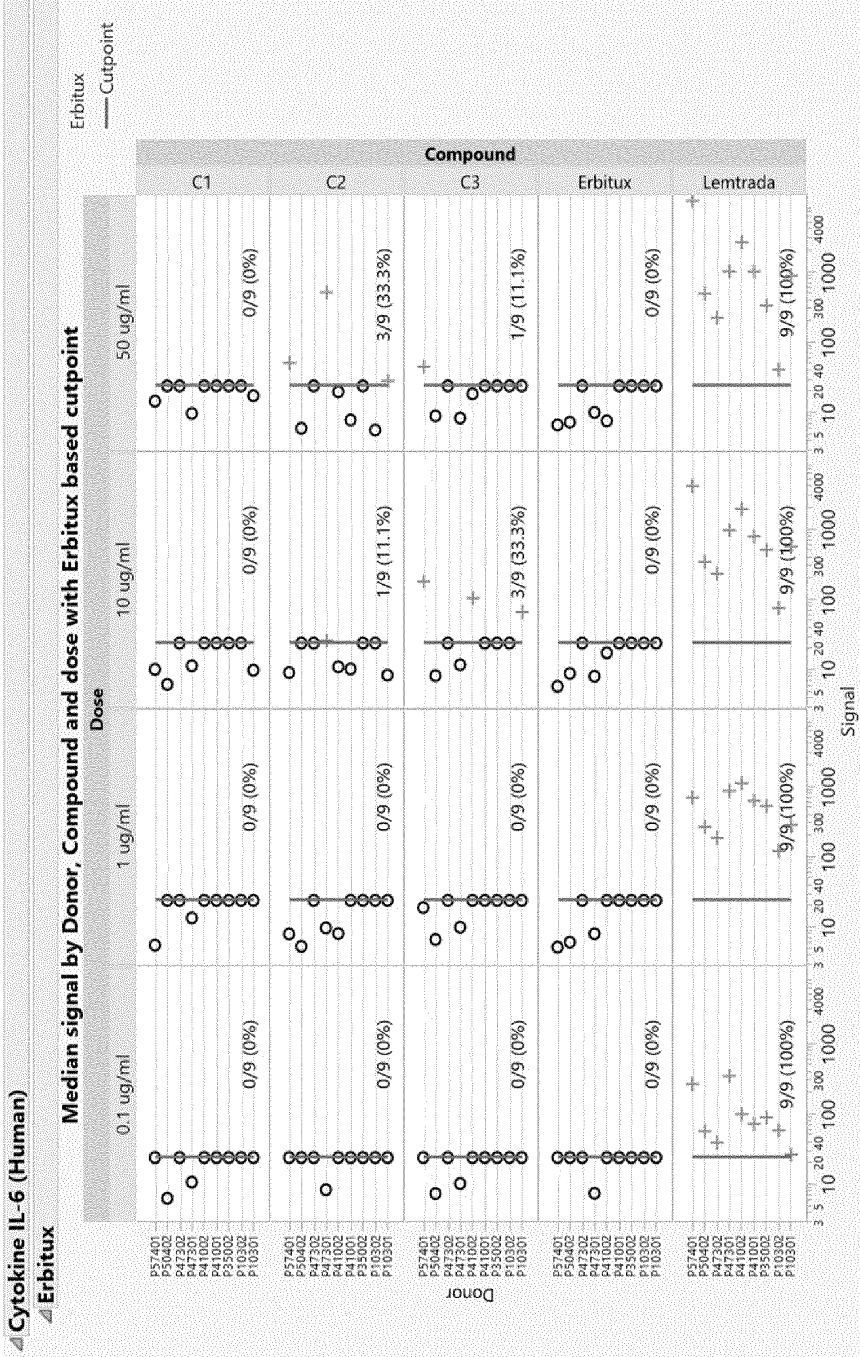


Fig. 43B

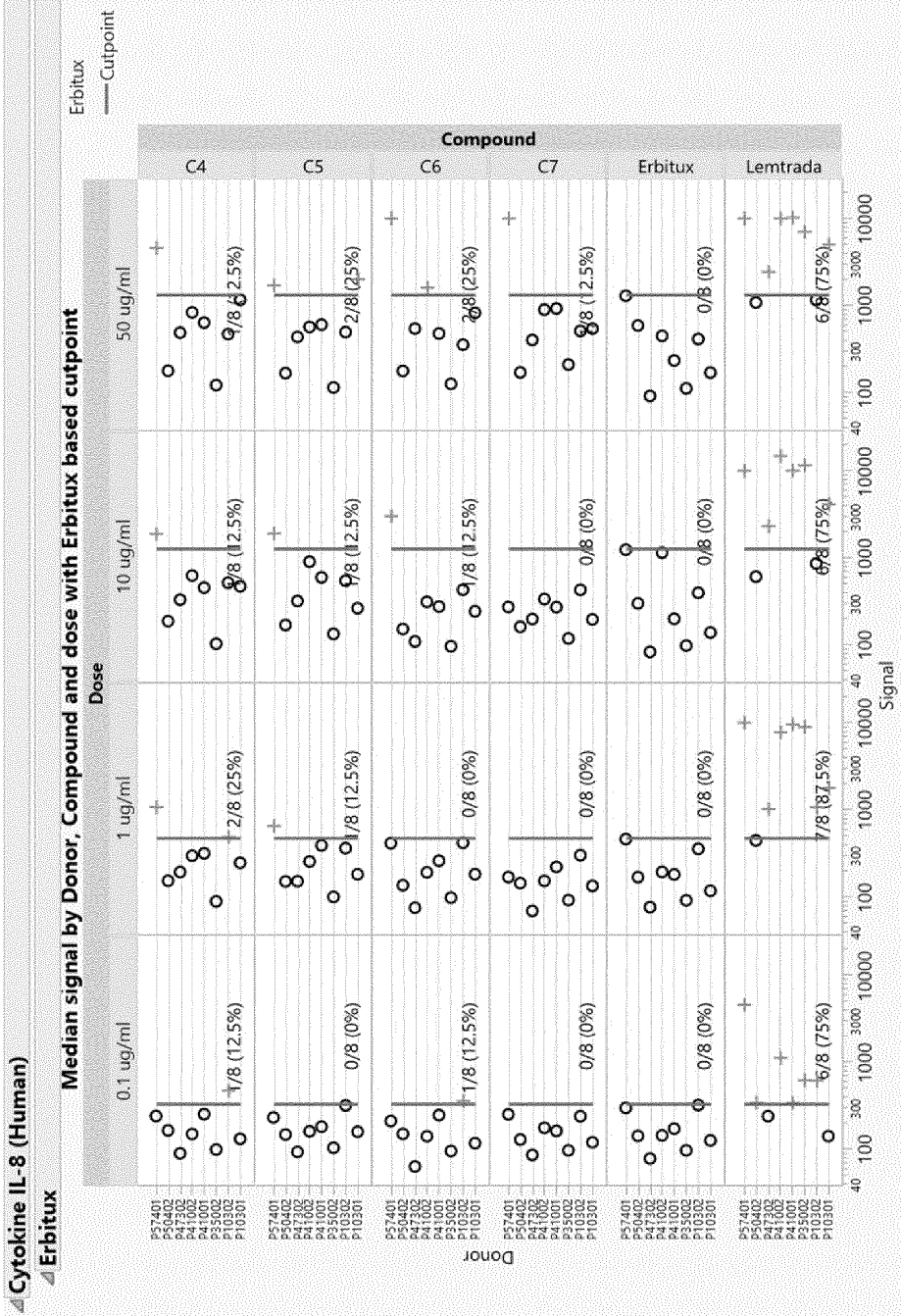


Fig. 44A

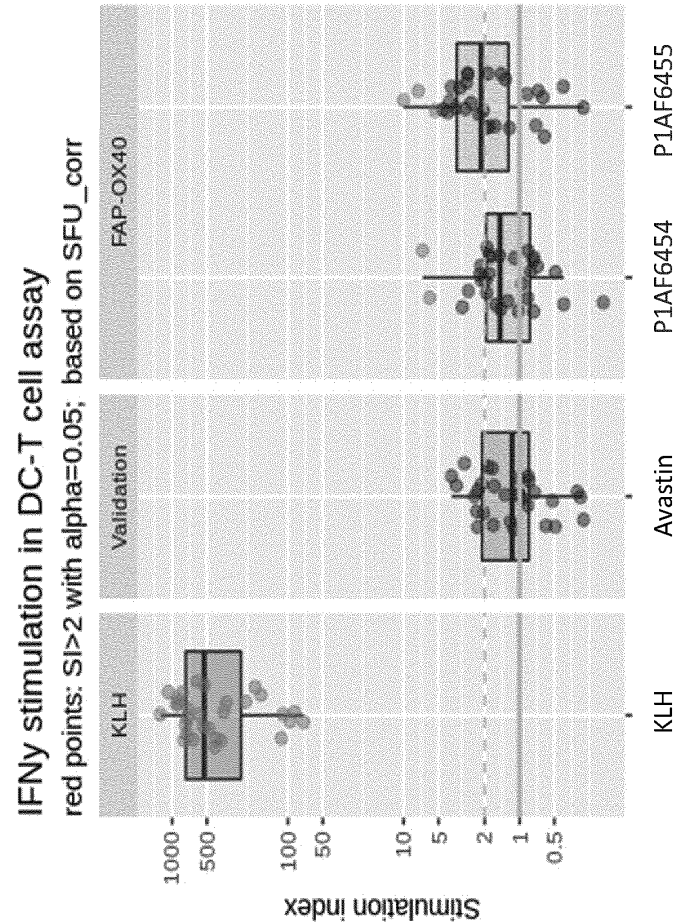


Fig. 44B

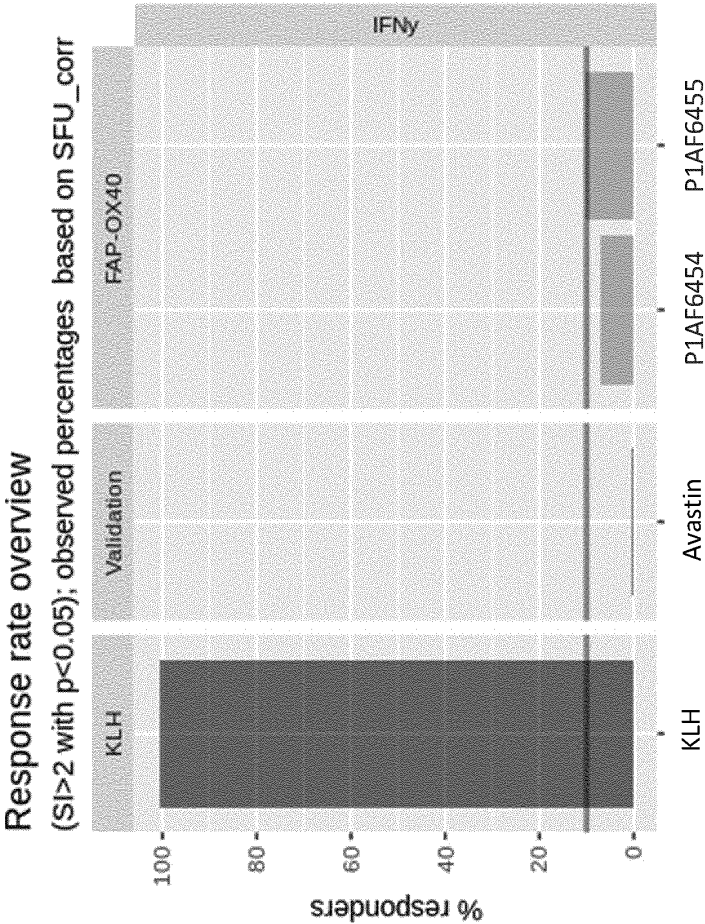


Fig. 45A

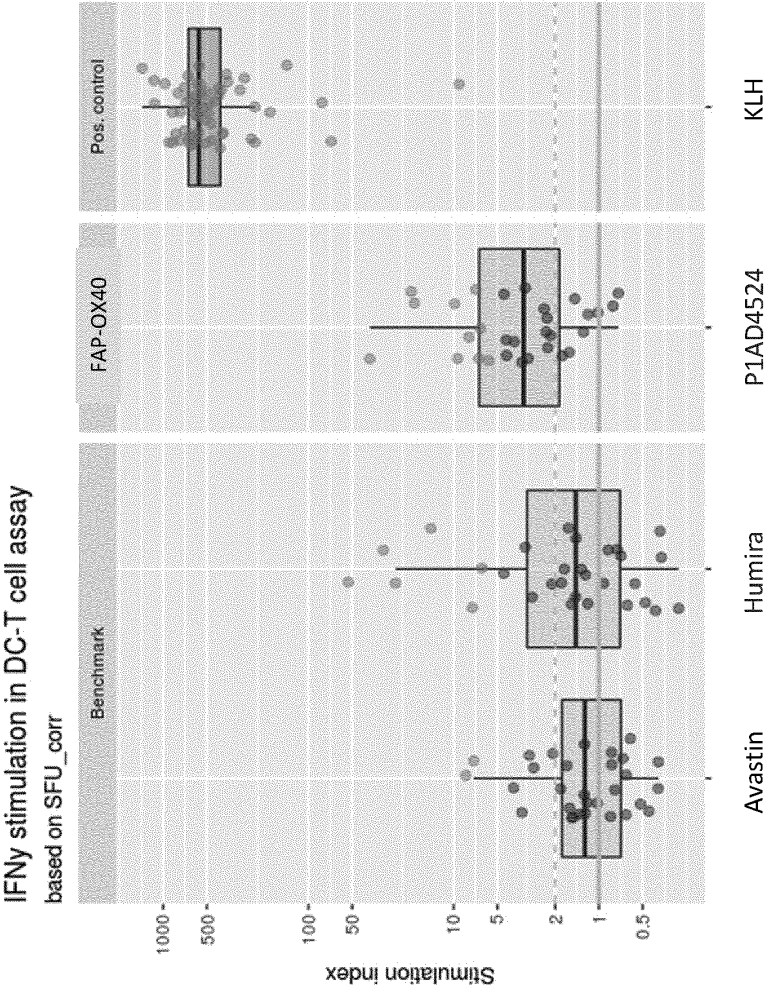


Fig. 45B

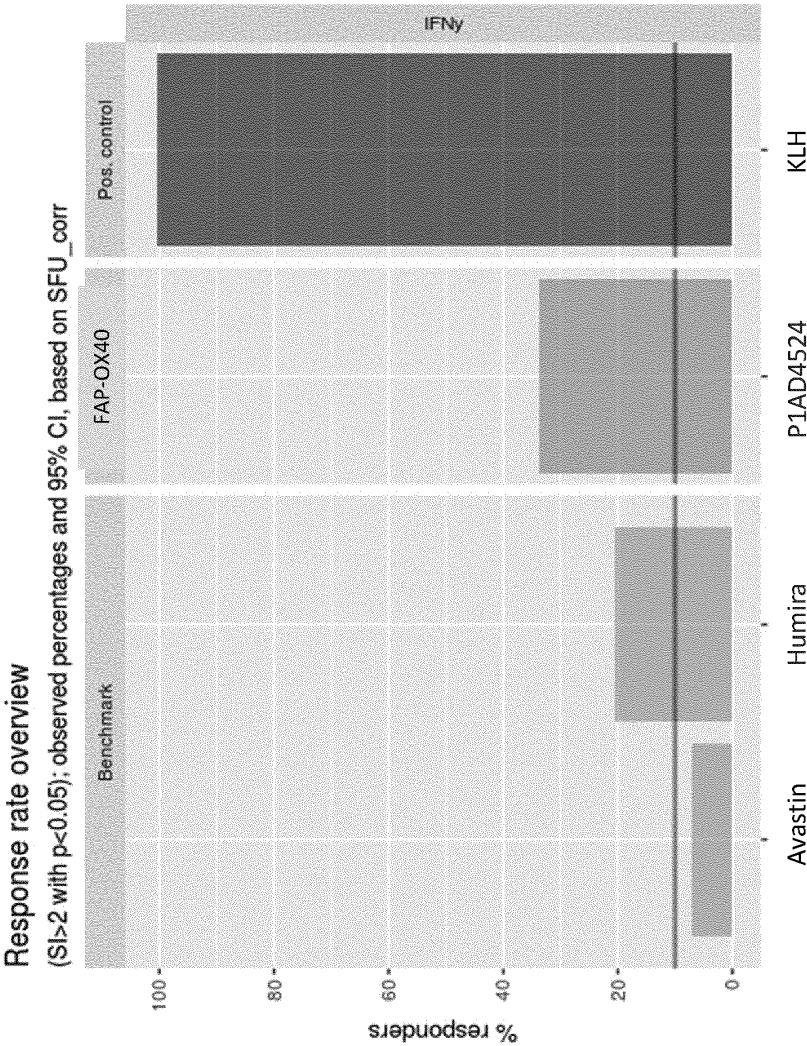


Fig. 46

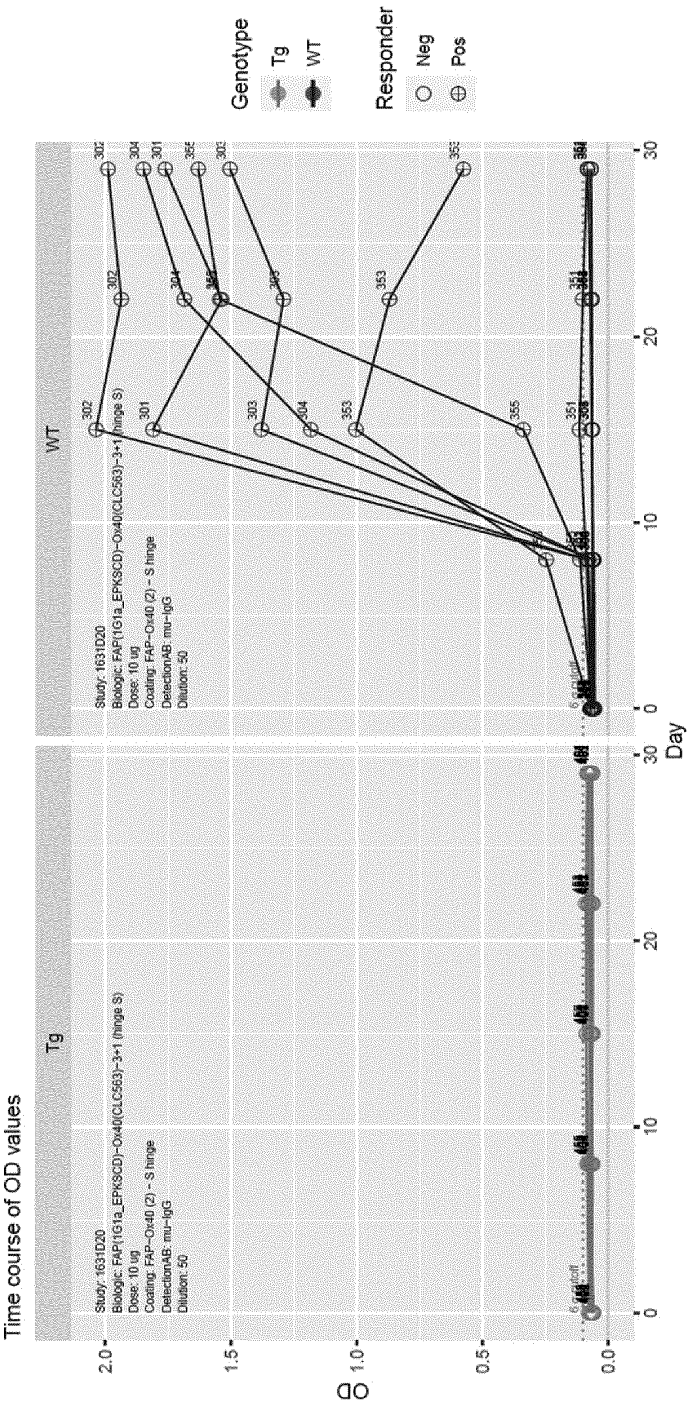


Fig. 47

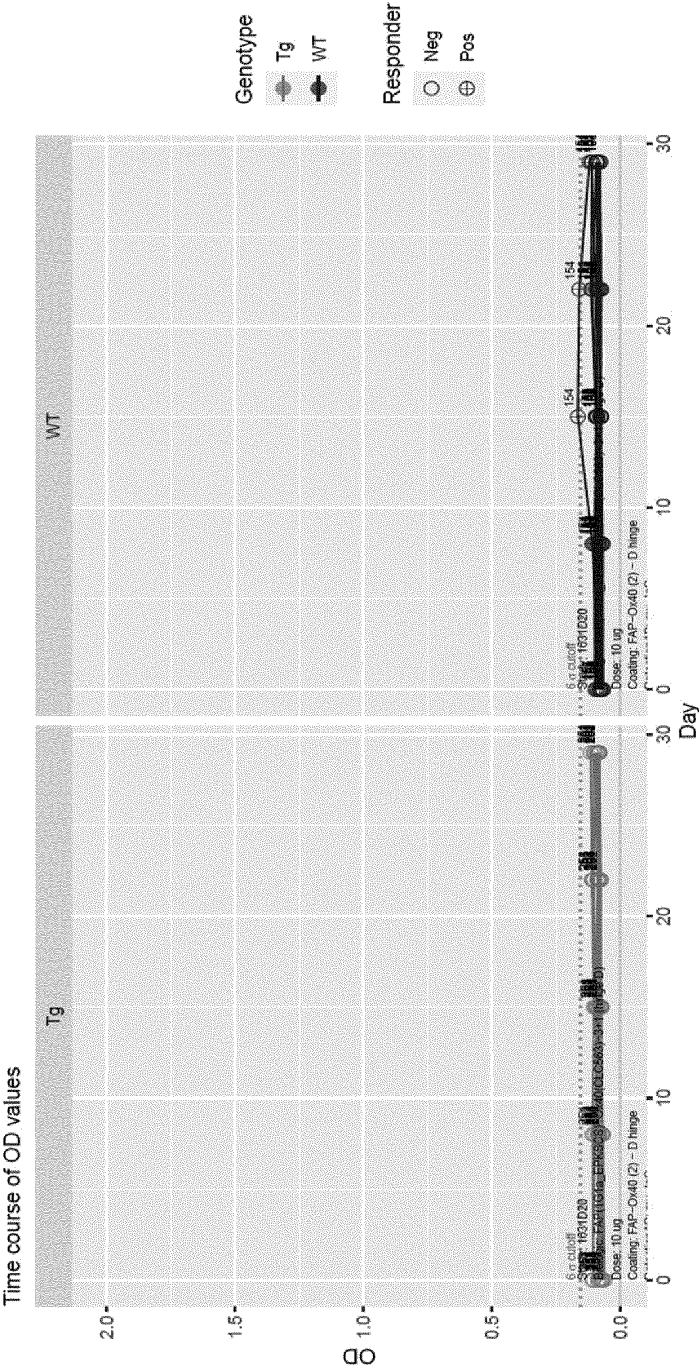


Fig. 48

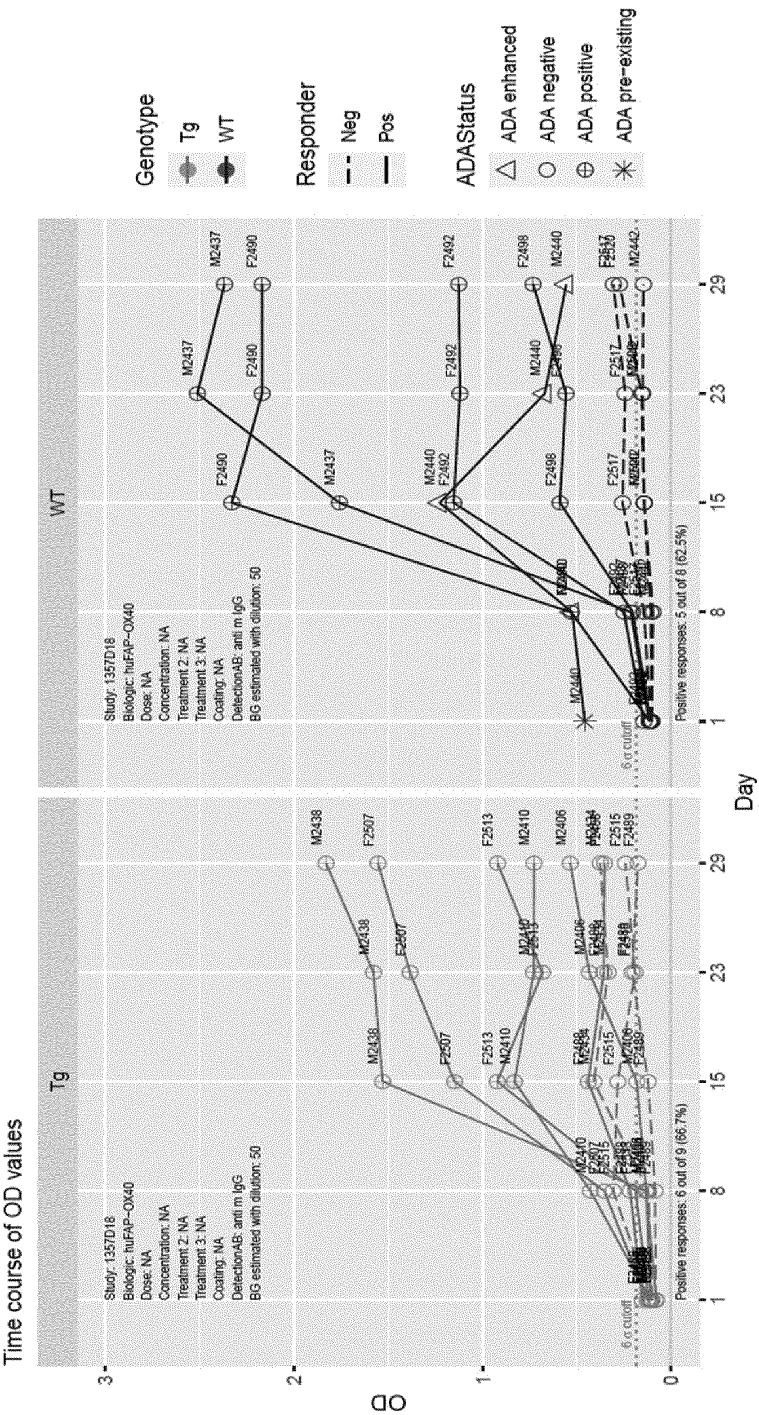
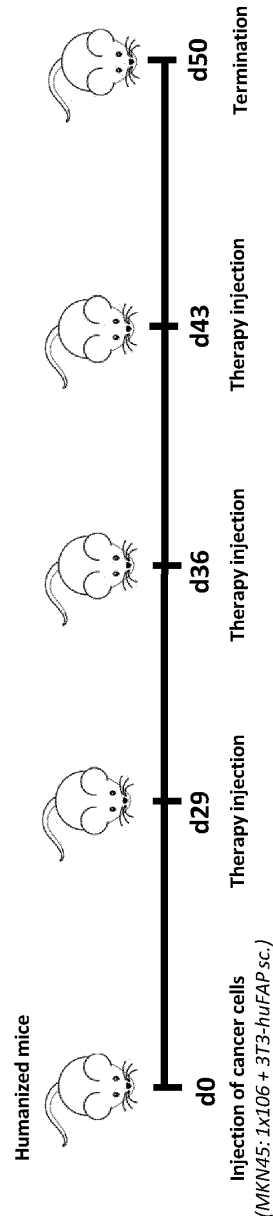


Fig. 49



Group	No. of animals	Compound	Dose (mg/kg)	Therapy	No. of treatments
1	9	Vehicle	-	i.v.	3 (once weekly)
2	9	CEACAM5-TCB	0.5	i.v.	3 (once weekly)
3	9	CEACAM5-TCB + FAP-OX40 (49B4 CPV; 4+1)	0.5 13.3	i.v.	3 (once weekly)
4	9	CEACAM5-TCB + FAP-OX40 (CLC563; 3+1)	0.44 12	i.v.	3 (once weekly)
5	9	CEACAM5-TCB + FAP-OX40 (8H9; 2+1)	0.5 9	i.v.	3 (once weekly)
6	9	CEACAM5-TCB + FAP-OX40 (49B4)	0.5 12	i.v.	3 (once weekly)

Fig. 50

FIG. 50A

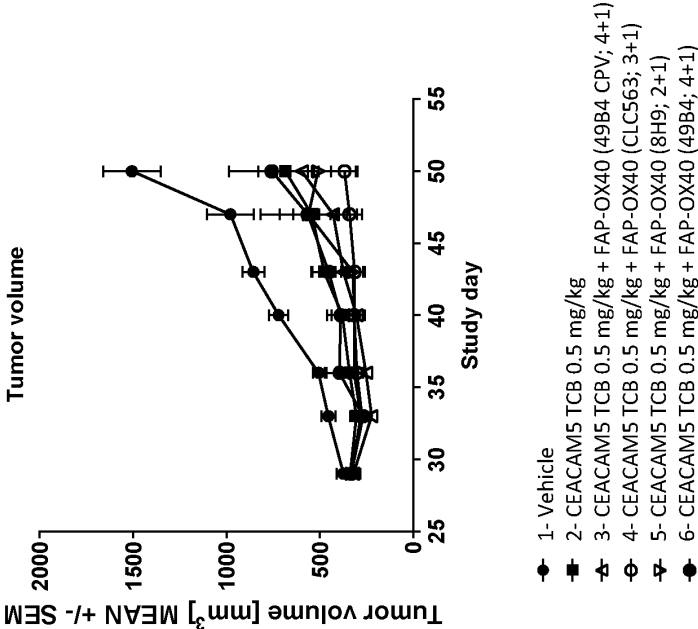


FIG. 50B

