

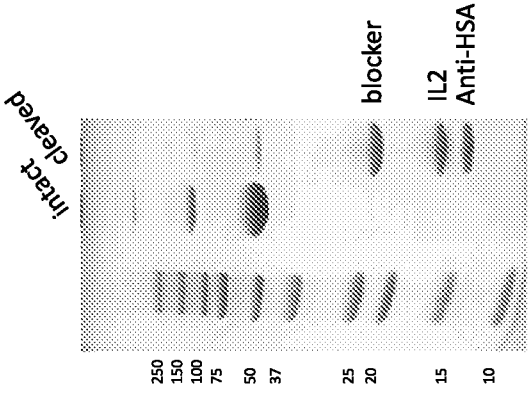
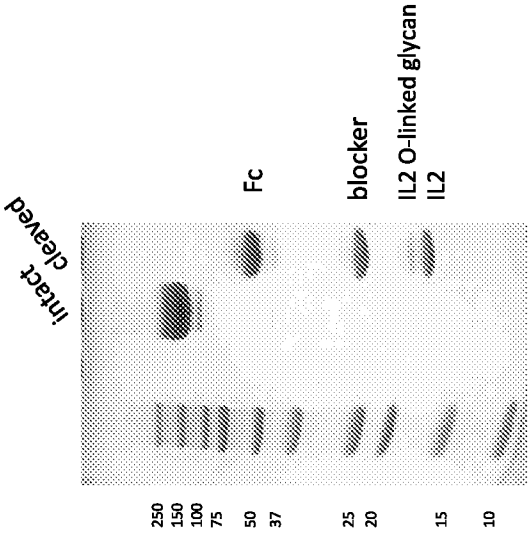


FIG. 1A

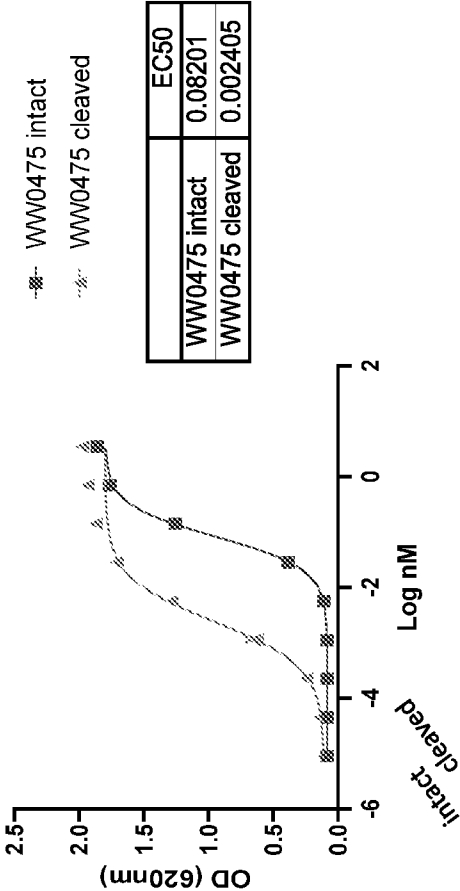
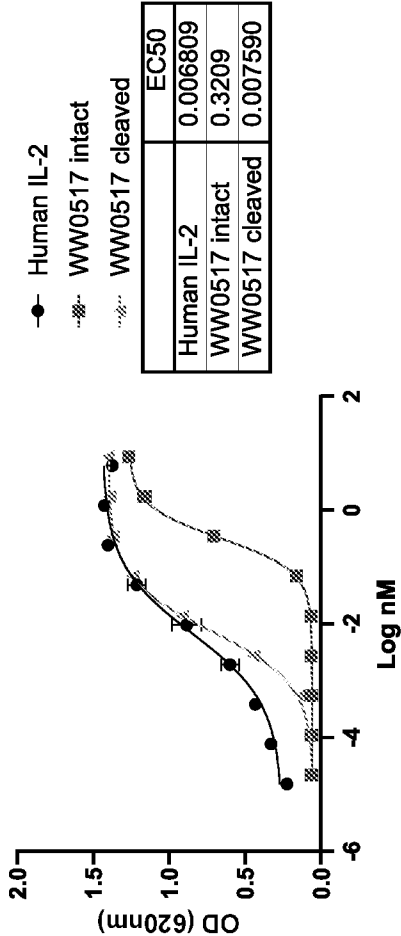


FIG. 1B



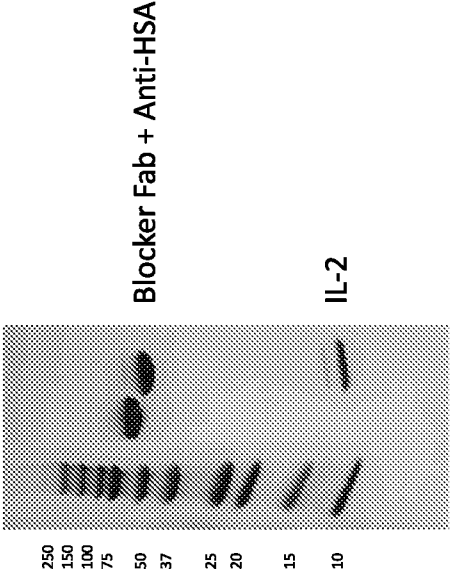
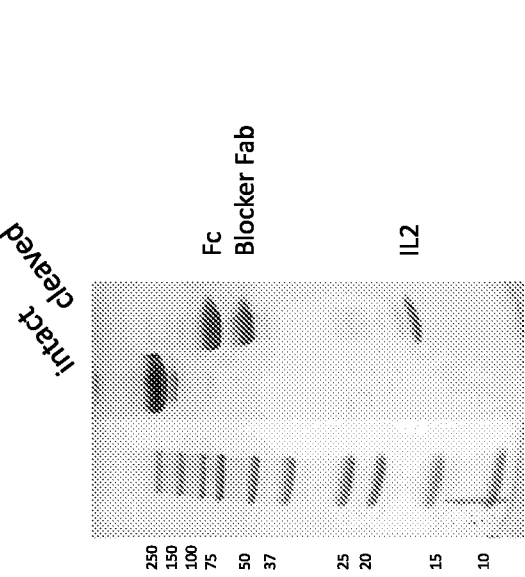


FIG. 1C

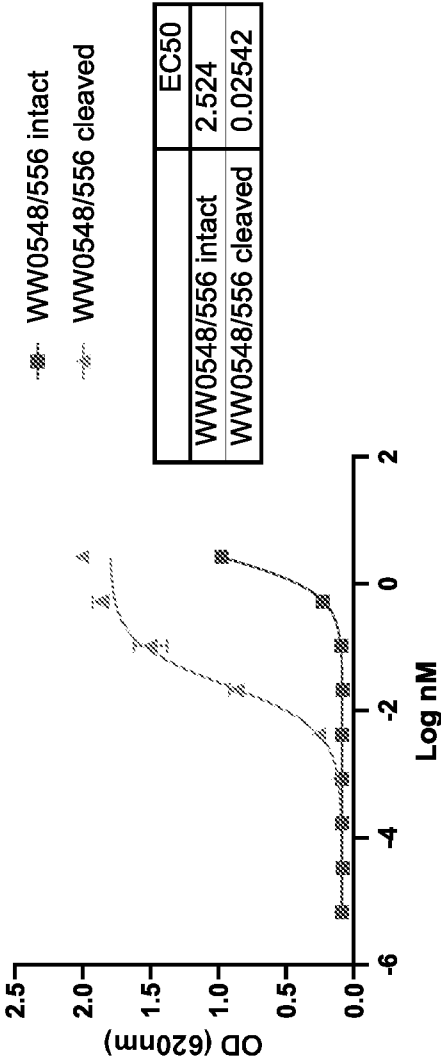
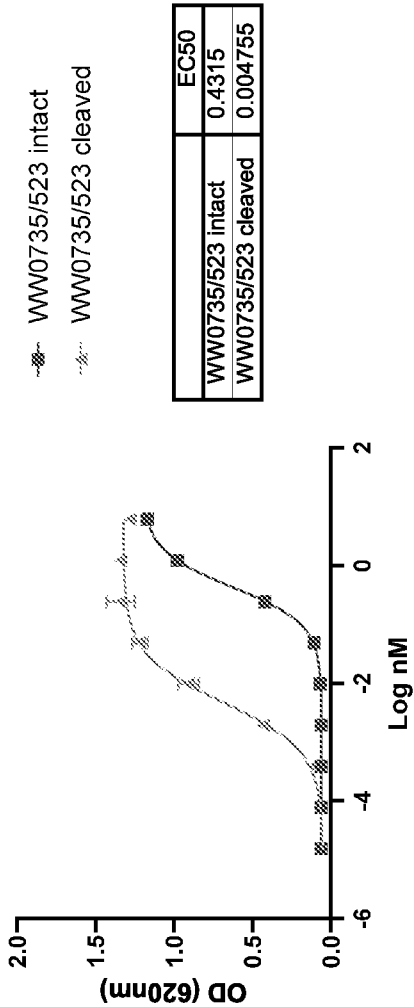


FIG. 1D



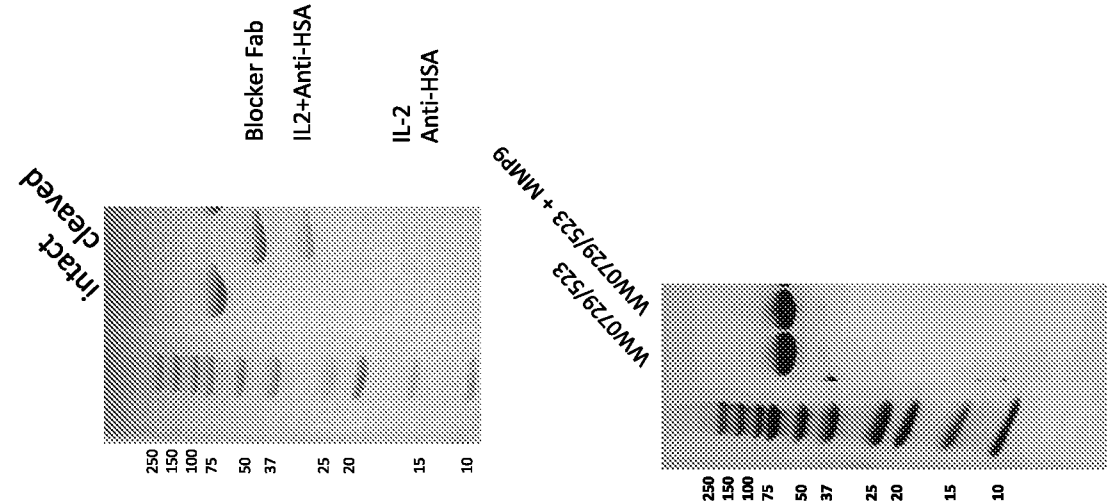


FIG. 1E

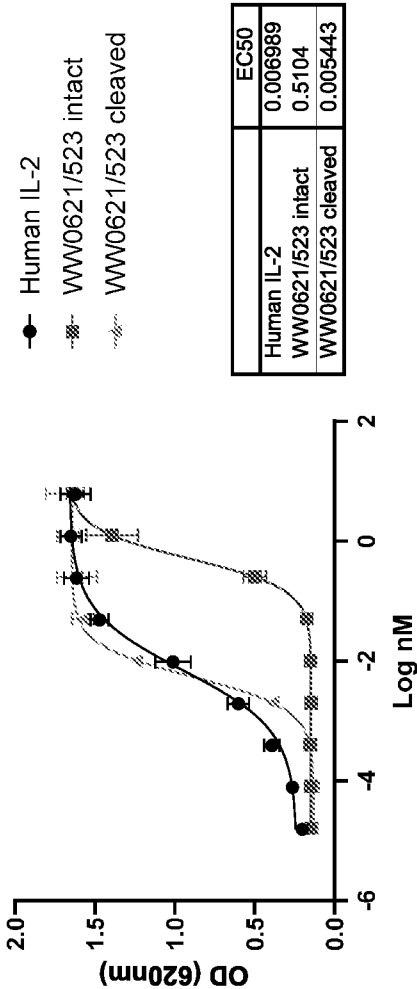
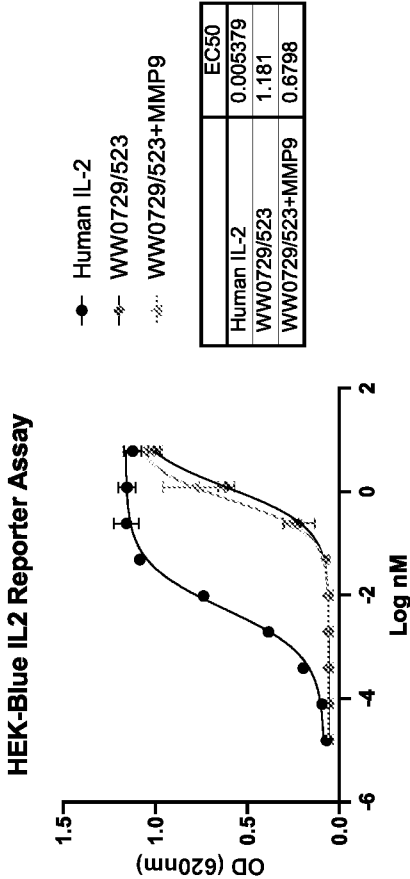


FIG. 1F



HEK-Blue IL2 Reporter Assay

FIG. 2A

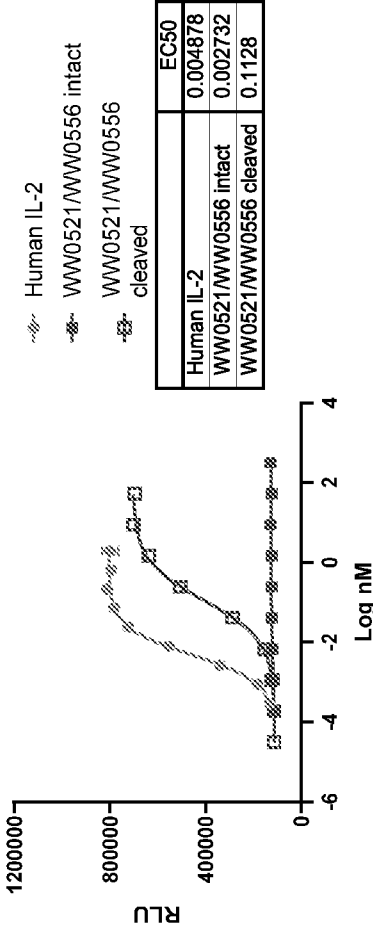


FIG. 2B

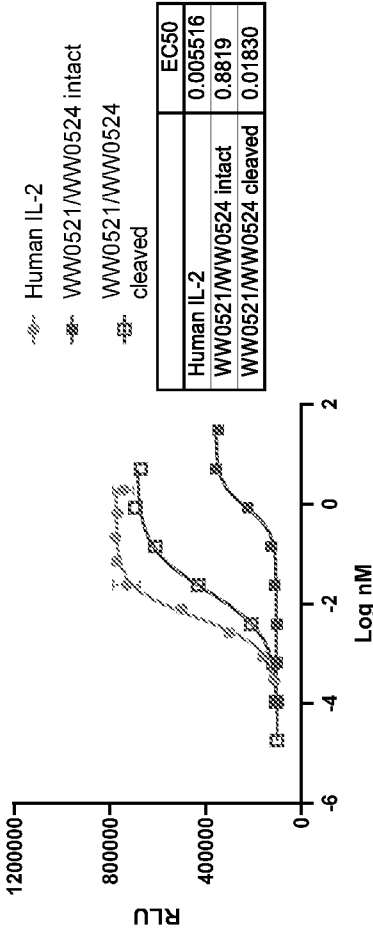


FIG. 2C

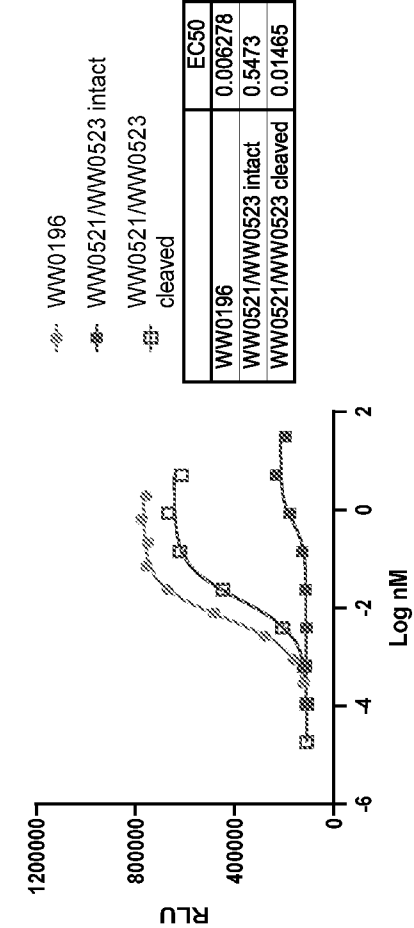


FIG. 2D

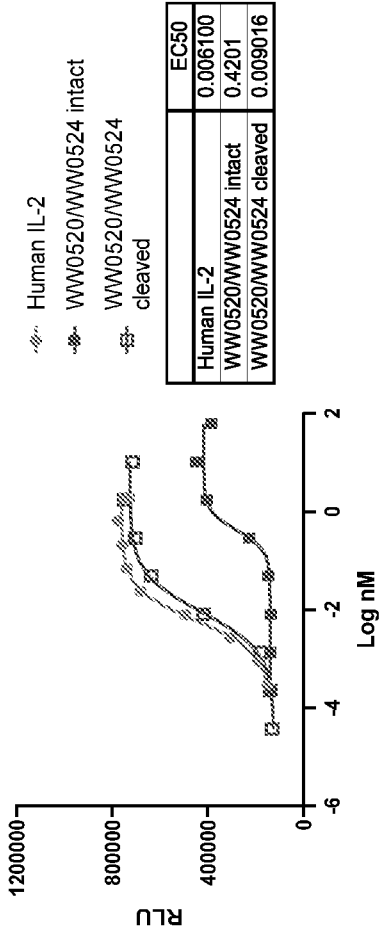


FIG. 2E

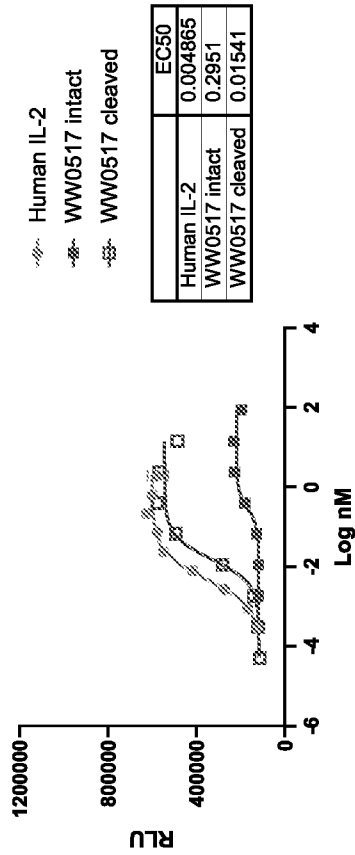


FIG. 2F

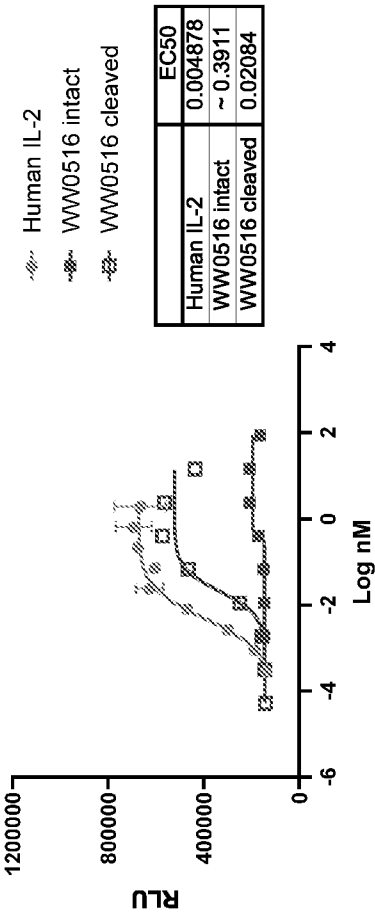


FIG. 2G

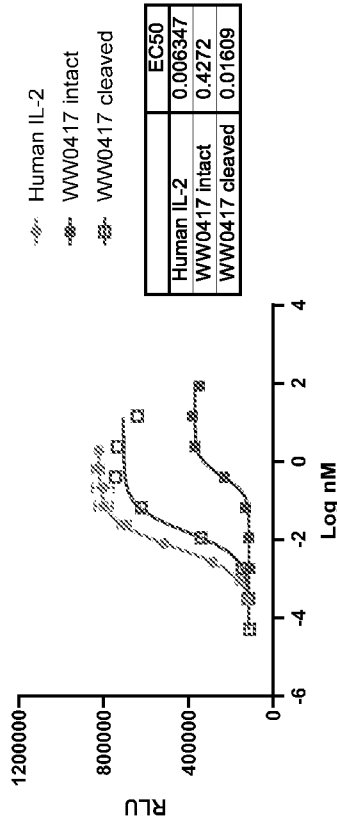


FIG. 2H

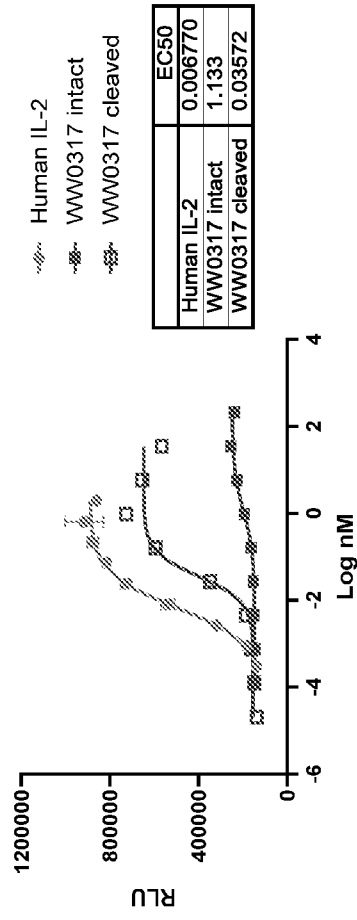


FIG. 2J

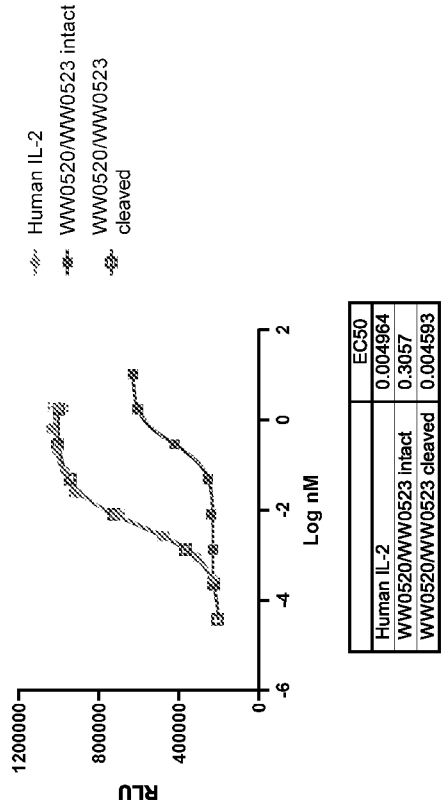


FIG. 2L

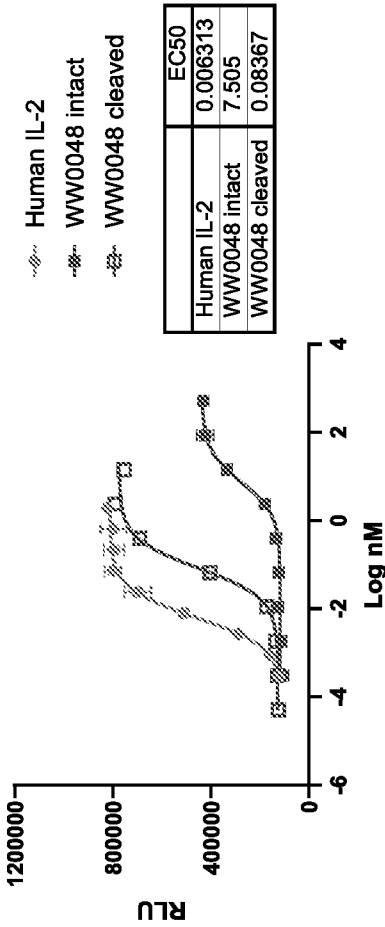


FIG. 2I

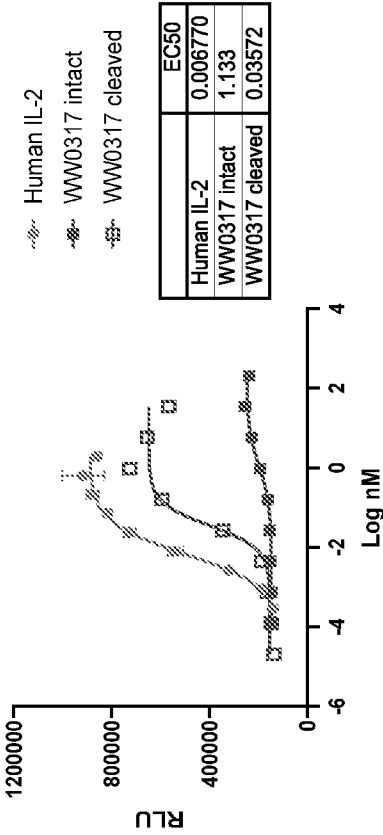


FIG. 2K

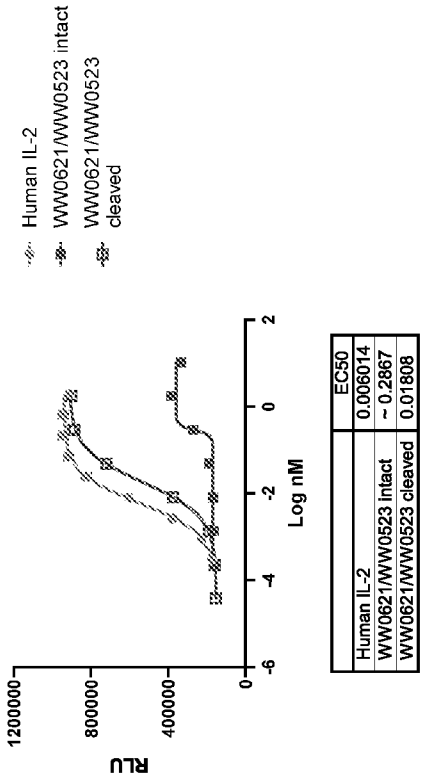


FIG. 3A

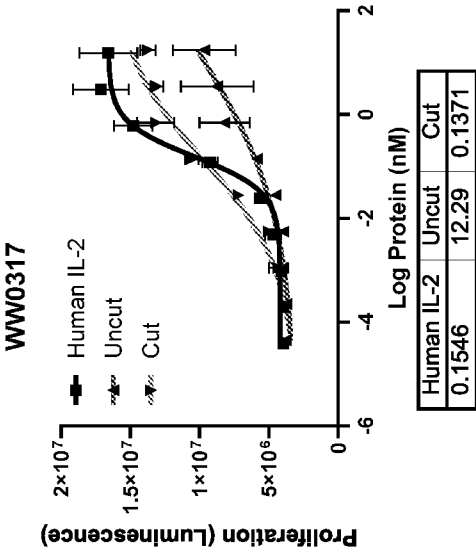


FIG. 3B

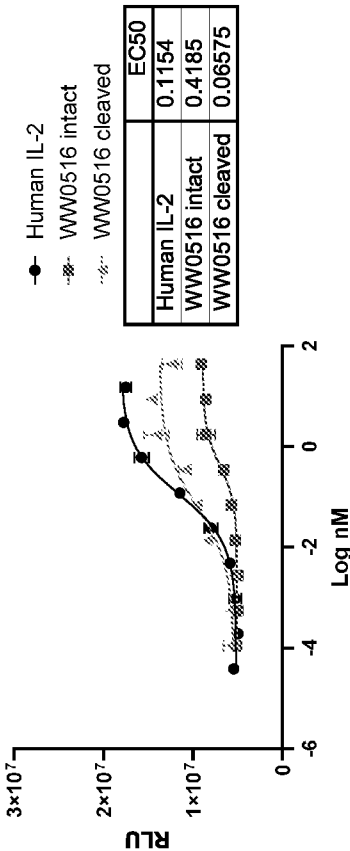


FIG. 3C

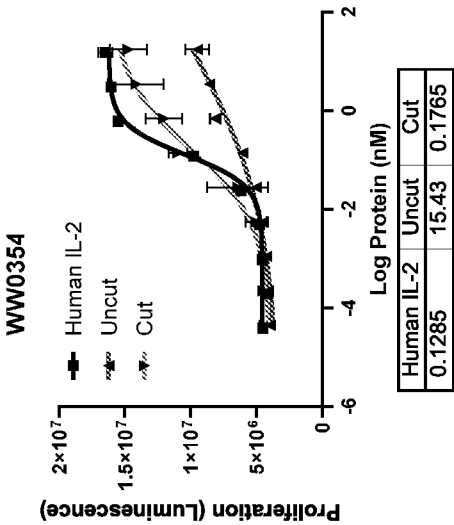


FIG. 3D

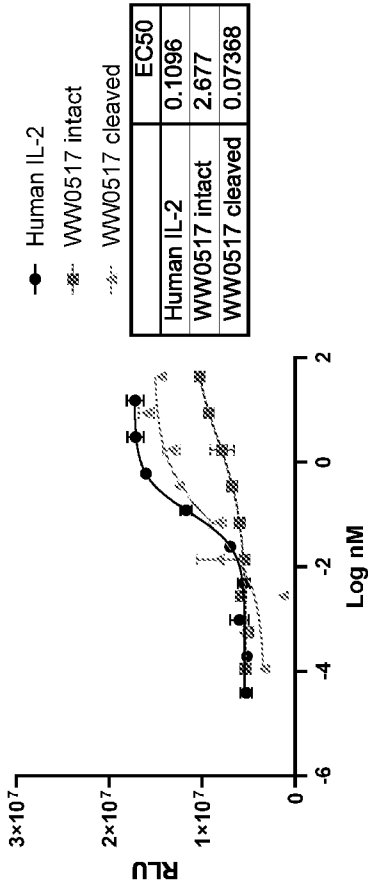


FIG. 3F

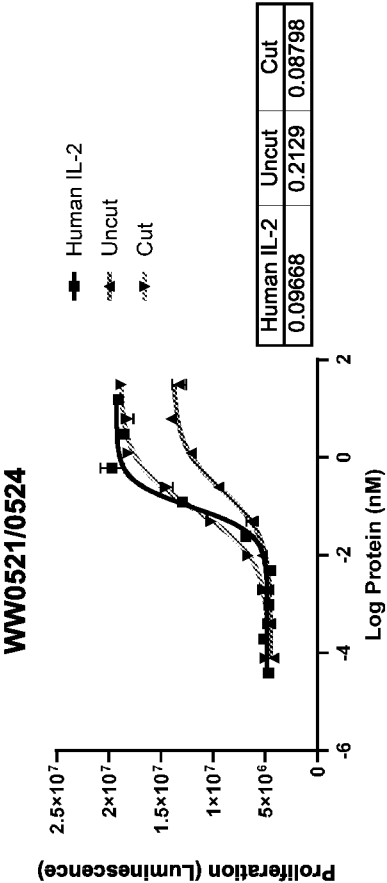


FIG. 3E

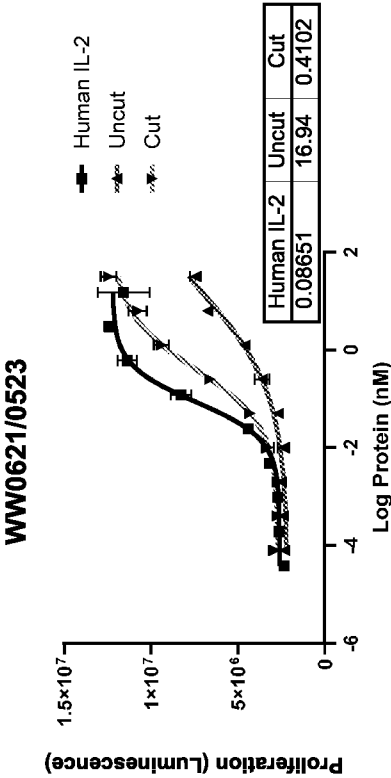


FIG. 3G

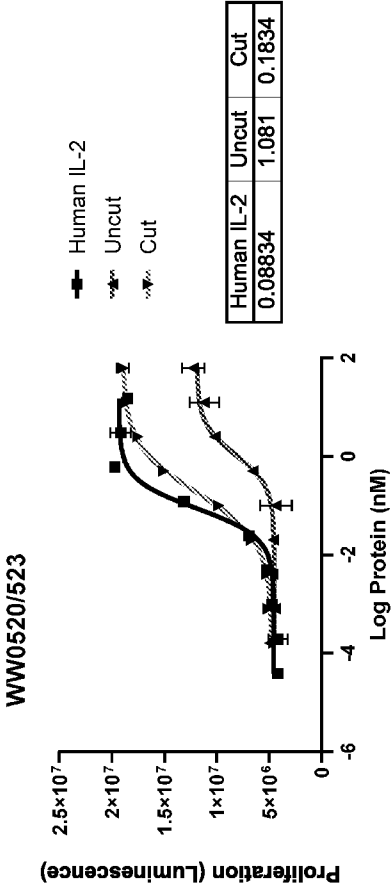


FIG. 3H

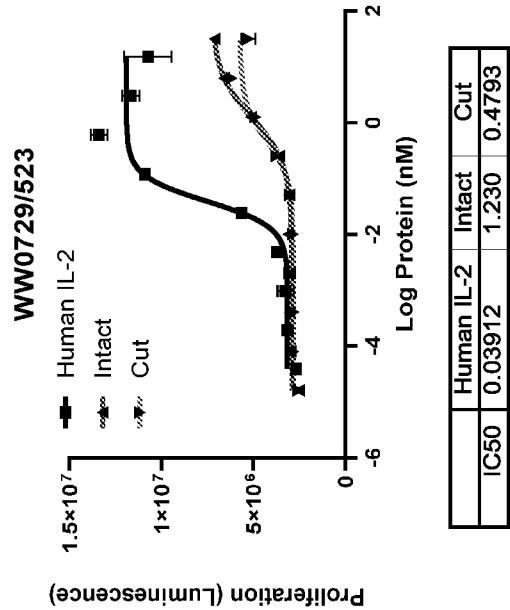


FIG. 3I

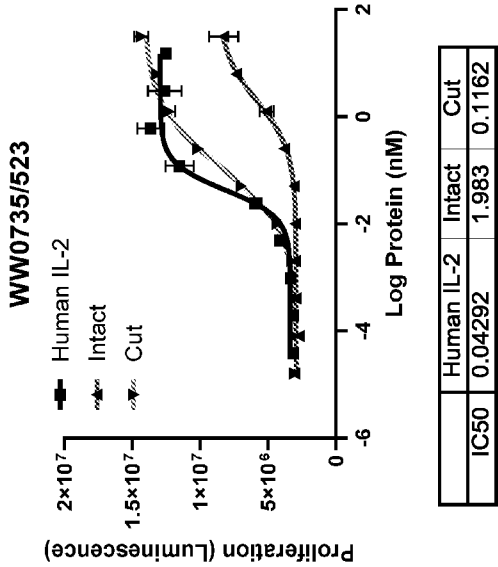


FIG. 3J

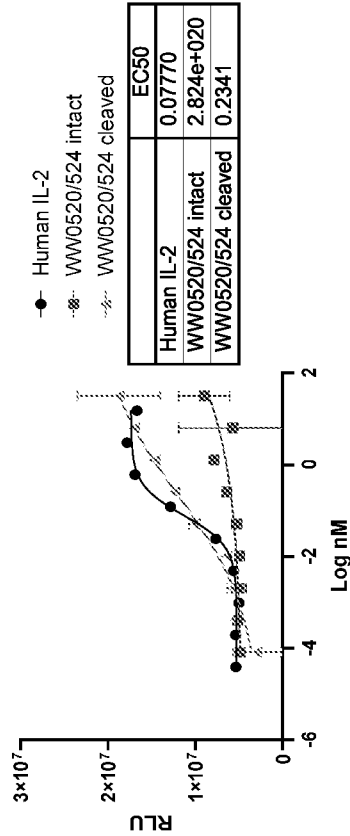


FIG. 4A

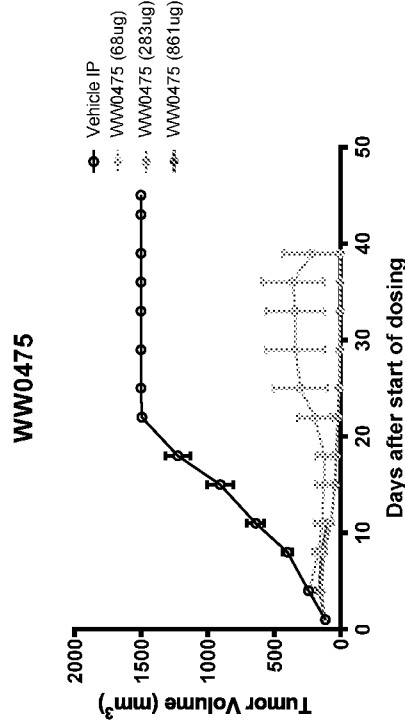


FIG. 4B

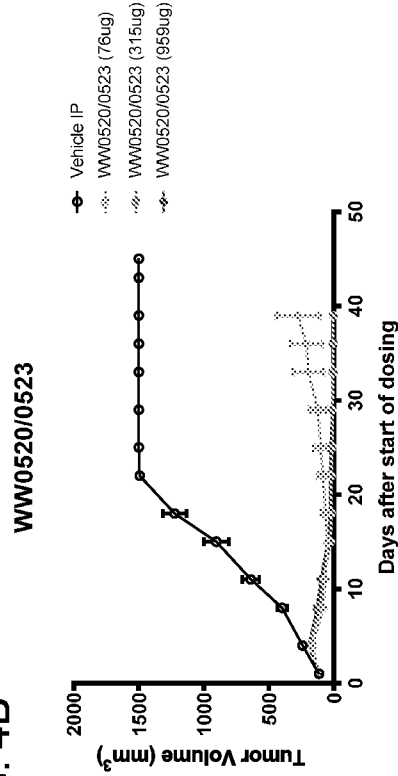


FIG. 4C

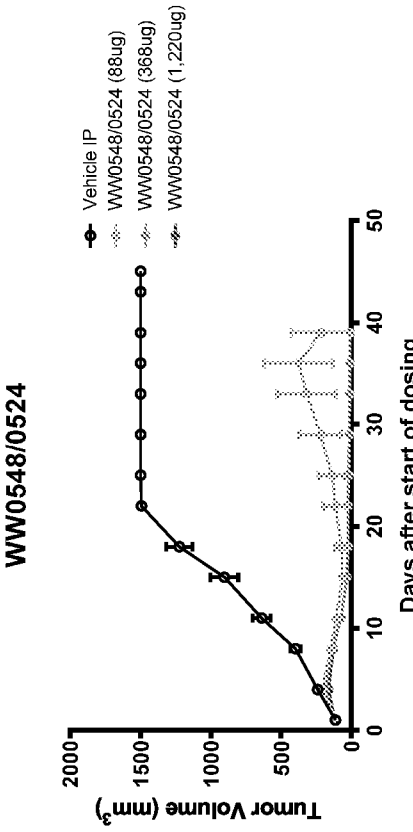


FIG. 4D

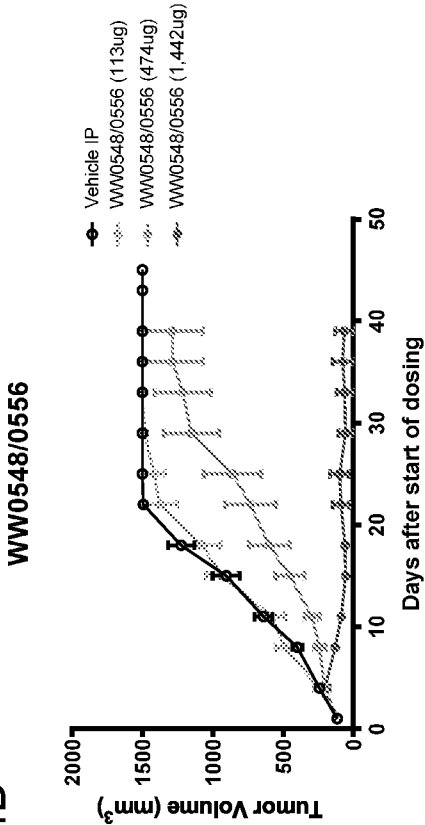


FIG. 4E

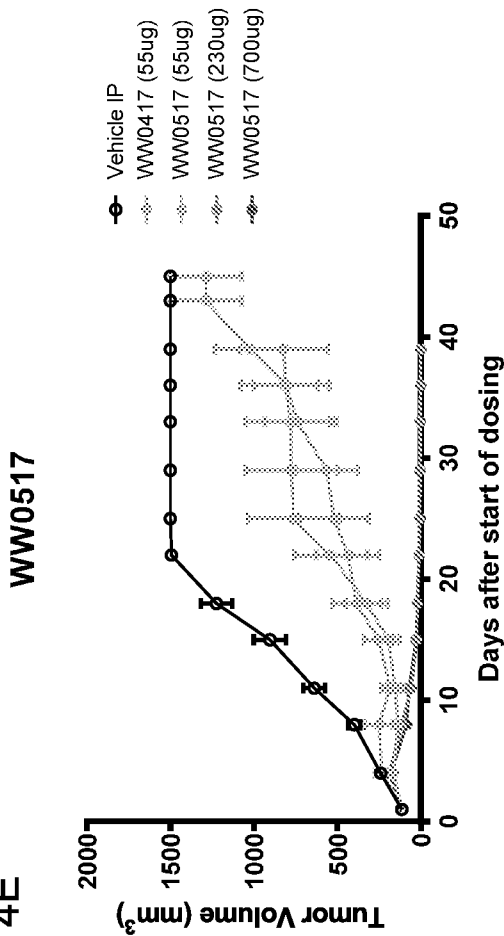


FIG. 4F

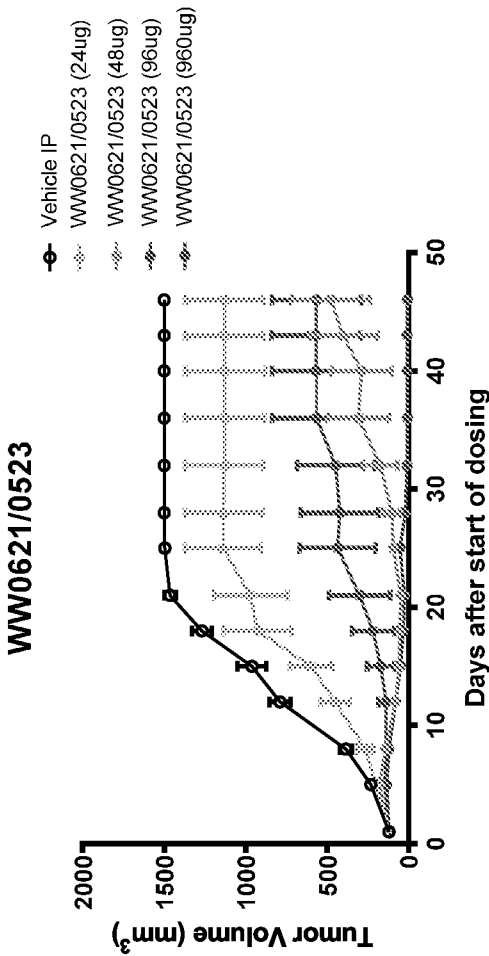


FIG. 4G

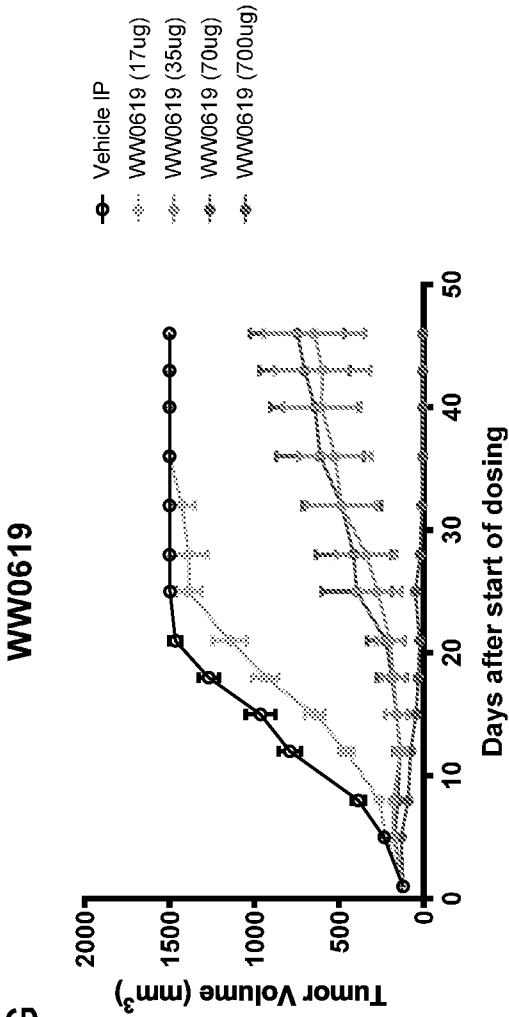


FIG. 5A

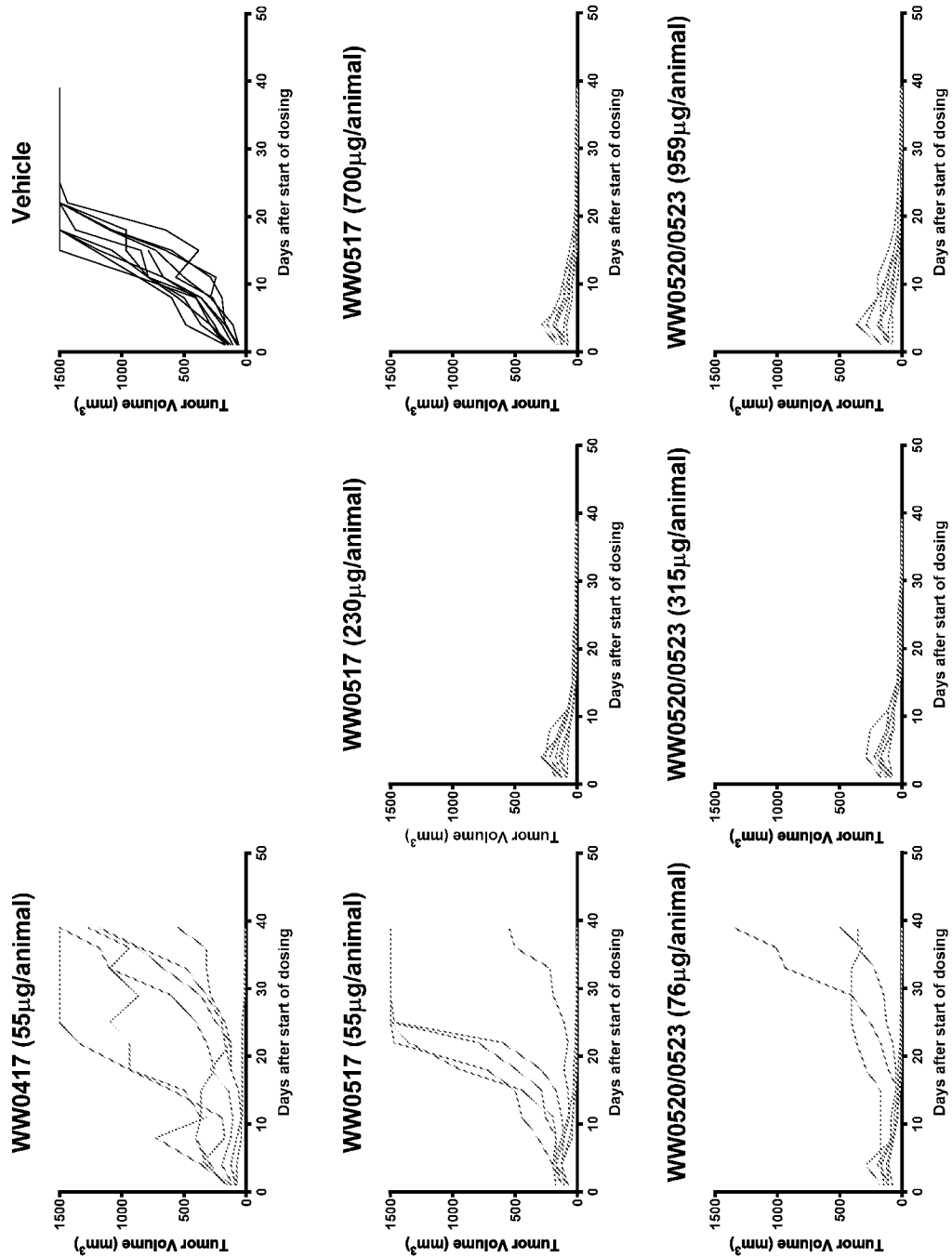


FIG. 5B

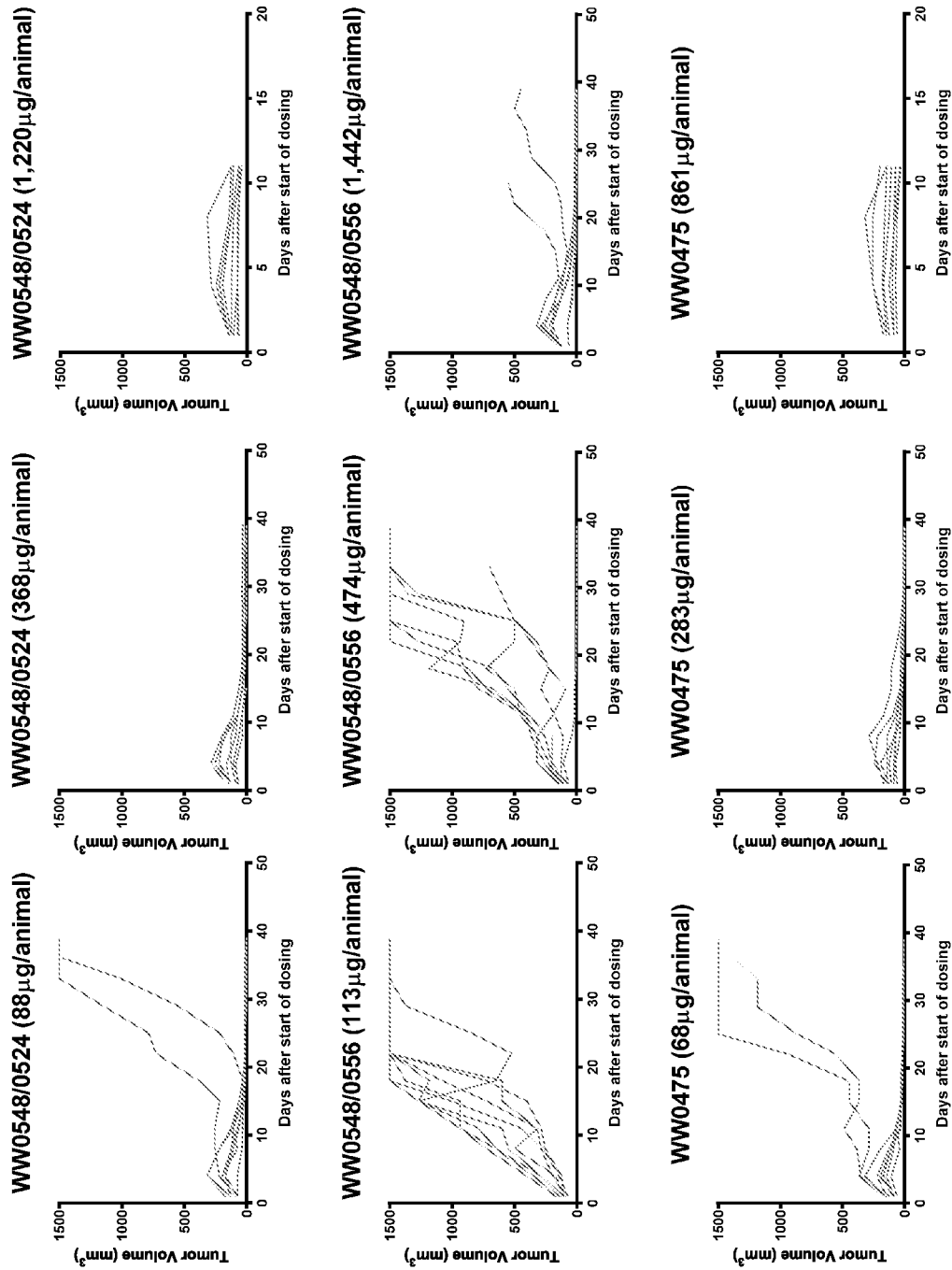


FIG. 5C

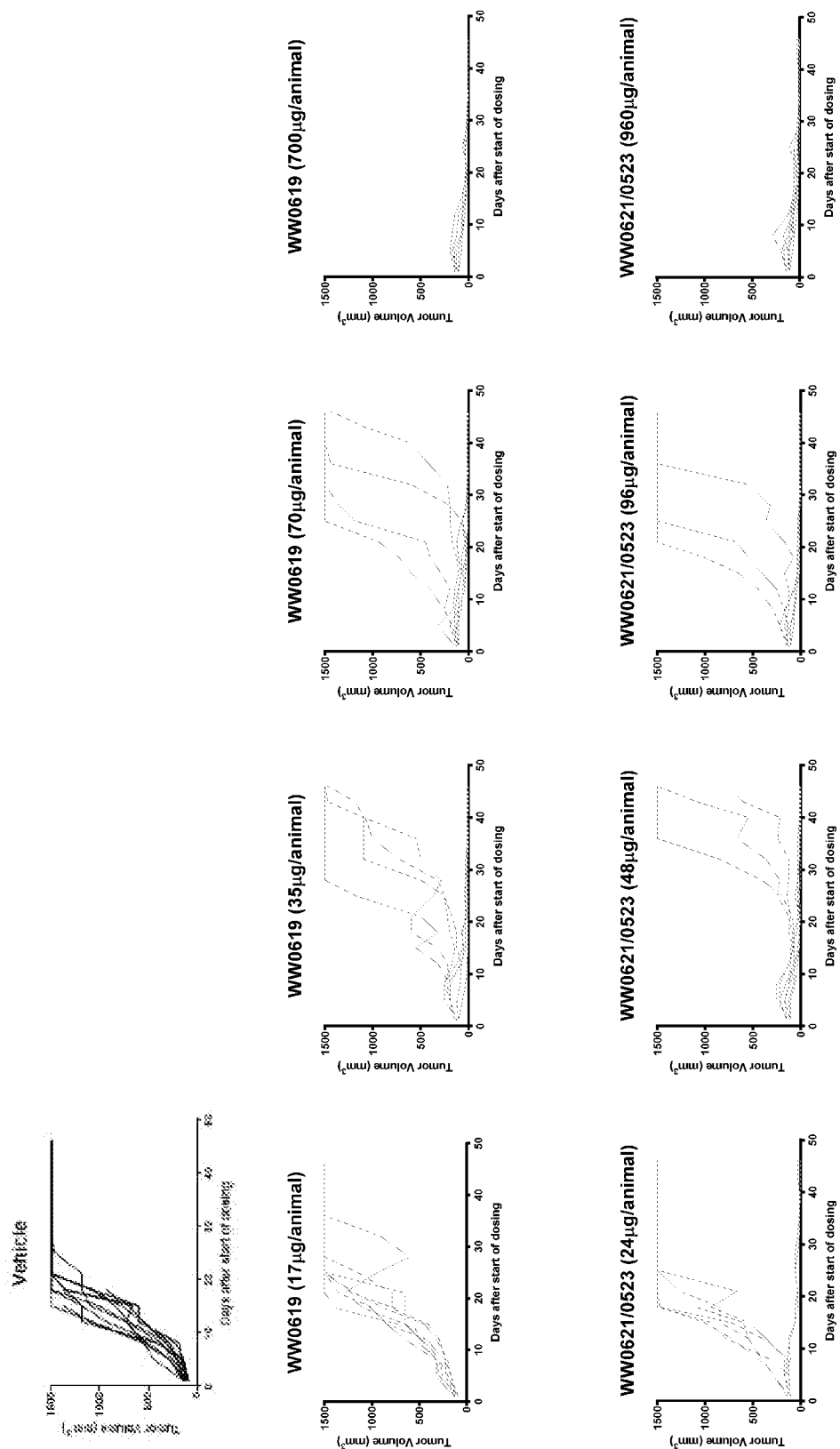


FIG. 6A

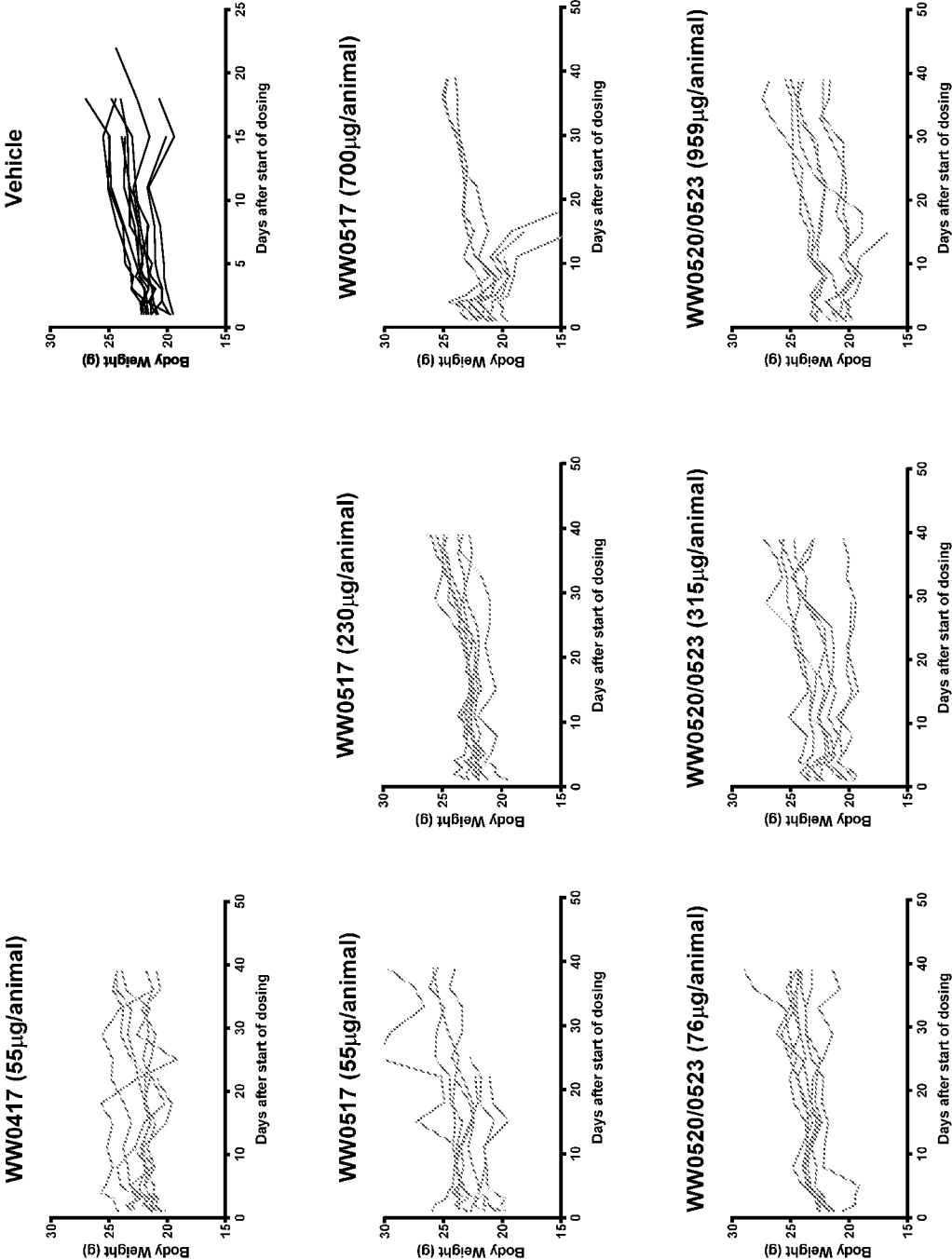


FIG. 6B

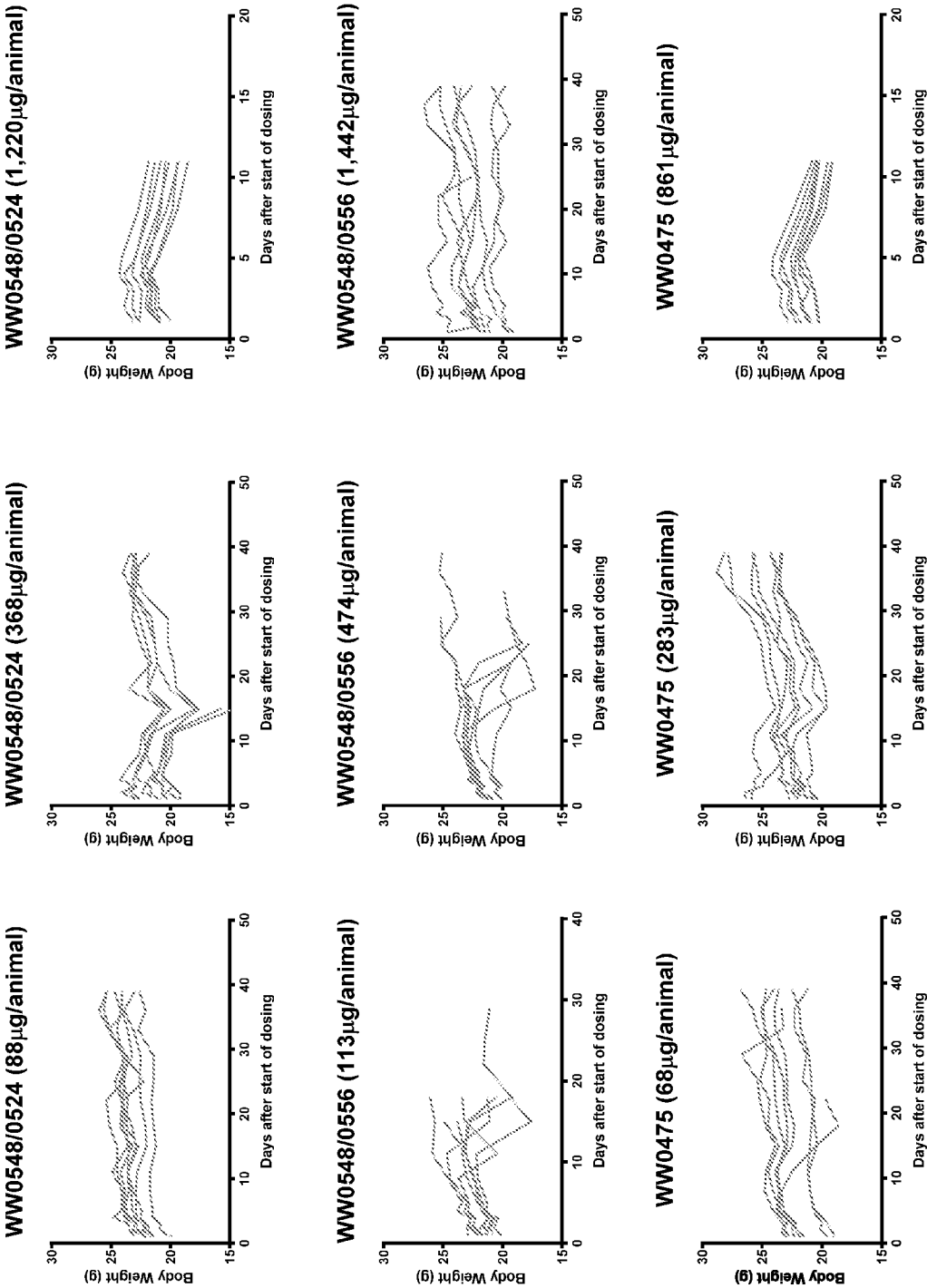


FIG. 6C

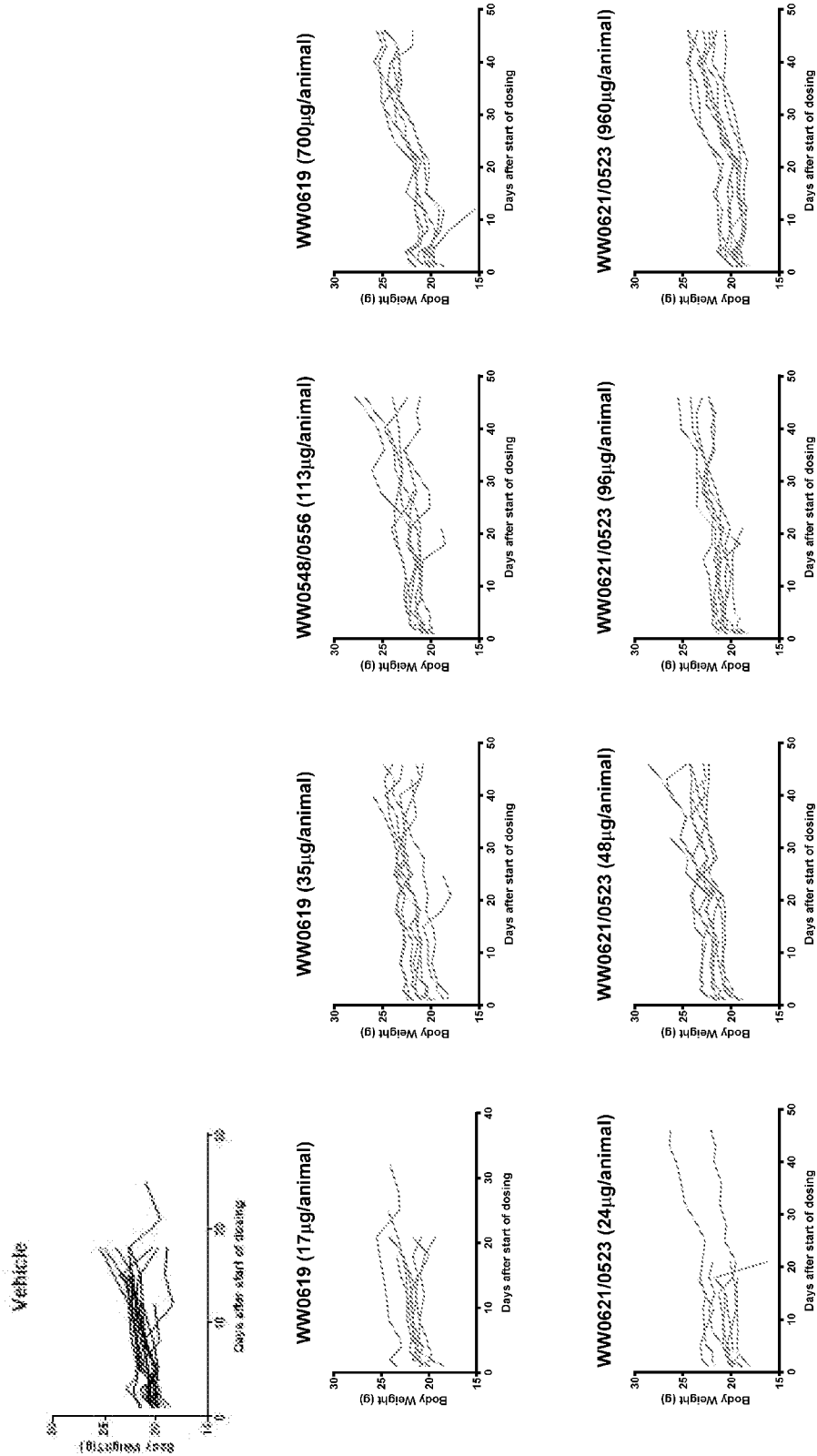


FIG. 7A

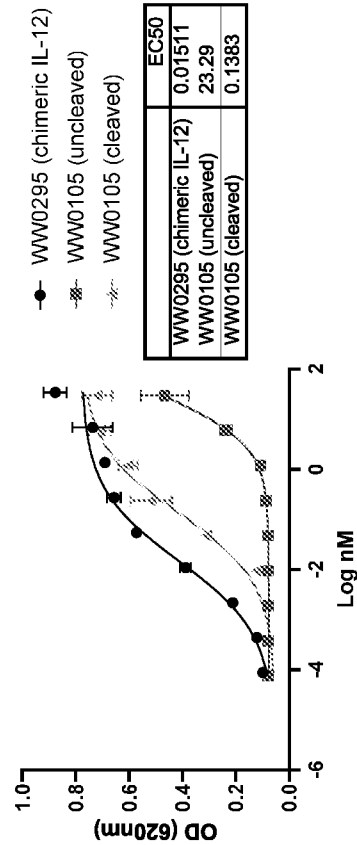


FIG. 7B

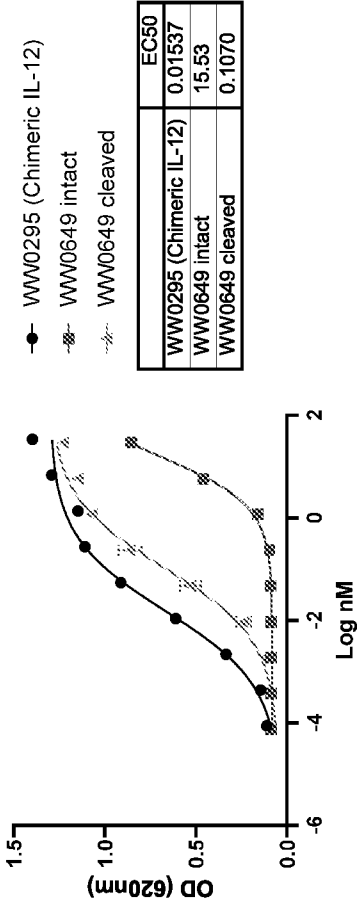


FIG. 7C

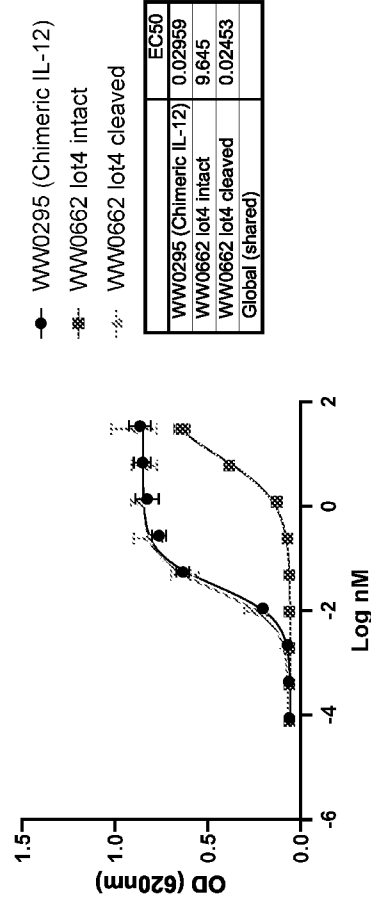


FIG. 7D

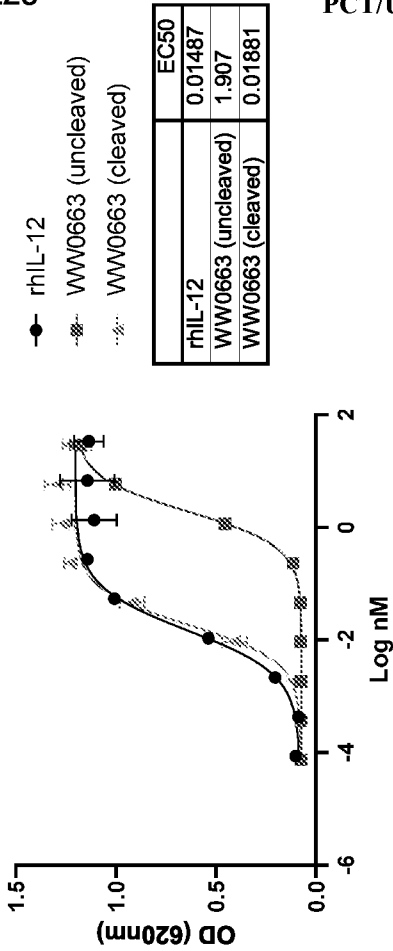


FIG. 7E

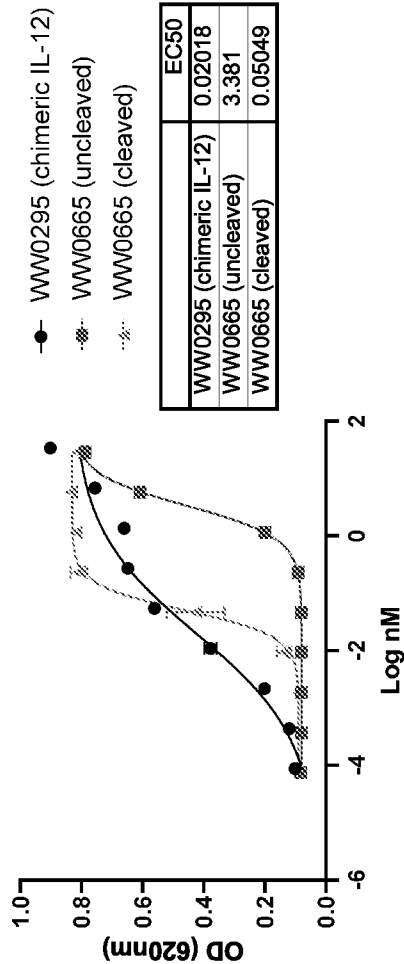


FIG. 7F

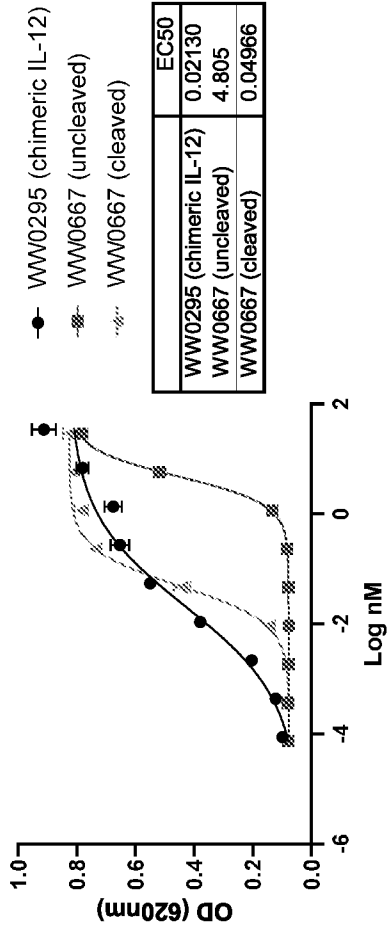


FIG. 7G

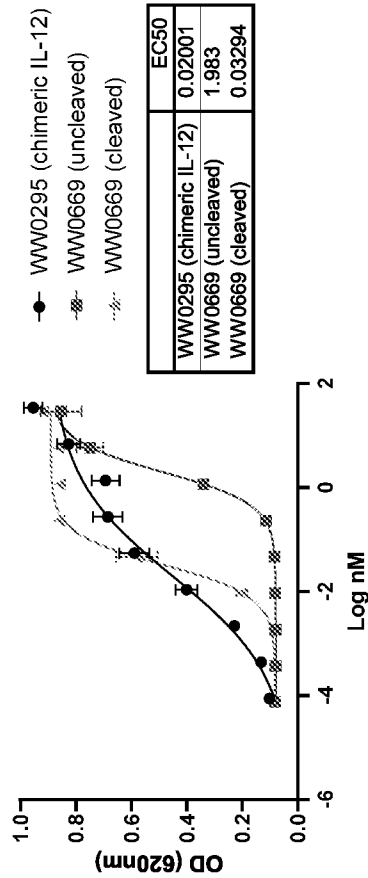


FIG. 7I

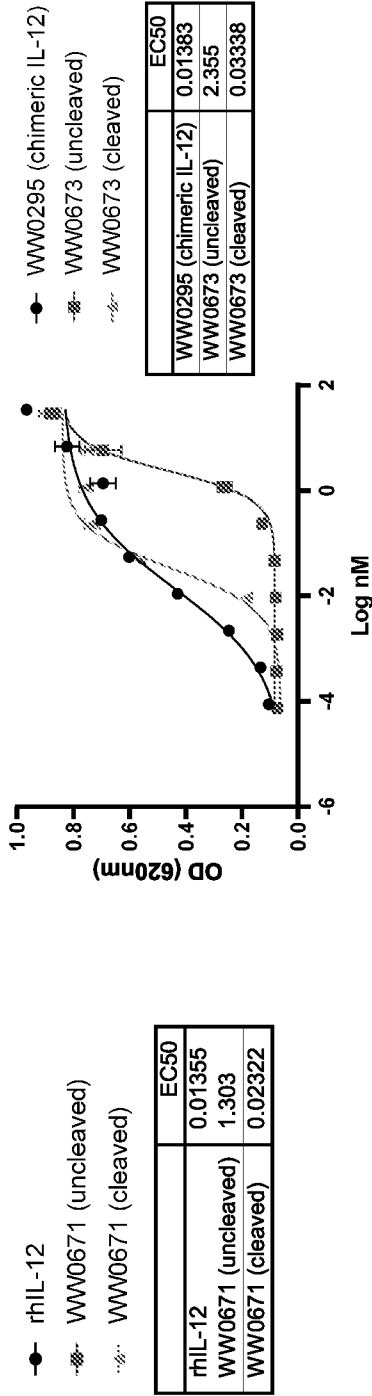


FIG. 7H

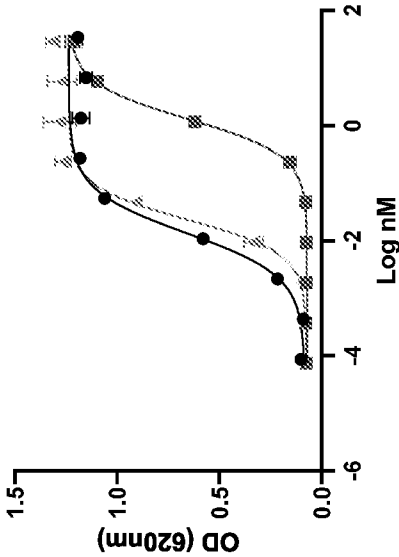


FIG. 7J

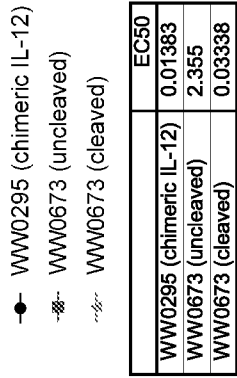
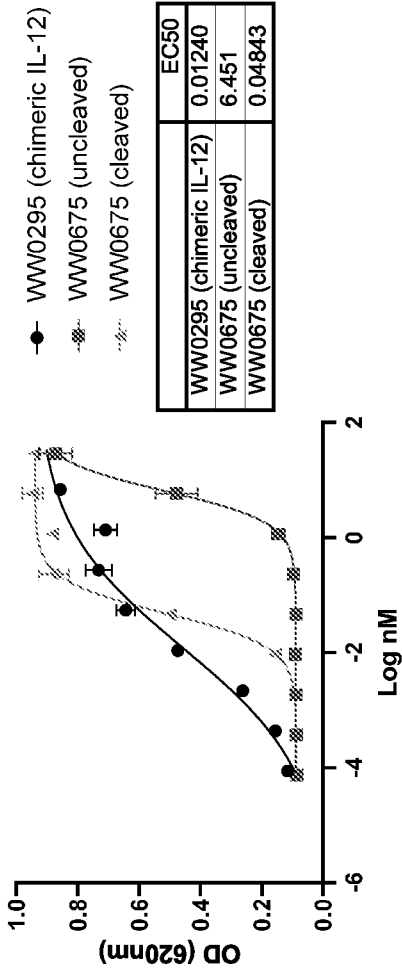


FIG. 7L

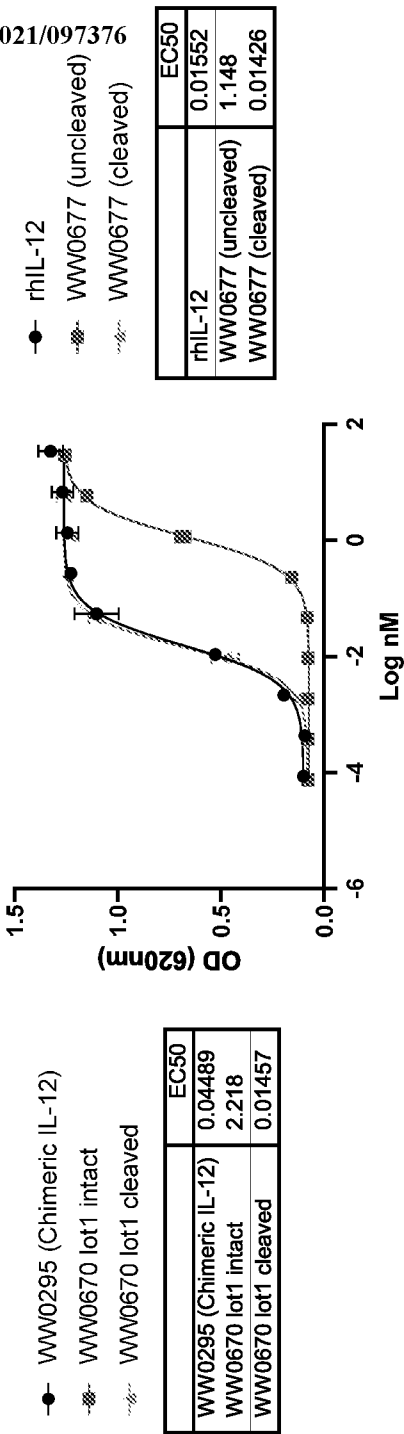


FIG. 7K

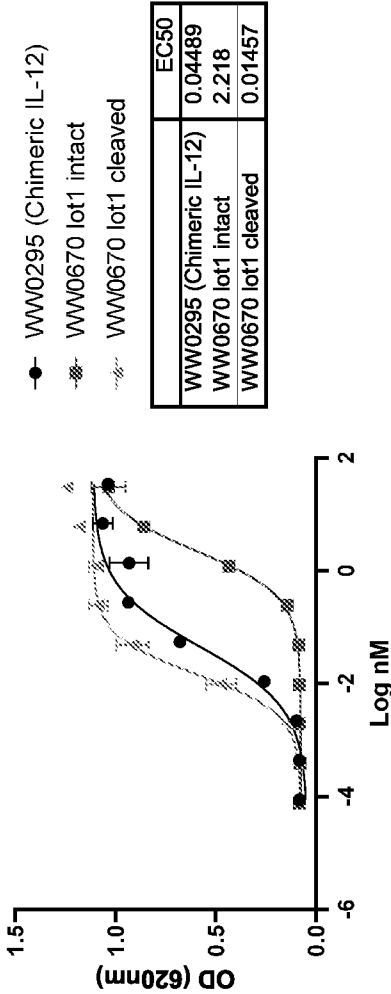


FIG. 7M

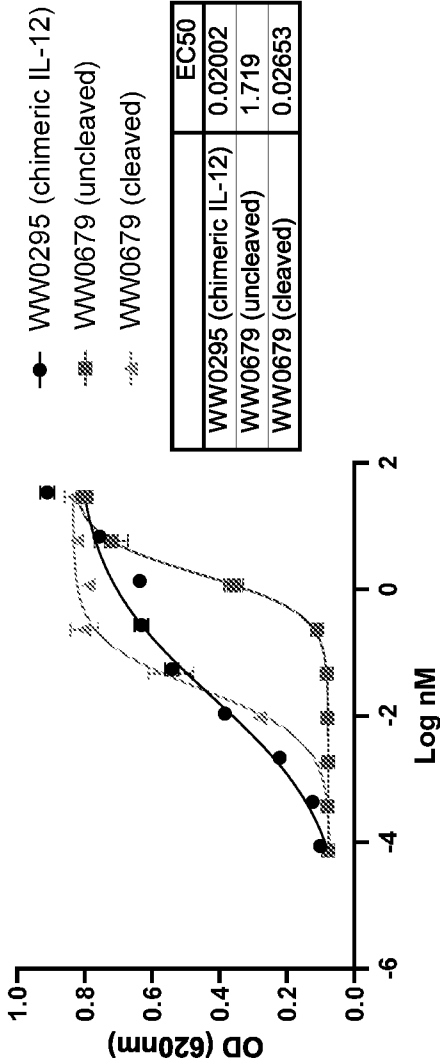


FIG. 7N

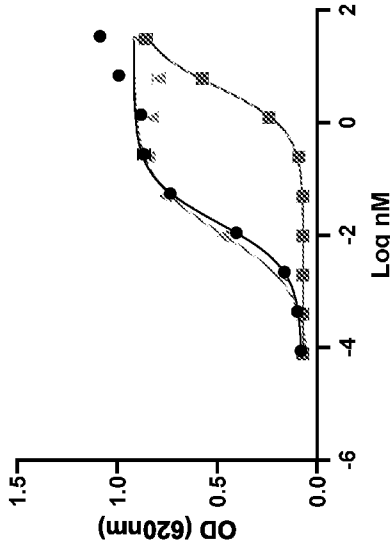


FIG. 7O

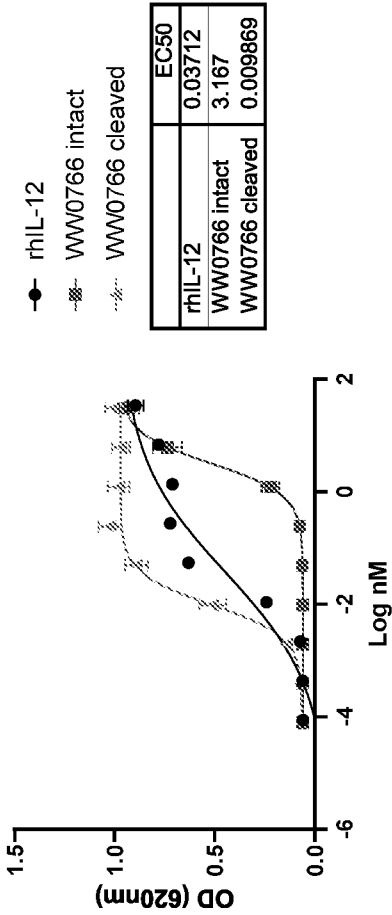


FIG. 7P

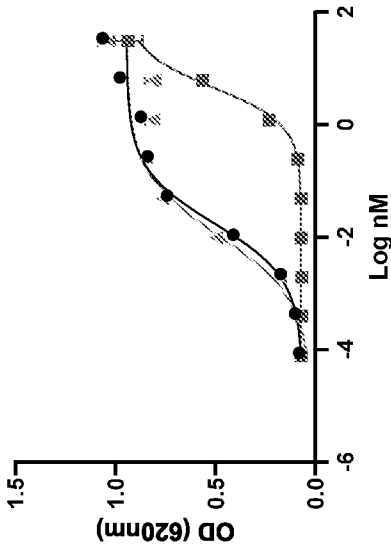


FIG. 7Q

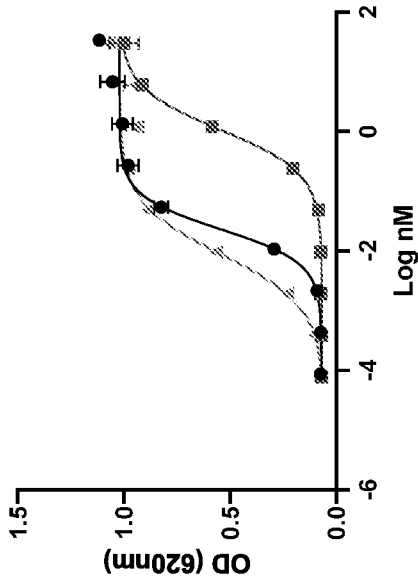


FIG. 8A

WW0663

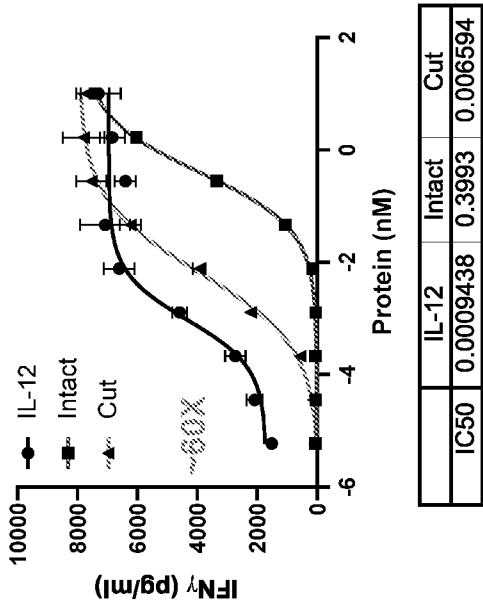


FIG. 8B

WW0671

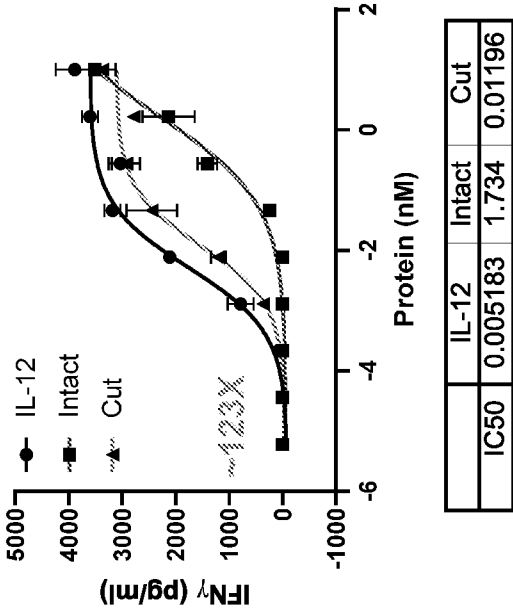


FIG. 8C

WW0766

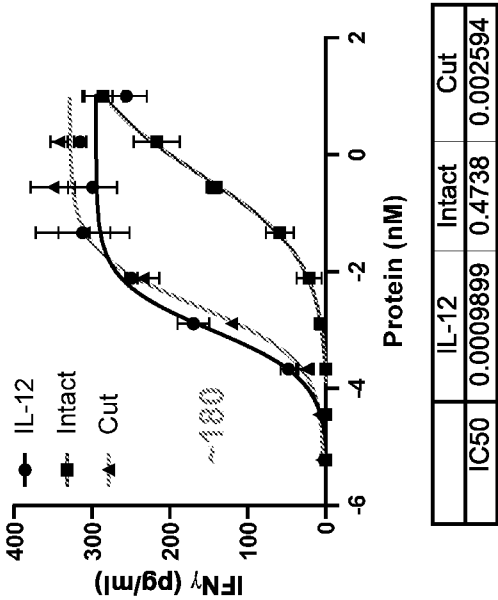


FIG. 8D

WW0765

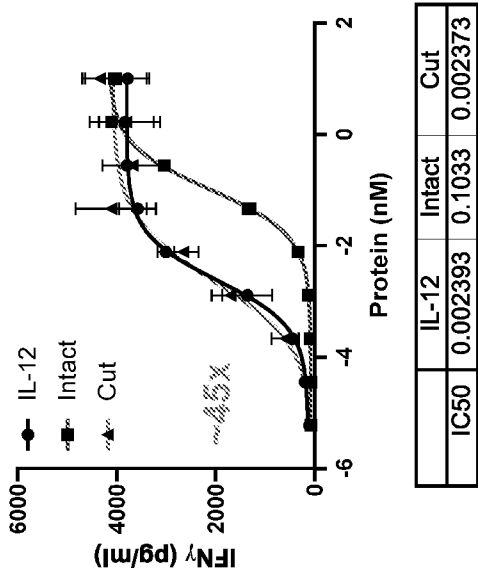


FIG. 9A

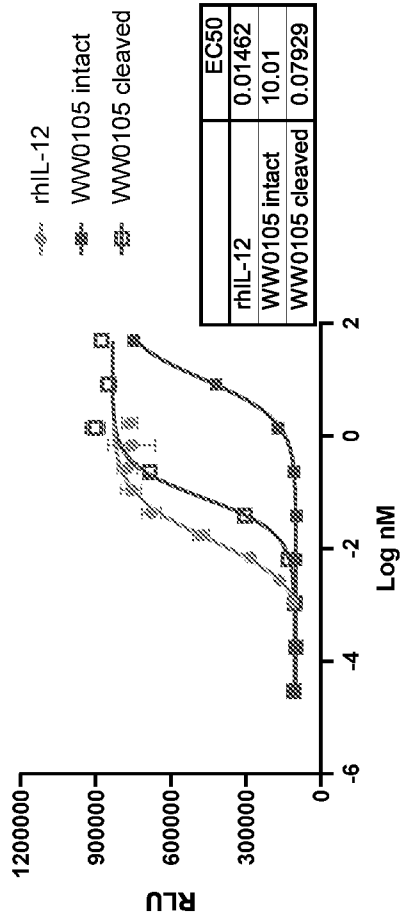


FIG. 9B

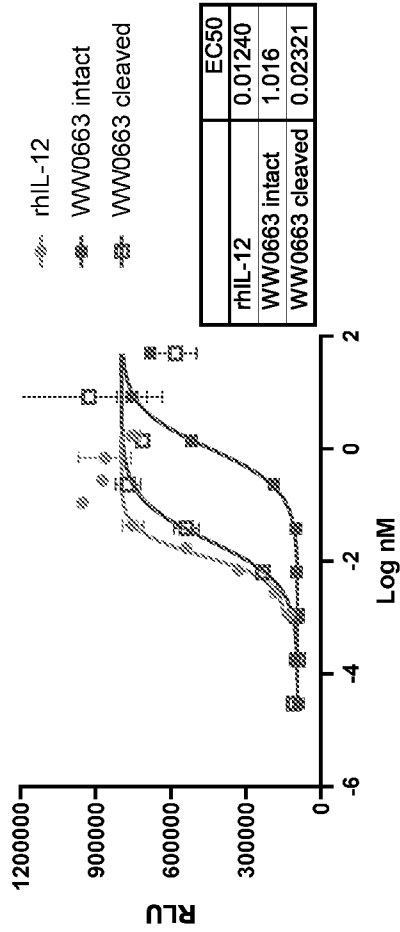


FIG. 9C

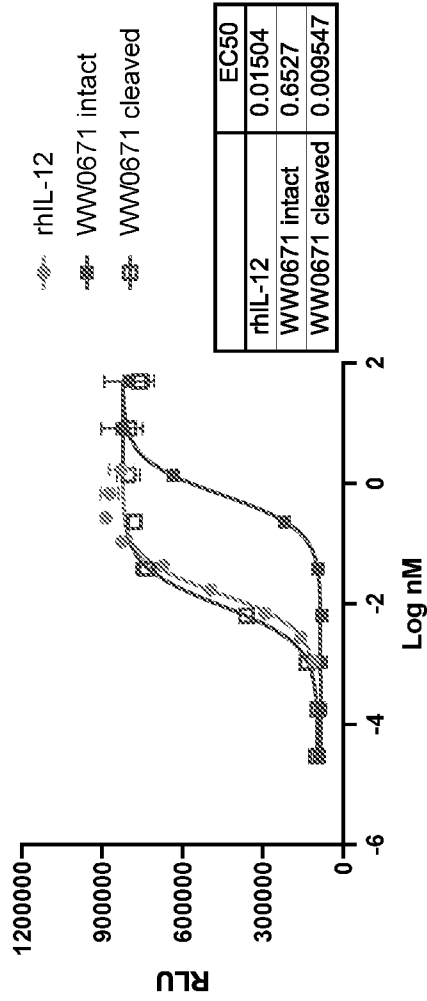


FIG. 10A

B16-Blue mIFNa/b Reporter Assay Using WW0609

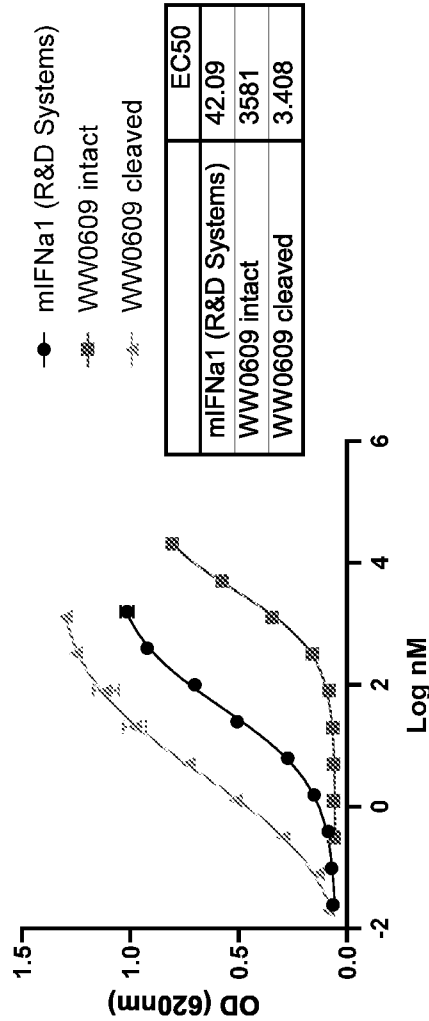


FIG. 10B

B16 Blue mIFNa/b Reporter assay WW0610

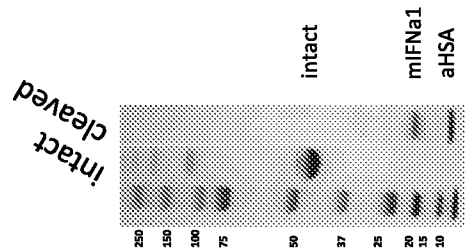
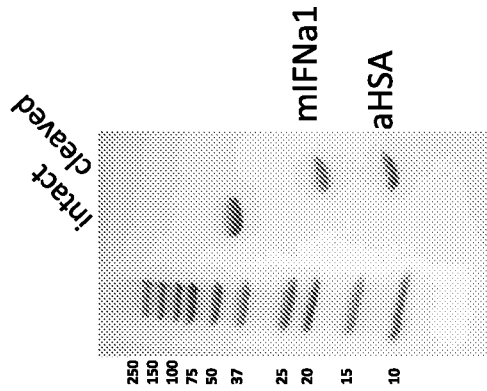
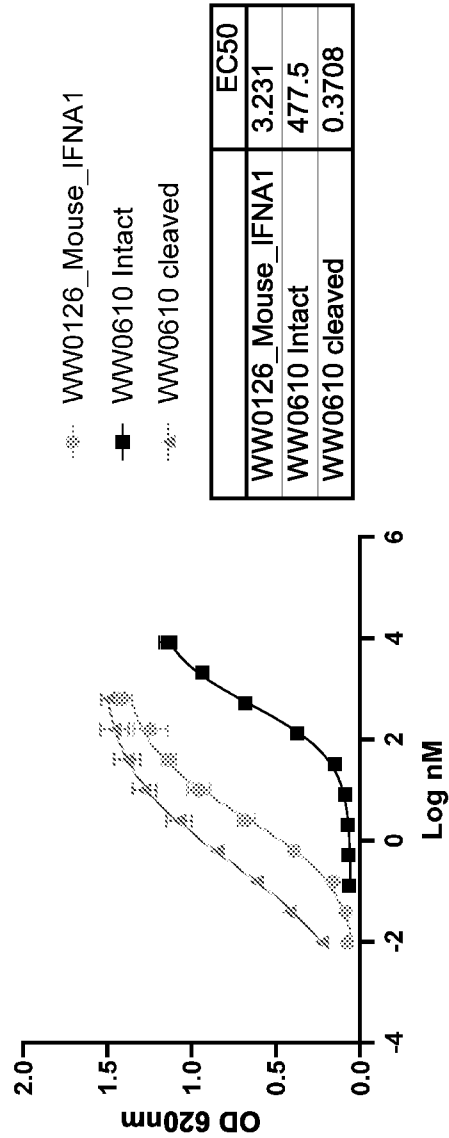


FIG. 10C

B16-Blue mIFNa/b Reporter Assay Using WW0643

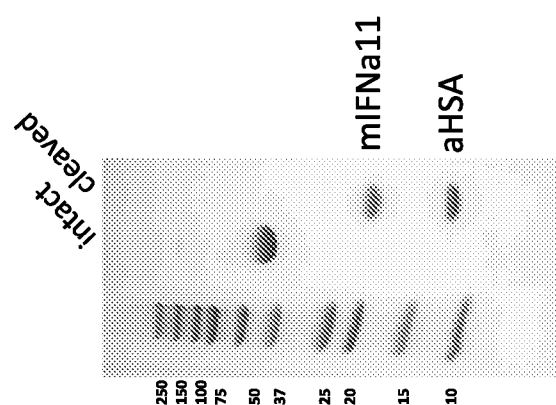
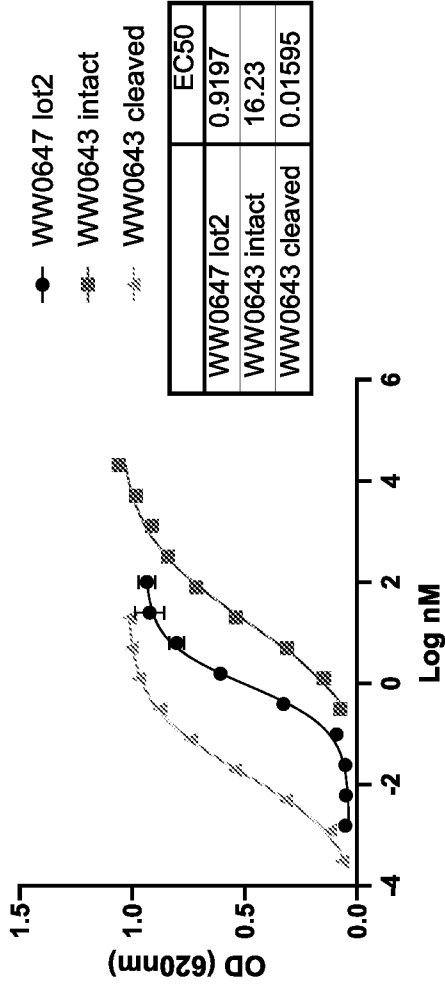


FIG. 10D

B16-Blue mIFNa/b Reporter Assay Using WW0644

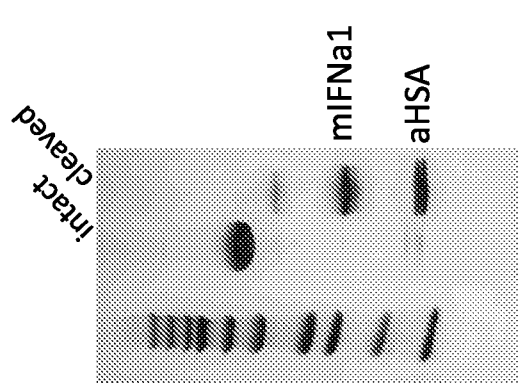
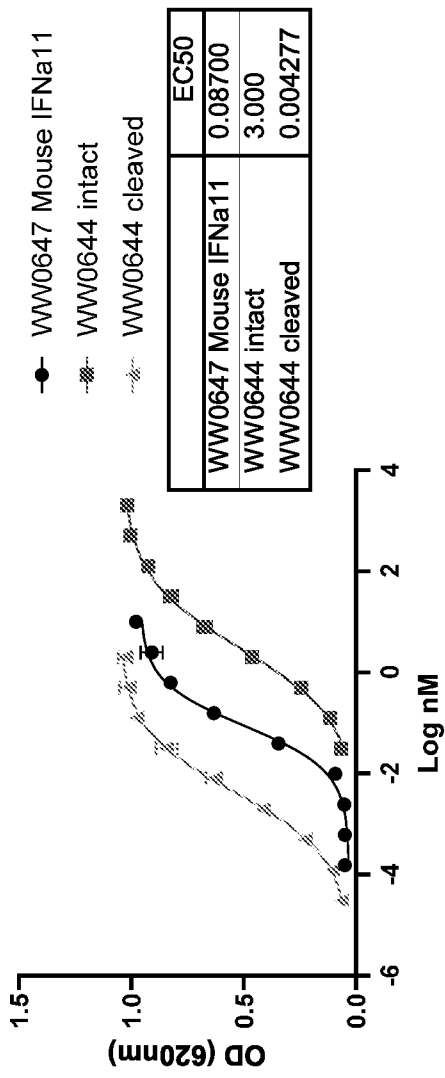


FIG. 10E

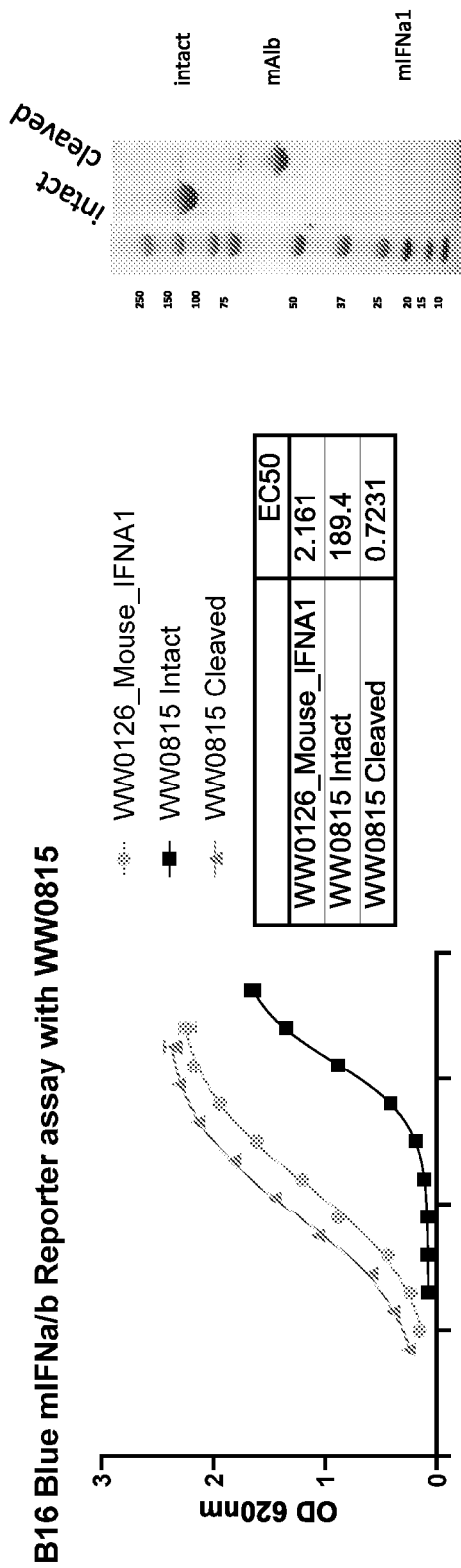


FIG. 10F

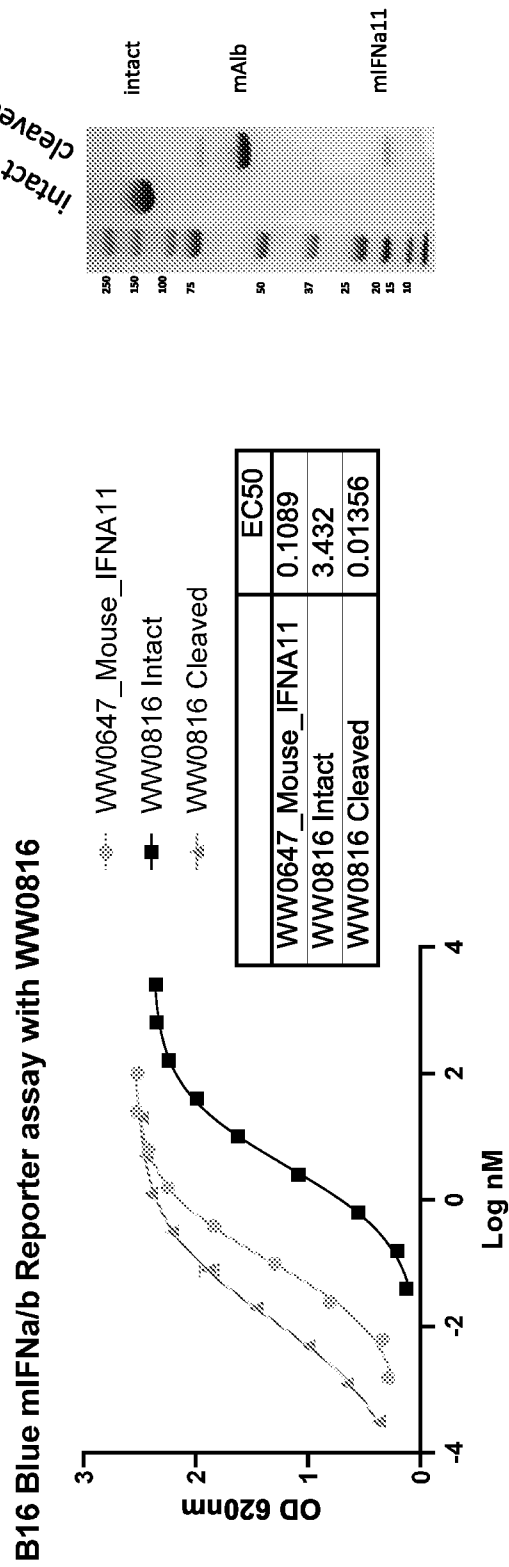


FIG. 10G

B16 mIFNa Reporter Assay for WW0817

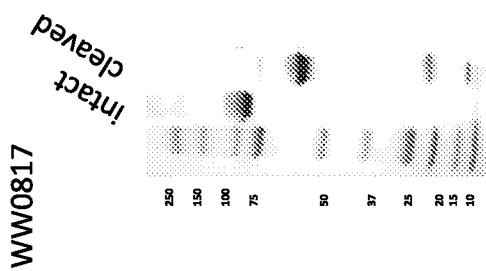
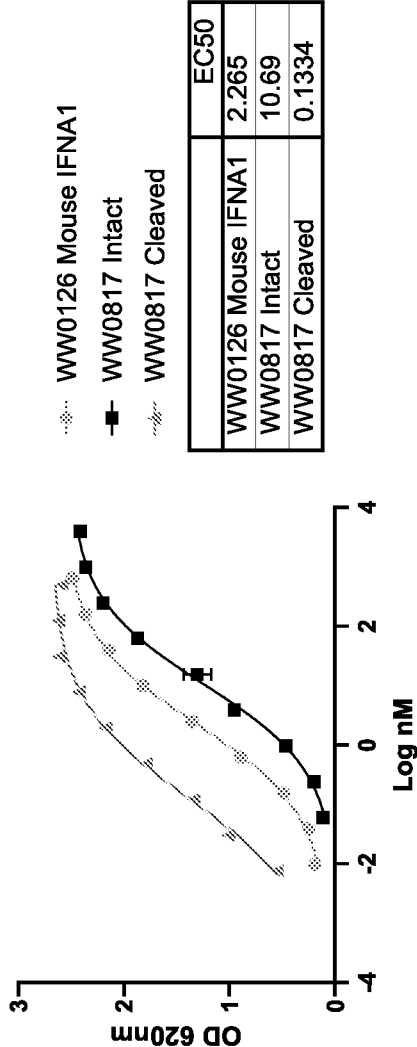


FIG. 10H

B16 mIFNa Reporter Assay for WW0819

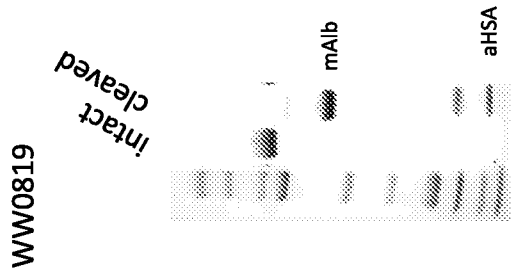
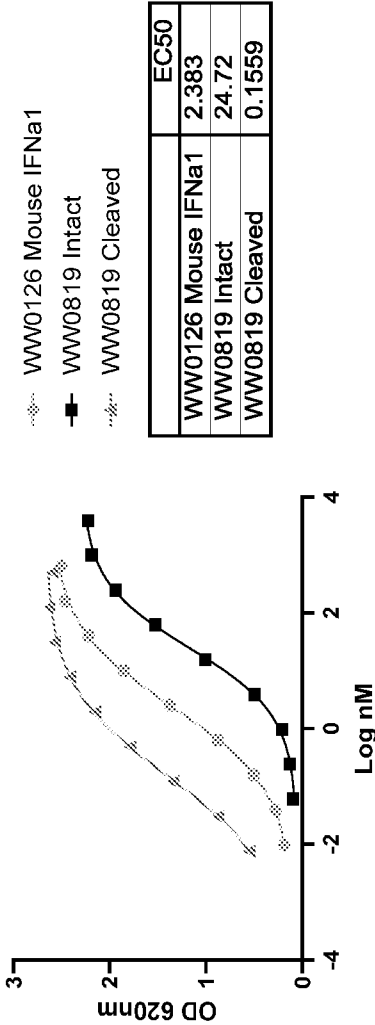


FIG. 10I

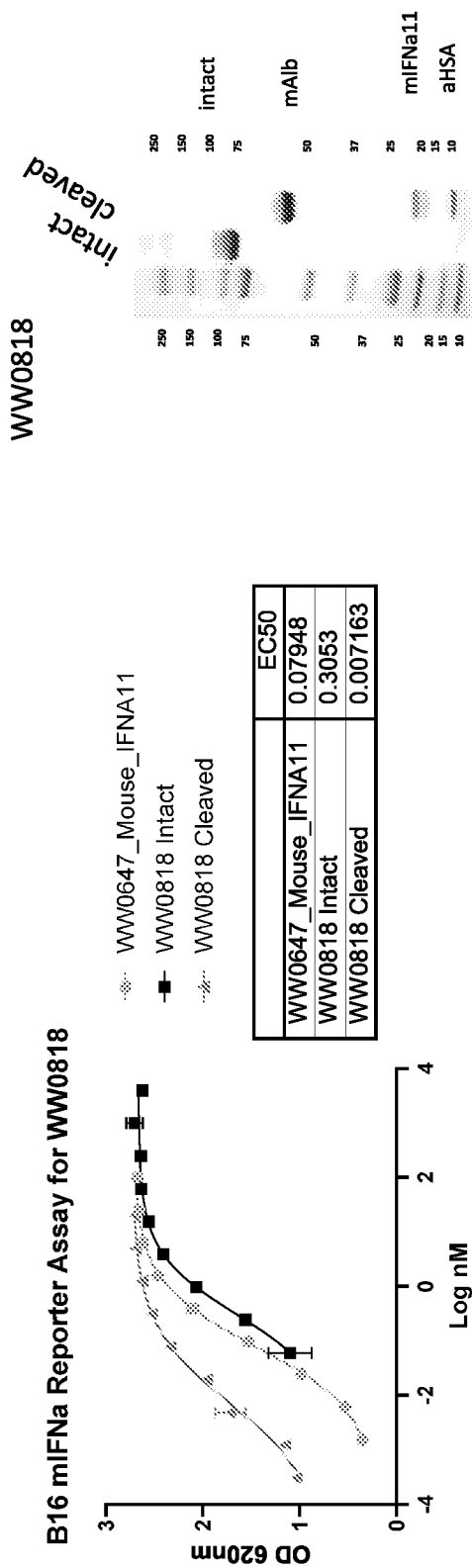


FIG. 10J

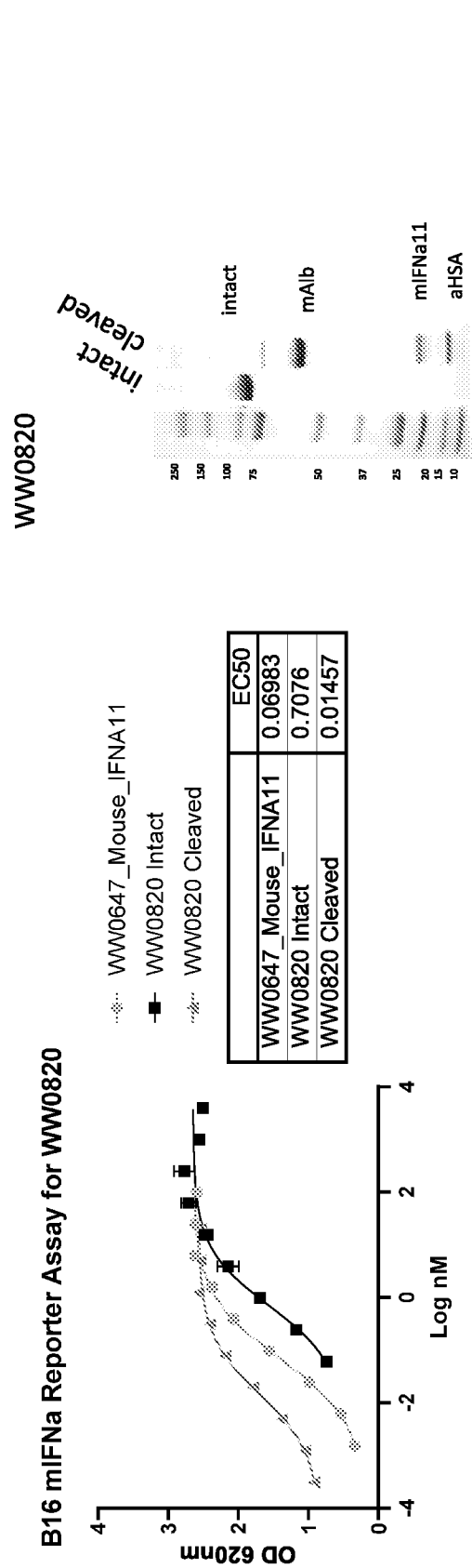


FIG. 11

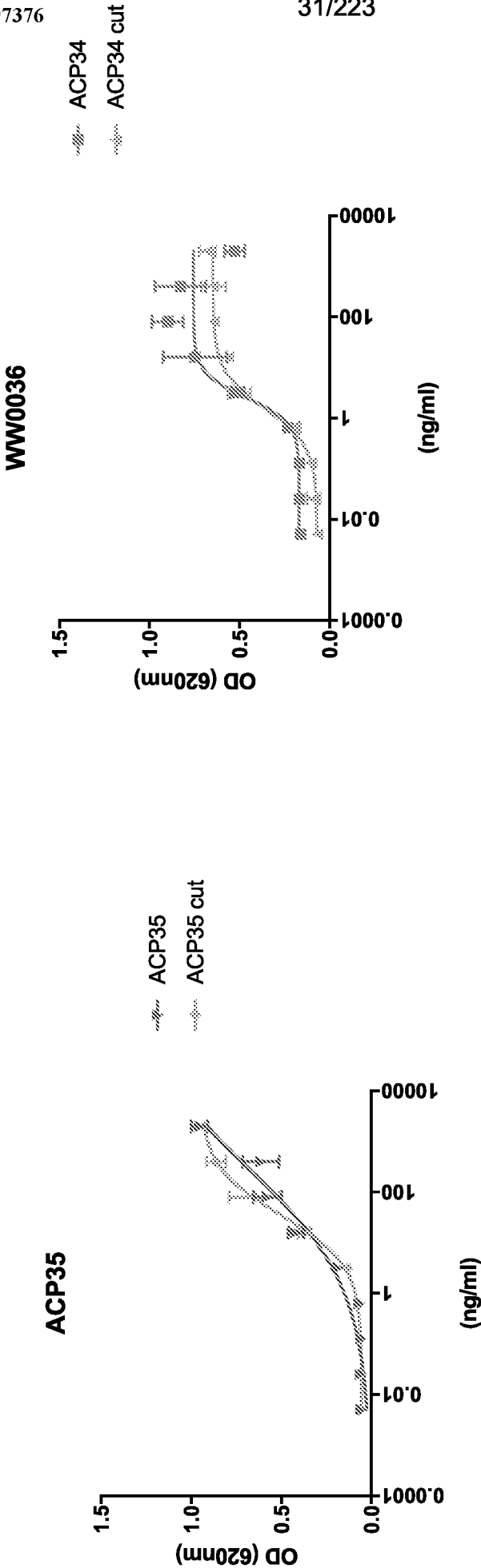


FIG. 11A

FIG. 11B

FIG. 12A-12D:

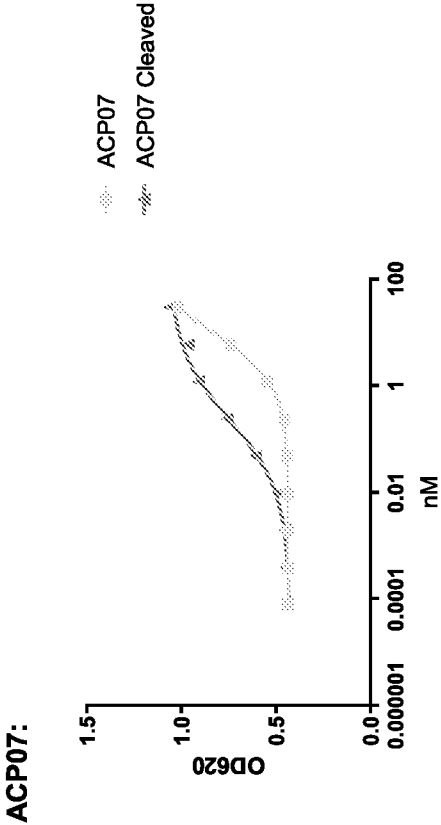


FIG. 12C

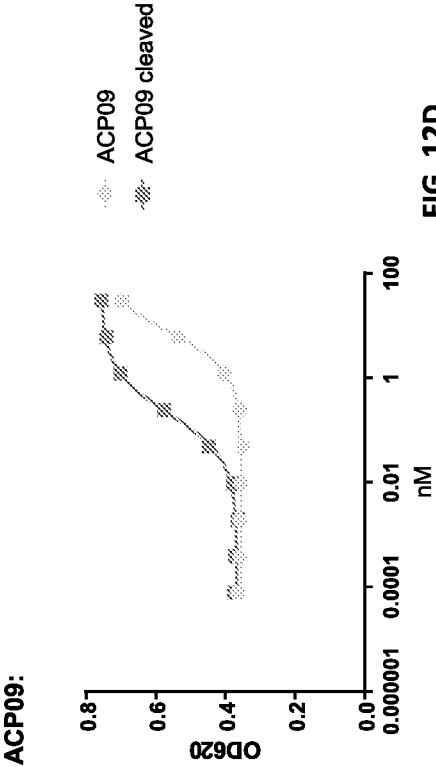


FIG. 12D

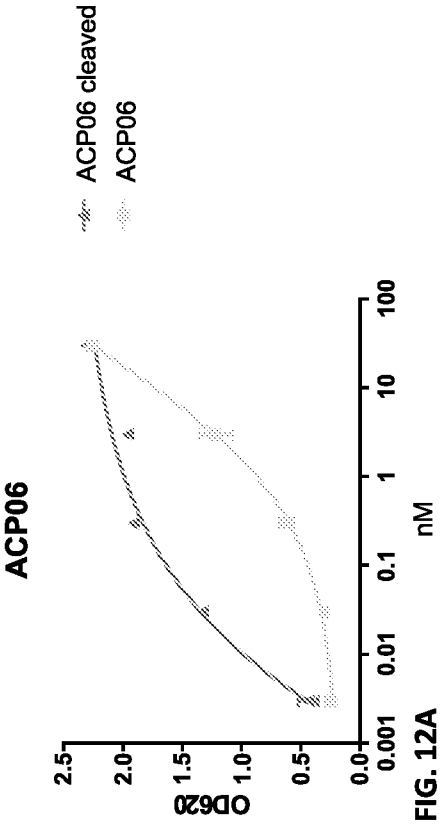


FIG. 12A

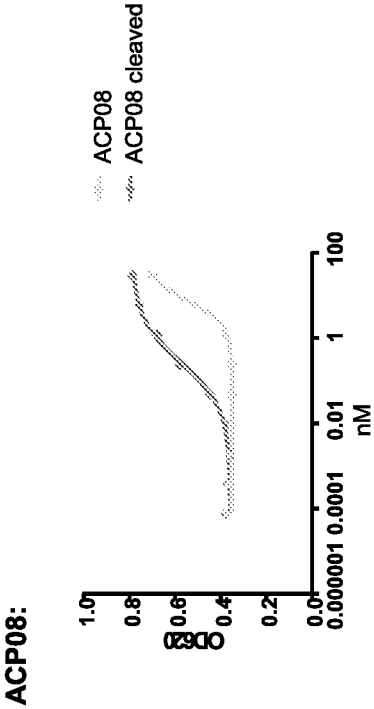


FIG. 12B

FIG. 12E-12F:

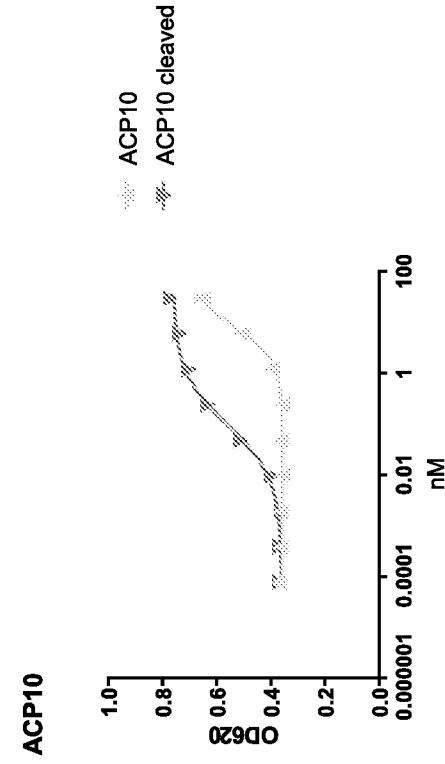


FIG. 12E

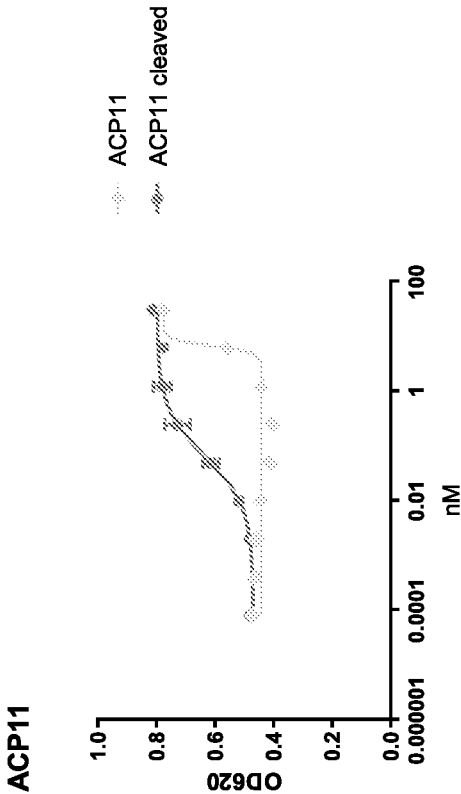


FIG. 12F

FIG. 14

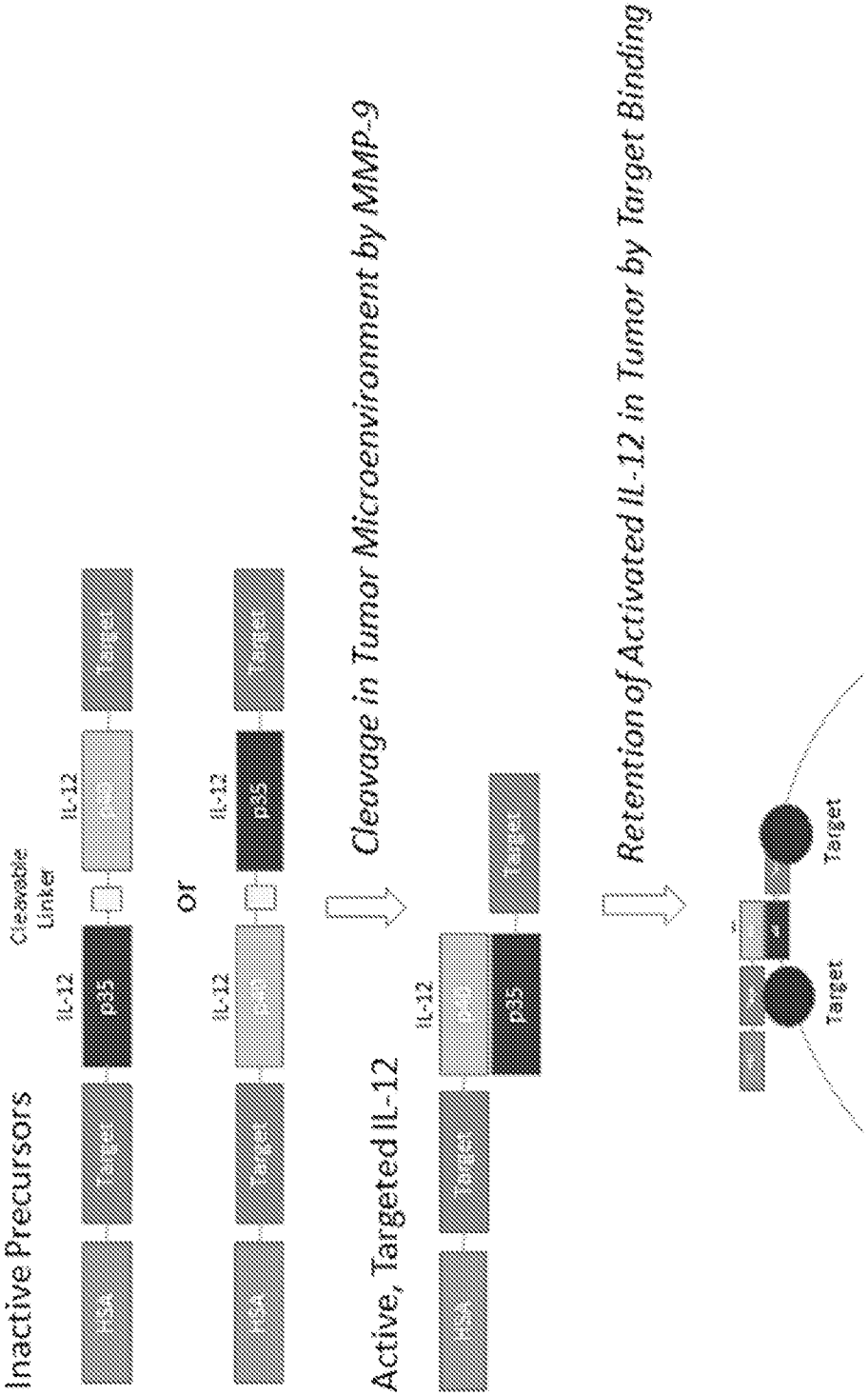


FIG. 15A

ACP36 - HSA

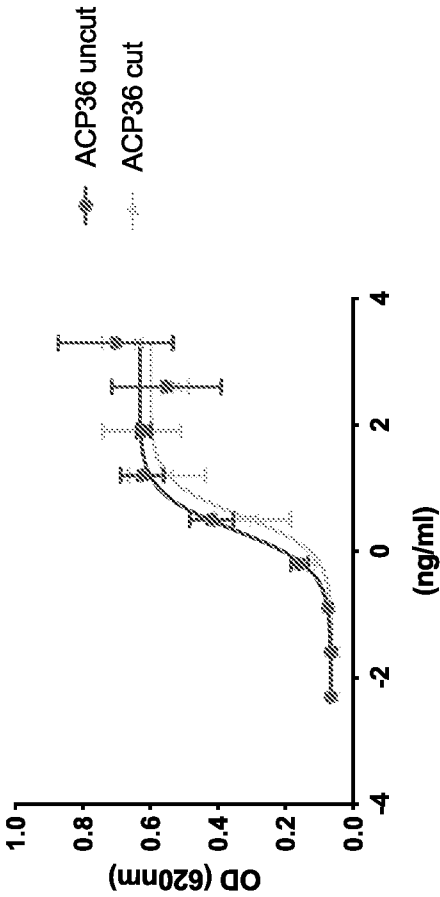


FIG. 15C

ACP37 - HSA

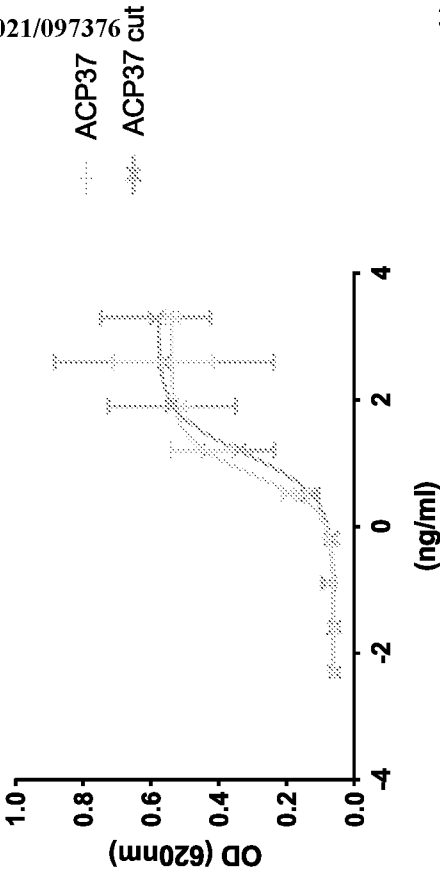


FIG. 15B

ACP36 + HSA

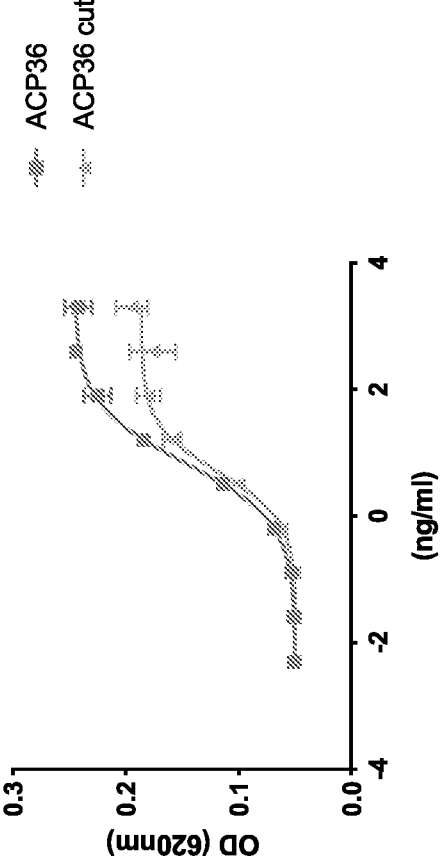


FIG. 15D

ACP37 + HSA

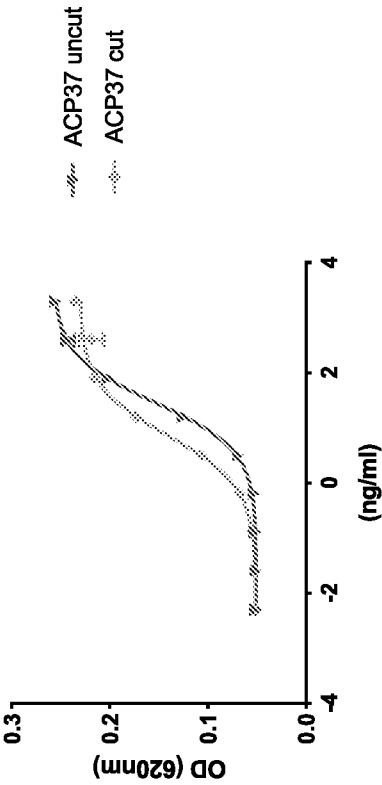


FIG. 16A-16F:

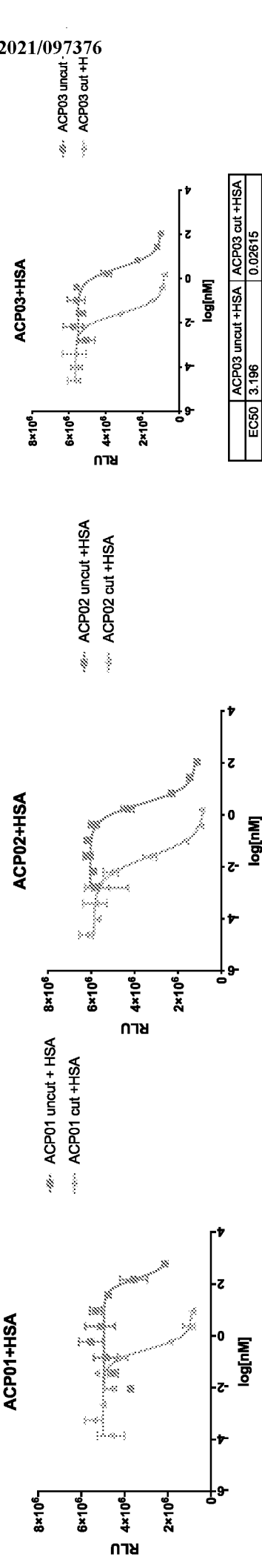


FIG. 16A

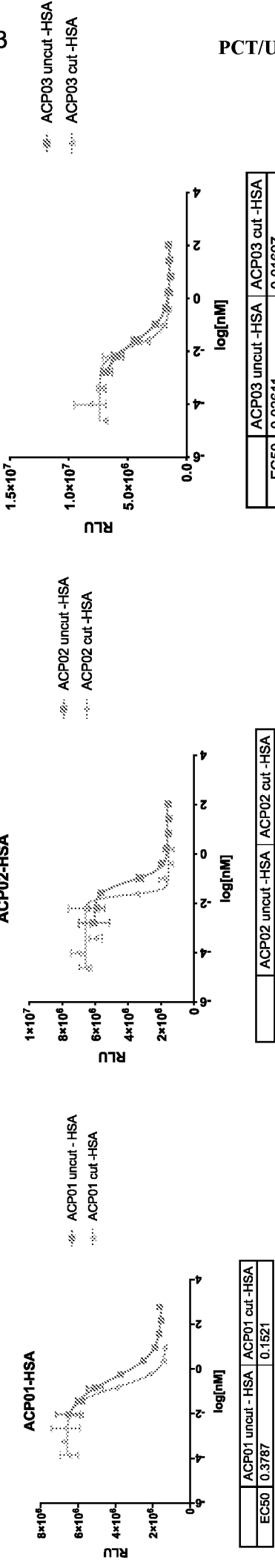


FIG. 16B

FIG. 16D

FIG. 16F

FIG. 17A-17F:

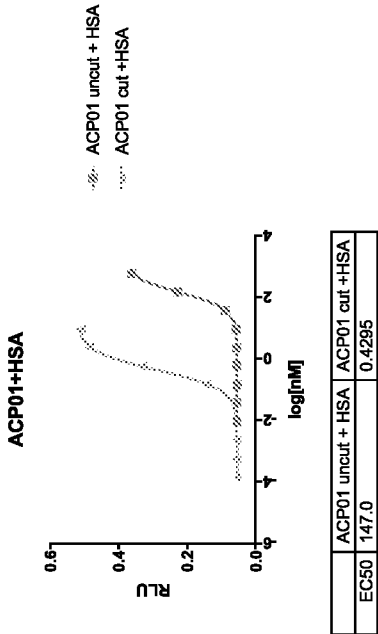
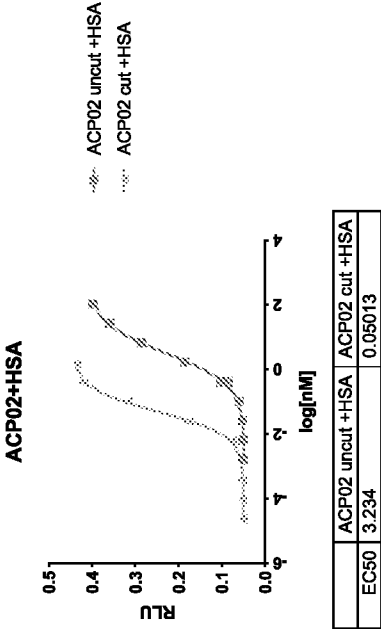
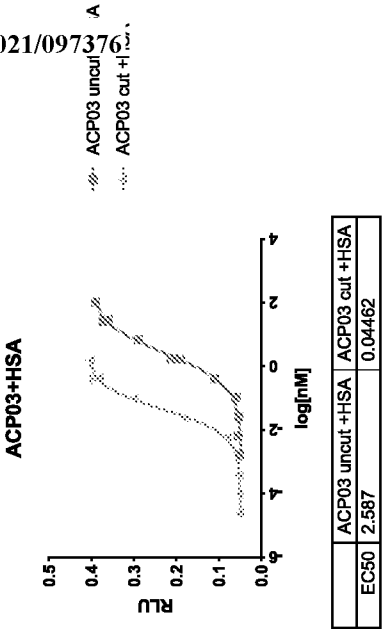


FIG. 17E

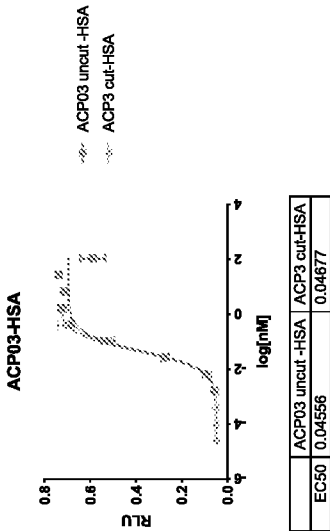


FIG. 17C

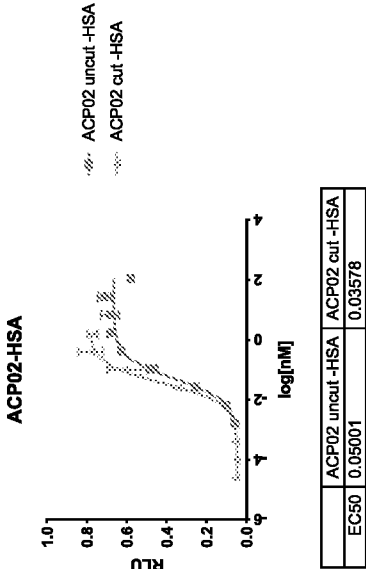


FIG. 17A

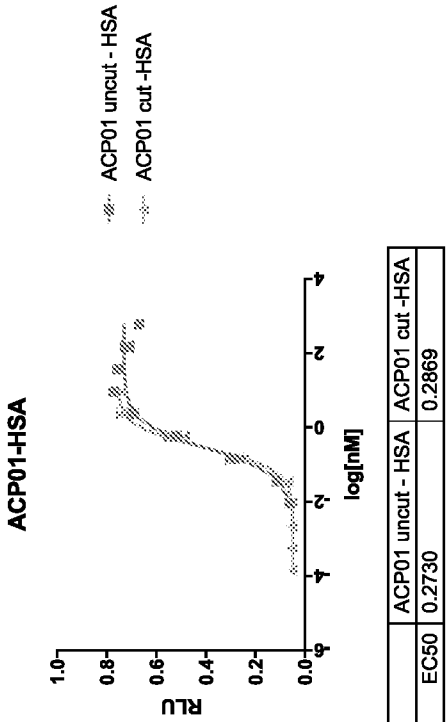


FIG. 17F

FIG. 17D

FIG. 17B

FIG. 18:

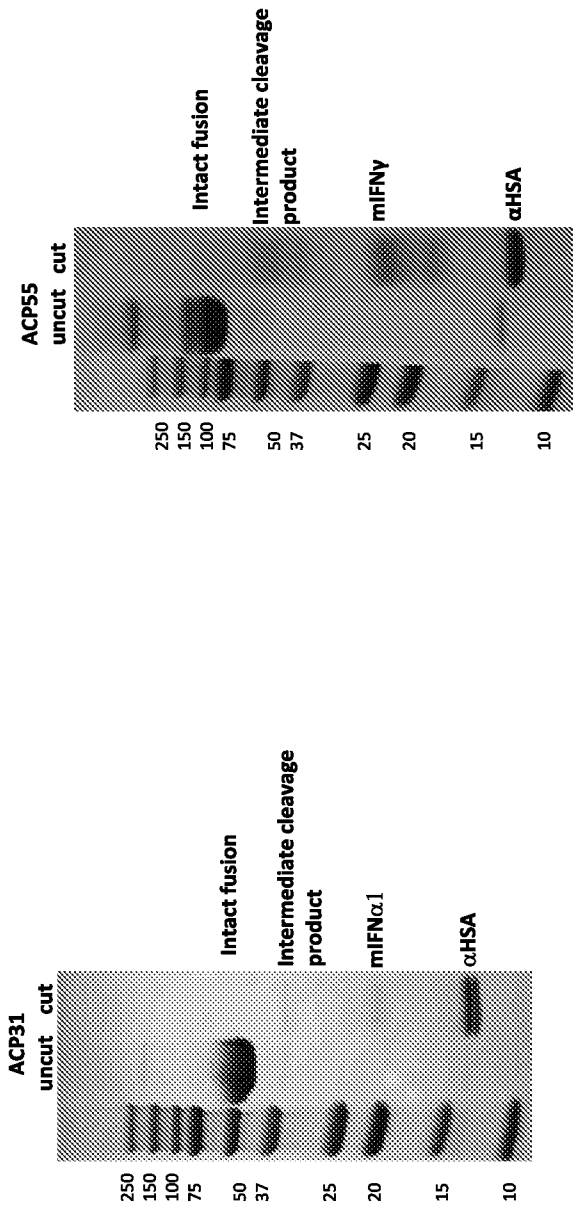


Fig. 18A

Fig. 18B

FIG 19.

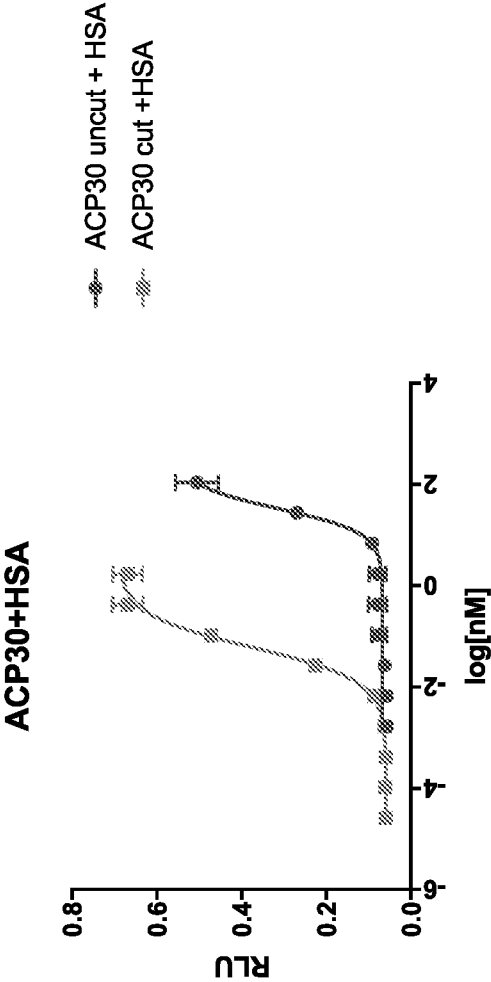


FIG. 19A

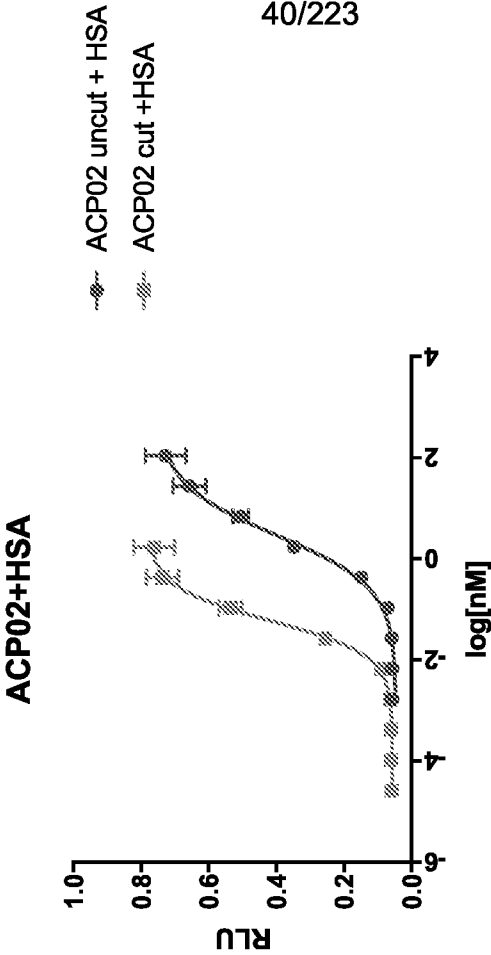


FIG. 19B

FIG. 20

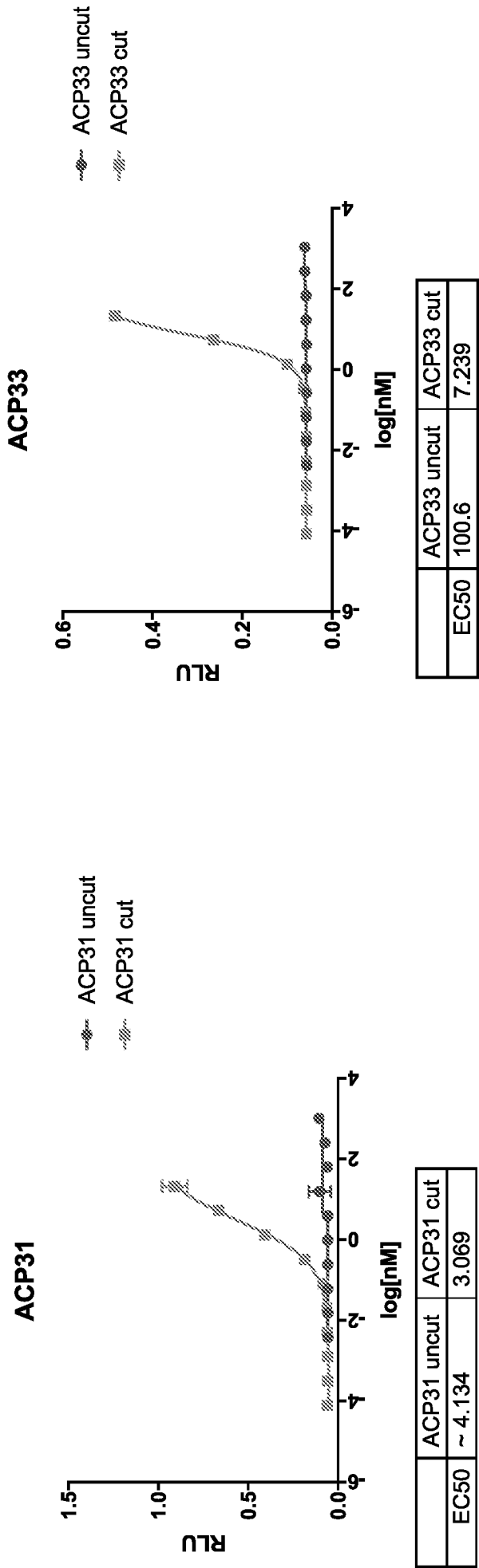


FIG. 20A

FIG. 20B

FIG. 21:

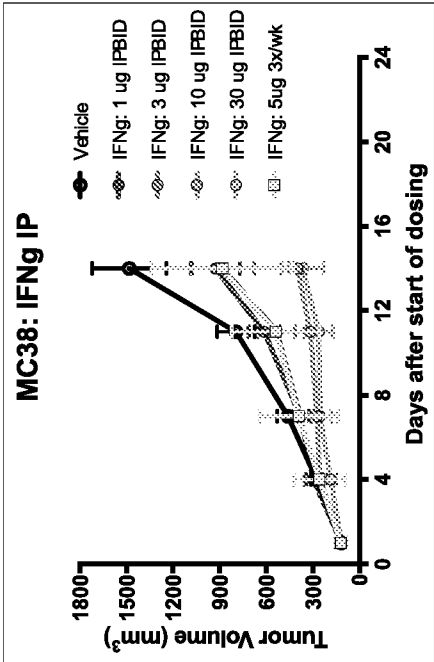


FIG. 21A

FIG. 21B

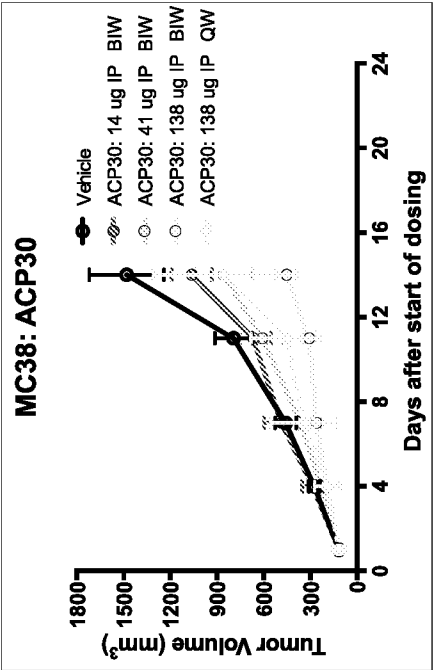
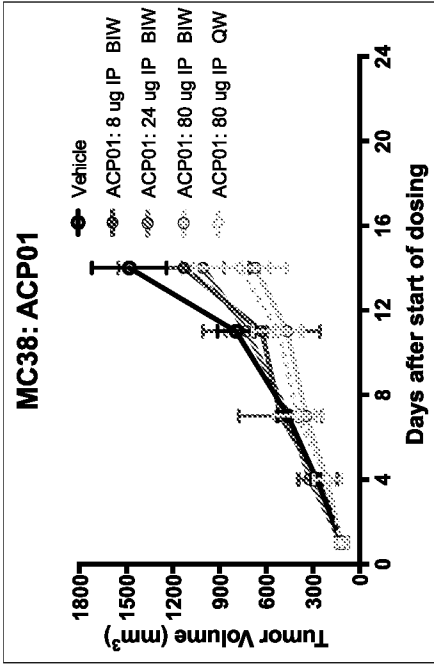


FIG. 21C

FIG. 21D

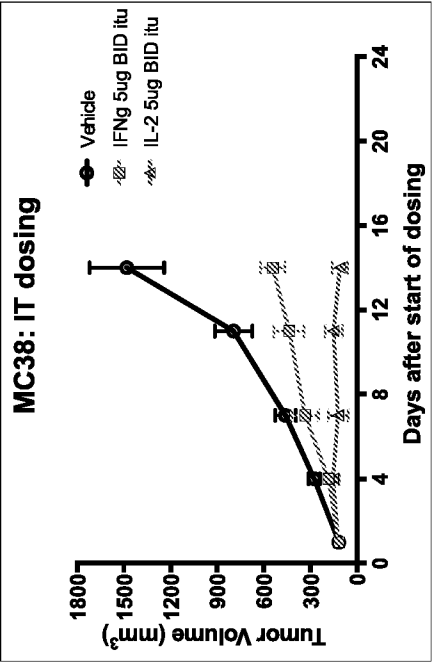


FIG. 22

FIG. 22A

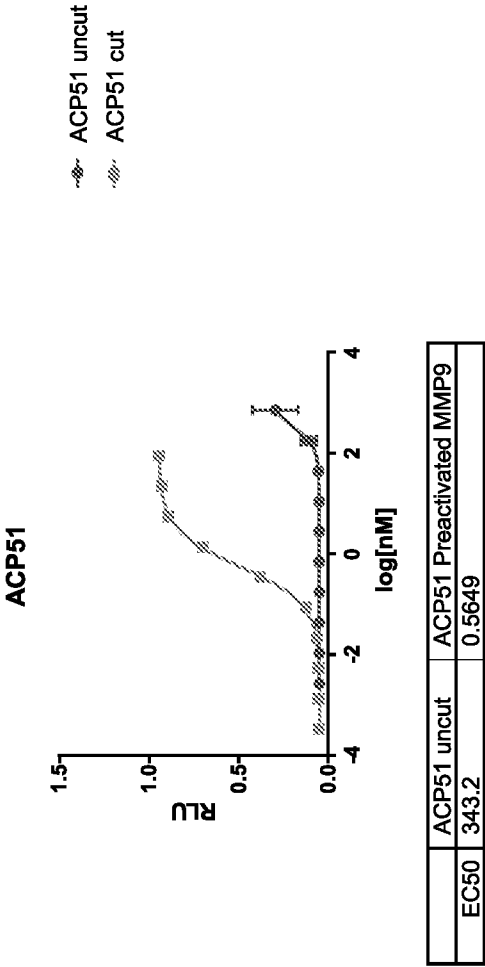


FIG. 22B

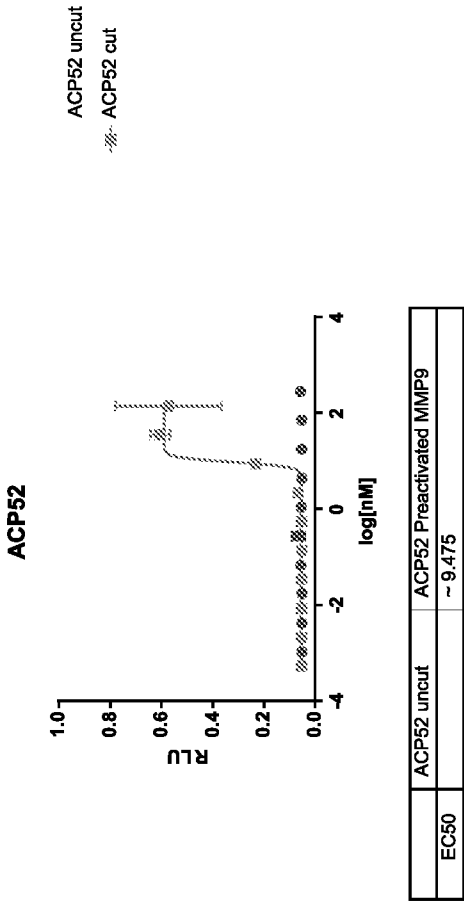


FIG. 23

FIG. 23A

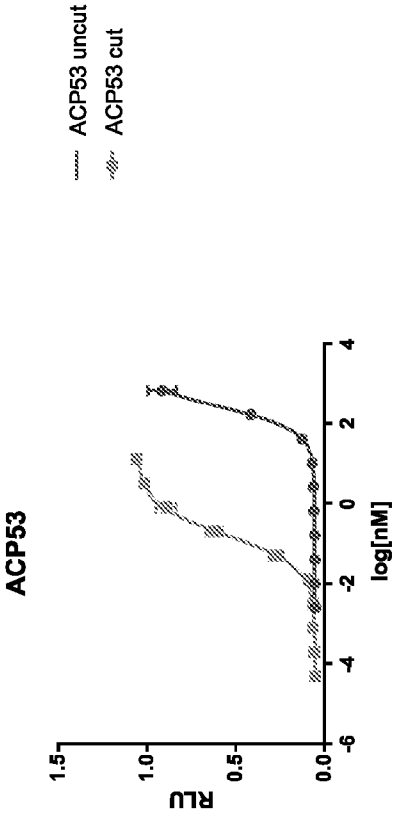


FIG. 23B

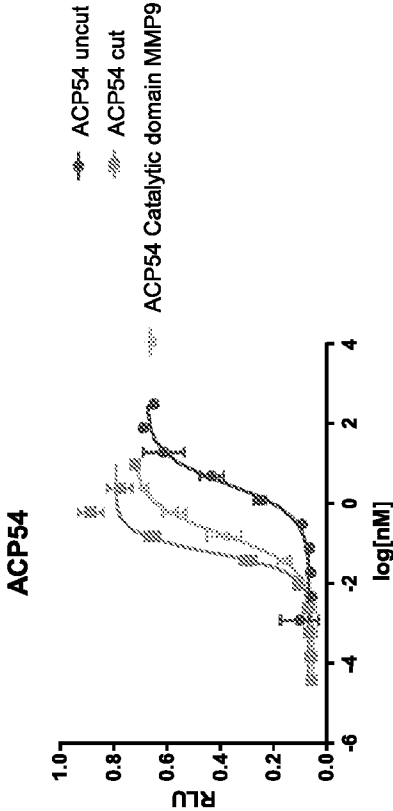


FIG. 24A

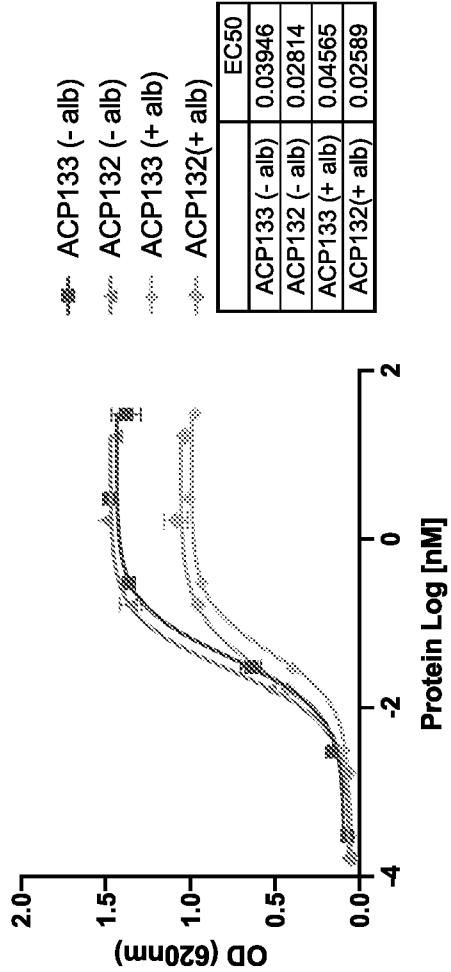


FIG. 24B

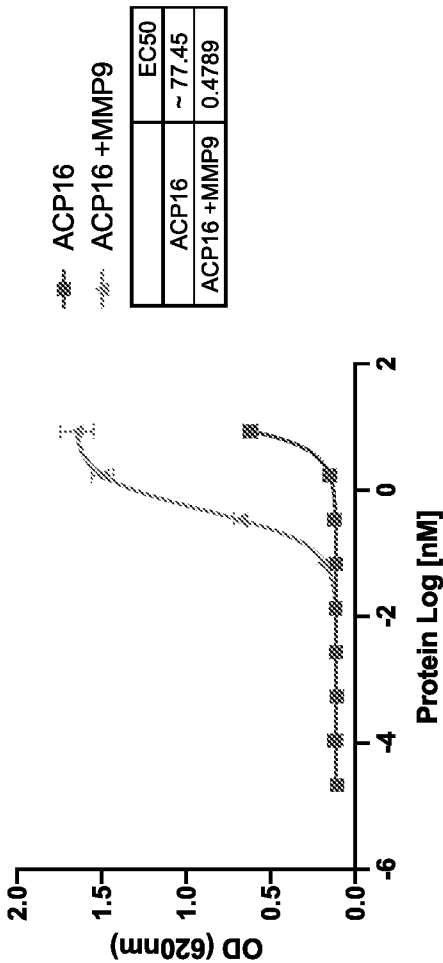
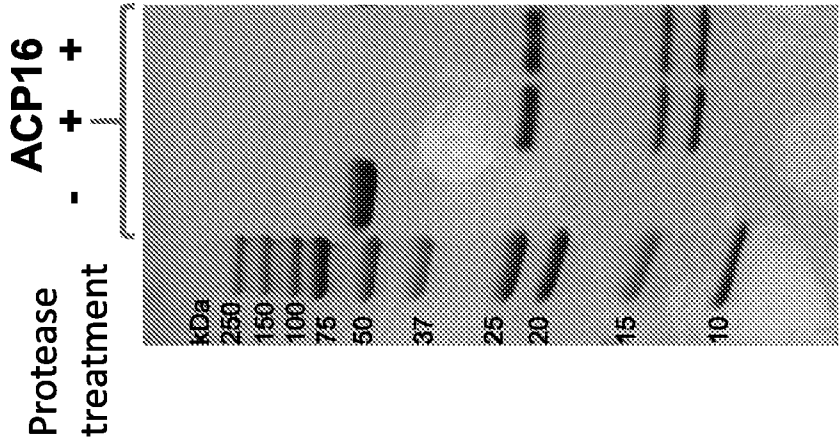


FIG. 24C

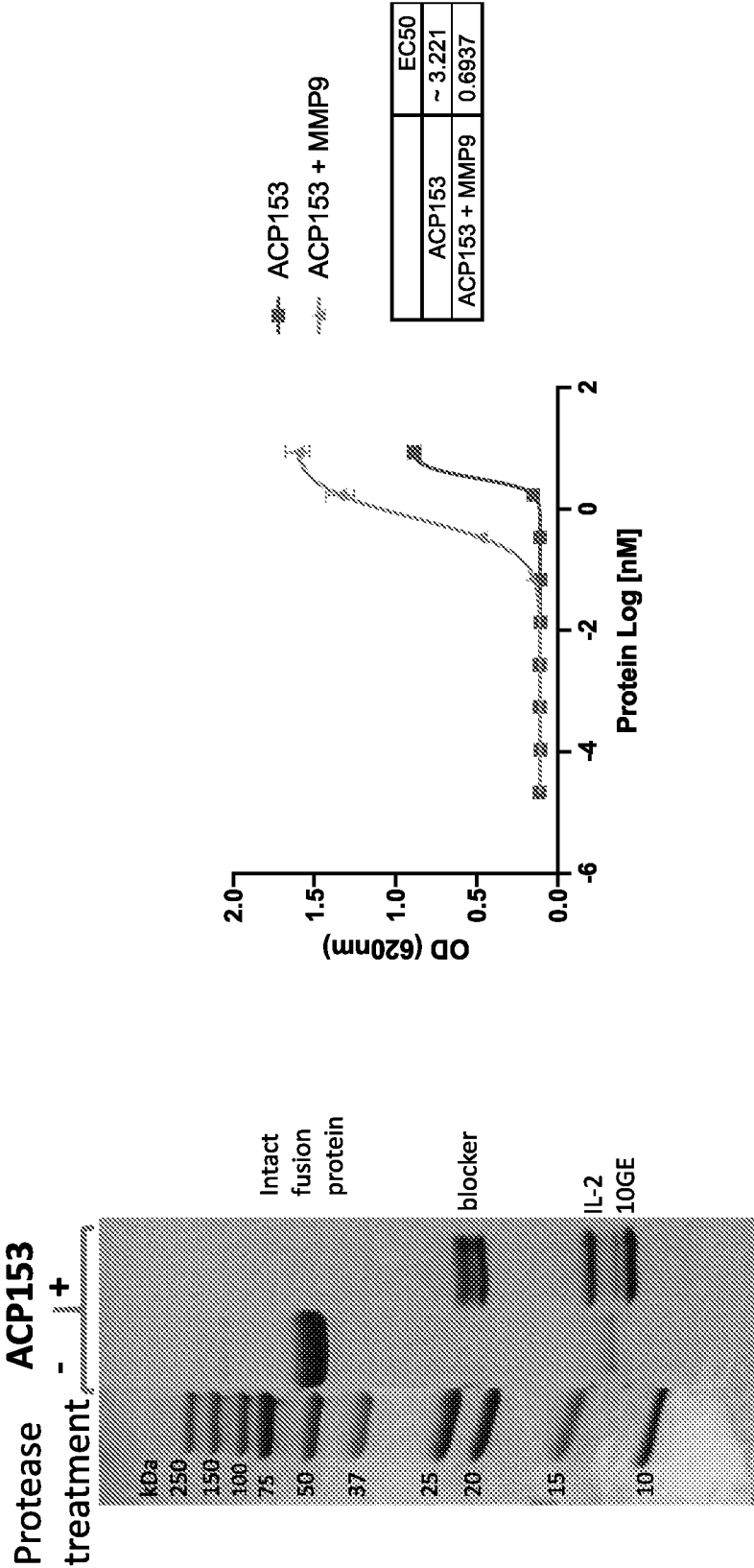


FIG. 24D

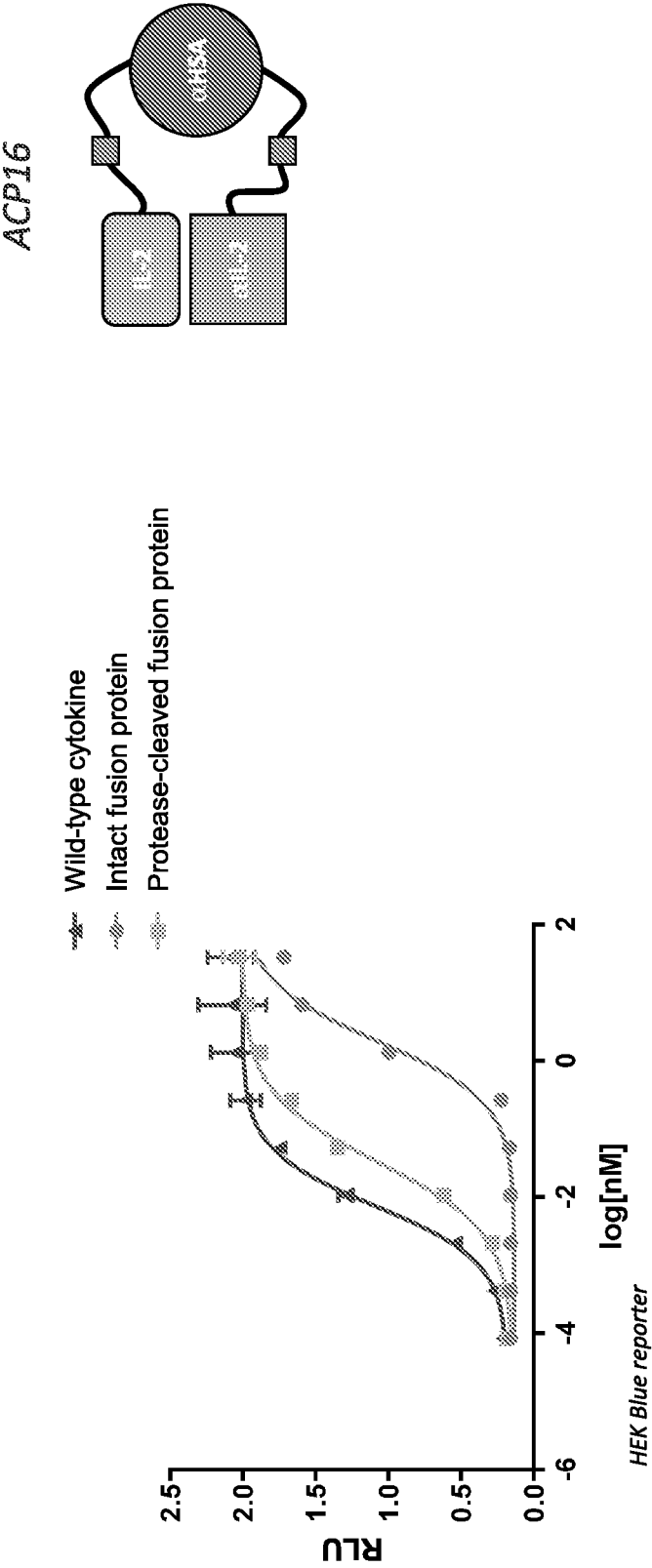


FIG. 25

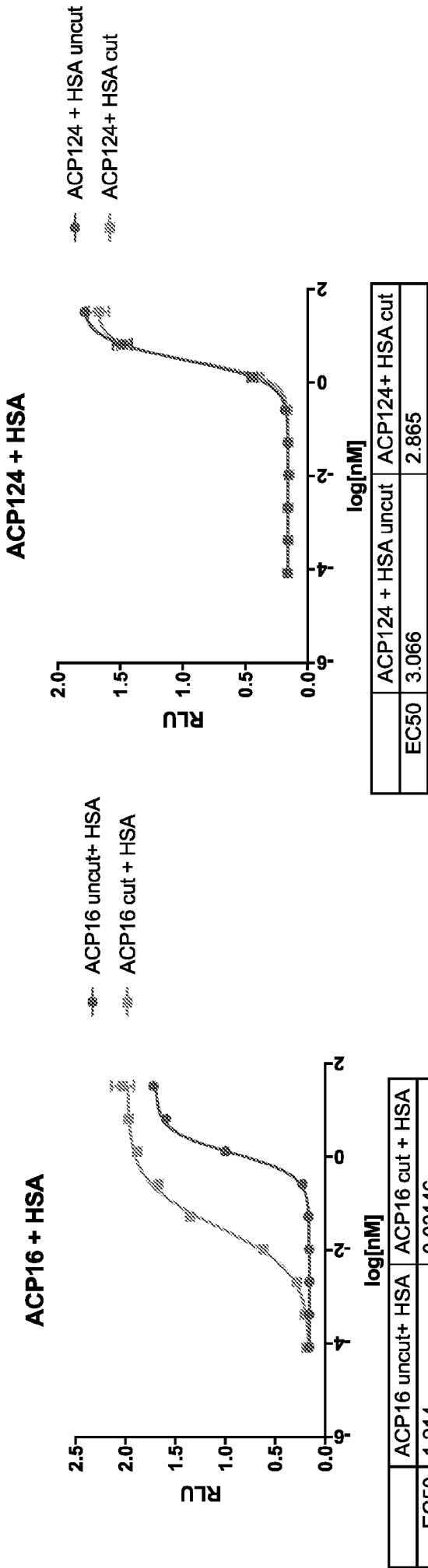


FIG. 25B

FIG. 25A

FIG. 25C

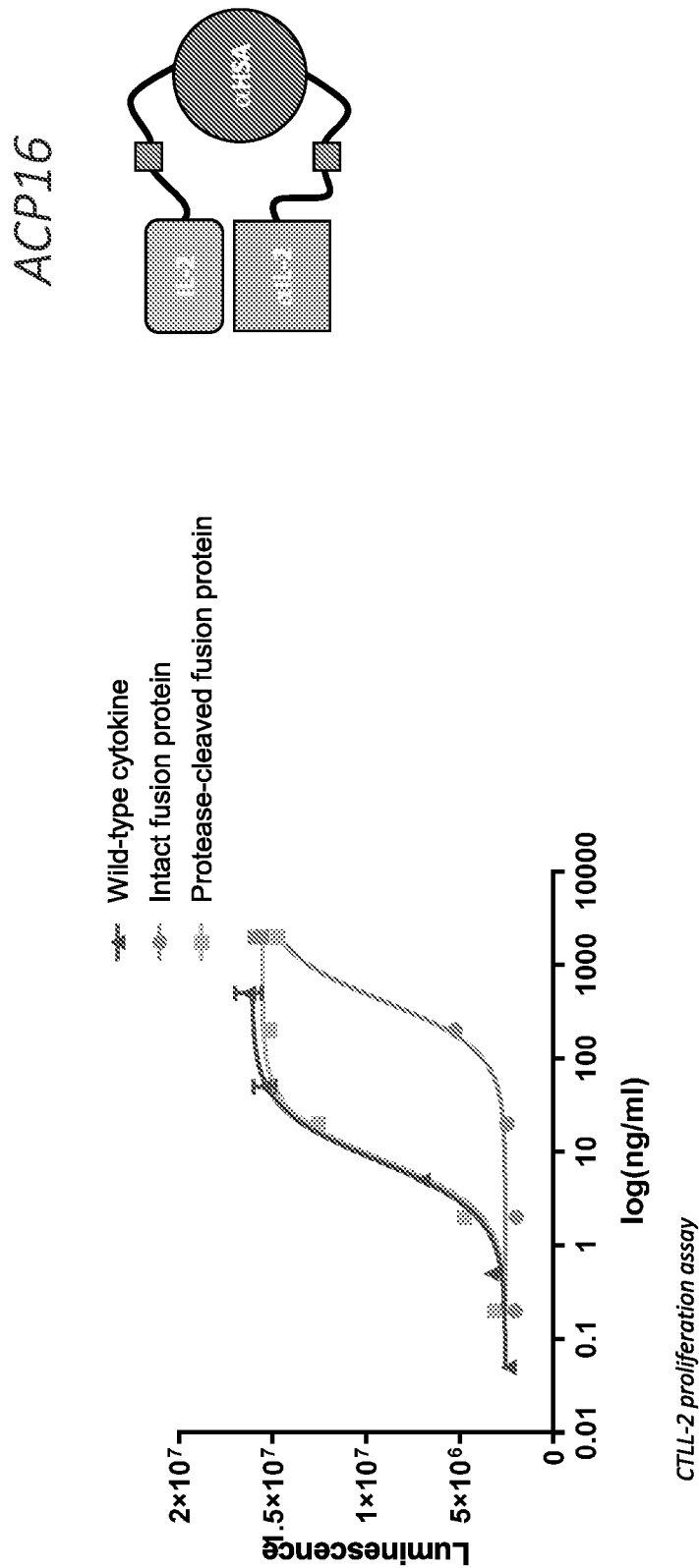


FIG. 26C

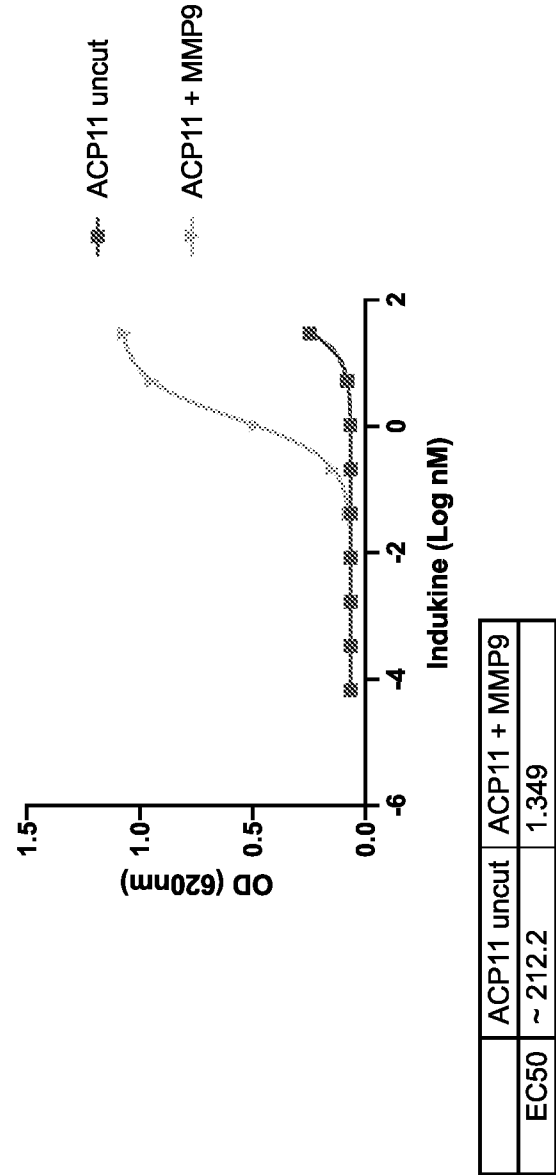


FIG. 26B

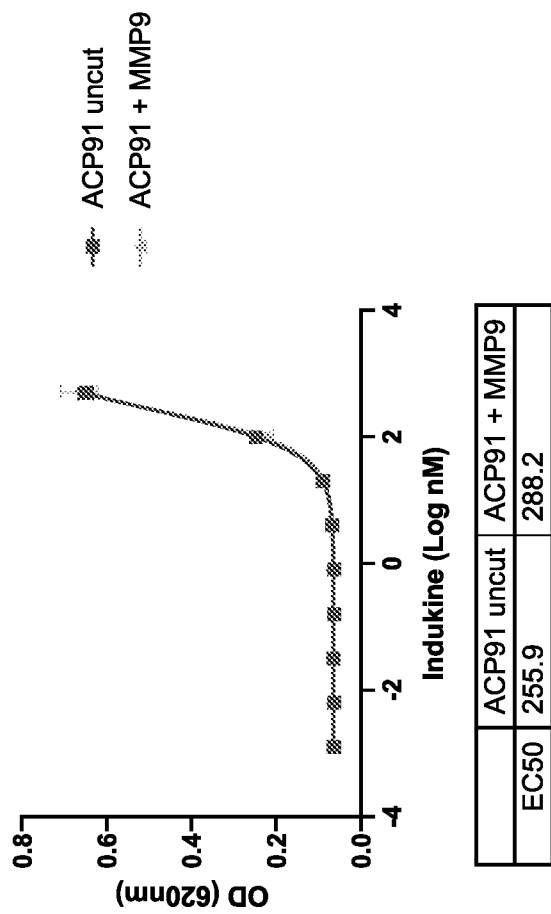


FIG. 26C

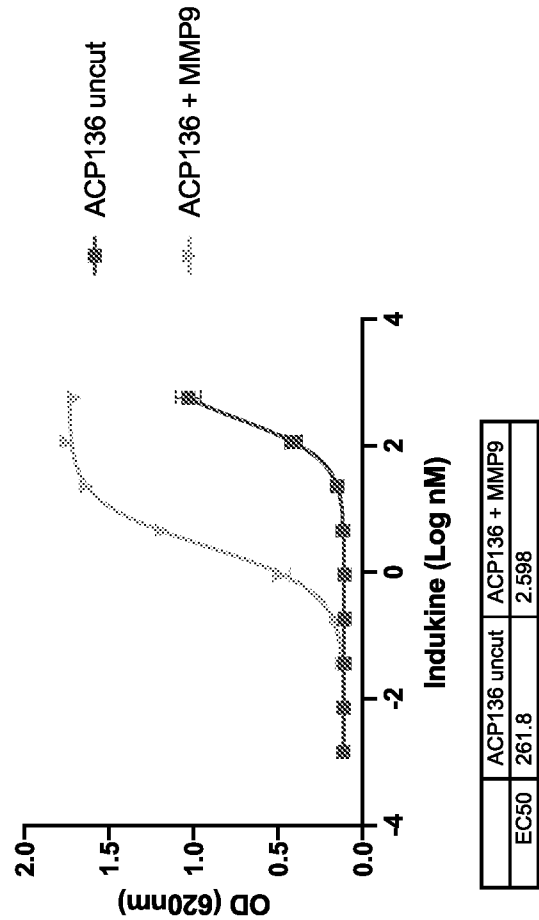


FIG. 27A

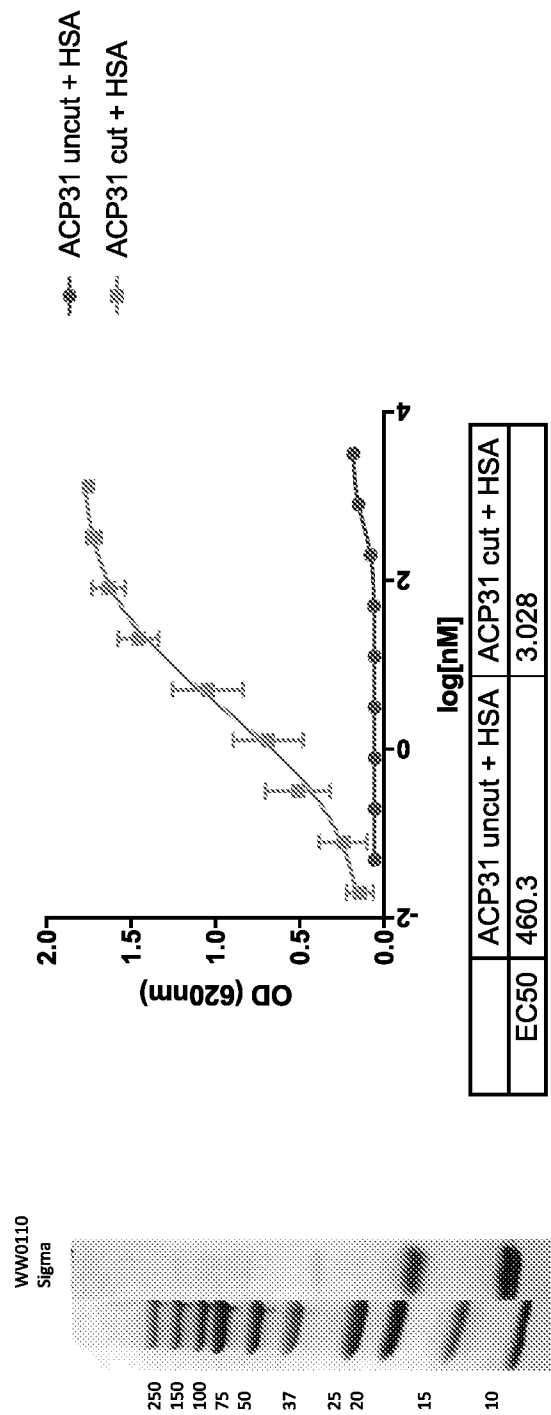


FIG. 27

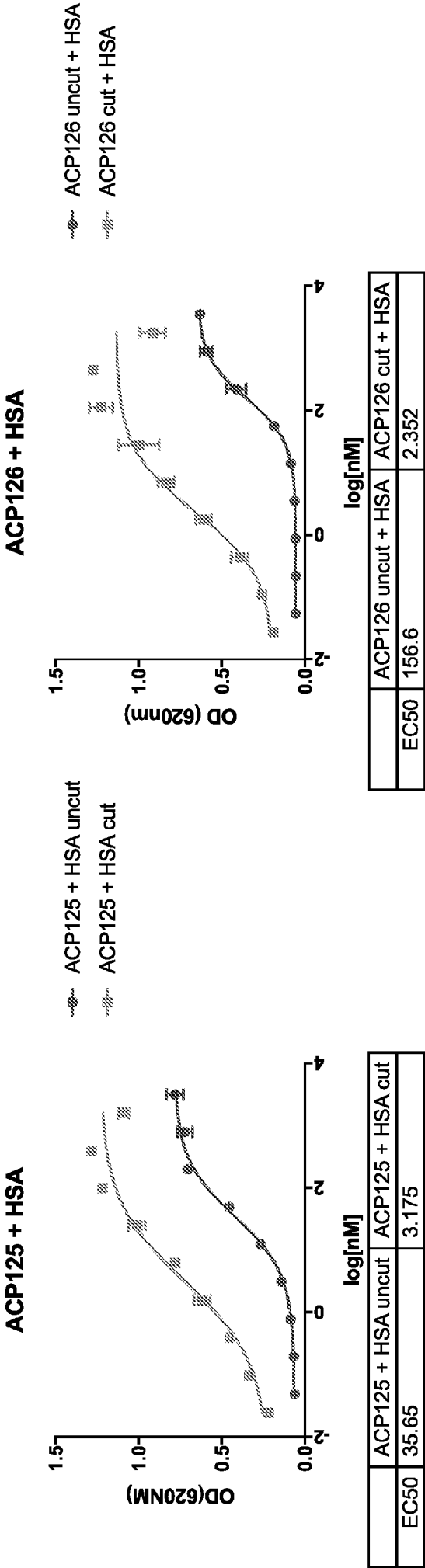


Fig 27C

Fig 27B

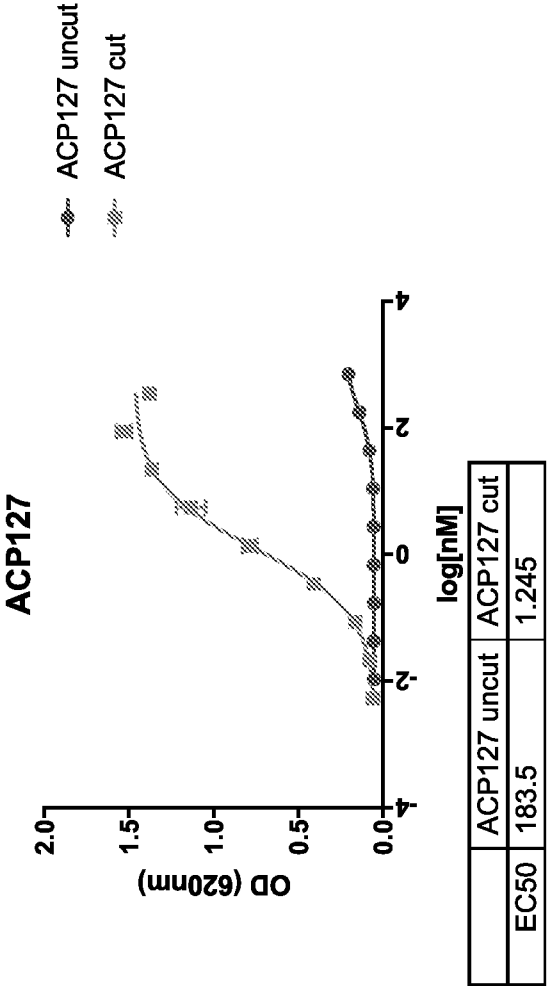
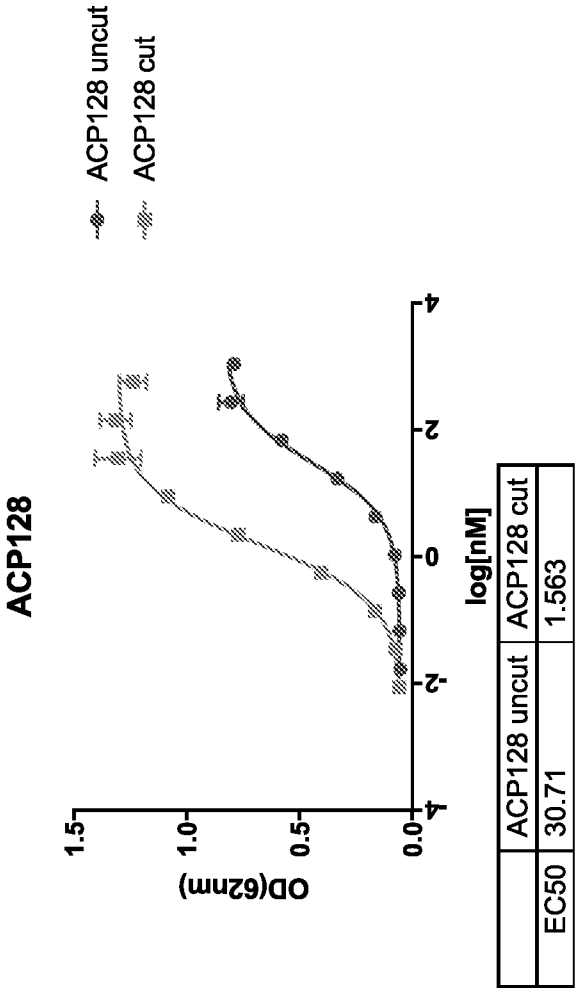


Fig 27D

FIG. 27

Fig 27E



ACP129

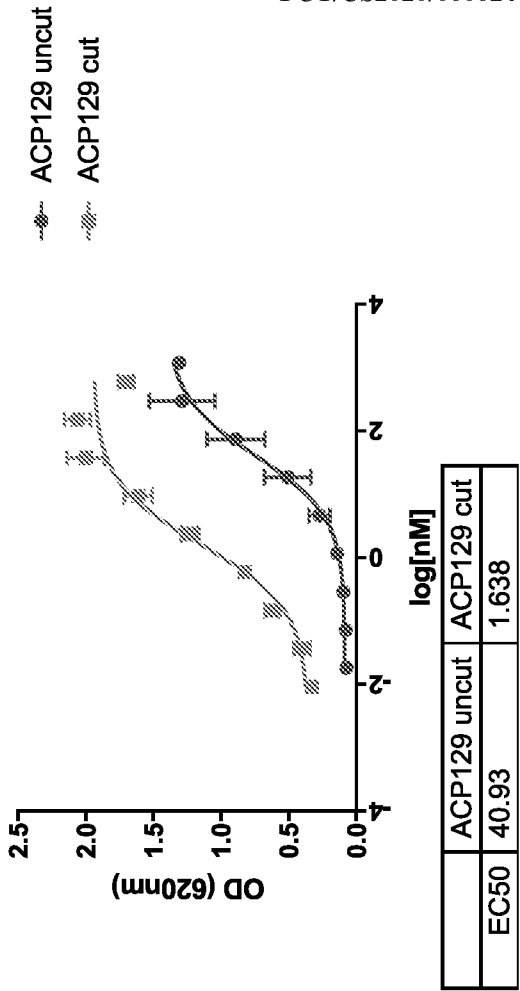


FIG. 28

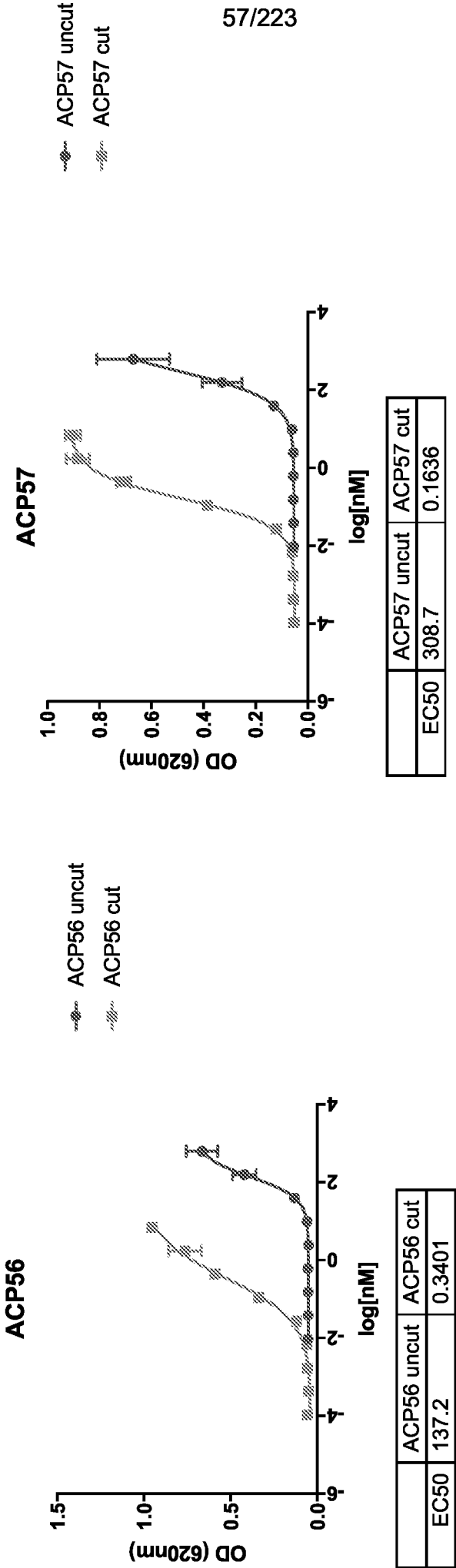


Fig 28A

Fig 28B

FIG. 28

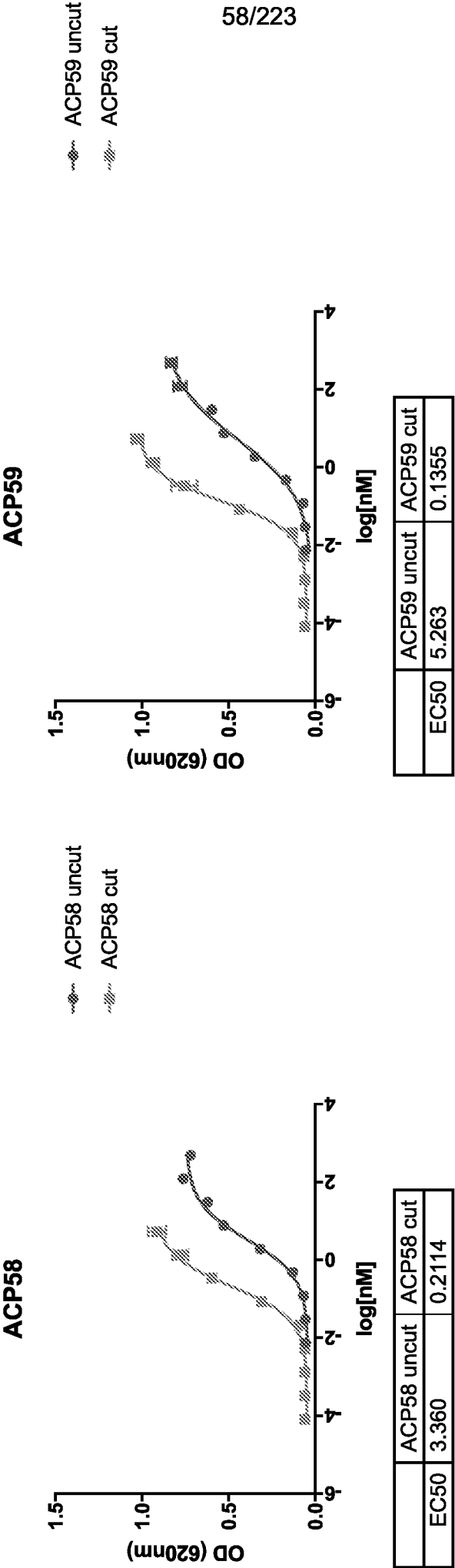


Fig 28D

Fig 28C

FIG. 28E

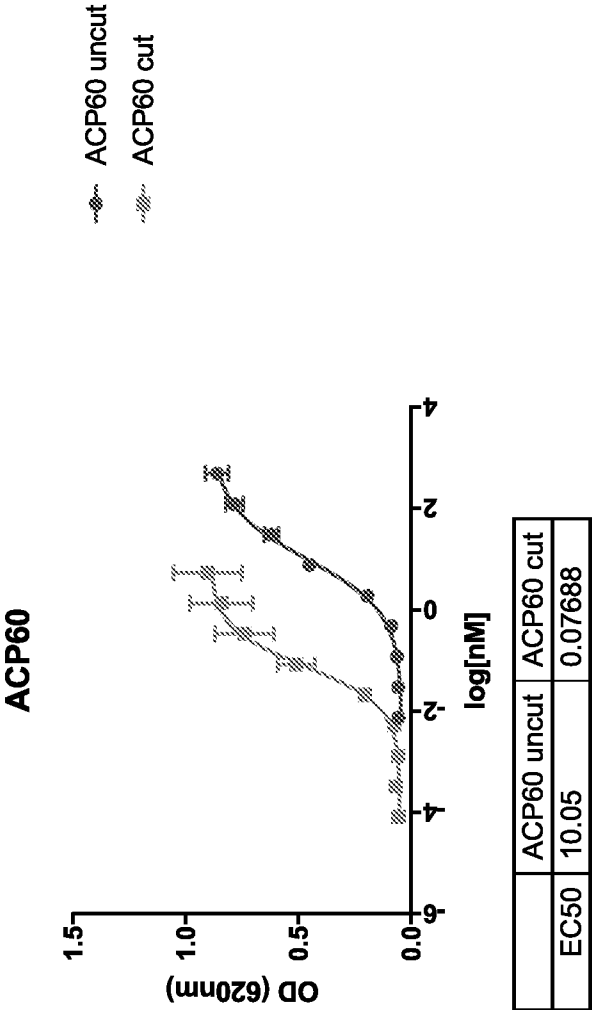


FIG. 28F

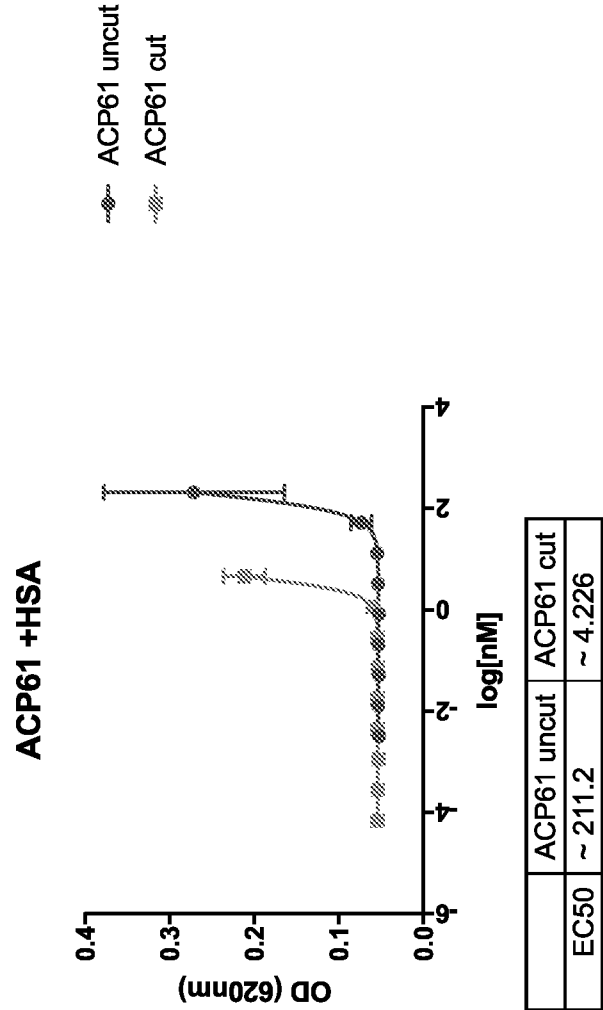
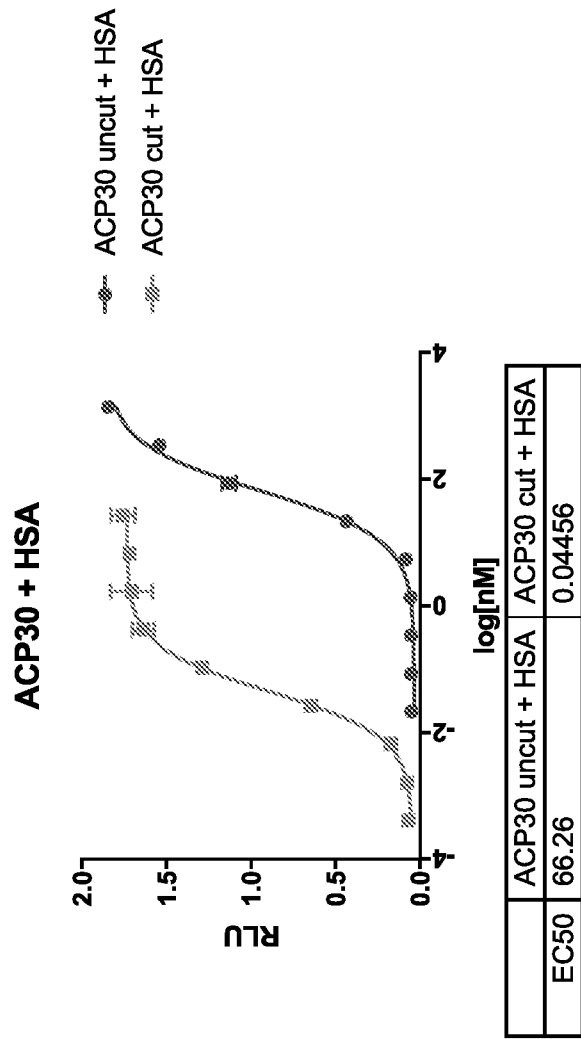


FIG. 28G



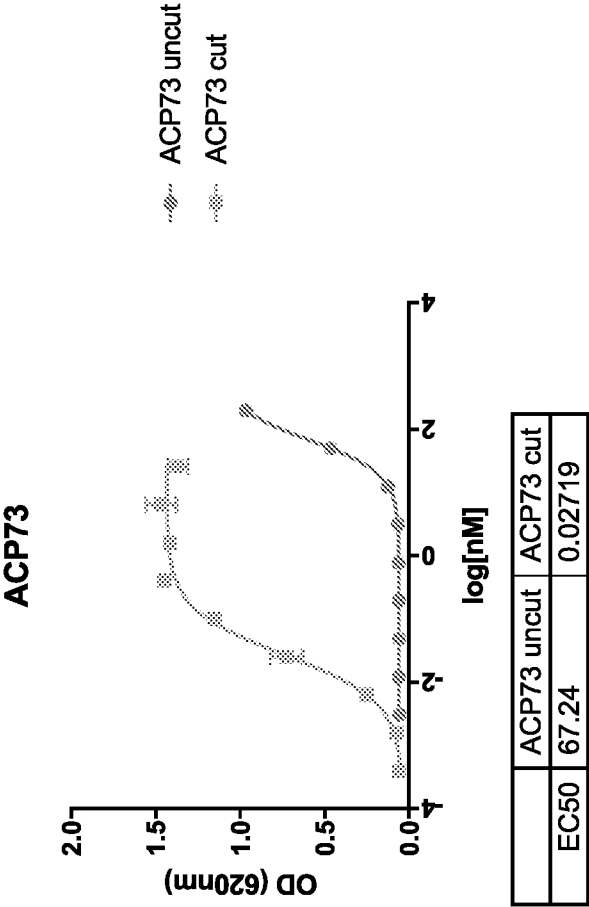


FIG. 28H

FIG. 28I

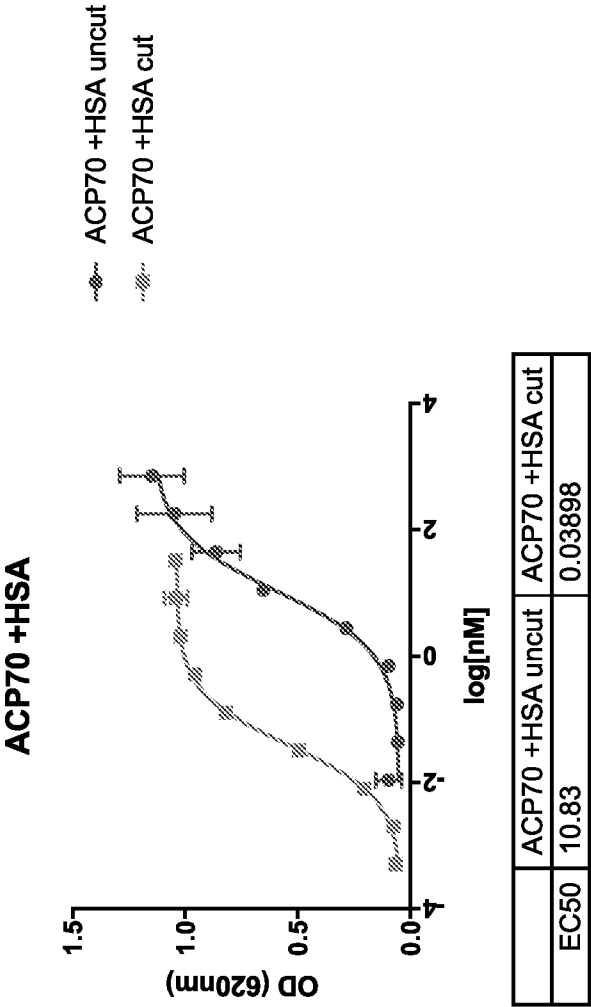


FIG. 28J

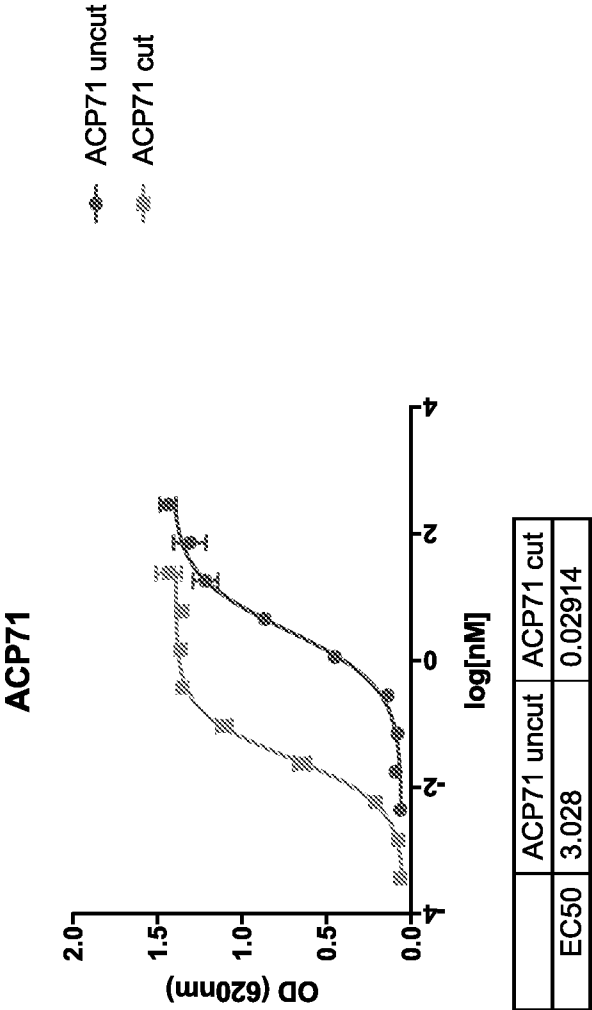


FIG. 28K

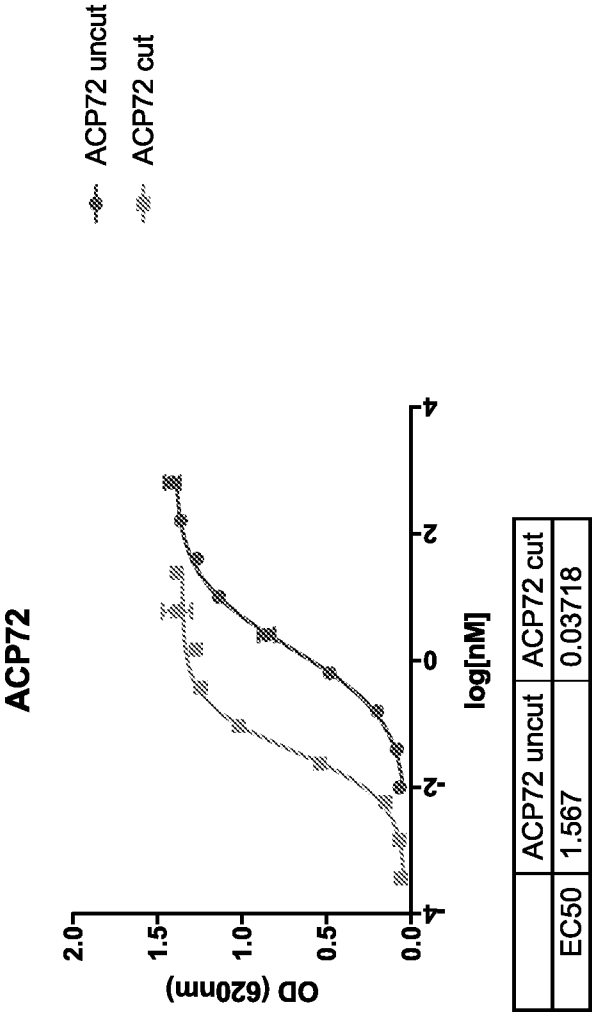


FIG. 28L

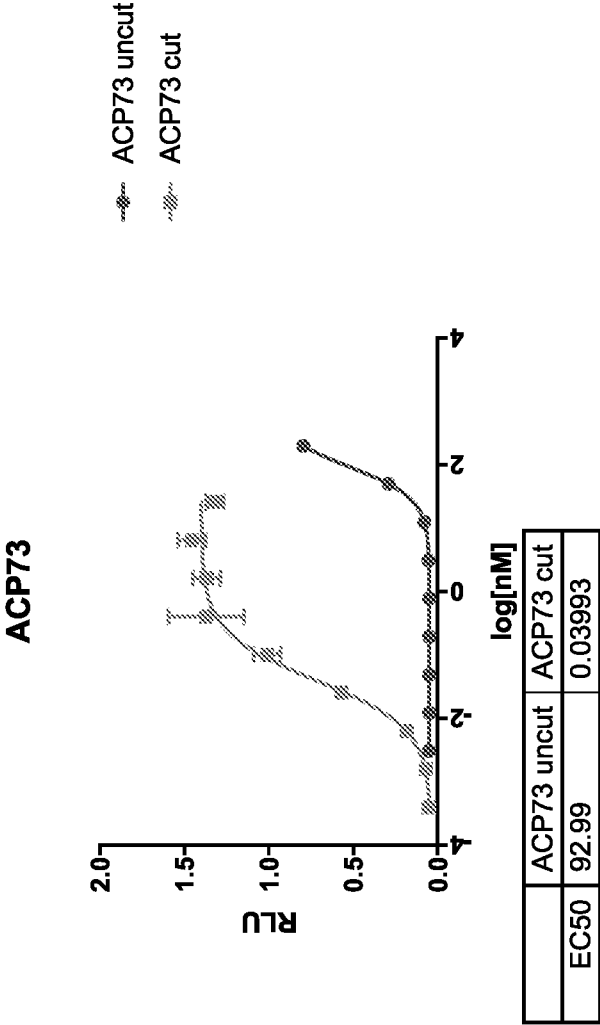


FIG. 28M

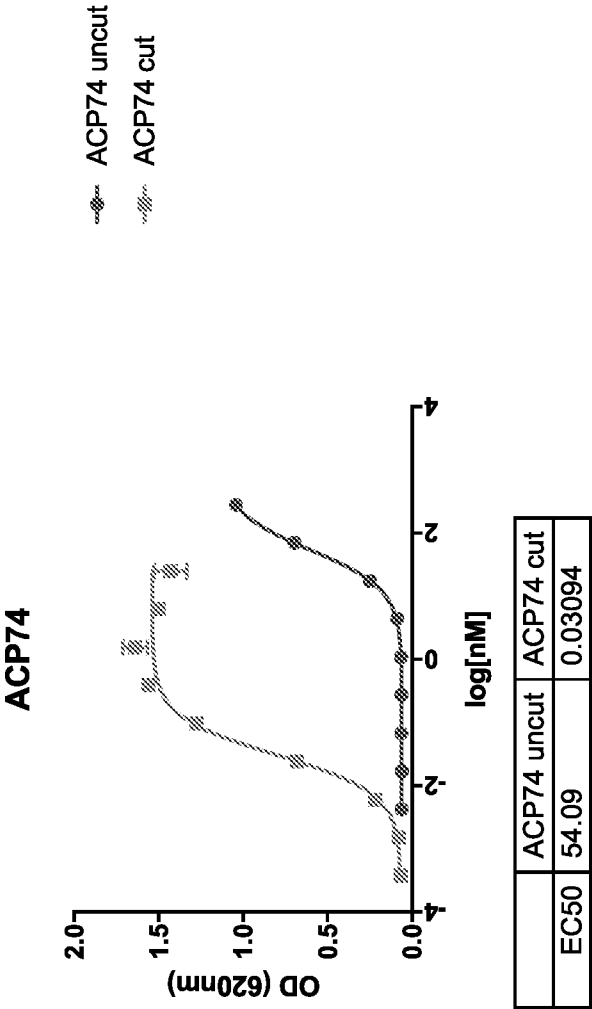


FIG. 28N

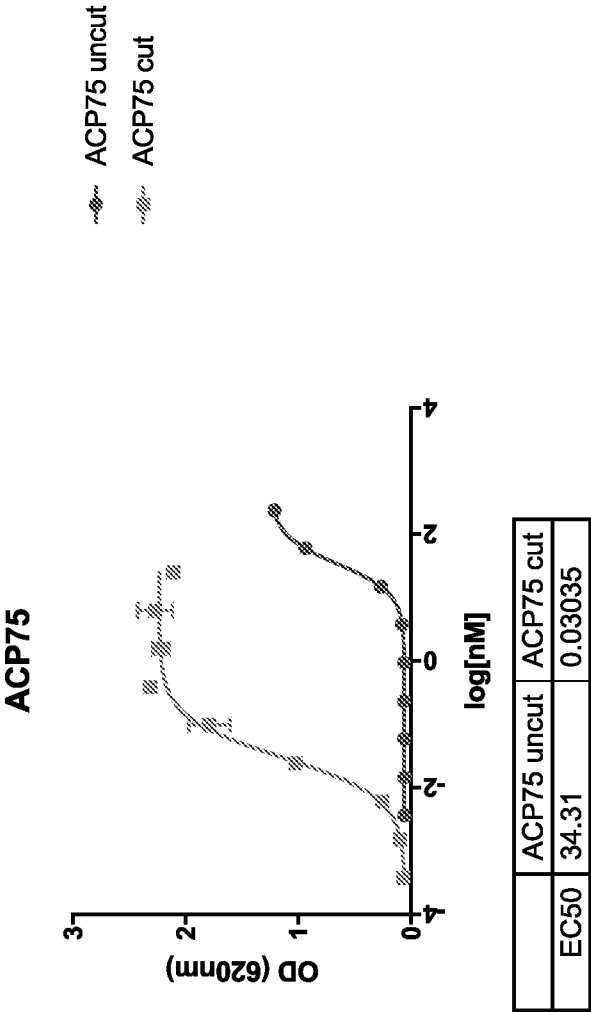
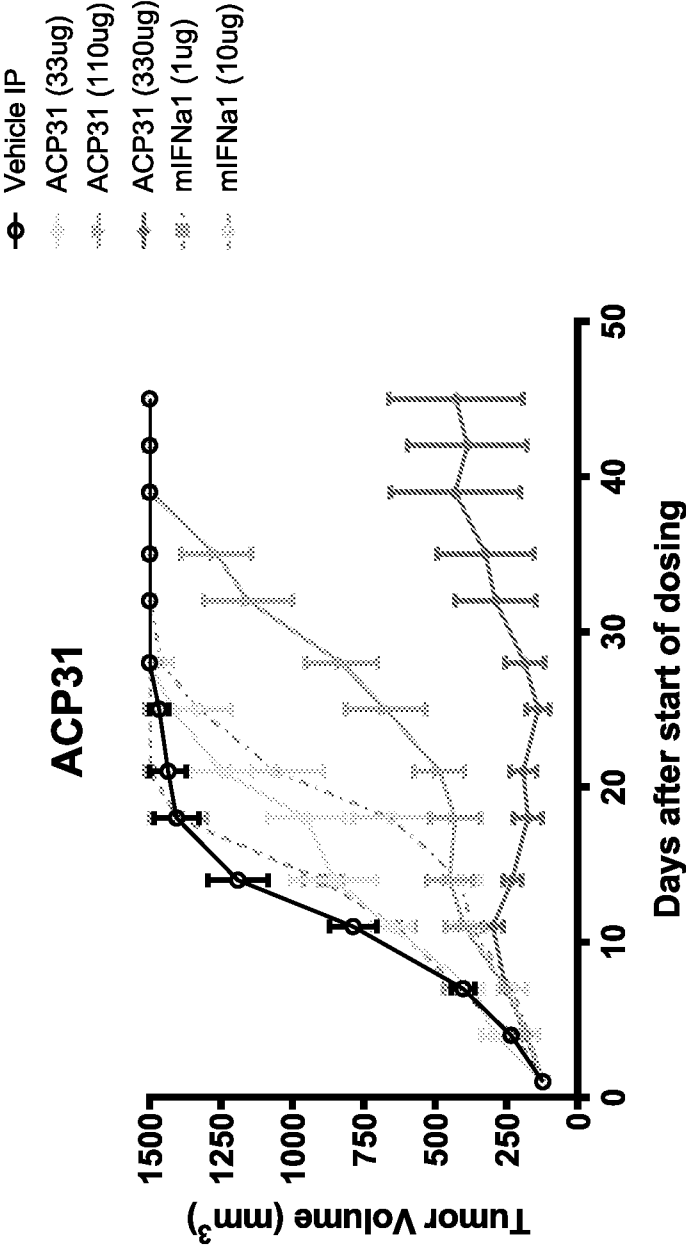


FIG. 29A



- Vehicle IP
- ACP11 (17.5ug)
- ACP11 (175ug)
- ACP11 (525ug)
- ACP04 (2ug)
- ACP04 (10ug)

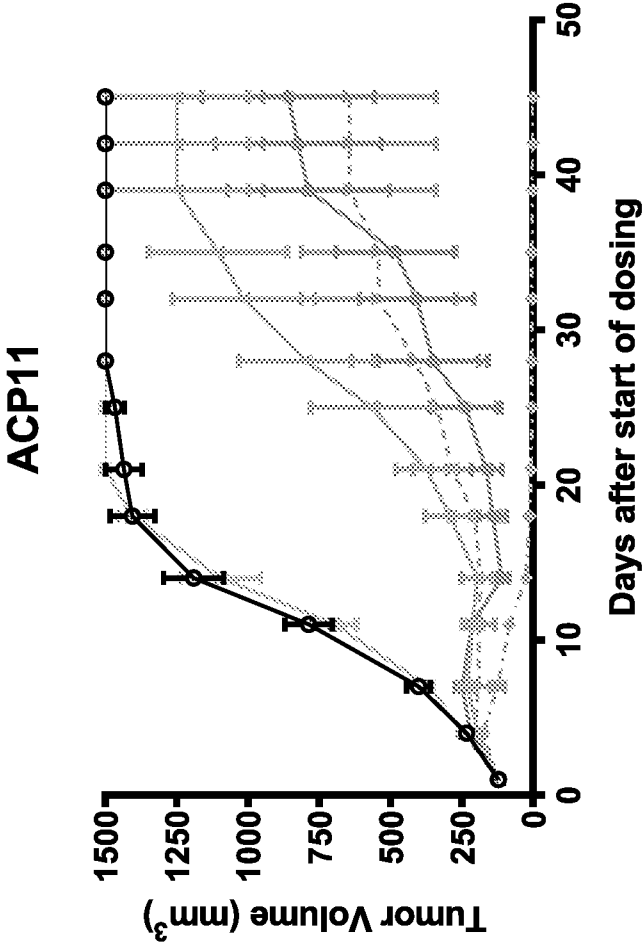


FIG. 29B

FIG. 30

FIG. 30A

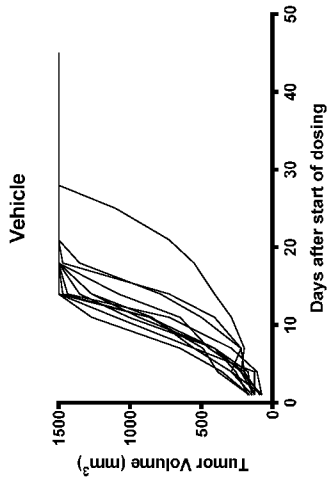


FIG. 30B

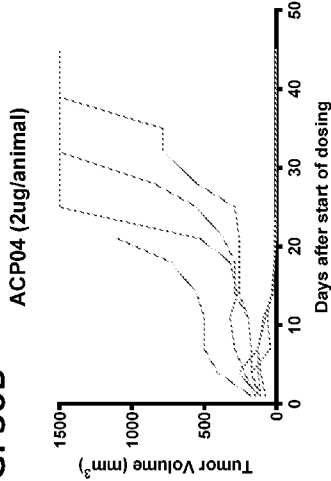


FIG. 30C

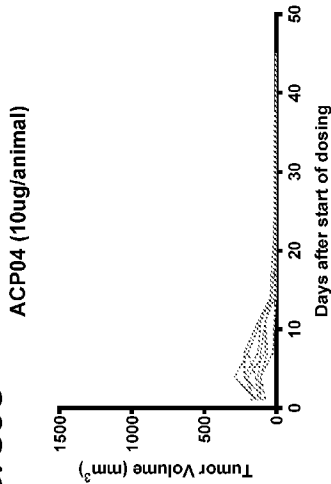


FIG. 30D

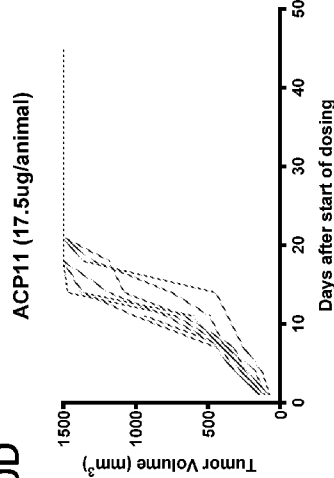


FIG. 30E

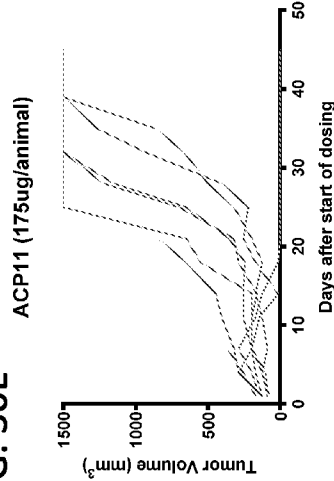


FIG. 30F

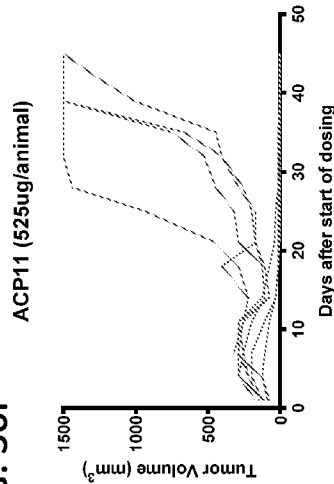
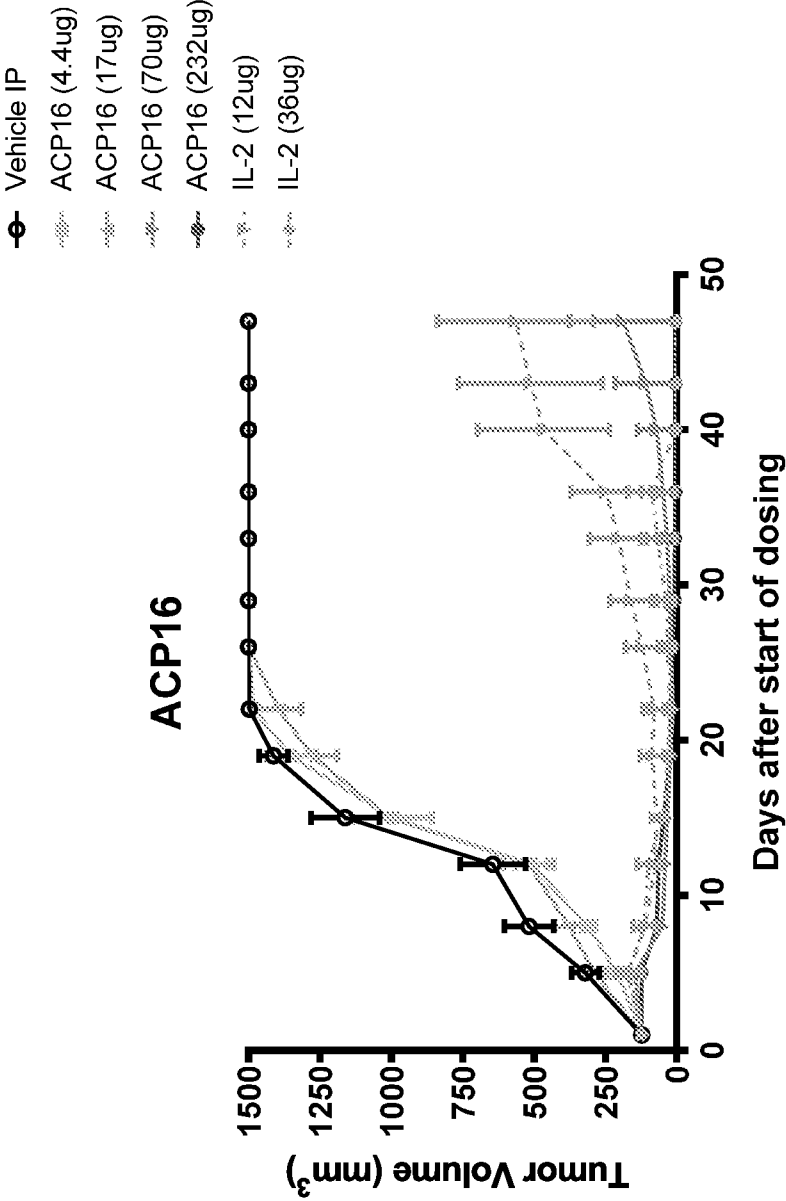


FIG. 31A



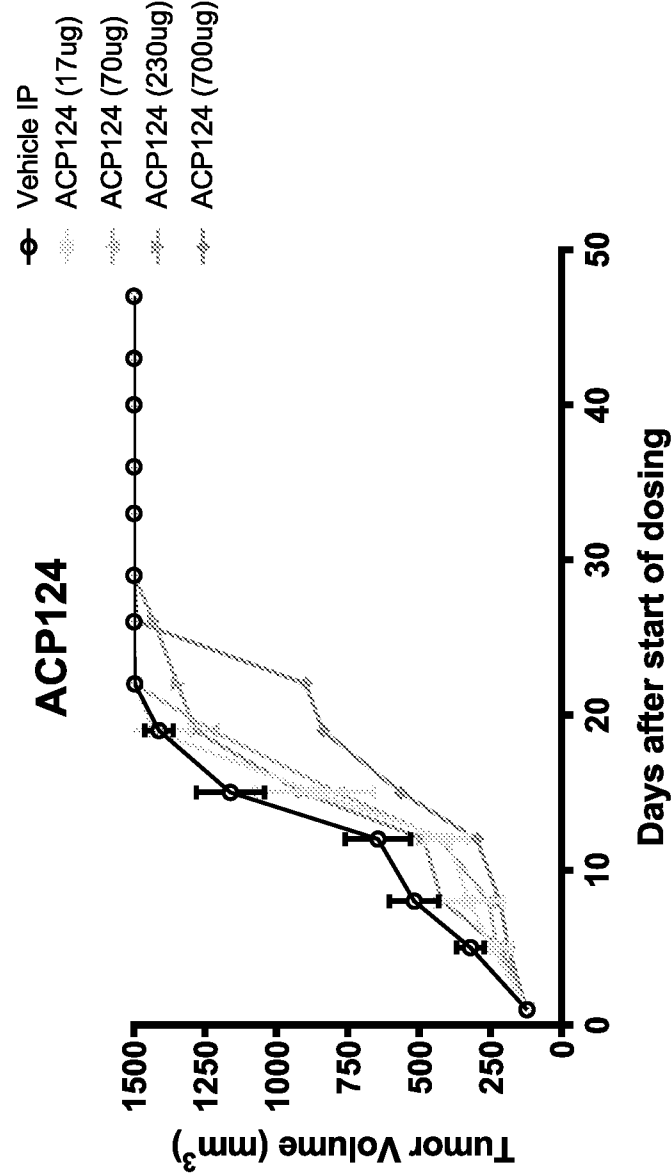


FIG. 31B

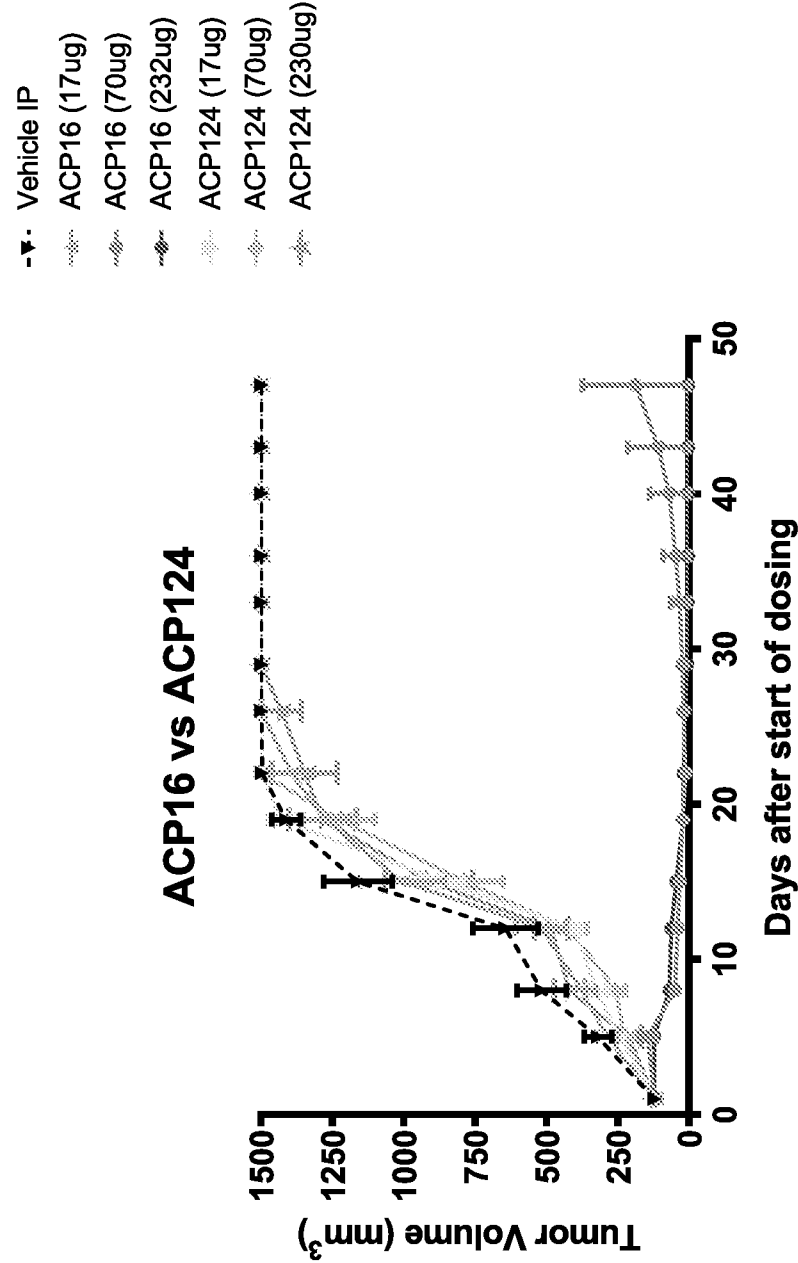


FIG. 31C

FIG. 32A

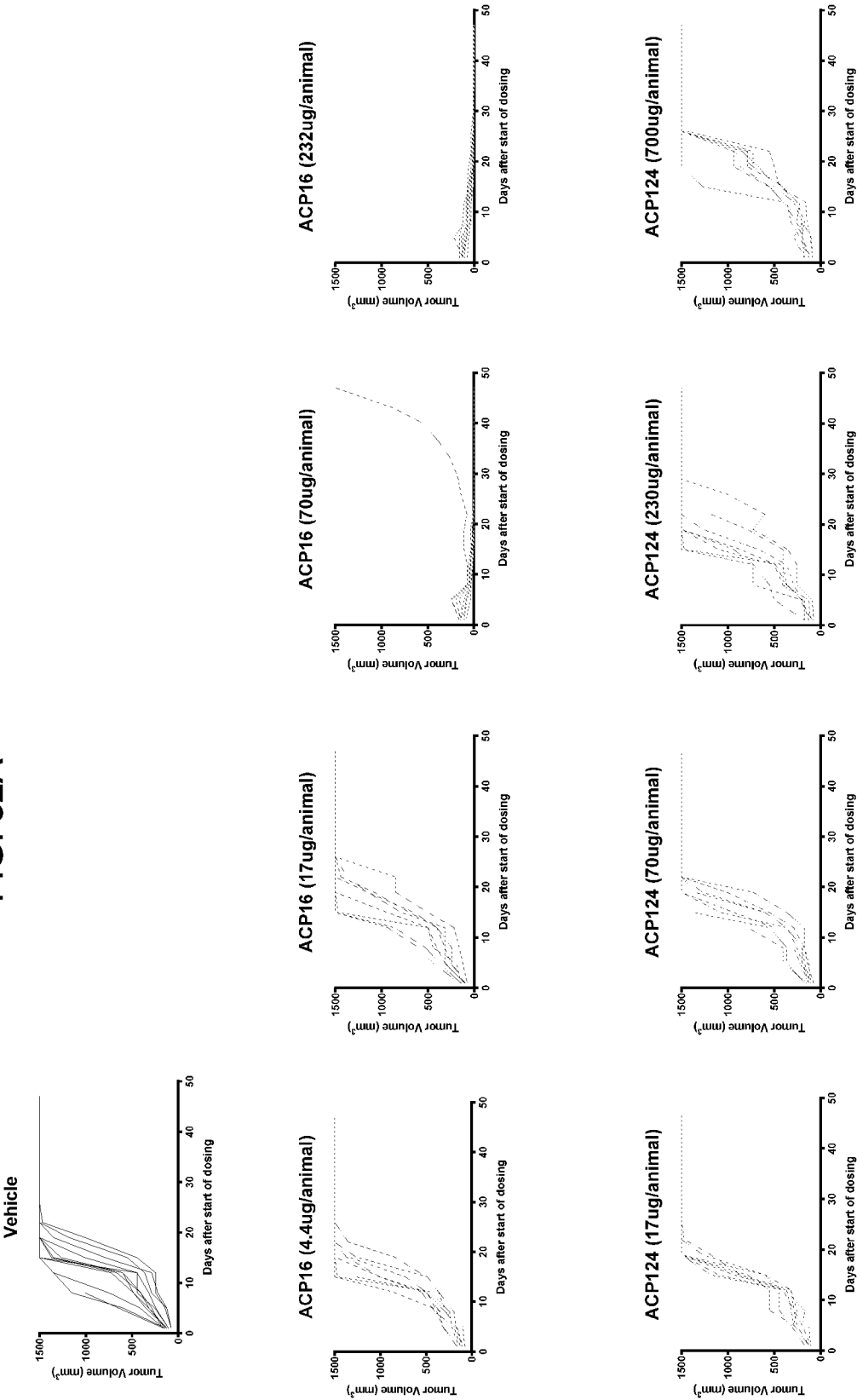


FIG. 32B

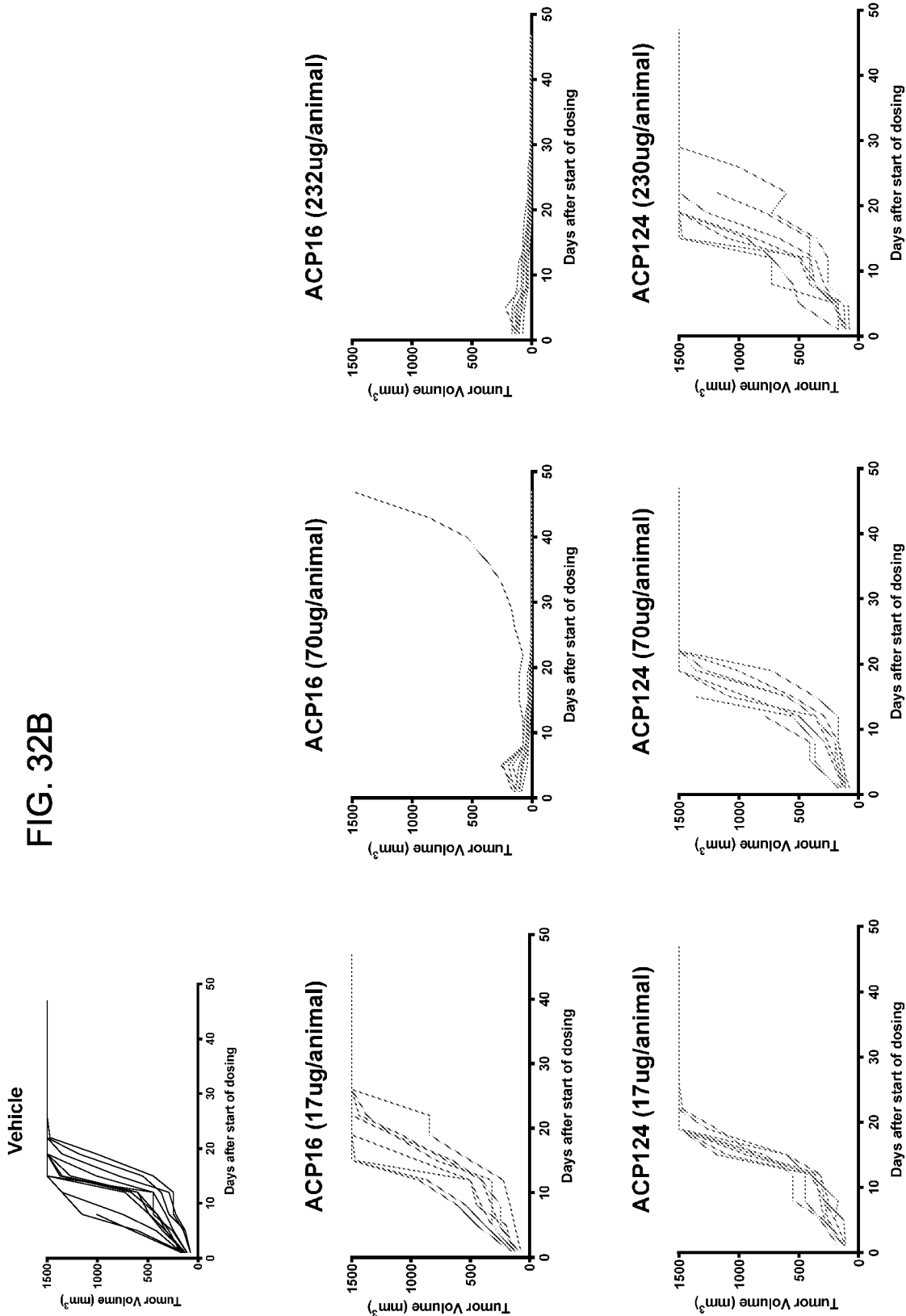


FIG. 33

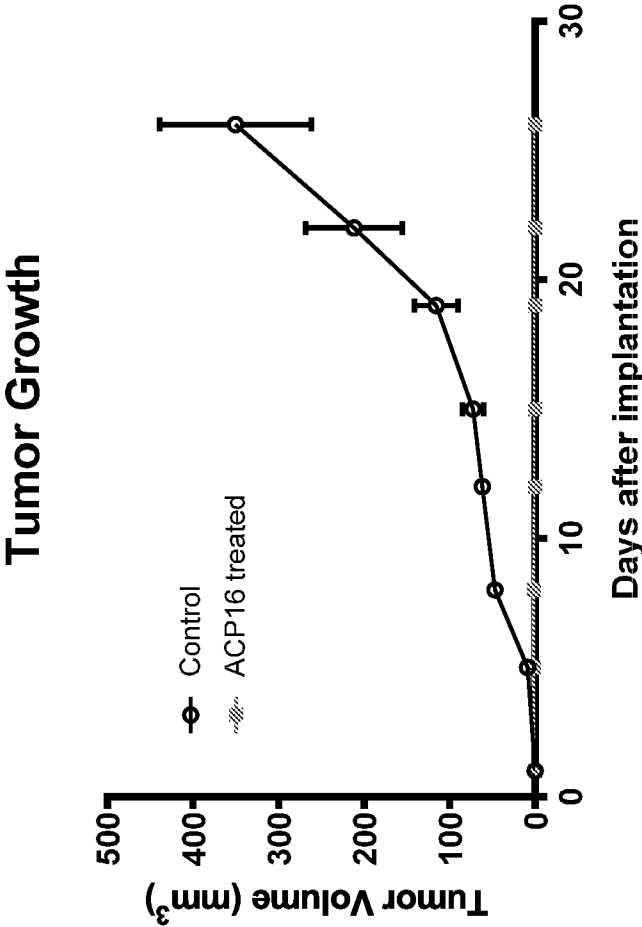


FIG. 34A

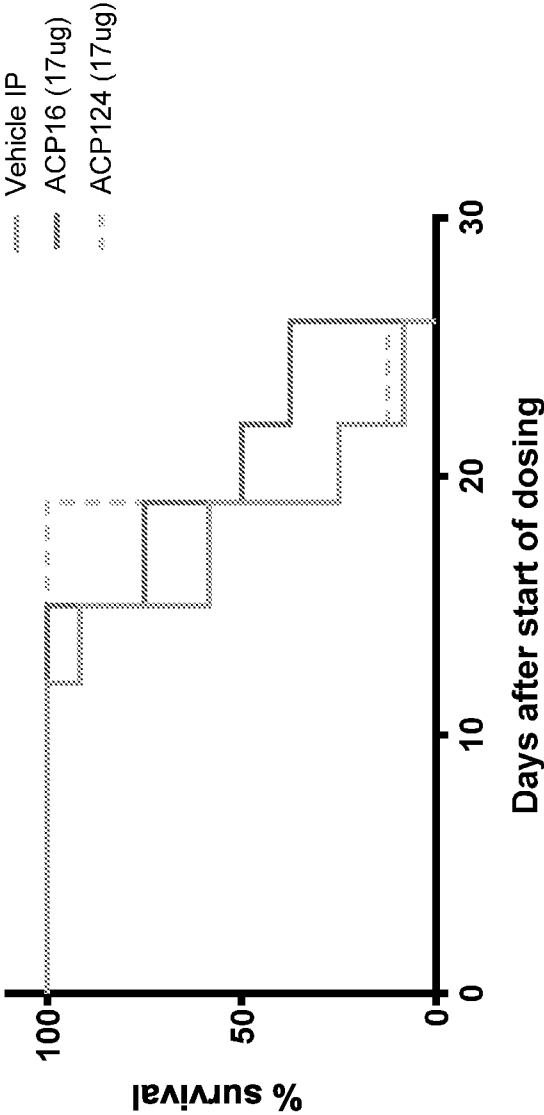


FIG. 34B

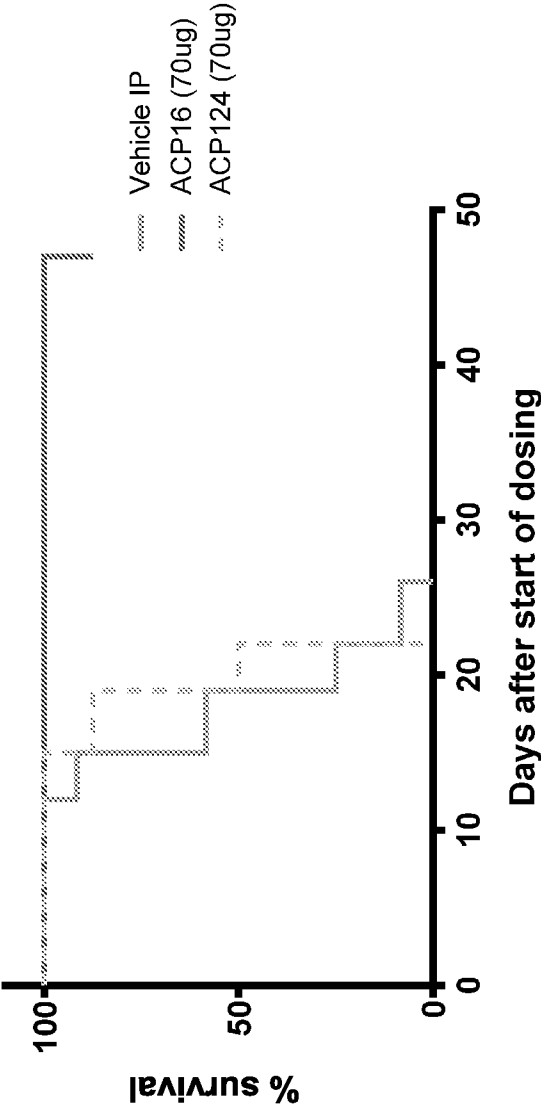


FIG. 34C

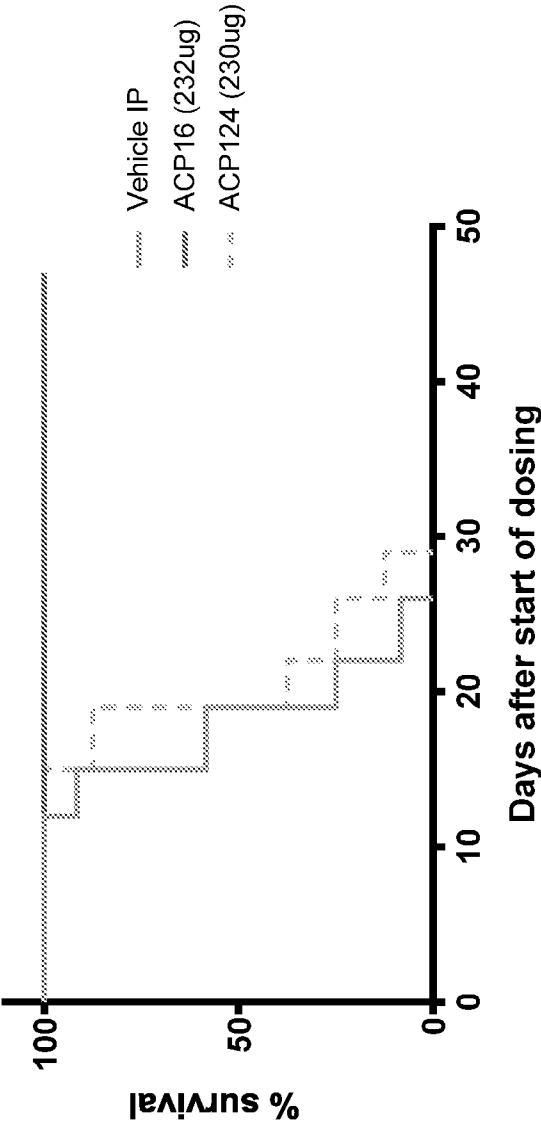
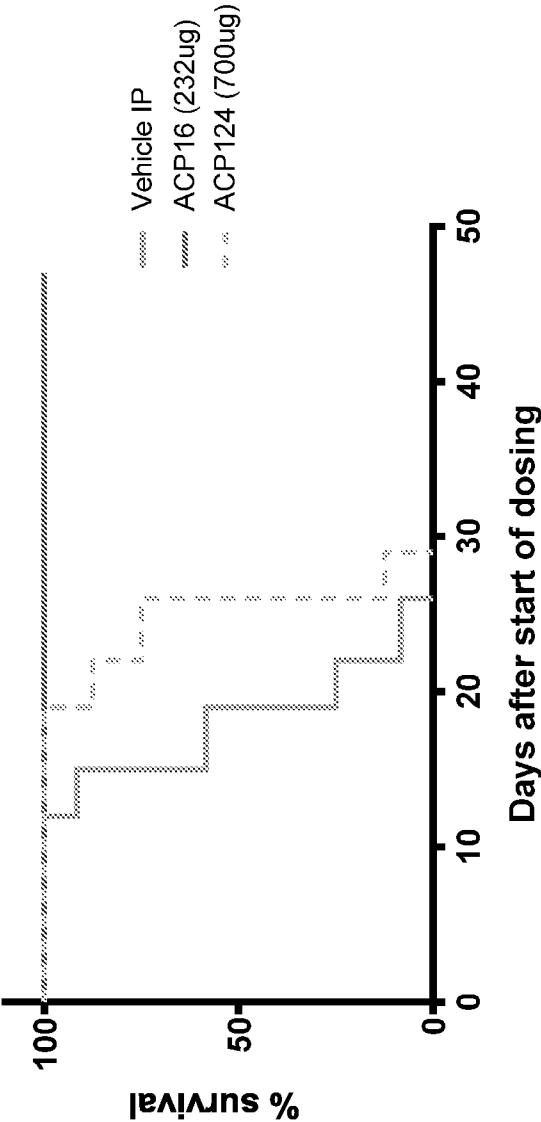
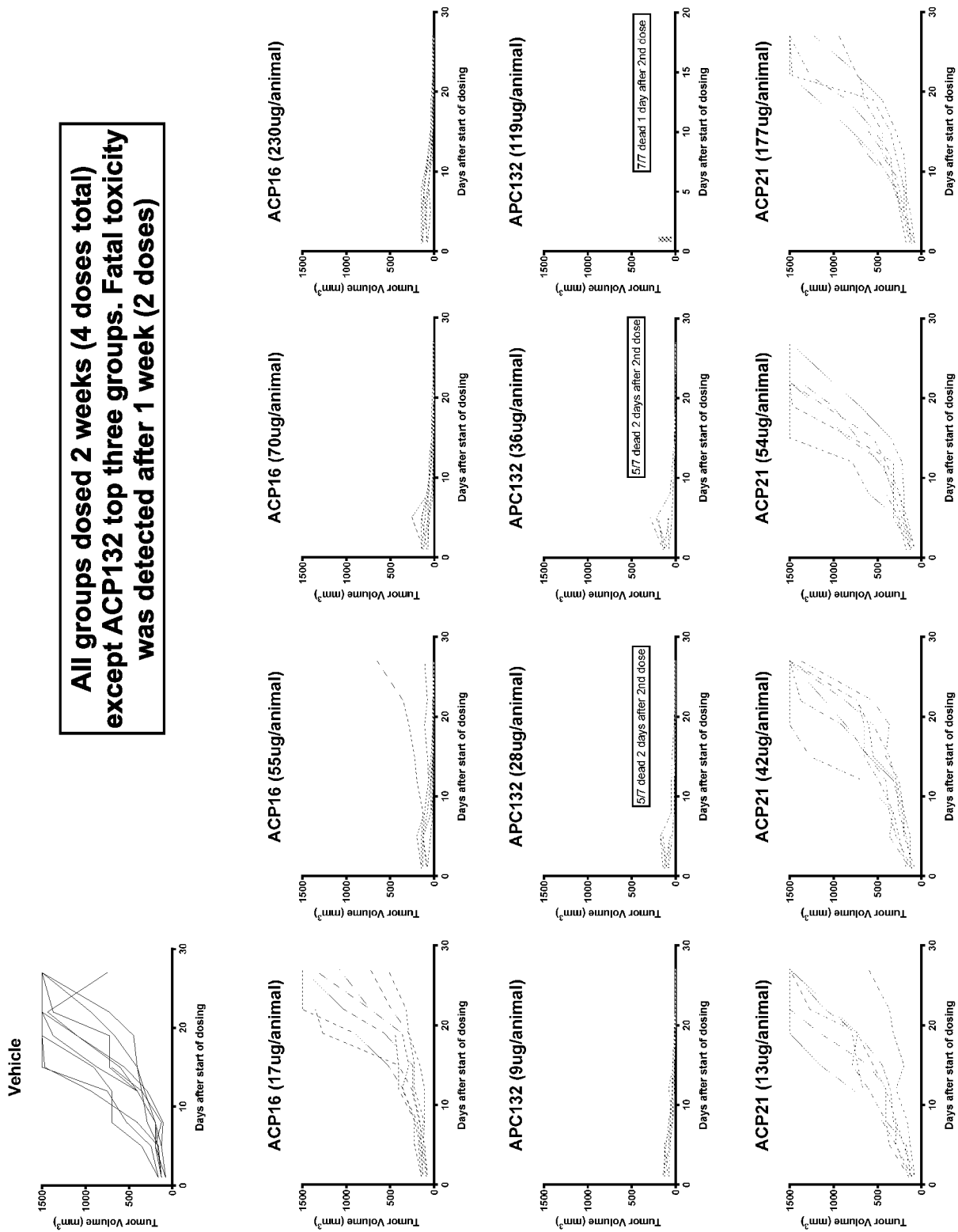


FIG. 34D





**All groups dosed 2 weeks (4 doses total)
except APC132 top three groups. Fatal toxicity
was detected after 1 week (2 doses)**

FIG. 35

FIG. 36A

HEK-Blue hIFNa/b Reporter Assay Using WW0612

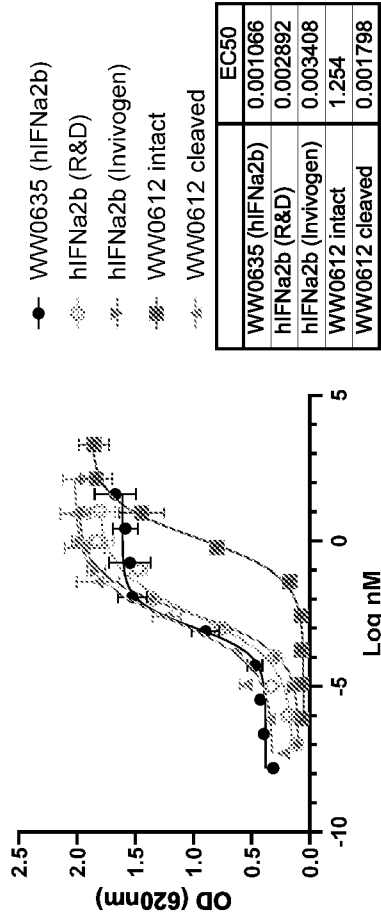


FIG. 36B

HEK-Blue hIFNa/b Reporter Assay Using WW0613

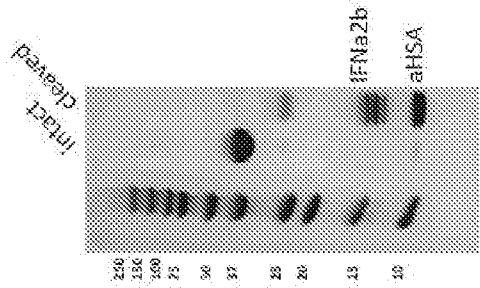
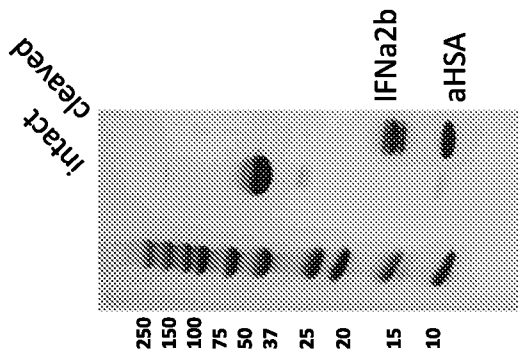
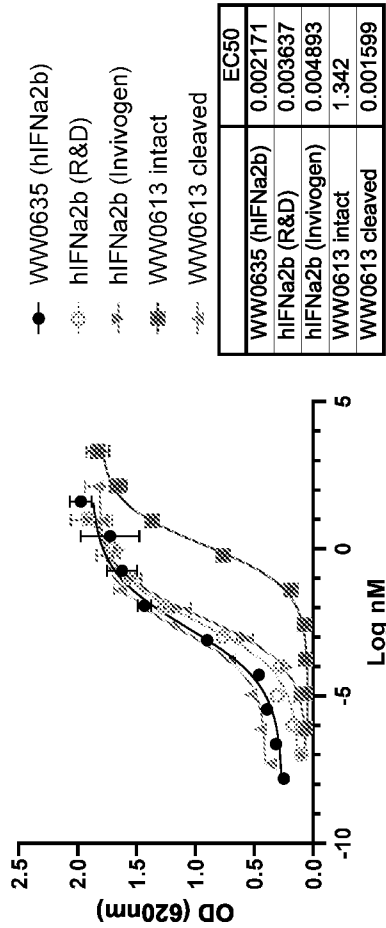


FIG. 36C HEK-Blue hIFNa/b Reporter Assay Using WW0645

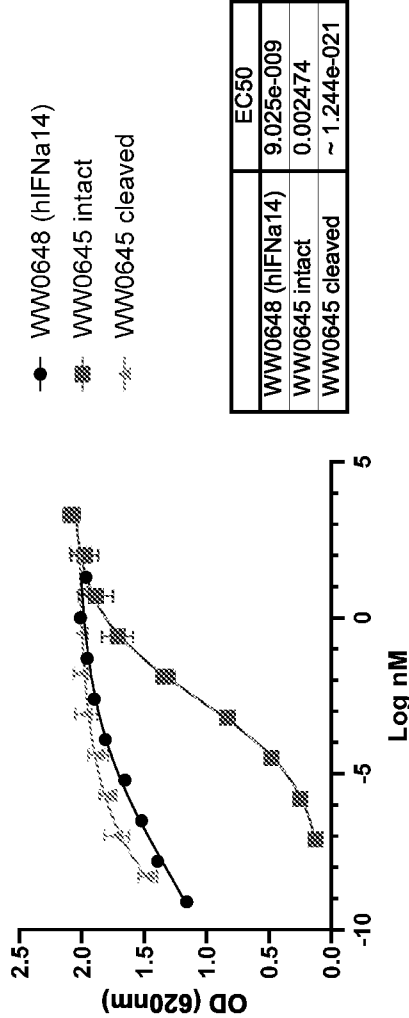


FIG. 36D

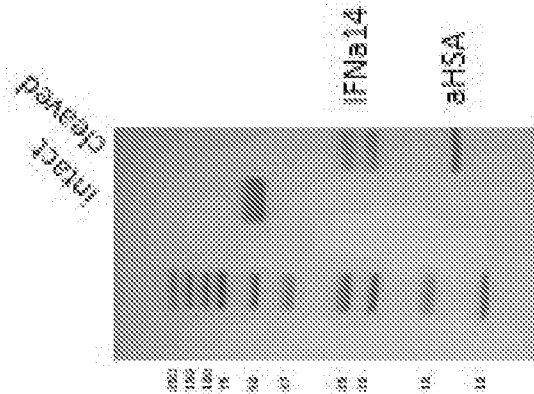
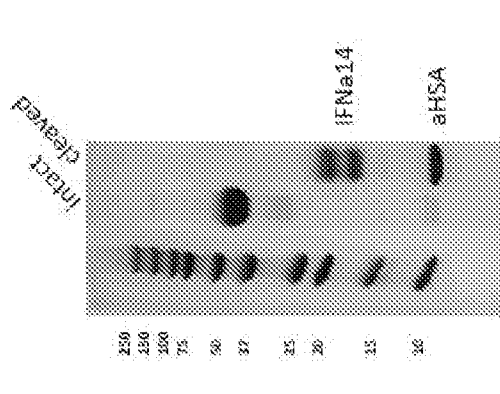
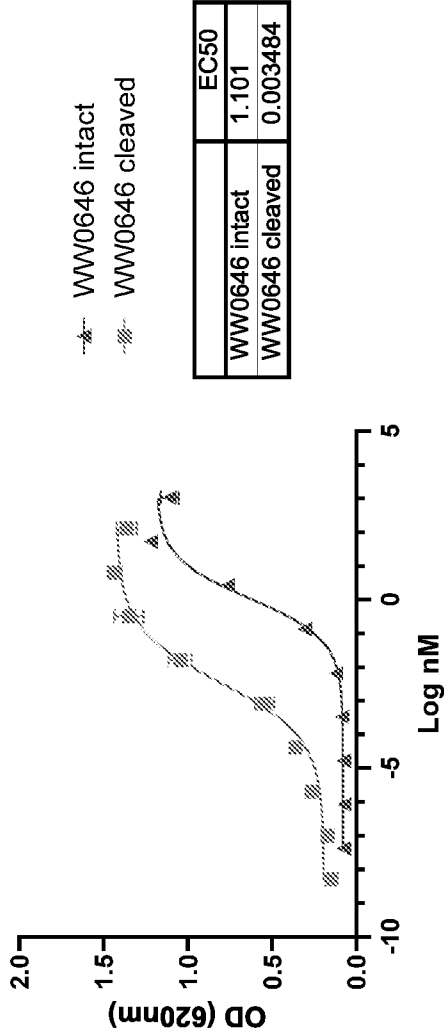


FIG. 36E

HEK-Blue hIFNa/b Reporter Assay Using WW0821

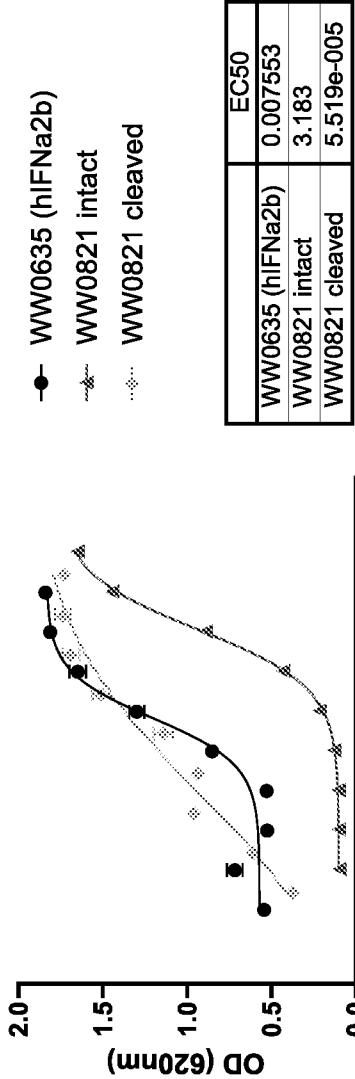


FIG. 36F

HEK-Blue hIFNa/b Reporter Assay Using WW0822

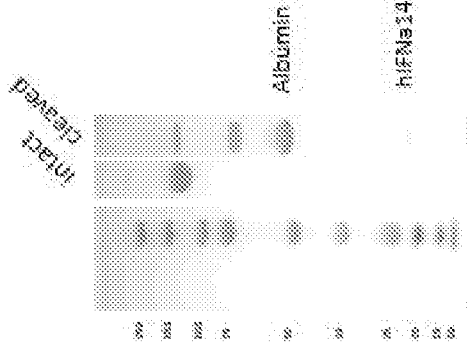
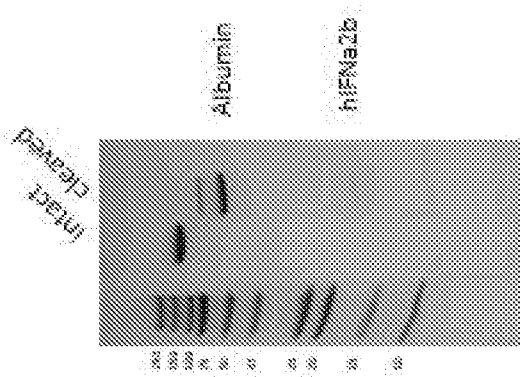
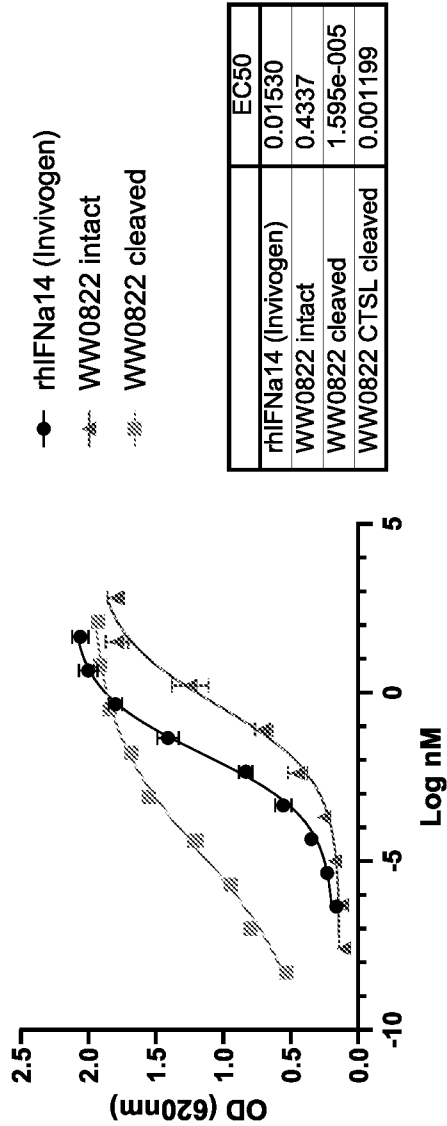


FIG. 36G

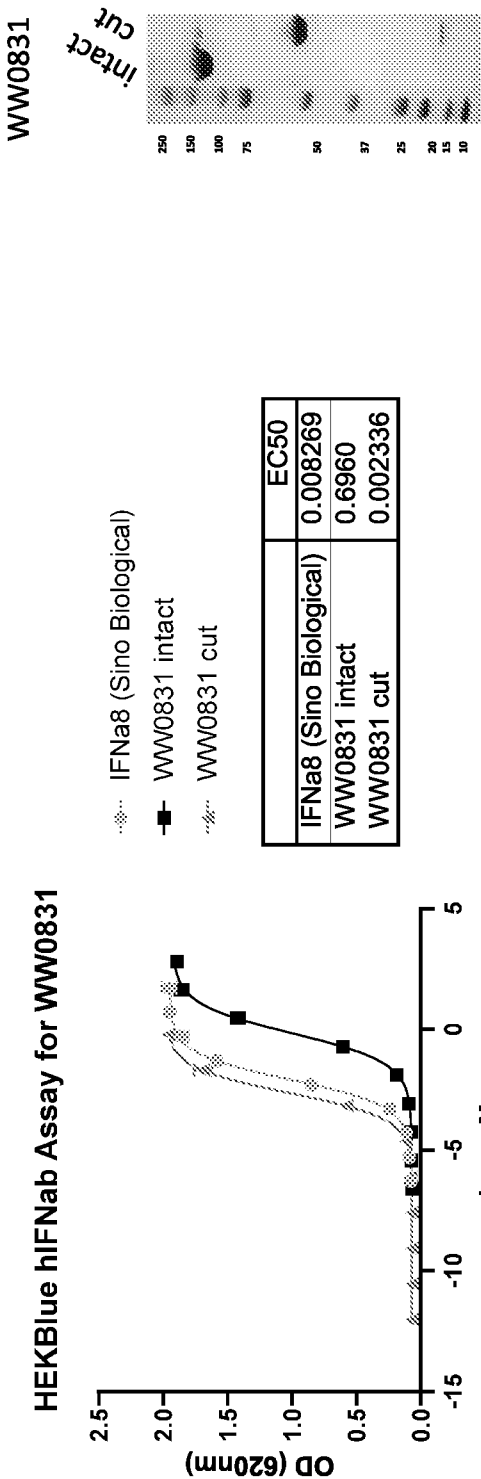


FIG. 36H

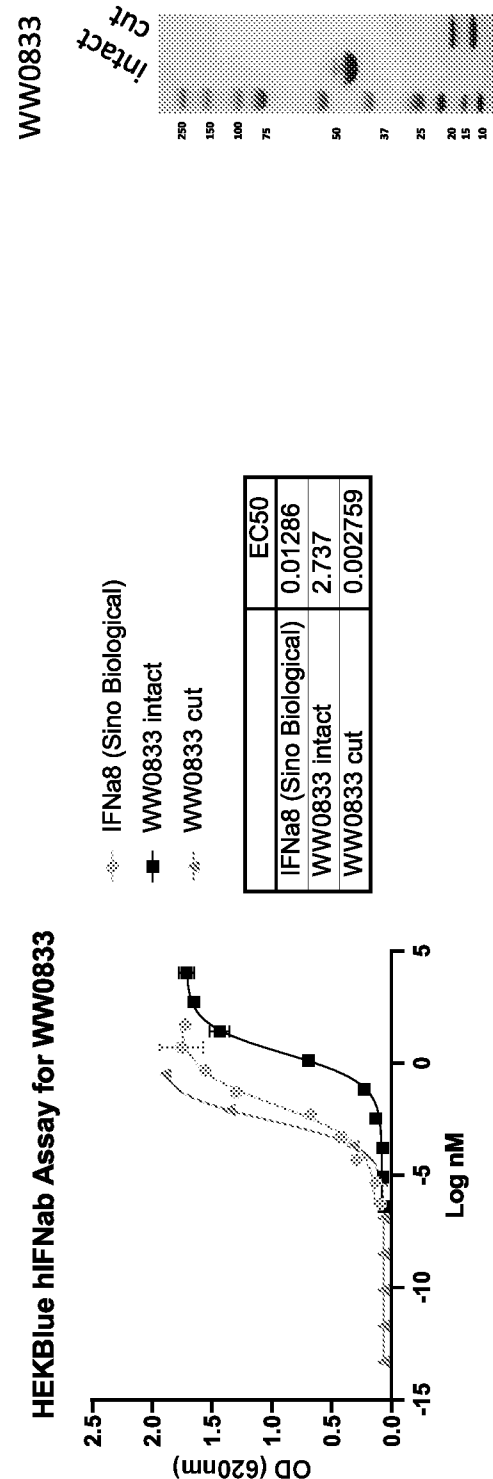


FIG. 37A

B16 Blue mIFN α /b Reporter assay WW0737

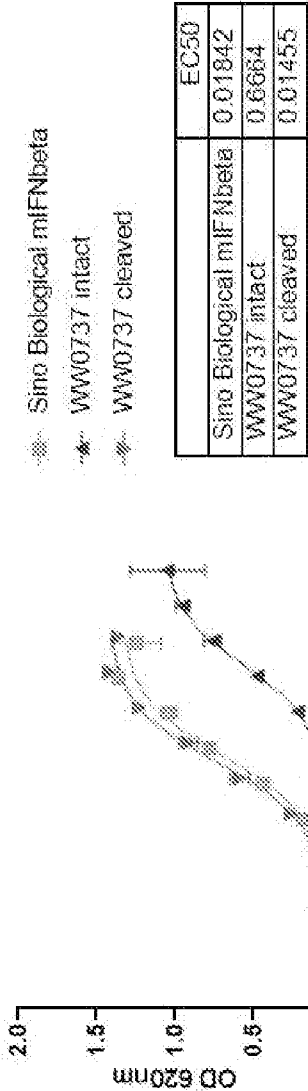


FIG. 37B

B16 Blue mIFN α /b Reporter assay WW0739

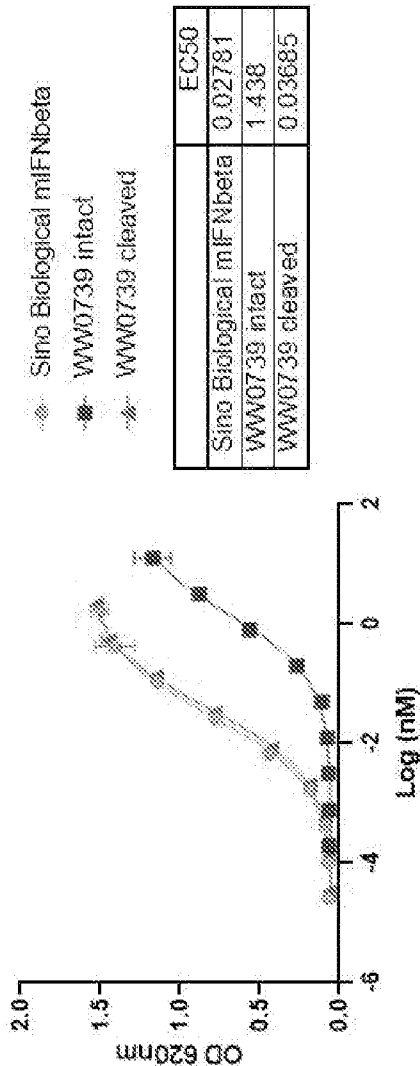


FIG. 37C

HEK-Blue hIFN α /b Reporter Assay Using WW0741

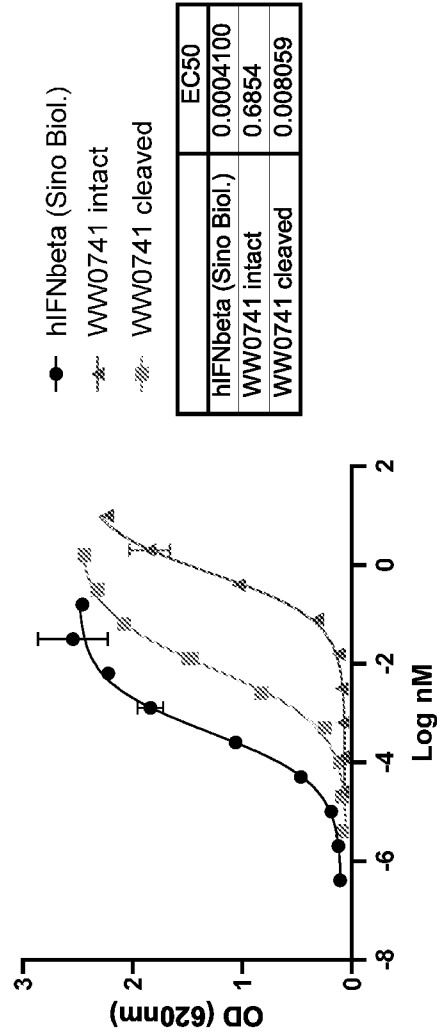


FIG. 37D

HEK-Blue hIFN α /b Reporter Assay Using WW0743

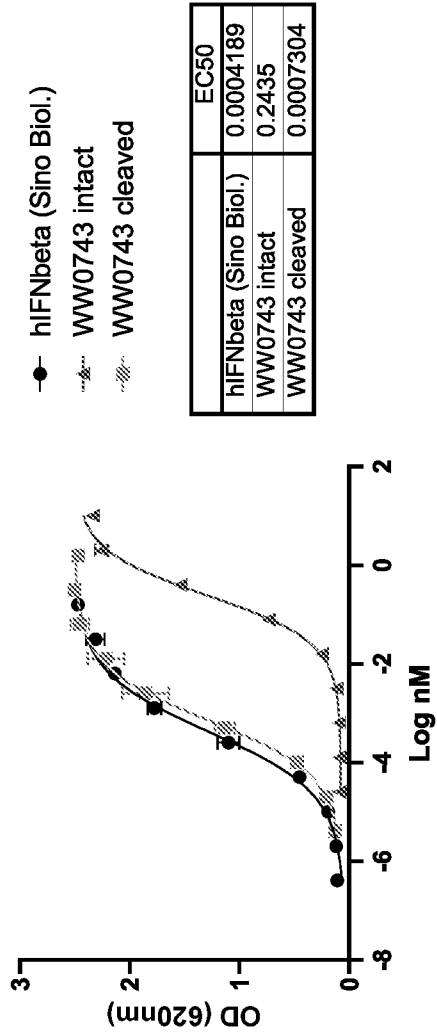
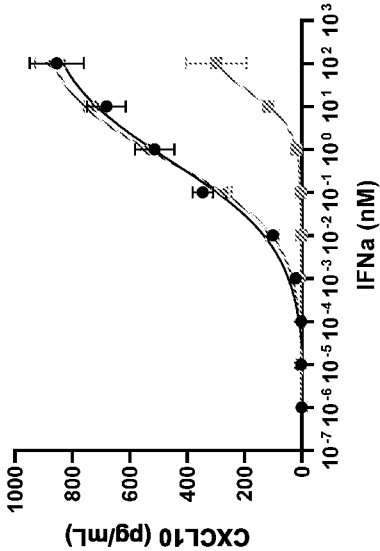


FIG. 38A

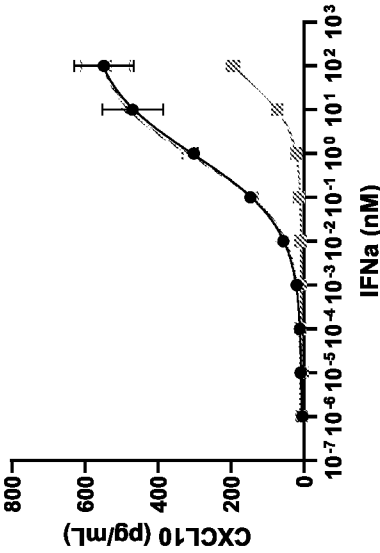
WW0612



WW0635	Intact WW0612	Cut WW0612
EC50	0.5252	21.63
		0.5189

FIG. 38B

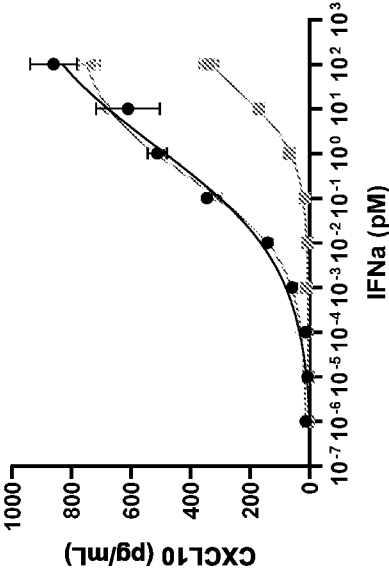
- WW0613



WW0635	Intact WW0613	Cut WW0613
EC50	0.9388	63.63
		0.7485

FIG. 38C

WW0645



WW0648	Intact WW0645	Cut WW0645
EC50	1.438	39.00
		0.2899

FIG. 39A

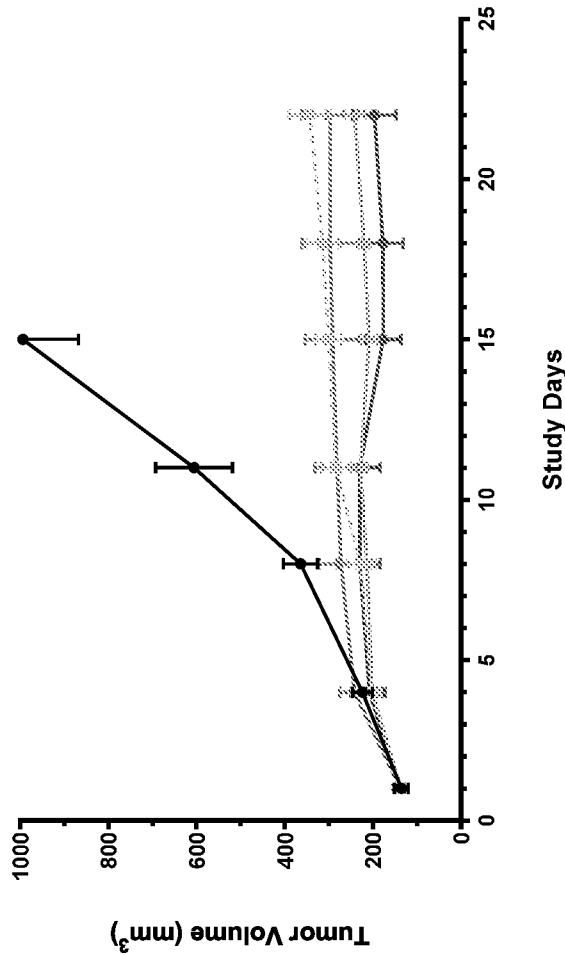


FIG. 39B

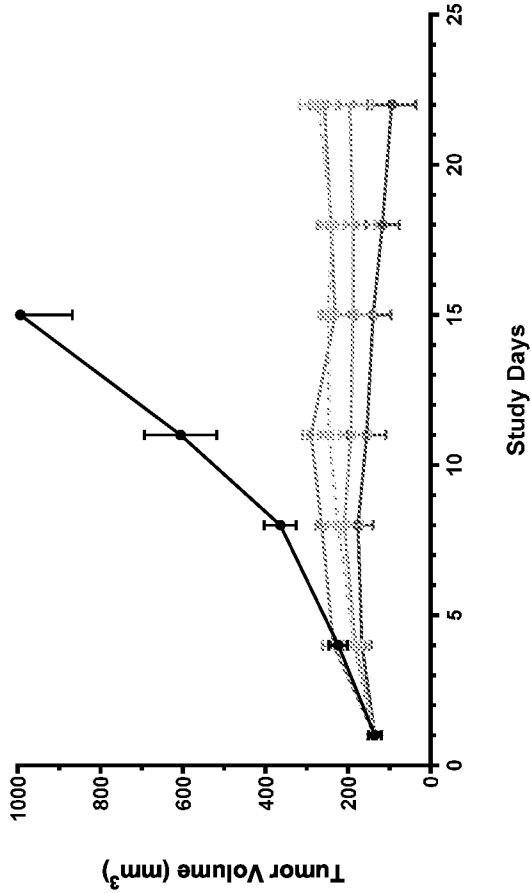
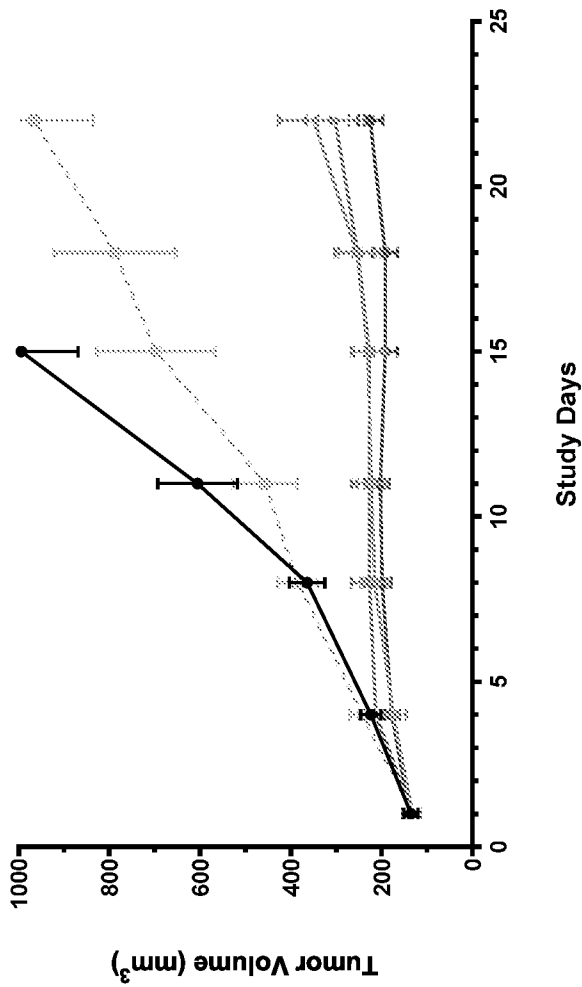


FIG. 39D



- Vehicle (x4)
- 816 (31 µg, x4)
- 816 (62 µg, x4)
- 816 (123 µg, x4)
- 816 (247 µg, x4)

Treatment	Dead	Day
644 (4.6 µg, x4)		
644 (9.3 µg, x4)		
644 (19 µg, x4)		
644 (37 µg, x4)	3	15, 18, 22

- Vehicle (x4)
- 644 (4.6 µg, x4)
- 644 (9.3 µg, x4)
- 644 (19 µg, x4)
- 644 (37 µg, x4)

FIG. 39C

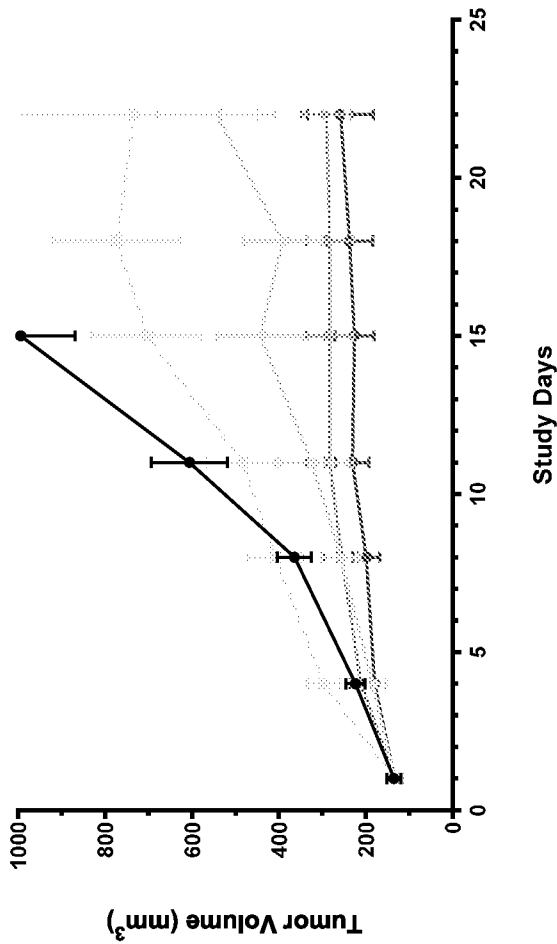


FIG. 39E

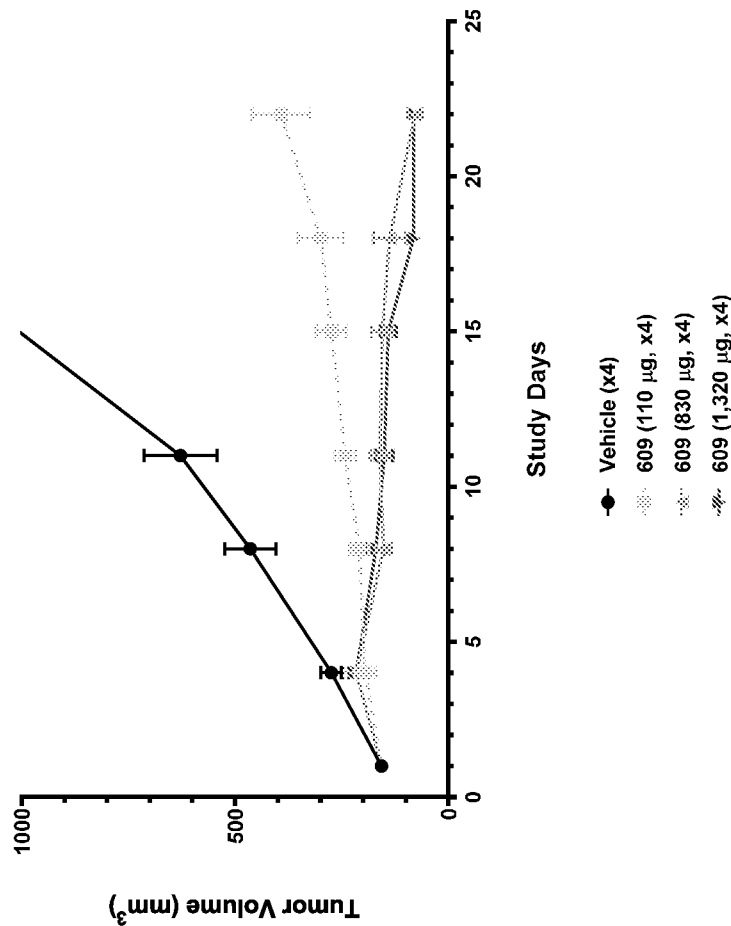


FIG. 39F

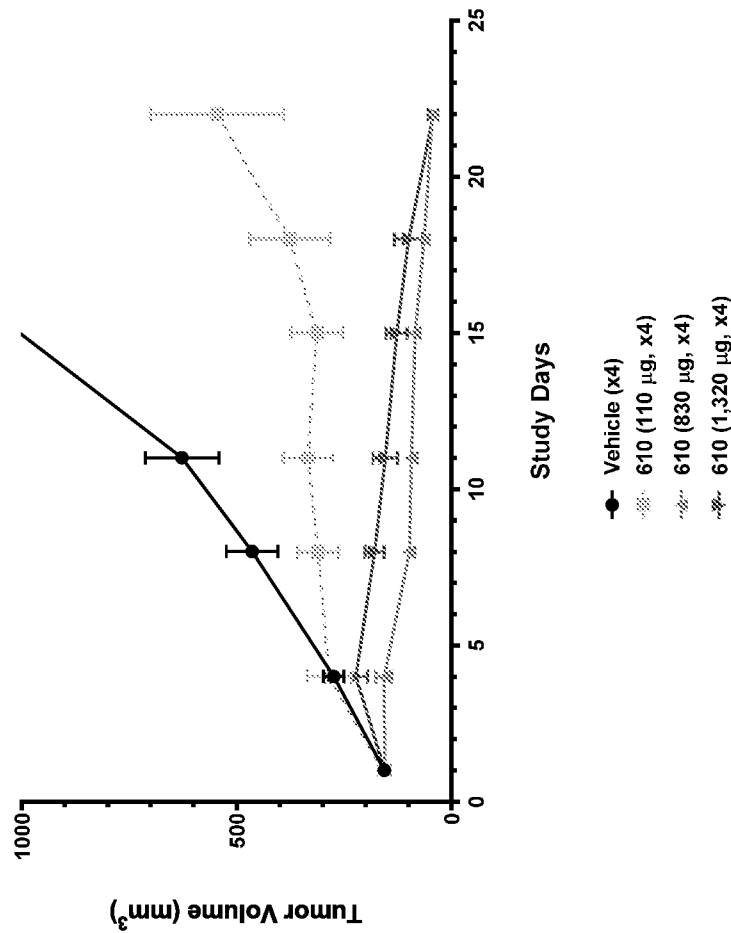


FIG. 39G

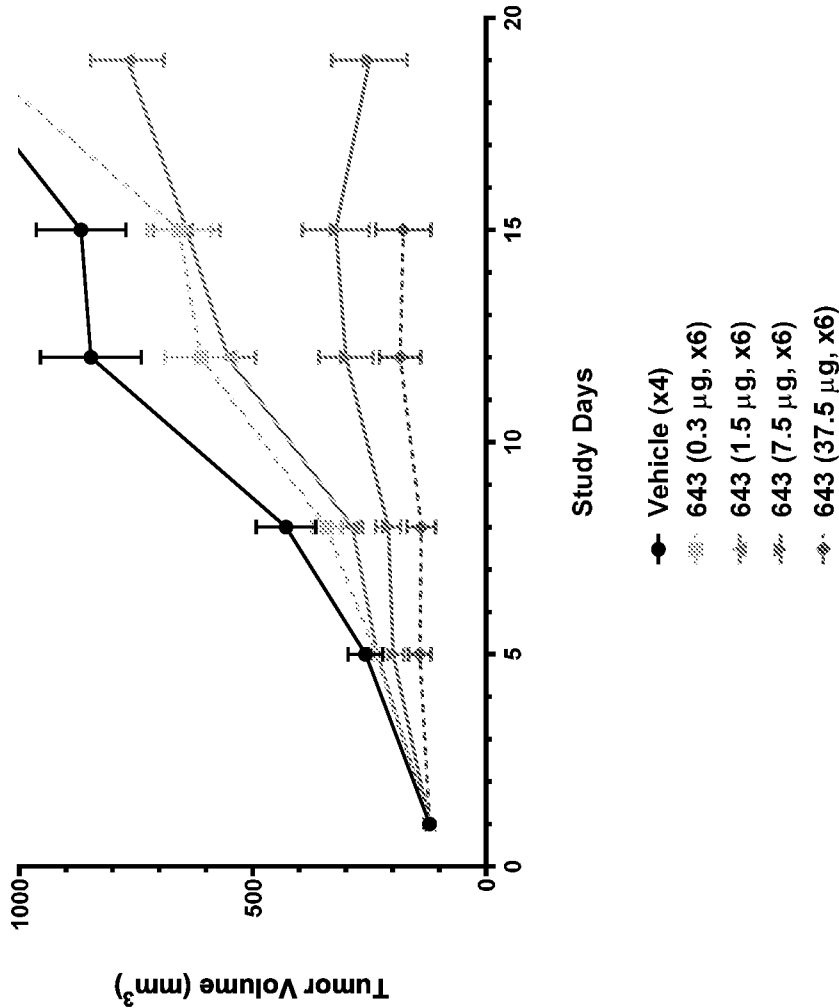


FIG. 40A

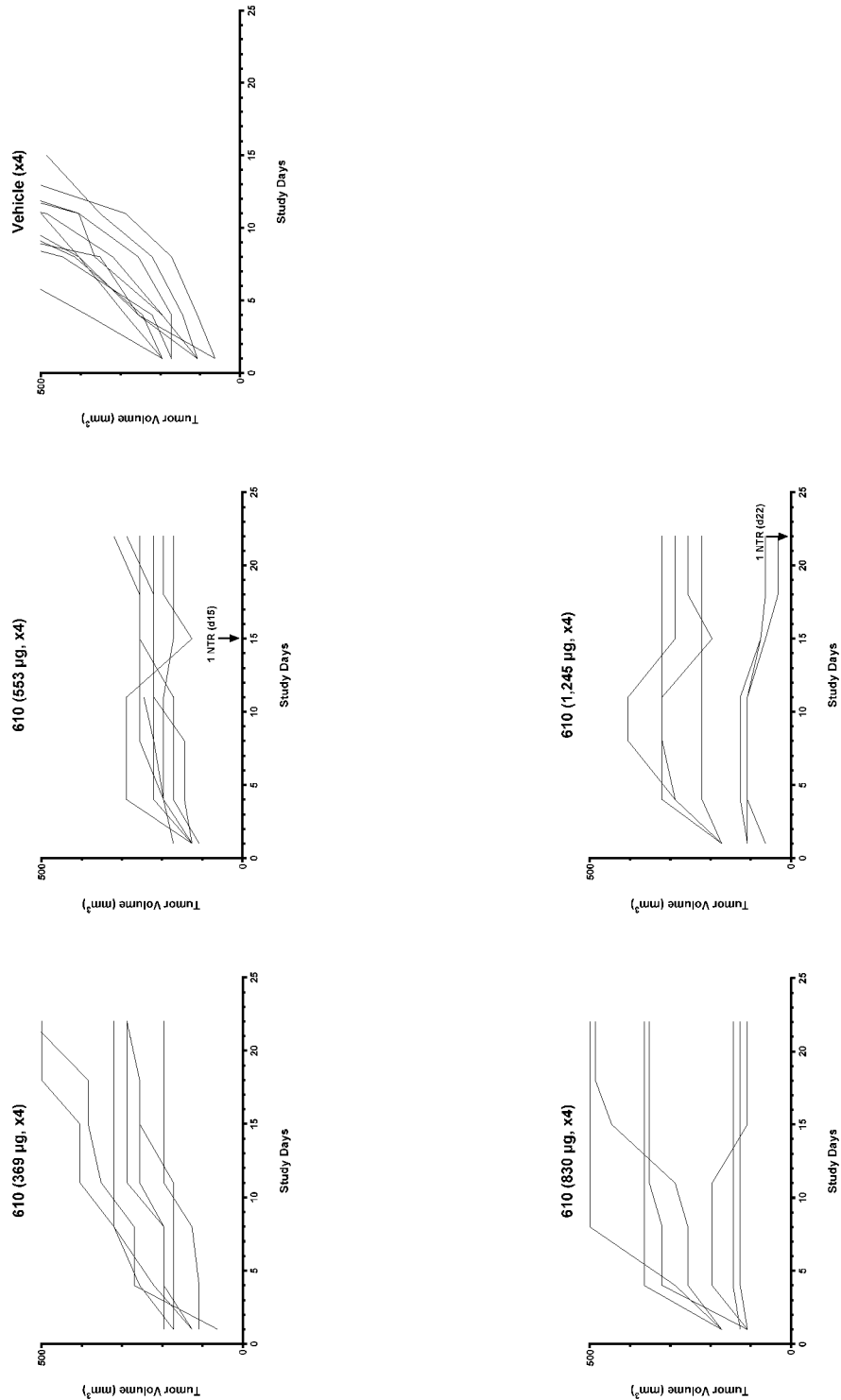


FIG. 40B

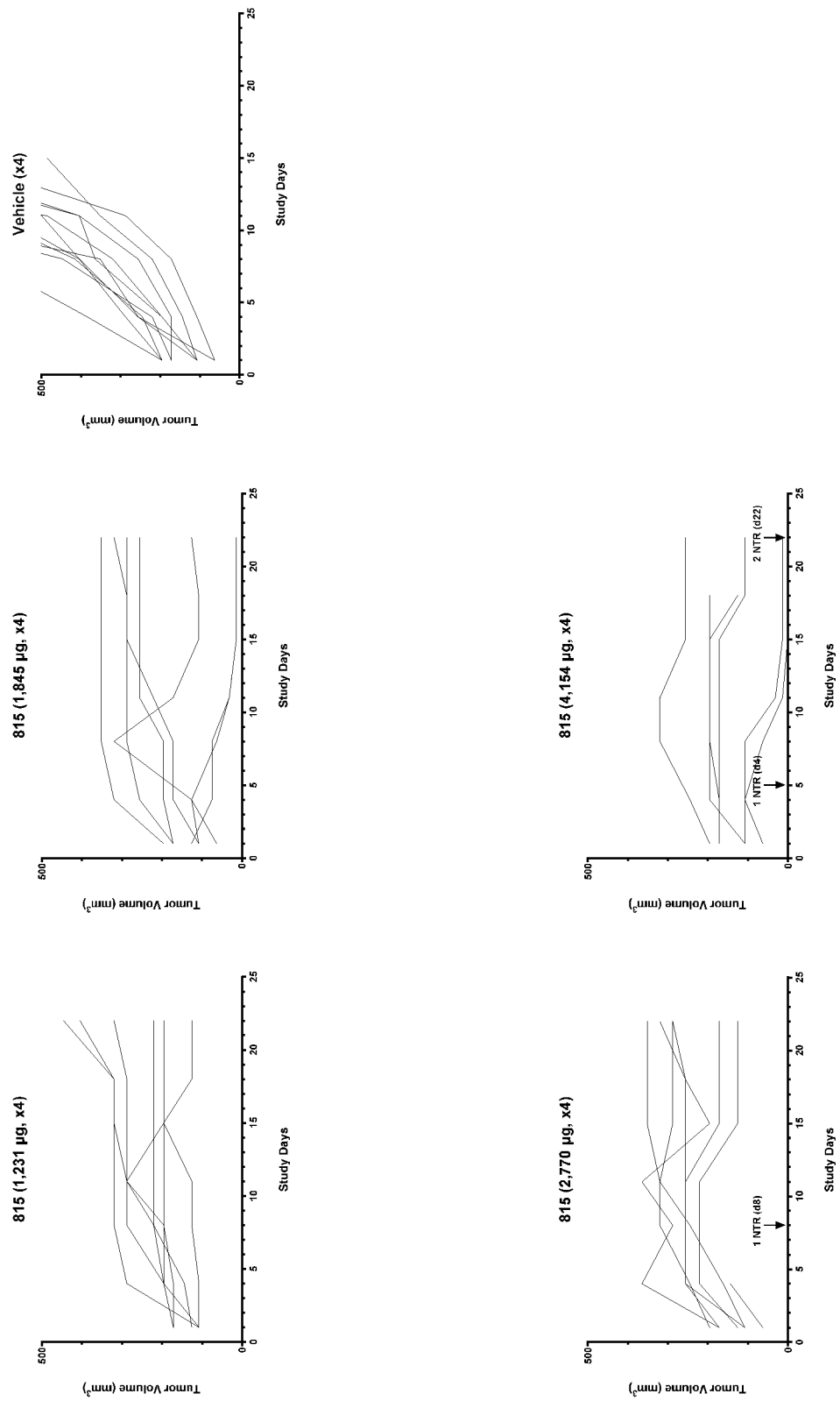


FIG. 40C

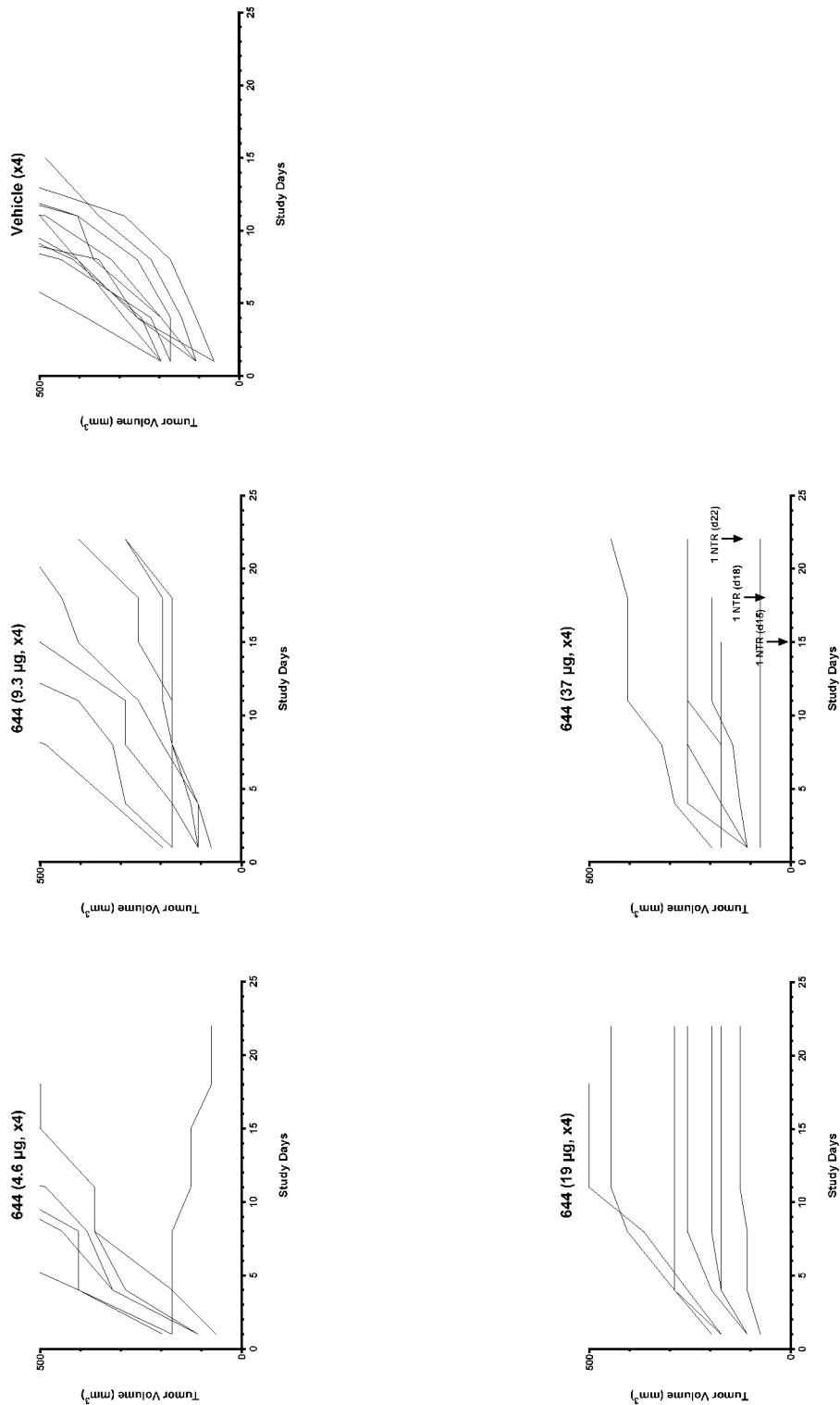


FIG. 40D

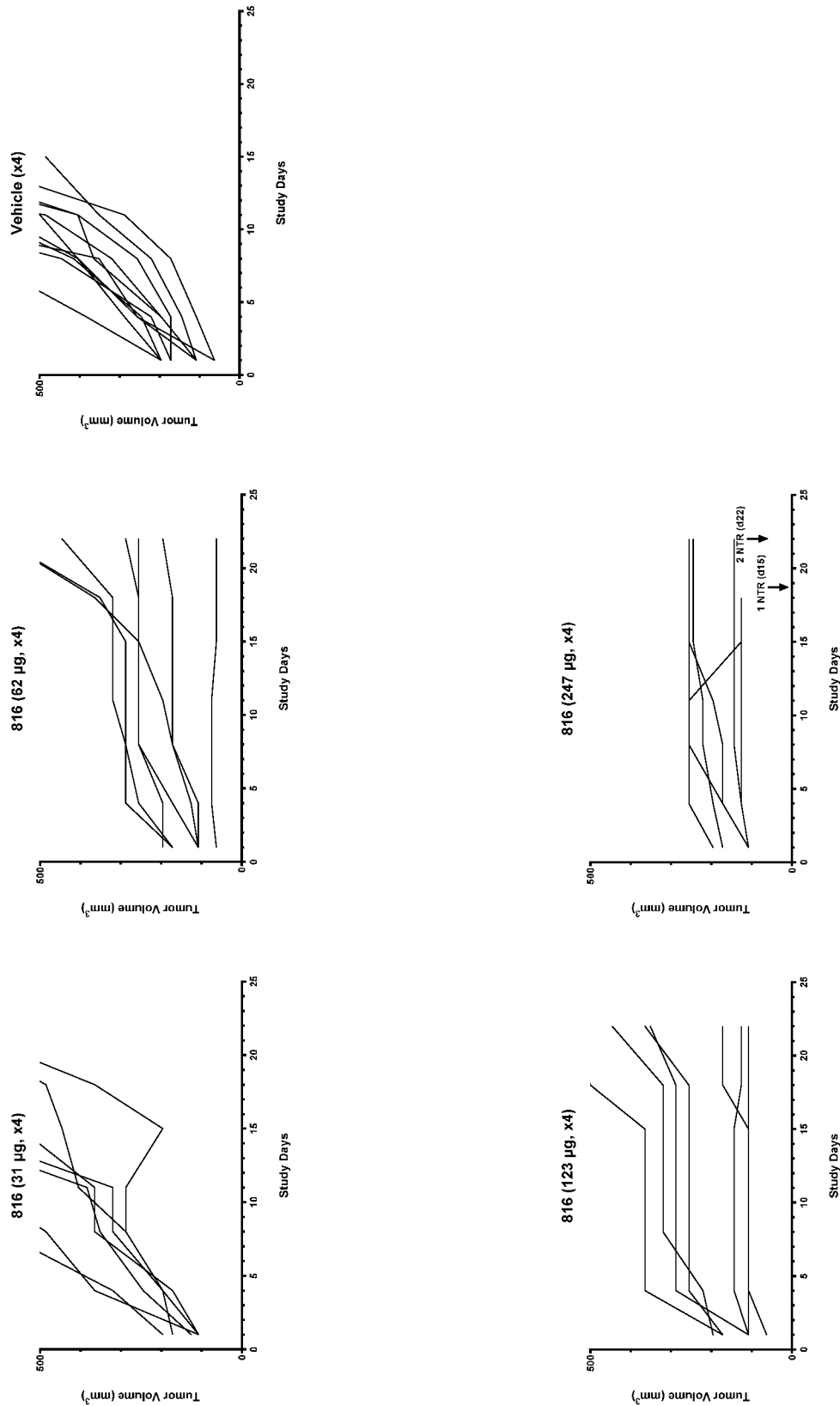


FIG. 40E

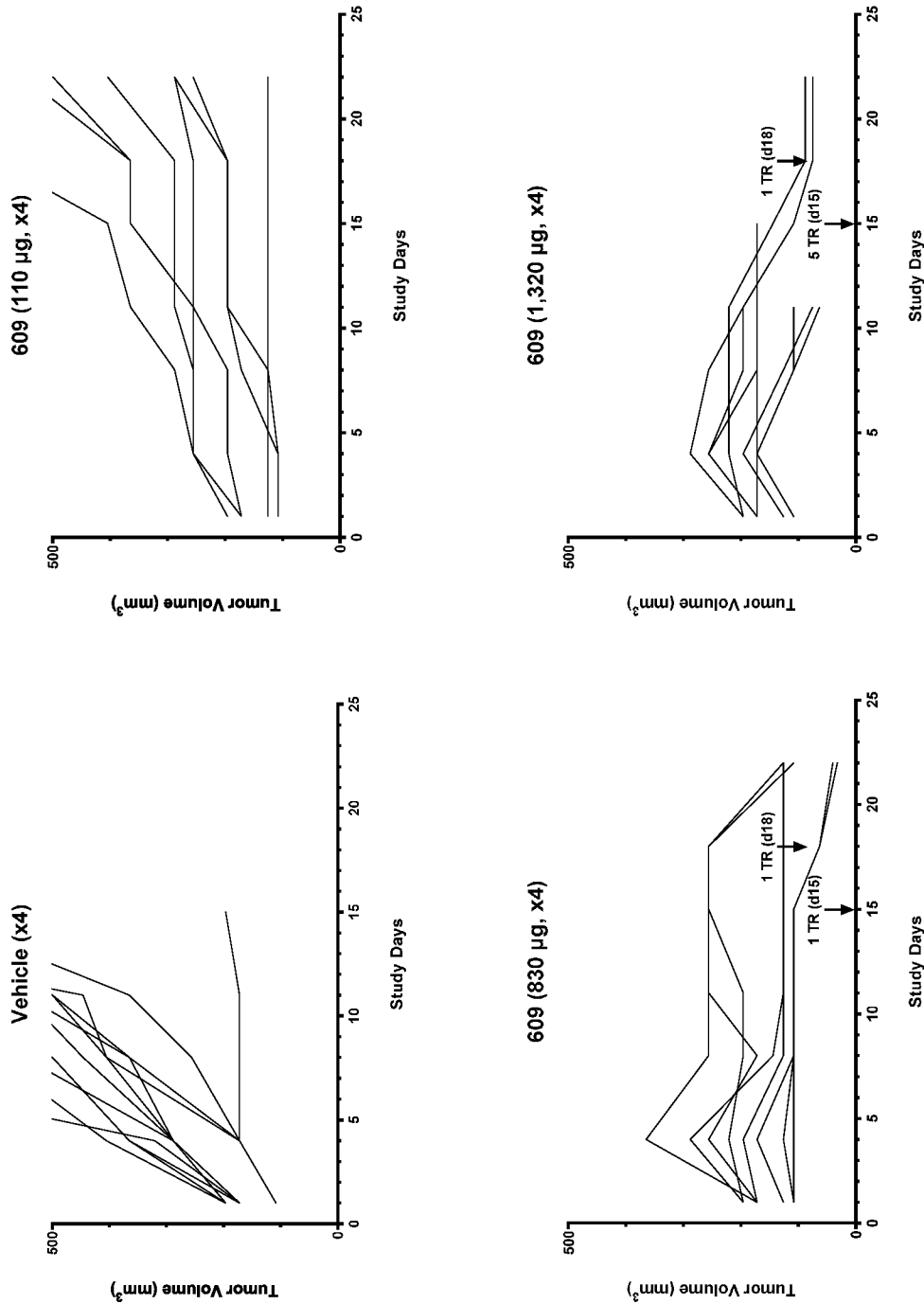


FIG. 40F

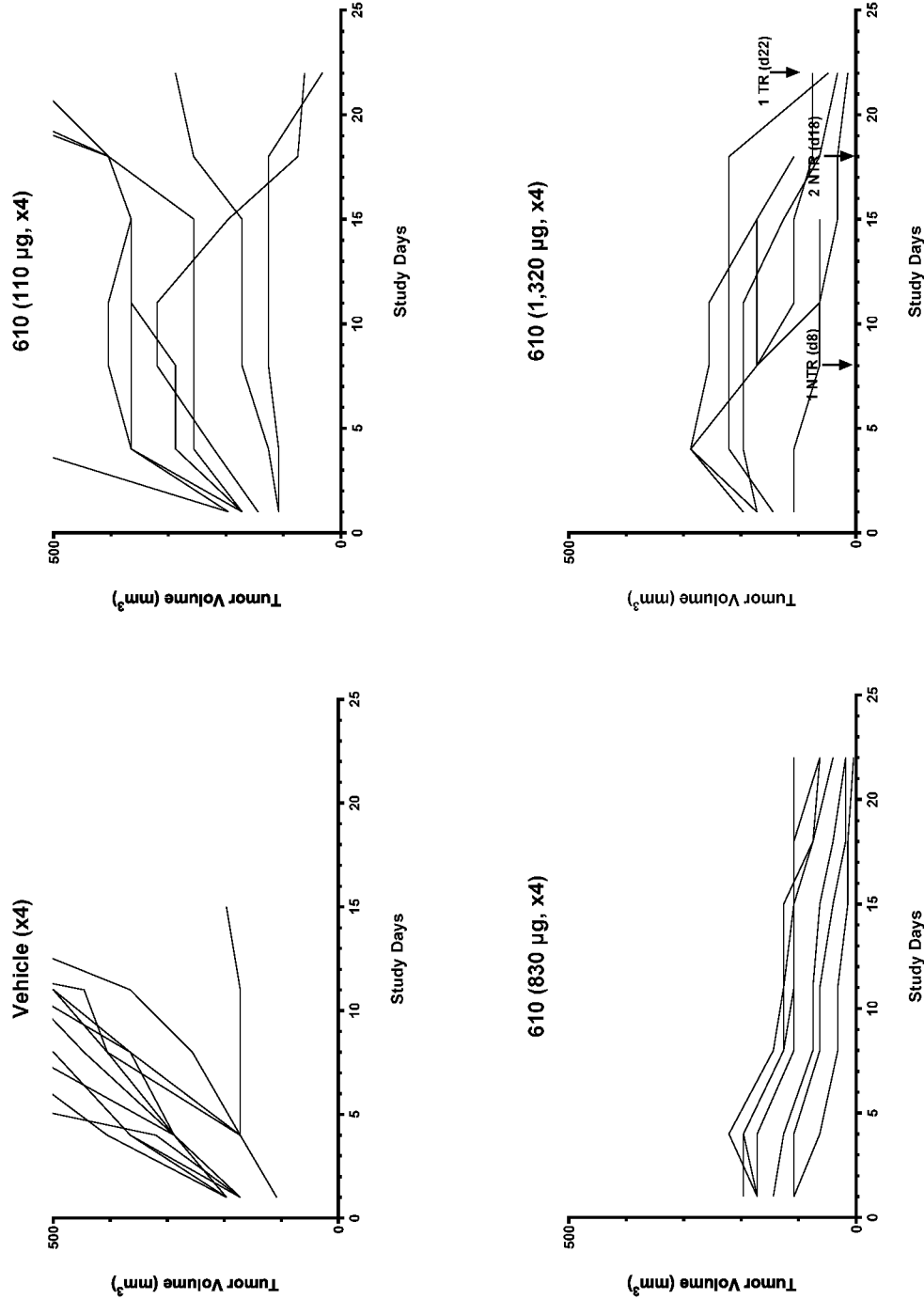


FIG. 40G

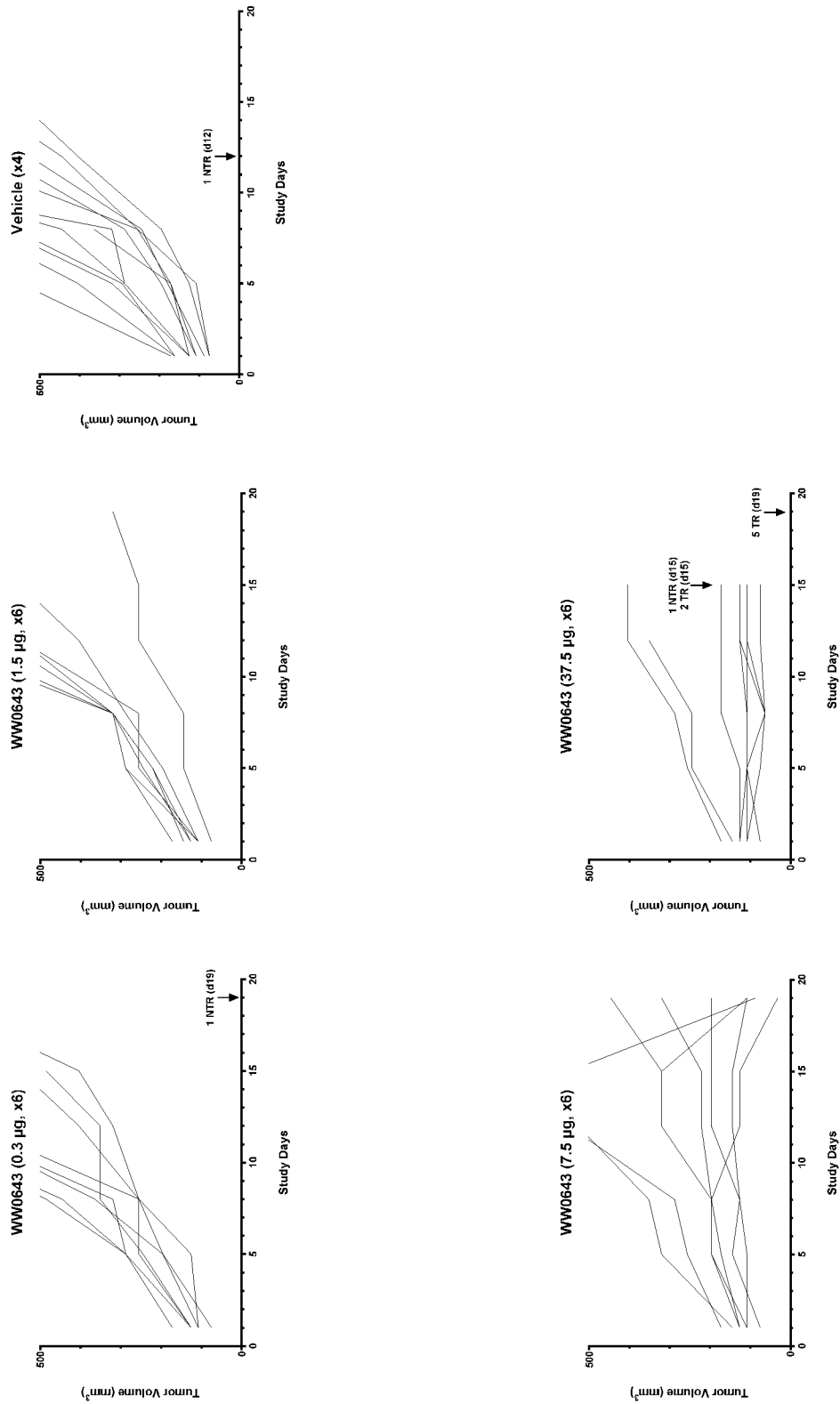


FIG. 41B

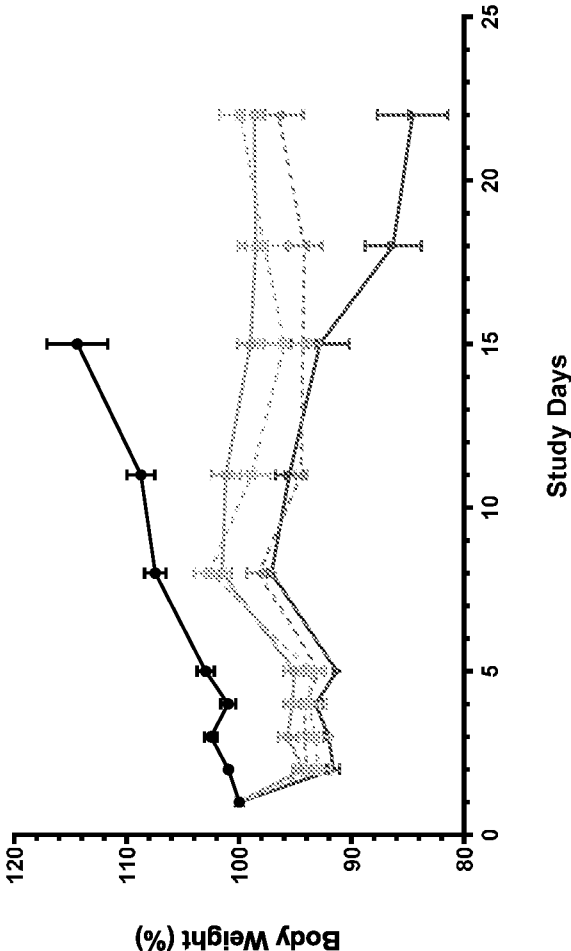


FIG. 41A

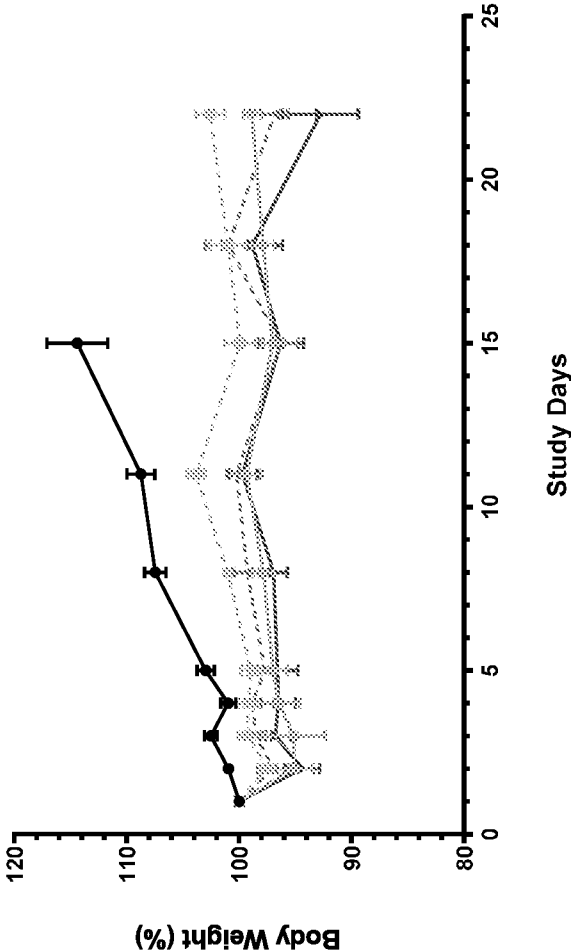
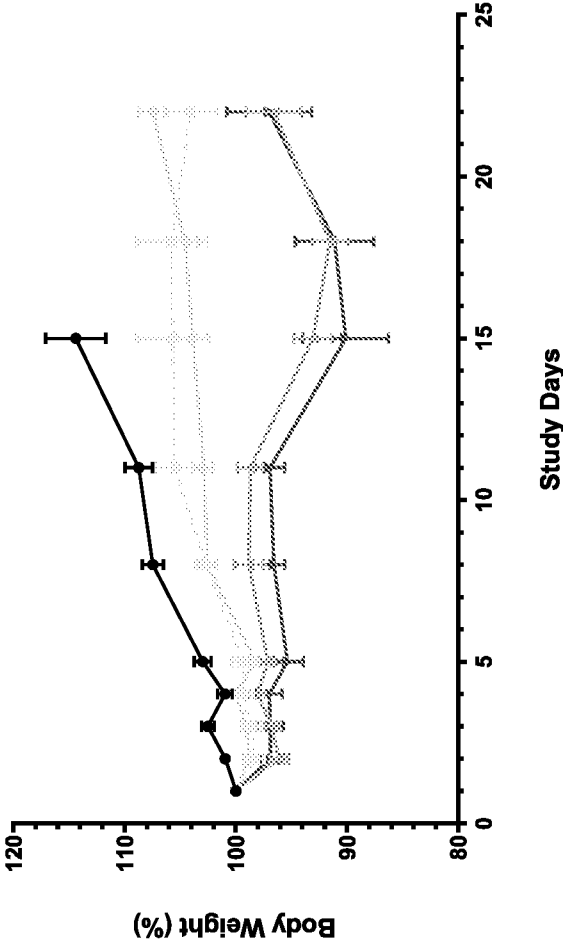


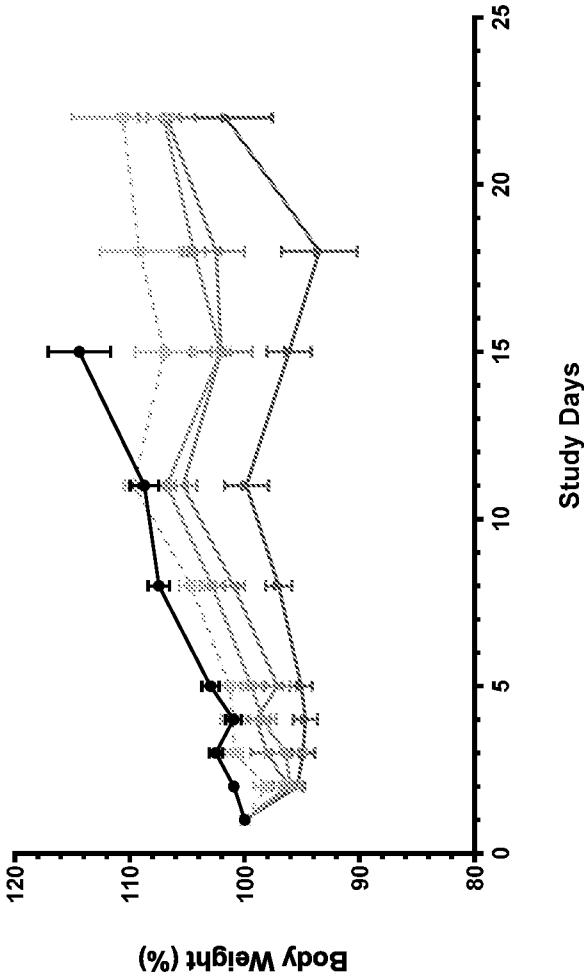
FIG. 41C



- Vehicle (x4)
- 644 (4.6 μg, x4)
- 644 (9.3 μg, x4)
- △ 644 (19 μg, x4)
- ◇ 644 (37 μg, x4)

Treatment	Dead	Day
644 (4.6 μg, x4)		
644 (9.3 μg, x4)		
644 (19 μg, x4)		
644 (37 μg, x4)	3	15, 18, 22

FIG. 41D



- Vehicle (x4)
- 816 (31 μg, x4)
- 816 (62 μg, x4)
- △ 816 (123 μg, x4)
- ◇ 816 (247 μg, x4)

Treatment	Dead	Day
816 (31 μg, x4)		
816 (62 μg, x4)		
816 (123 μg, x4)		
816 (247 μg, x4)	3	15, 22, 22

FIG. 41E

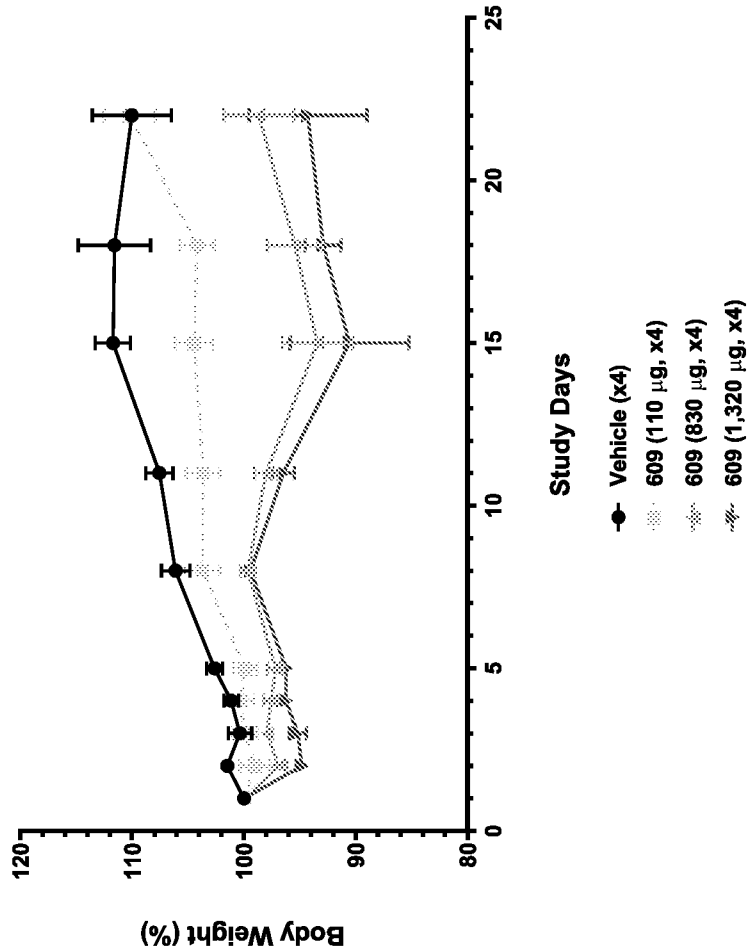


FIG. 41F

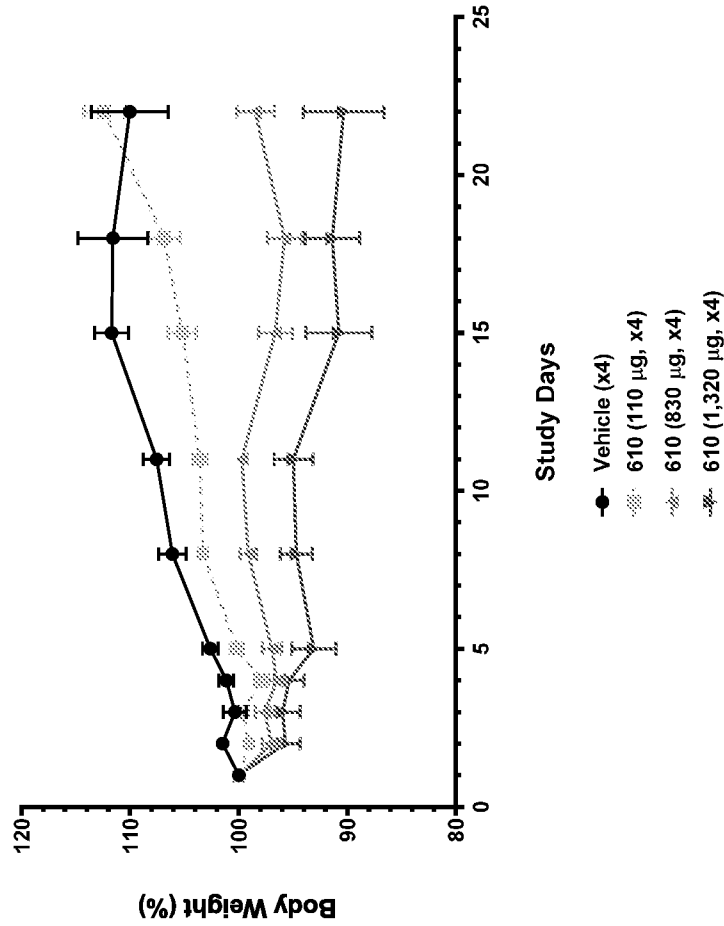
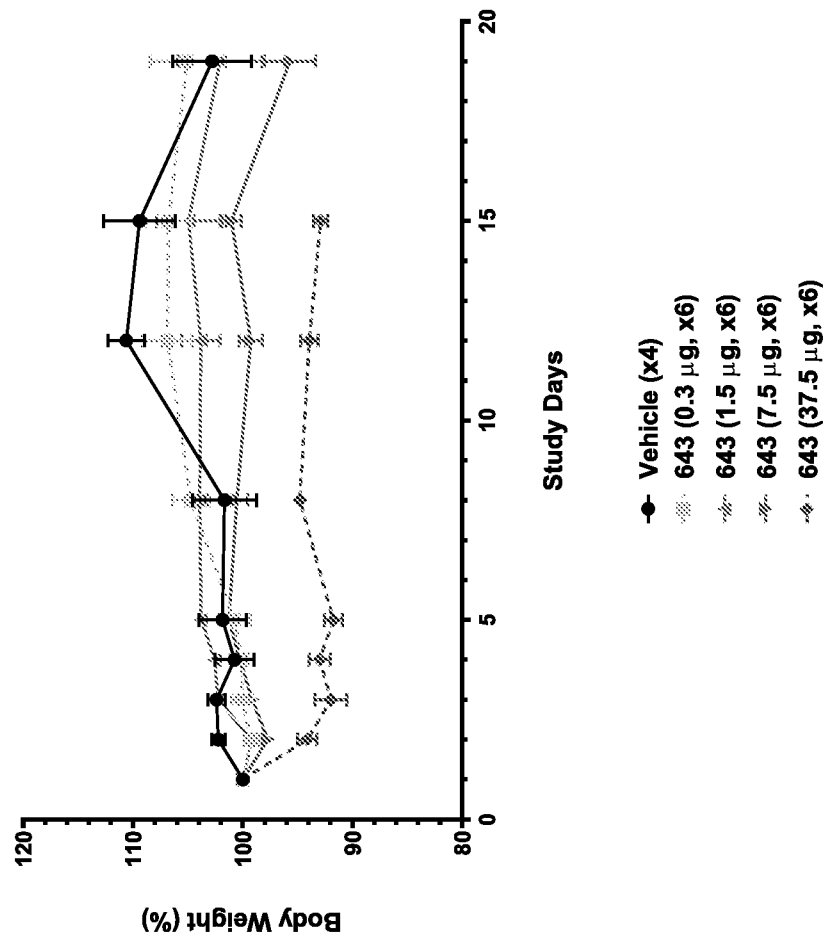


FIG. 41G



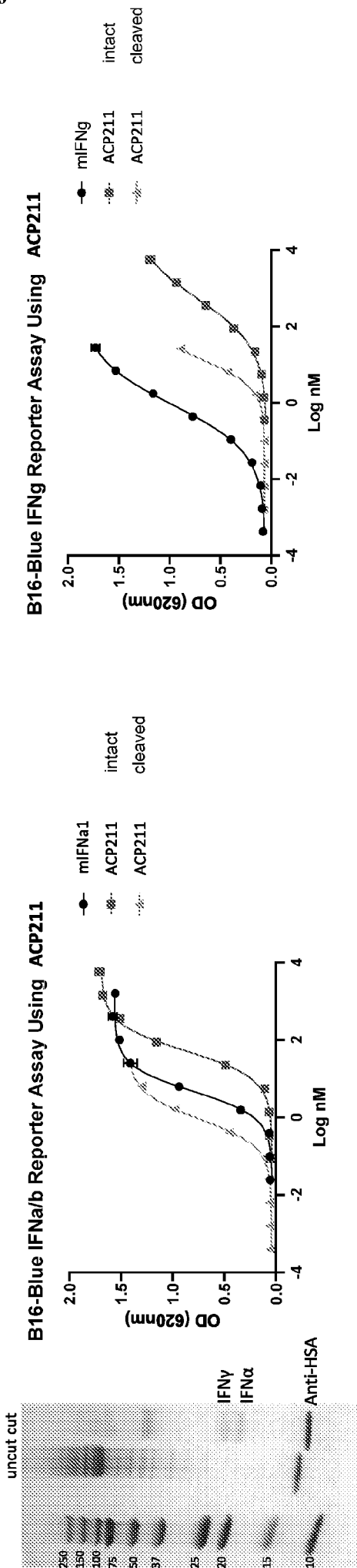


FIG. 42A

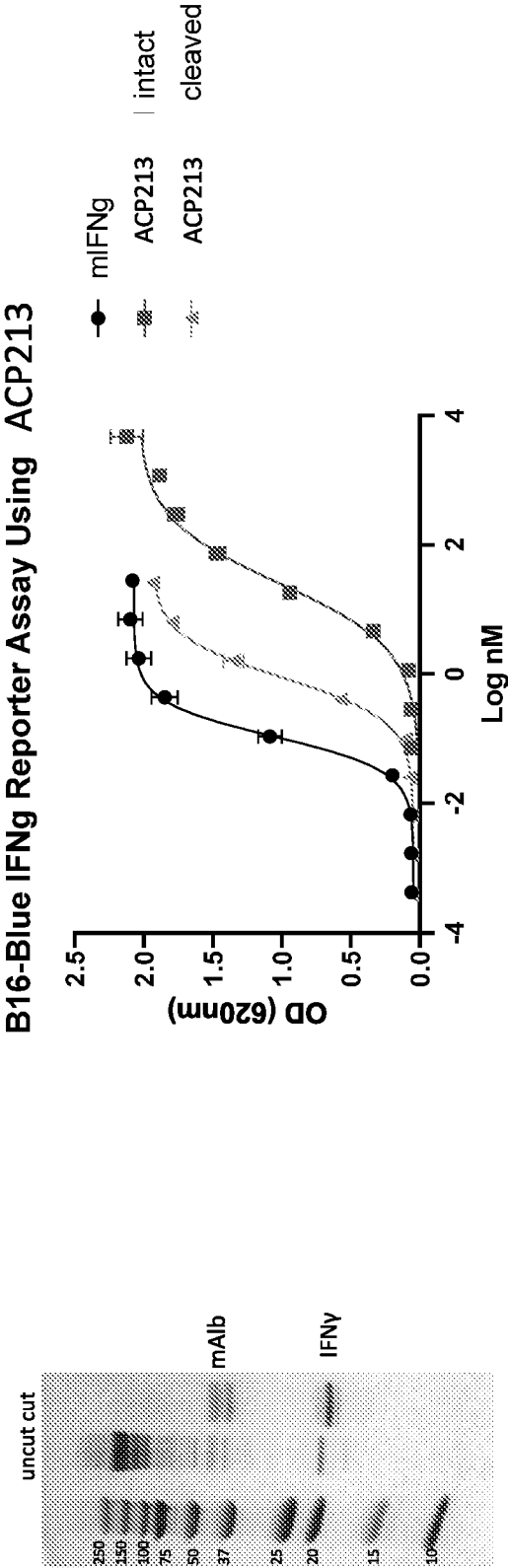


FIG. 42B

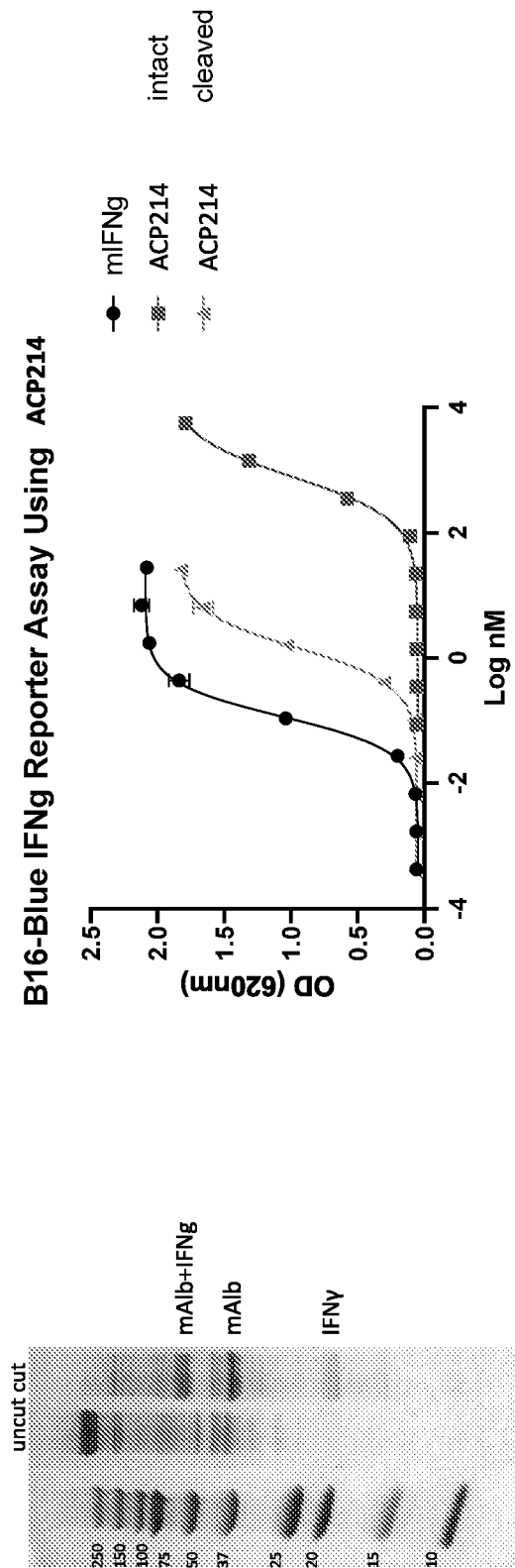


FIG. 42C

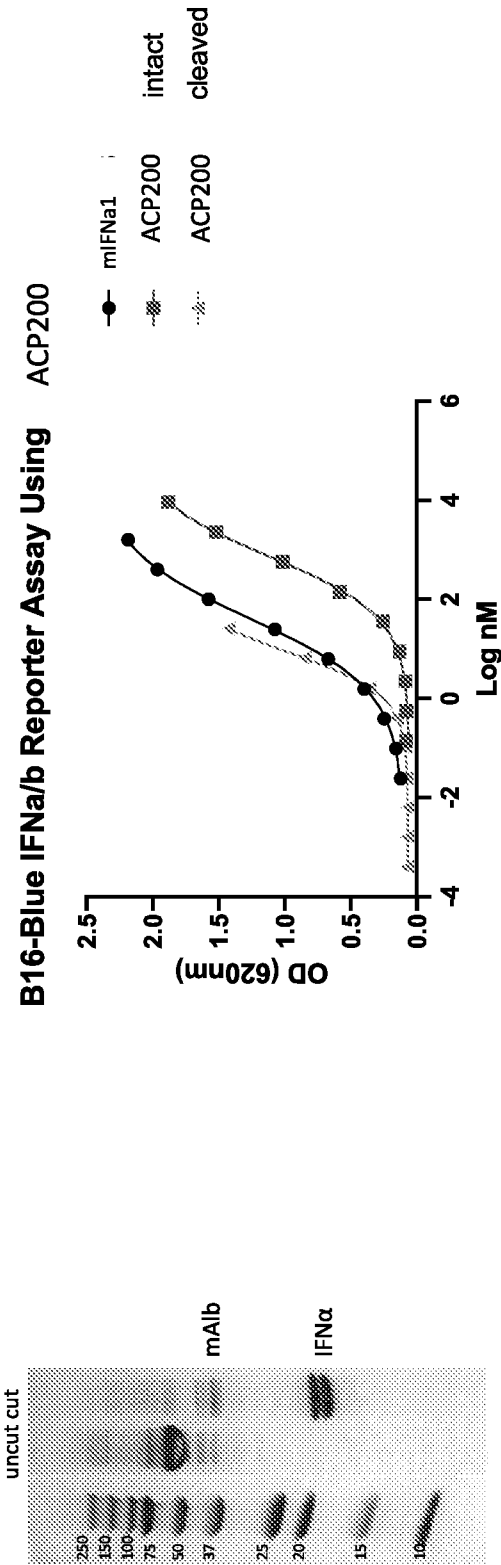


FIG. 42D

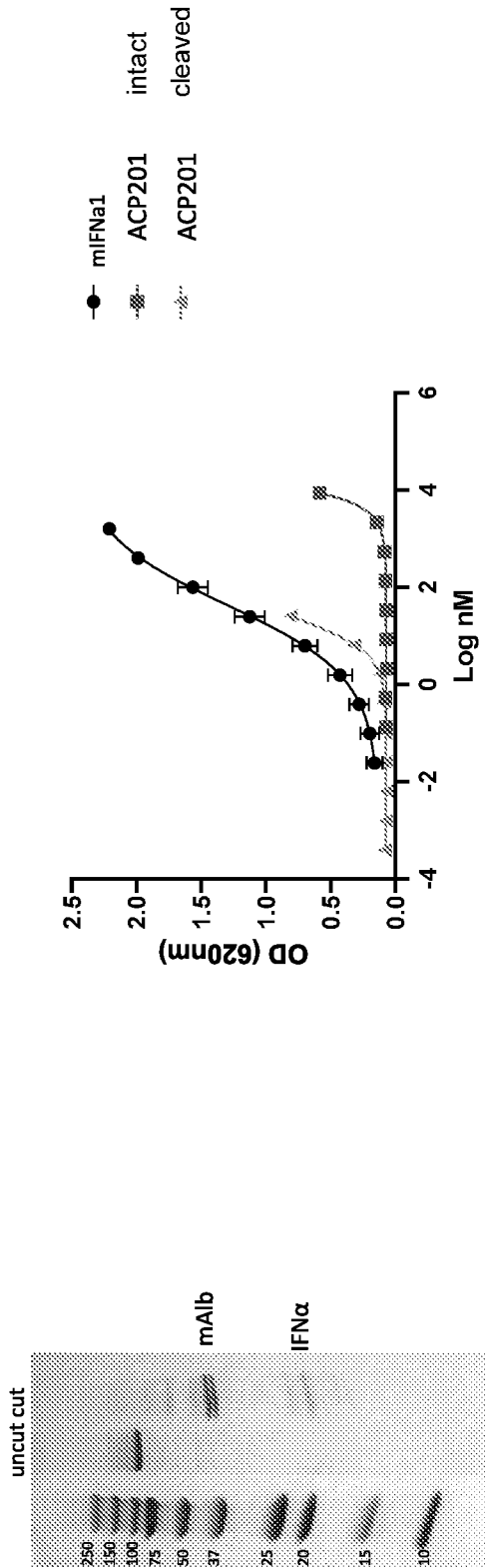


FIG. 42E

Mouse Serum Albumin	KD (nM)	kon(1/Ms)	kdis(1/s)
ACP16	2.6	4.3E+4	1.1E-4
ACP10	0.38	2.5E+5	9.7E-5
ACP11	No binding	No binding	No binding

Human Serum Albumin	KD (nM)	kon(1/Ms)	kdis(1/s)
ACP16	4.3	3.4E+5	1.5E-3
ACP10	4.4	2.8E+5	1.2E-3
ACP11	79	3.1E+4	2.5E-3

FIG. 43

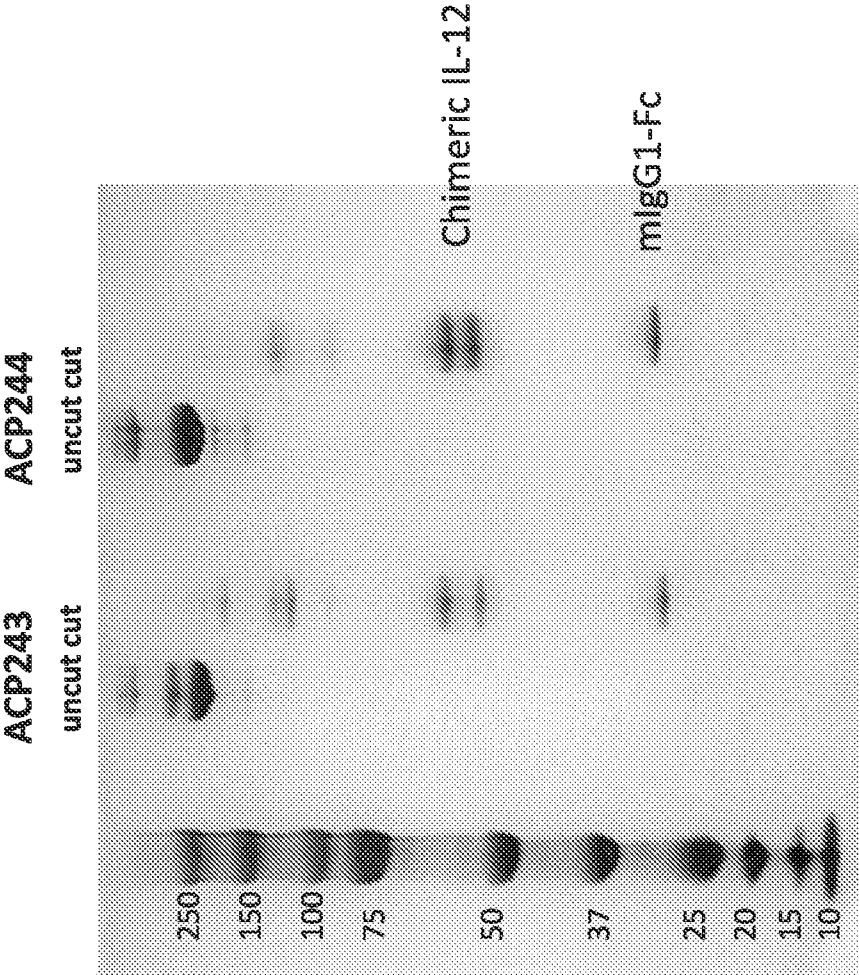
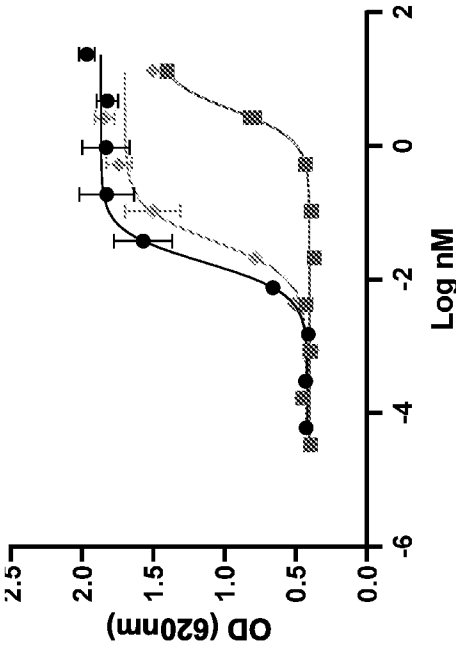
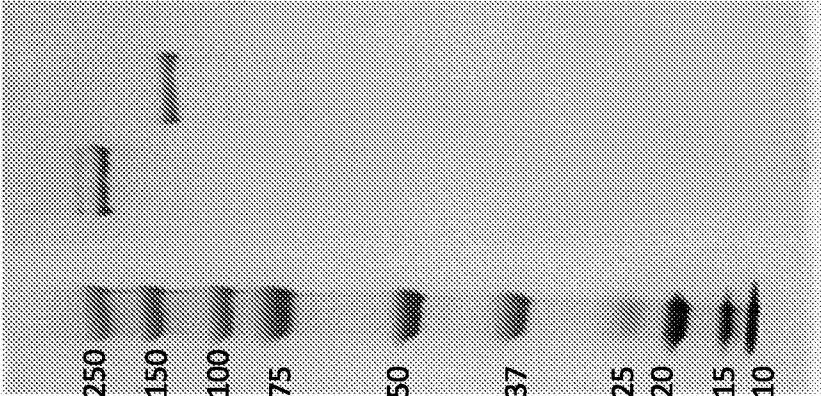


FIG. 44A

NR R



WW0205 (human p40/mouse p35)
IL12 (human p40/mouse p35)
ACP243 intact
ACP243 cleaved

FIG. 44B

NR R

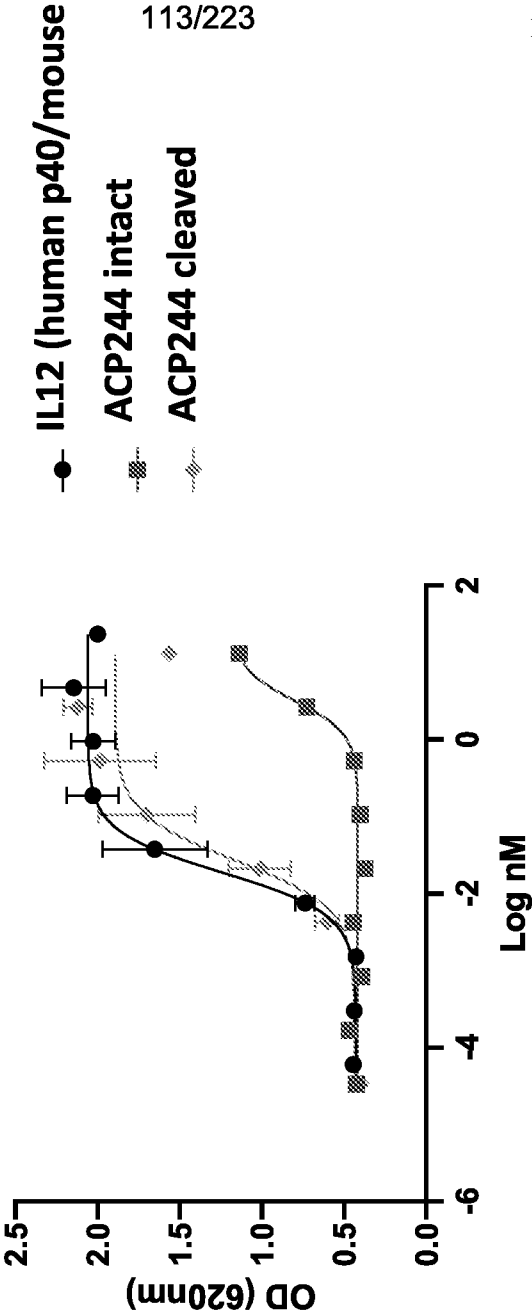
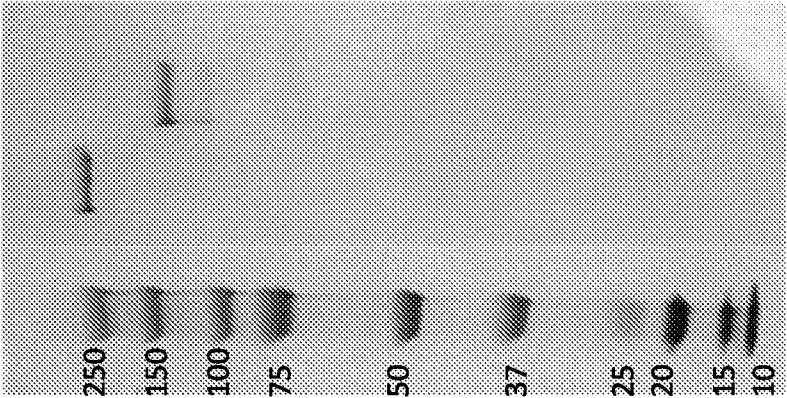
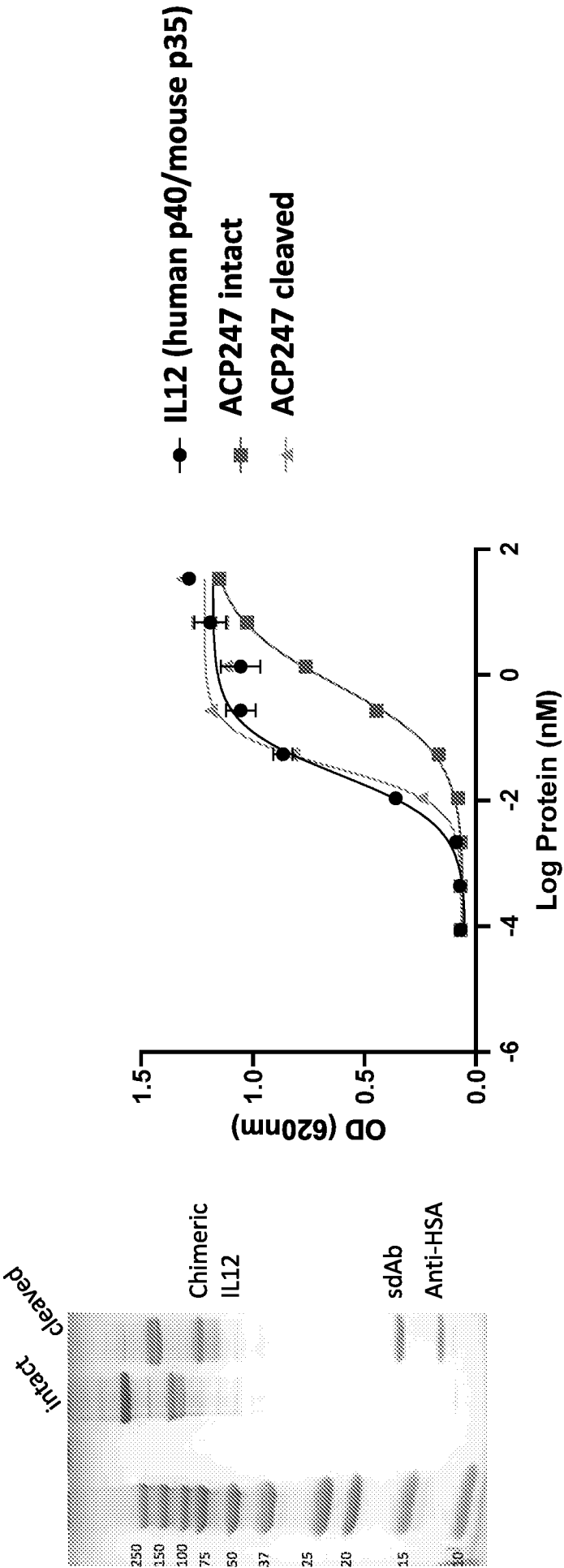


FIG. 44C

FIG. 44D



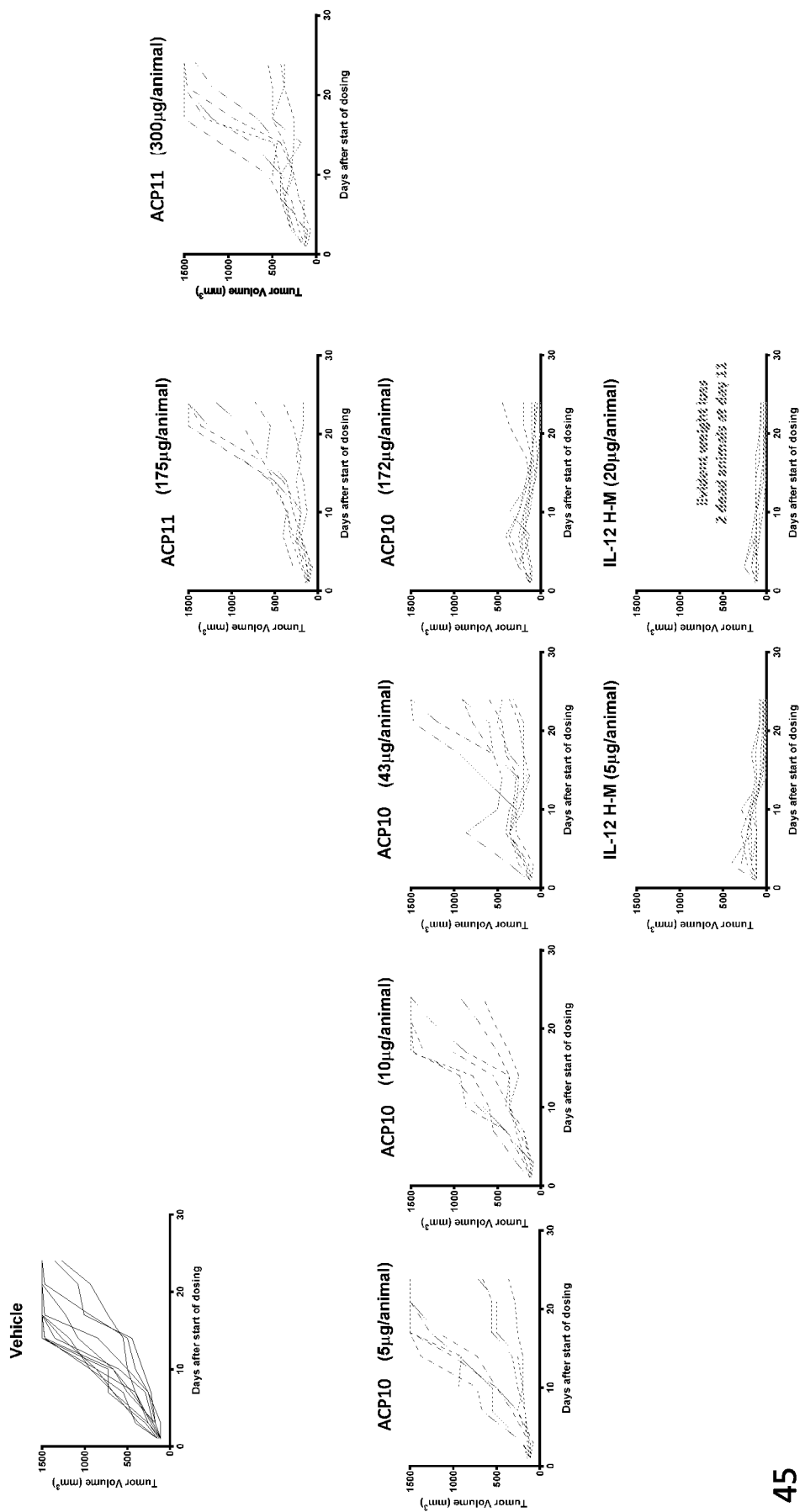


FIG. 45

FIG. 46A

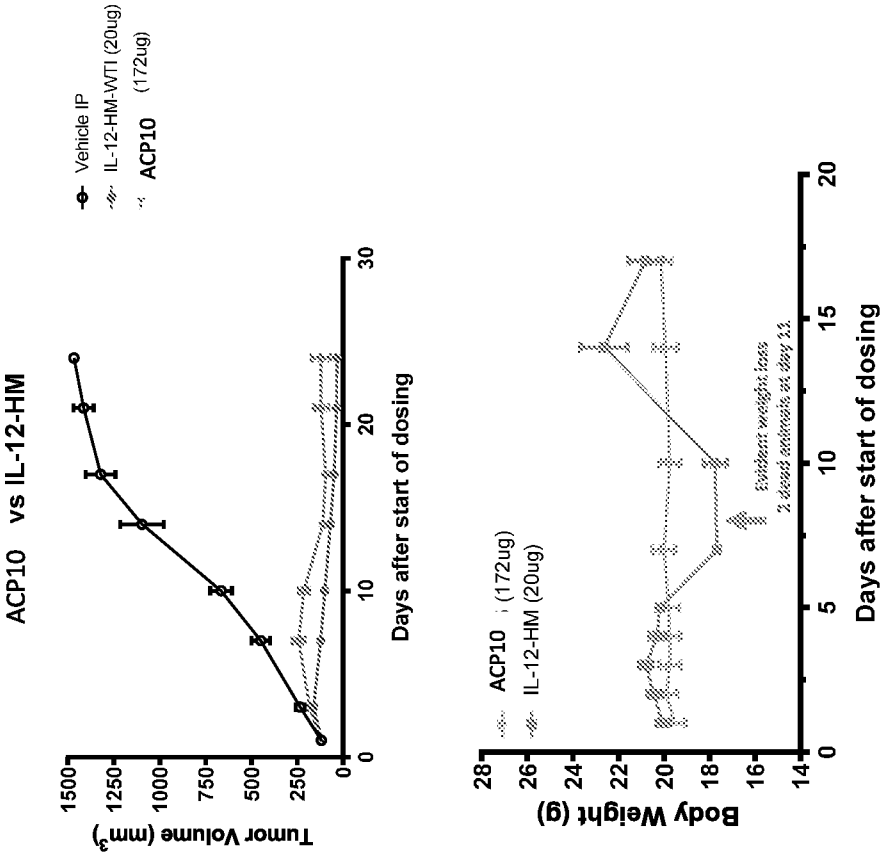


FIG. 46B

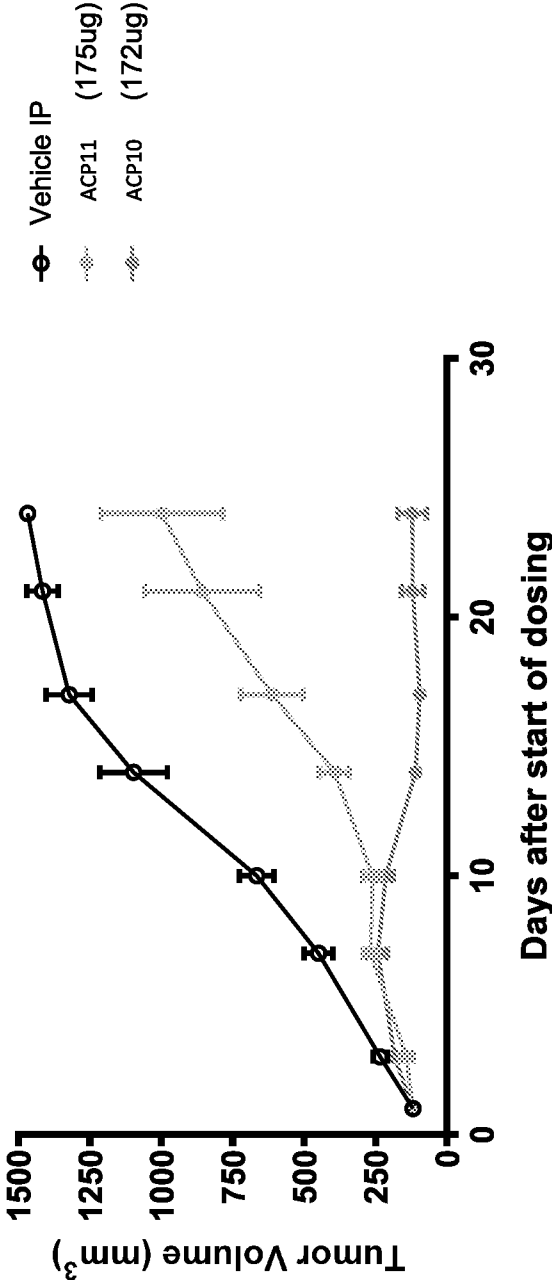


FIG. 46C

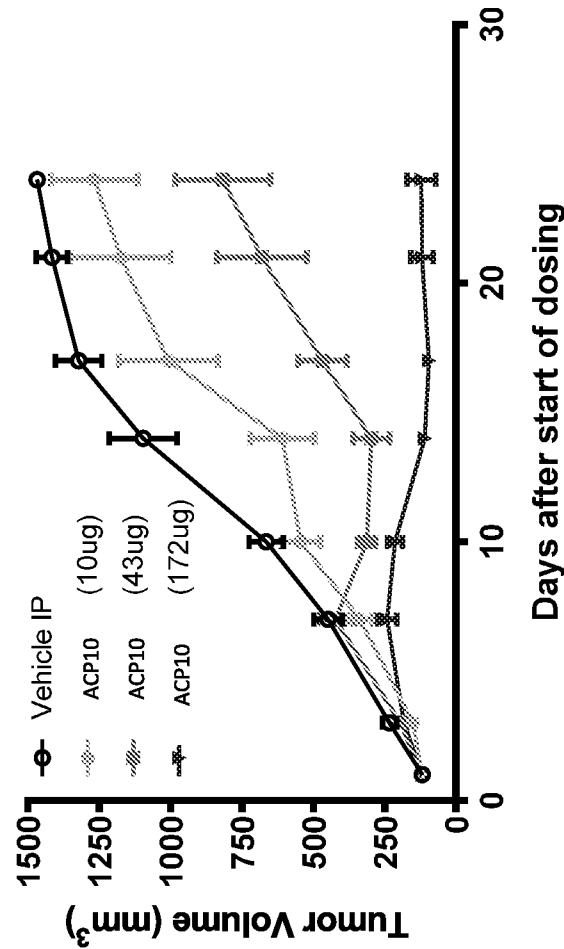


FIG. 46D

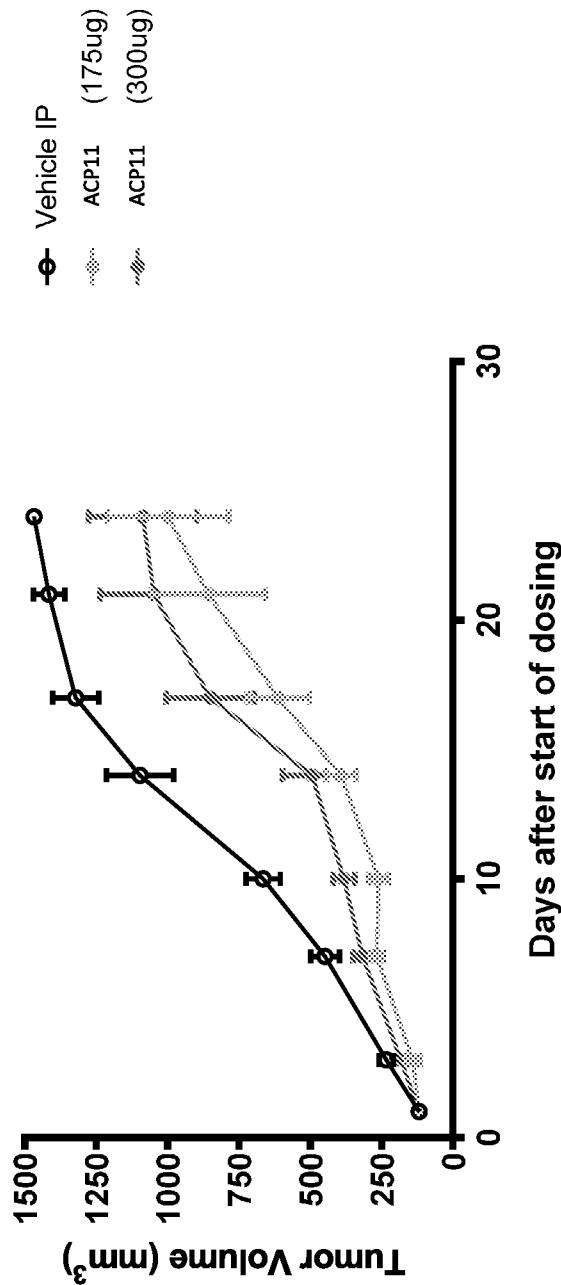
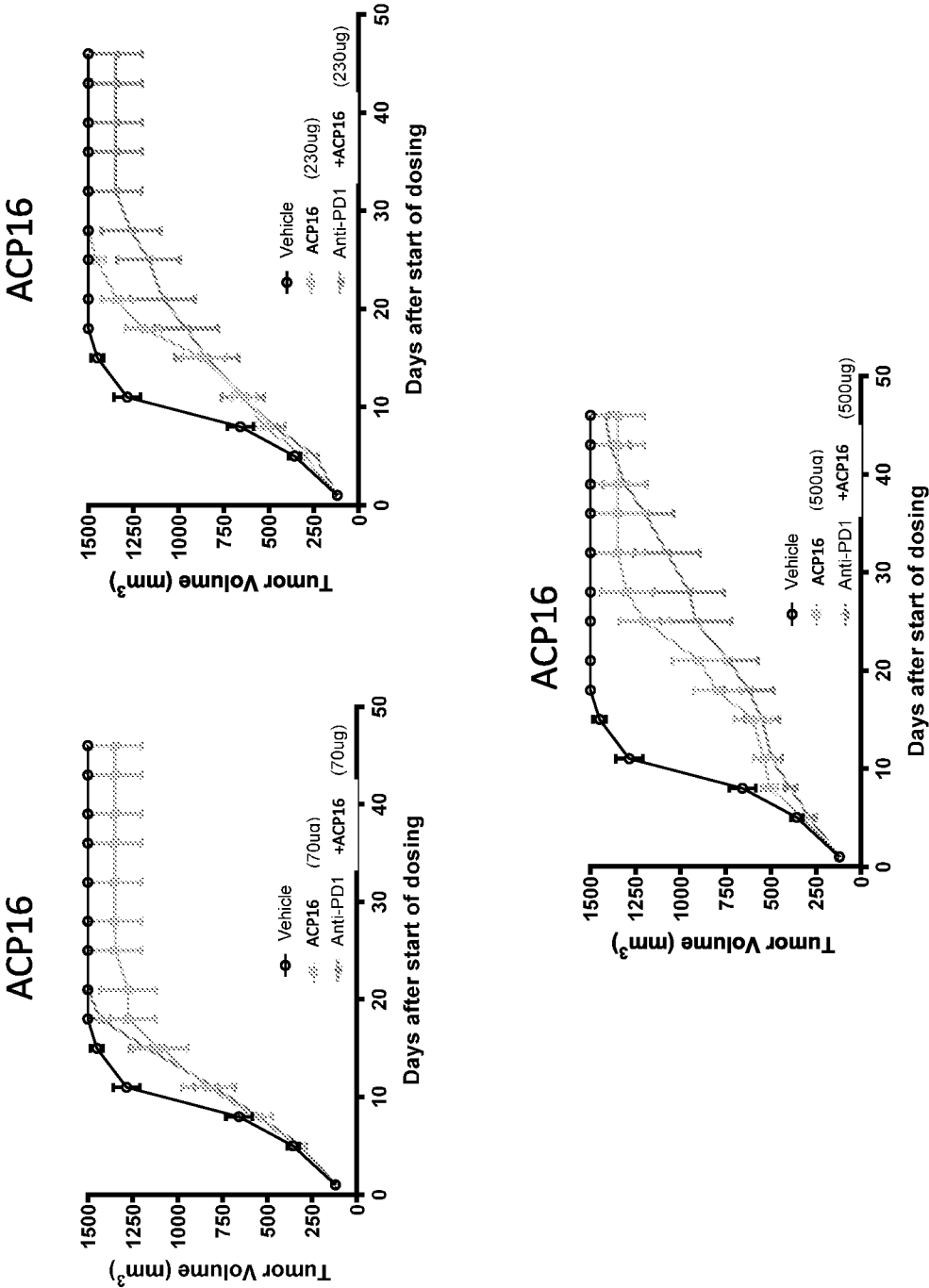


FIG. 47A



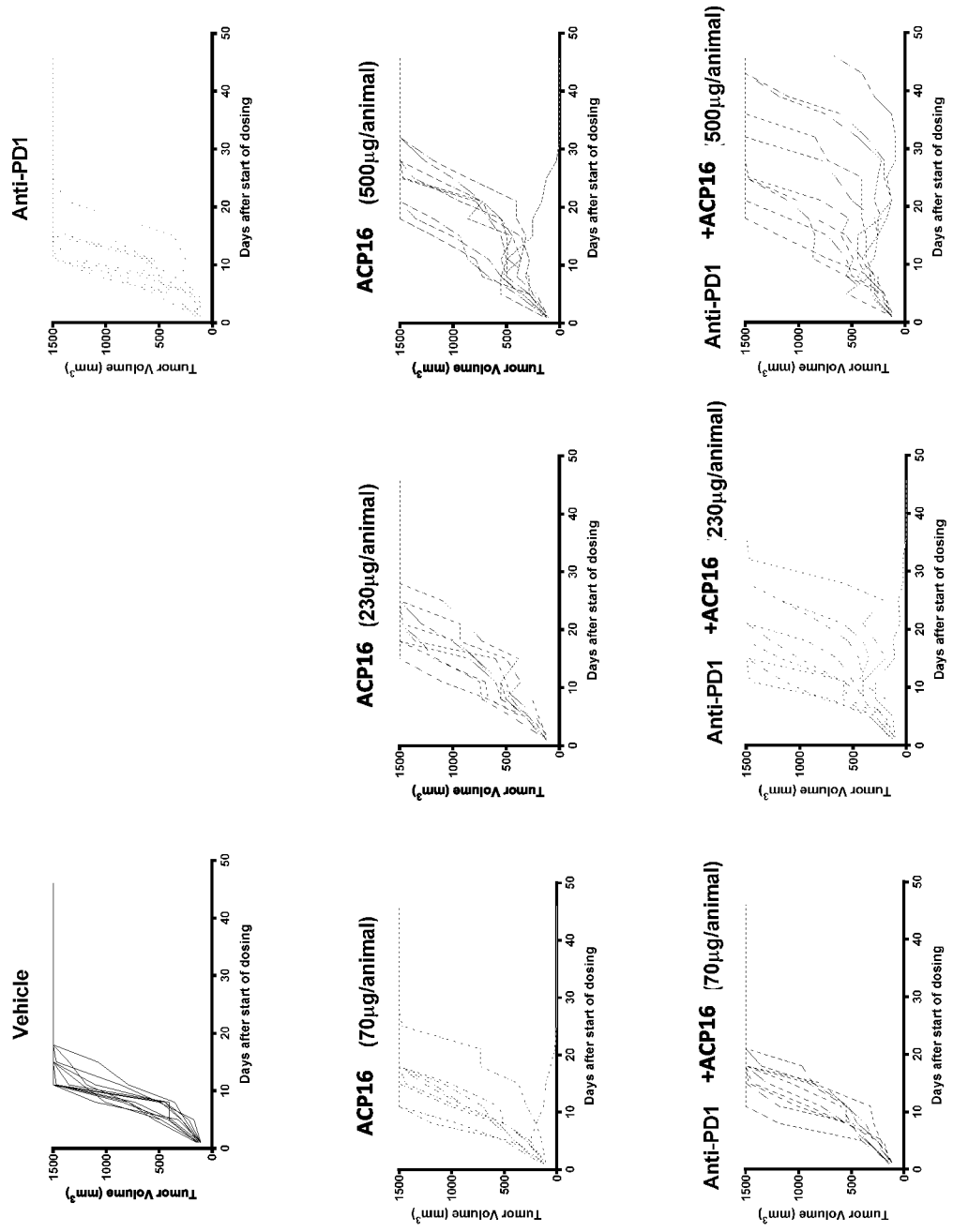


FIG. 47B

FIG. 47C

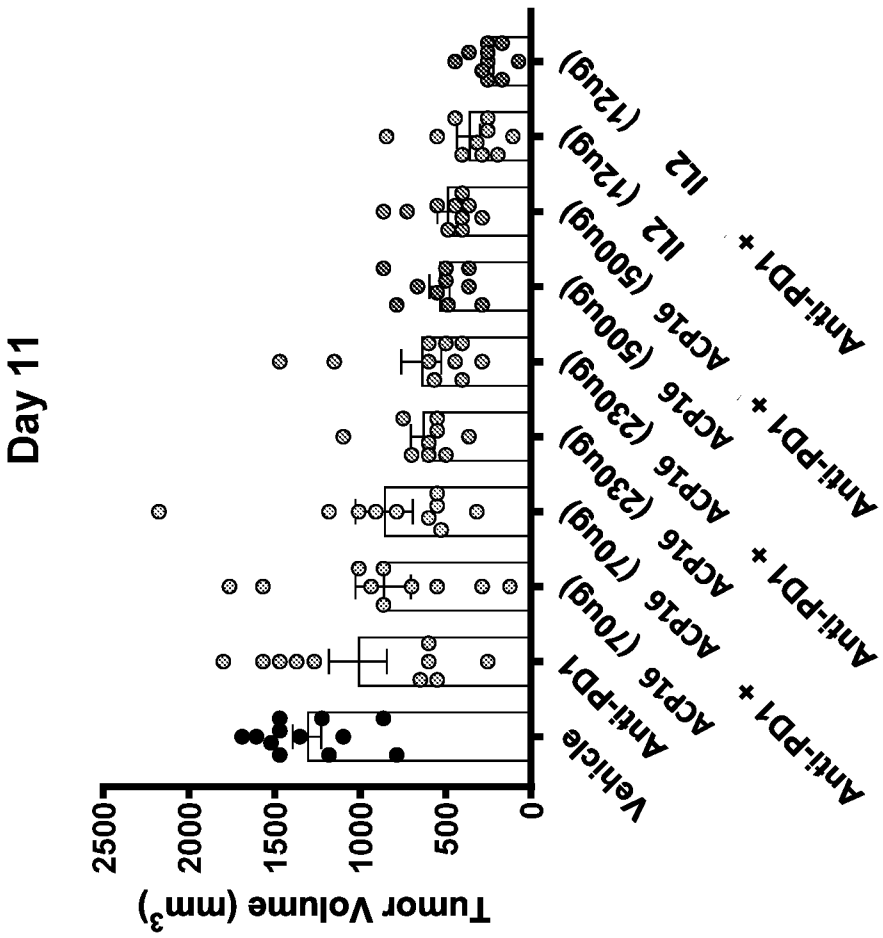
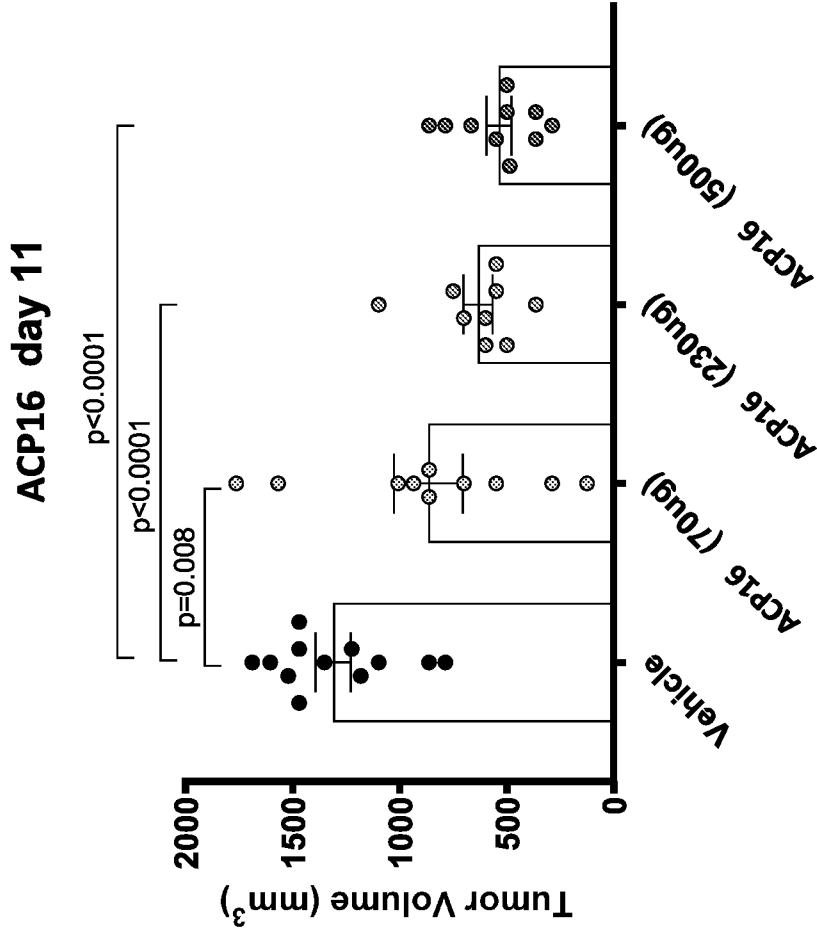


FIG. 47D



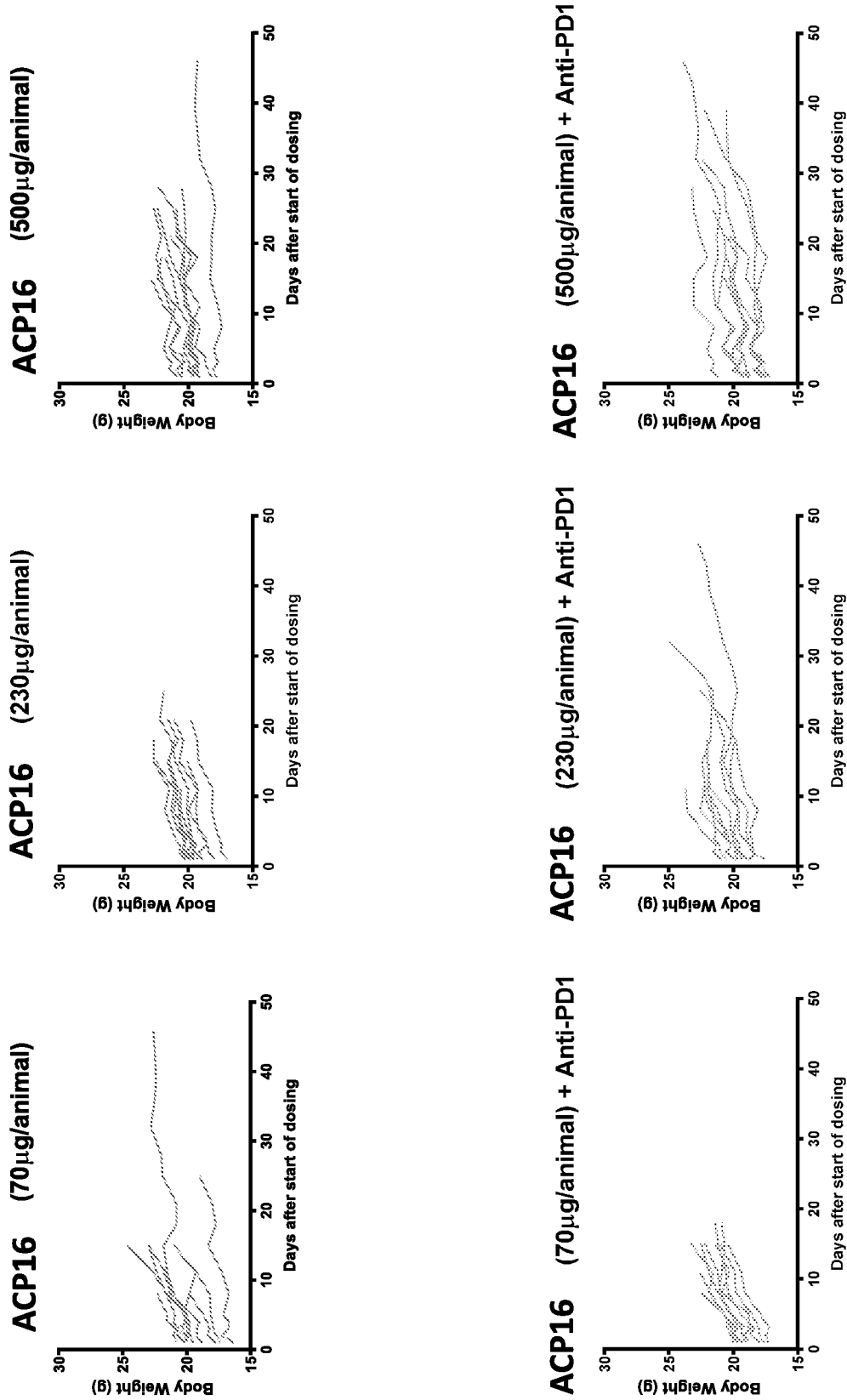
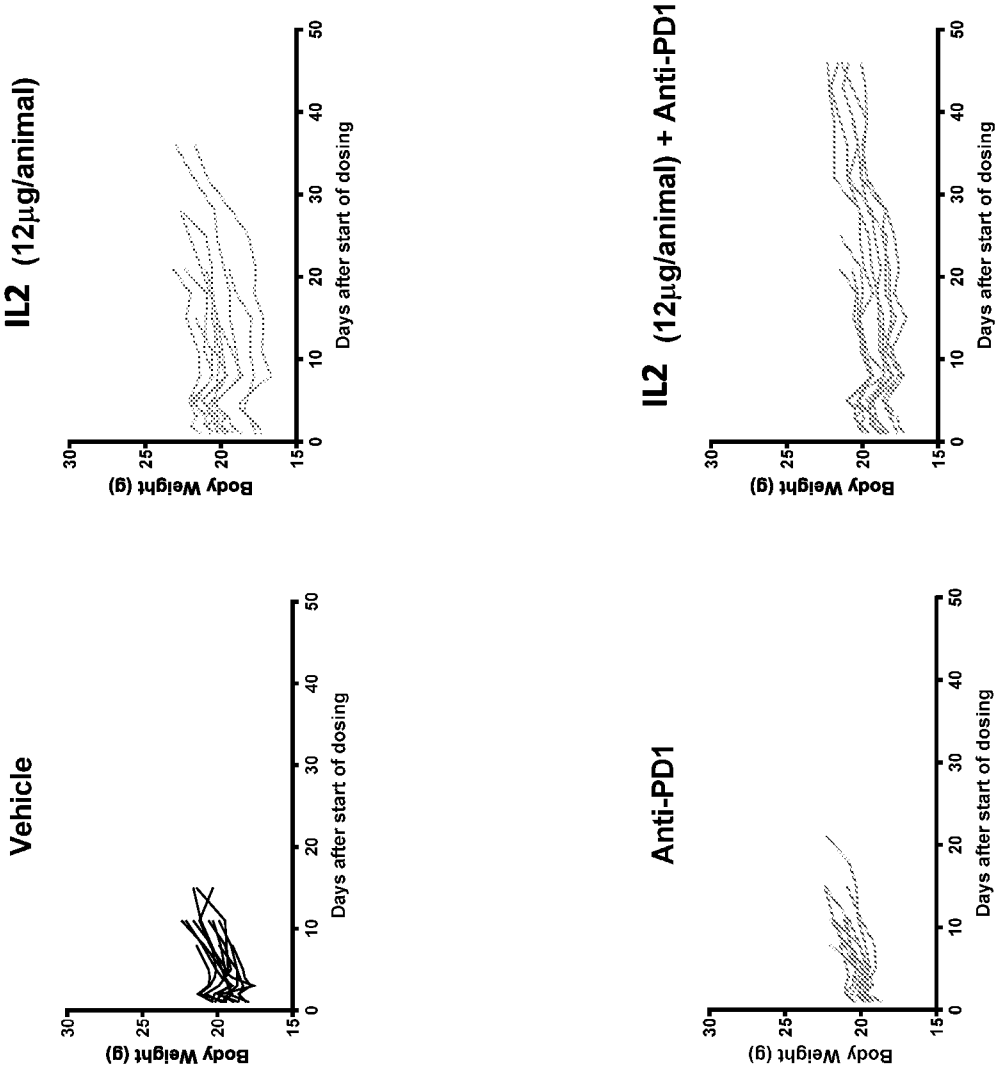


FIG. 48A

FIG. 48B



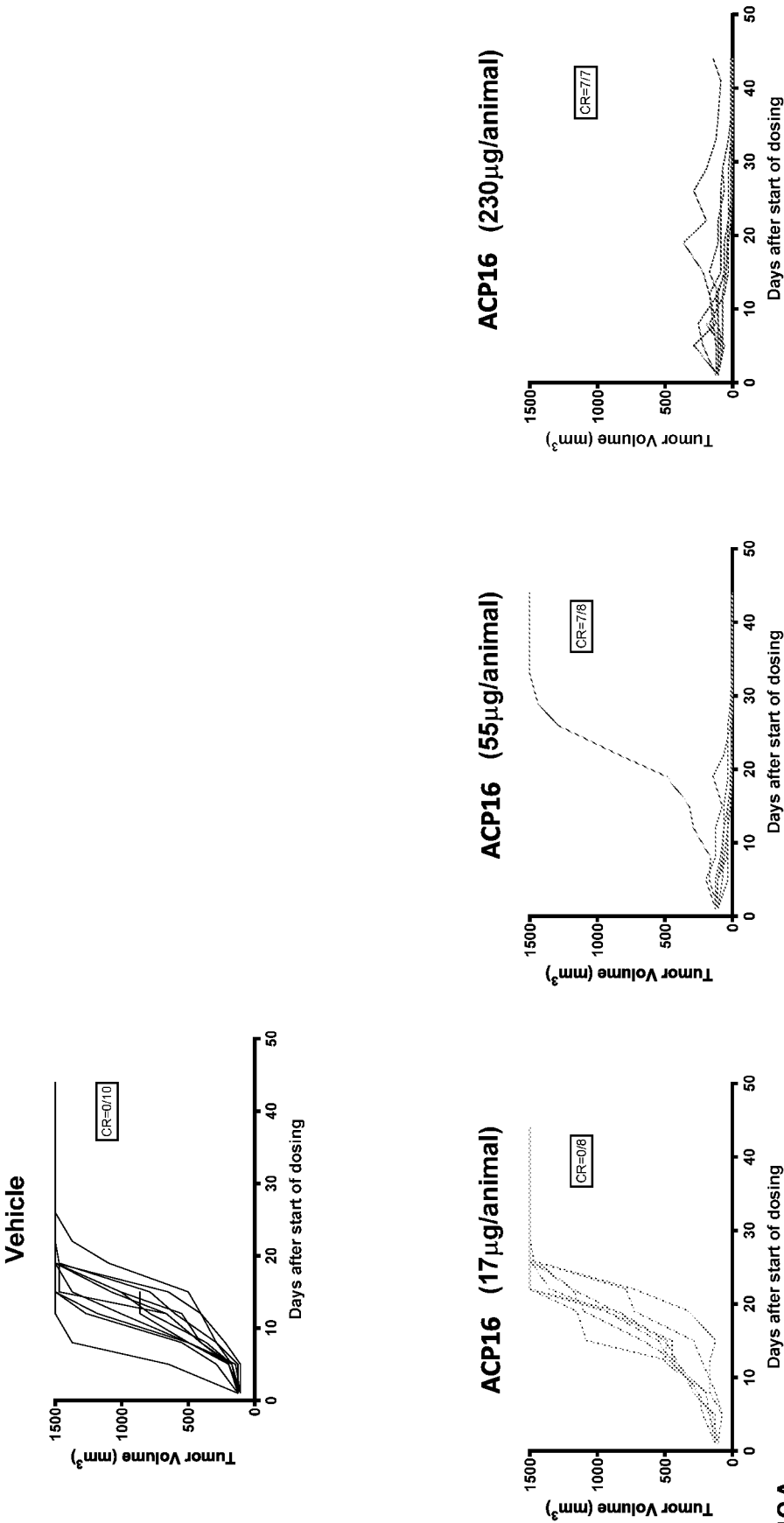


FIG. 49A

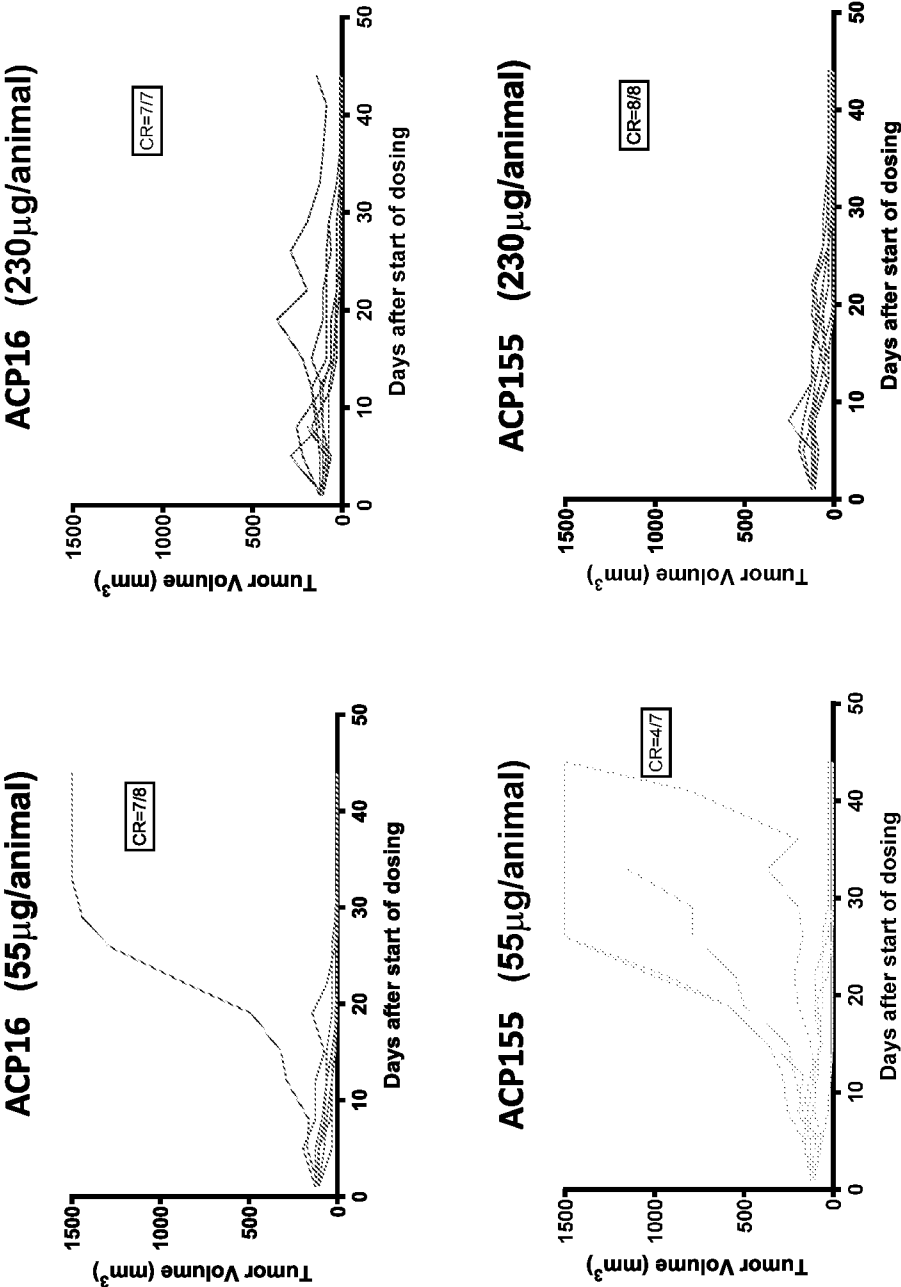


FIG. 49B

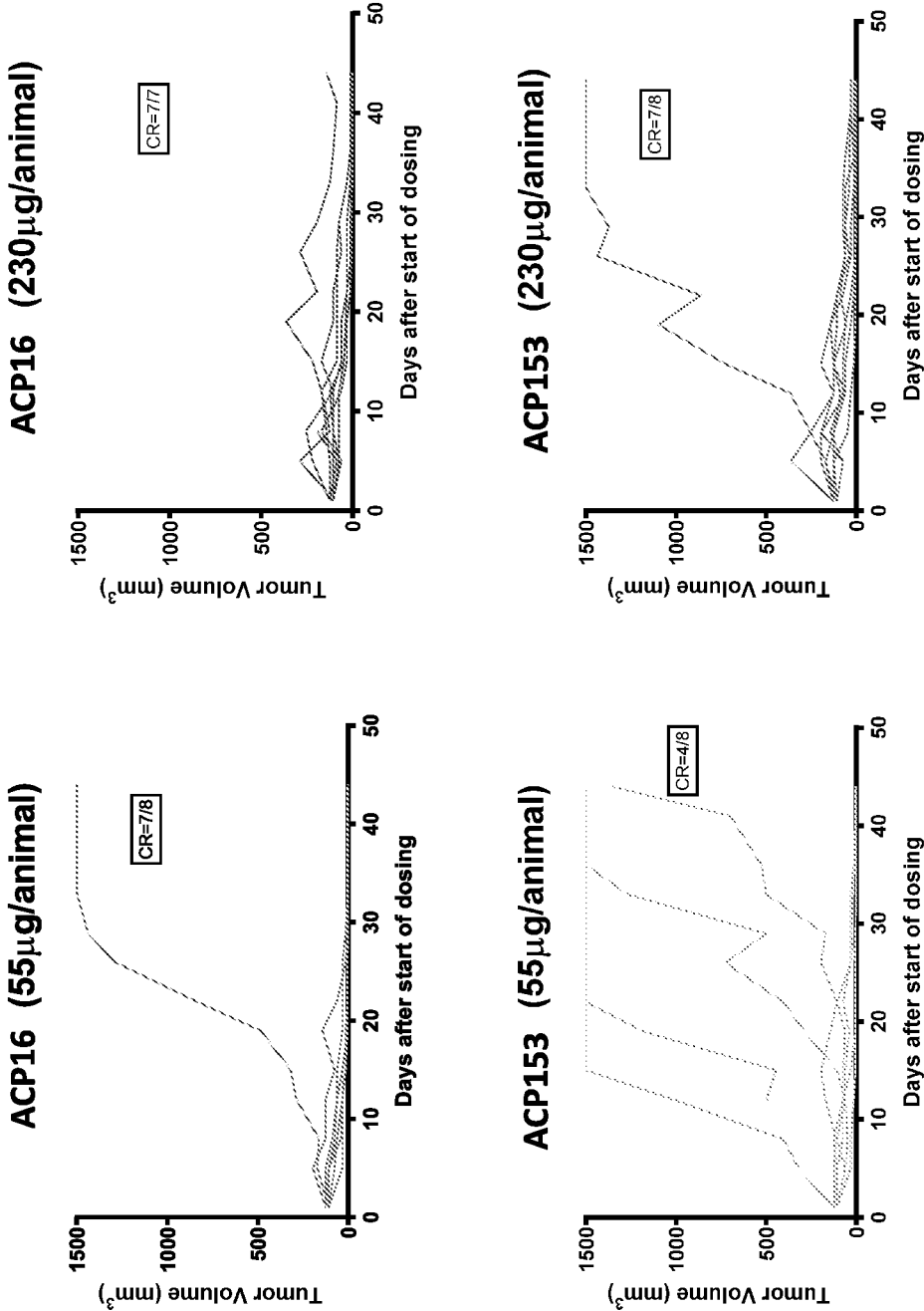


FIG. 49C

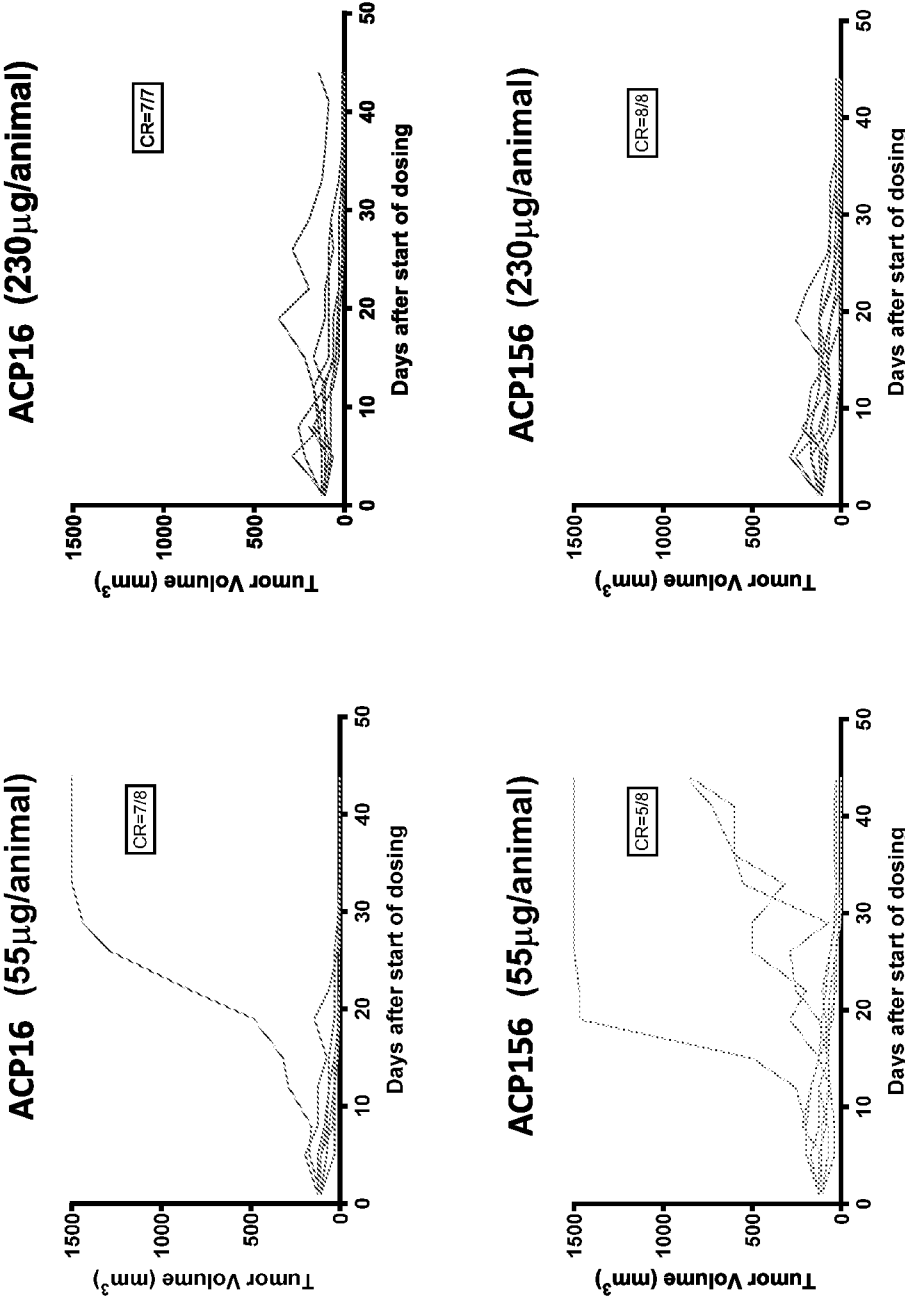


FIG. 49D

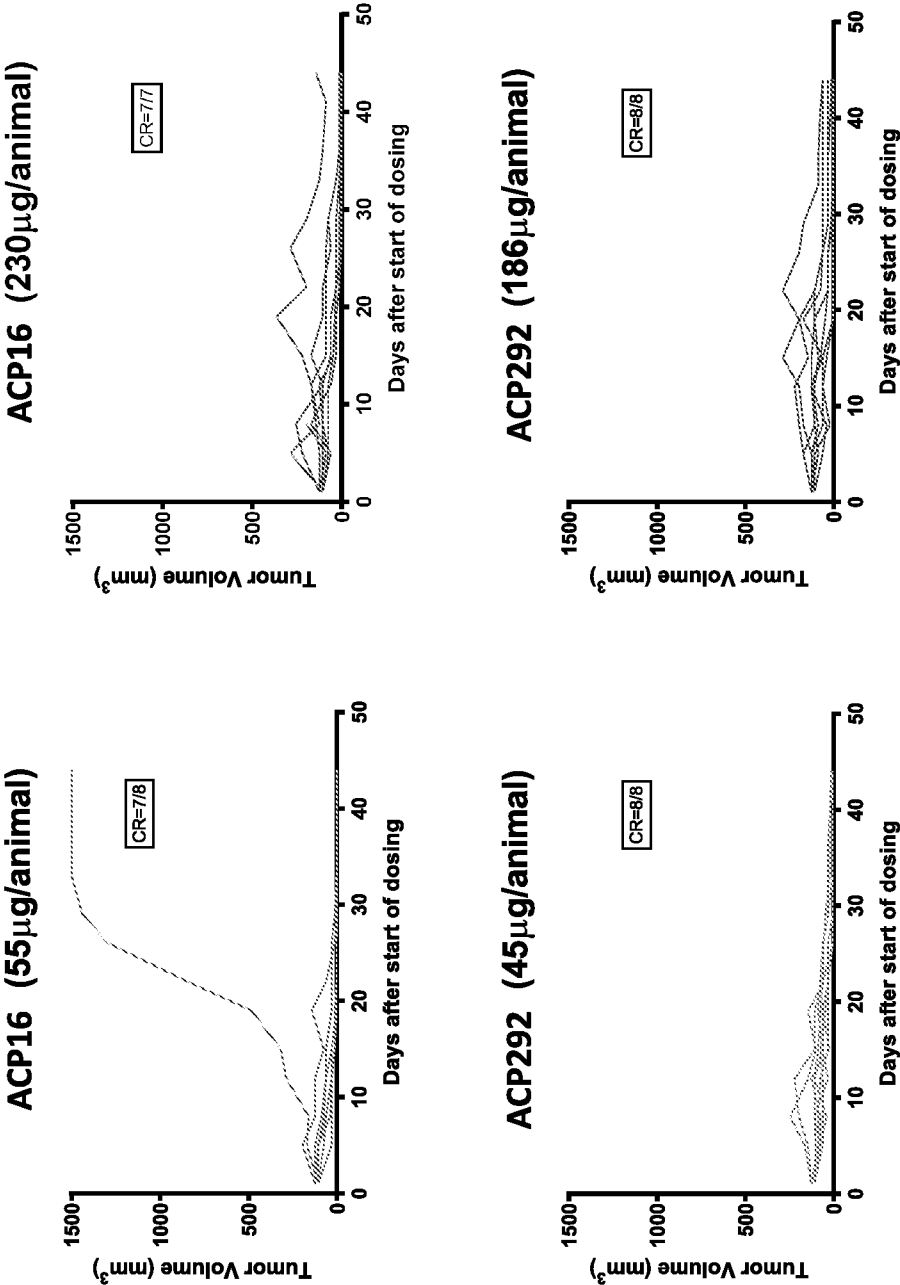


FIG. 49E

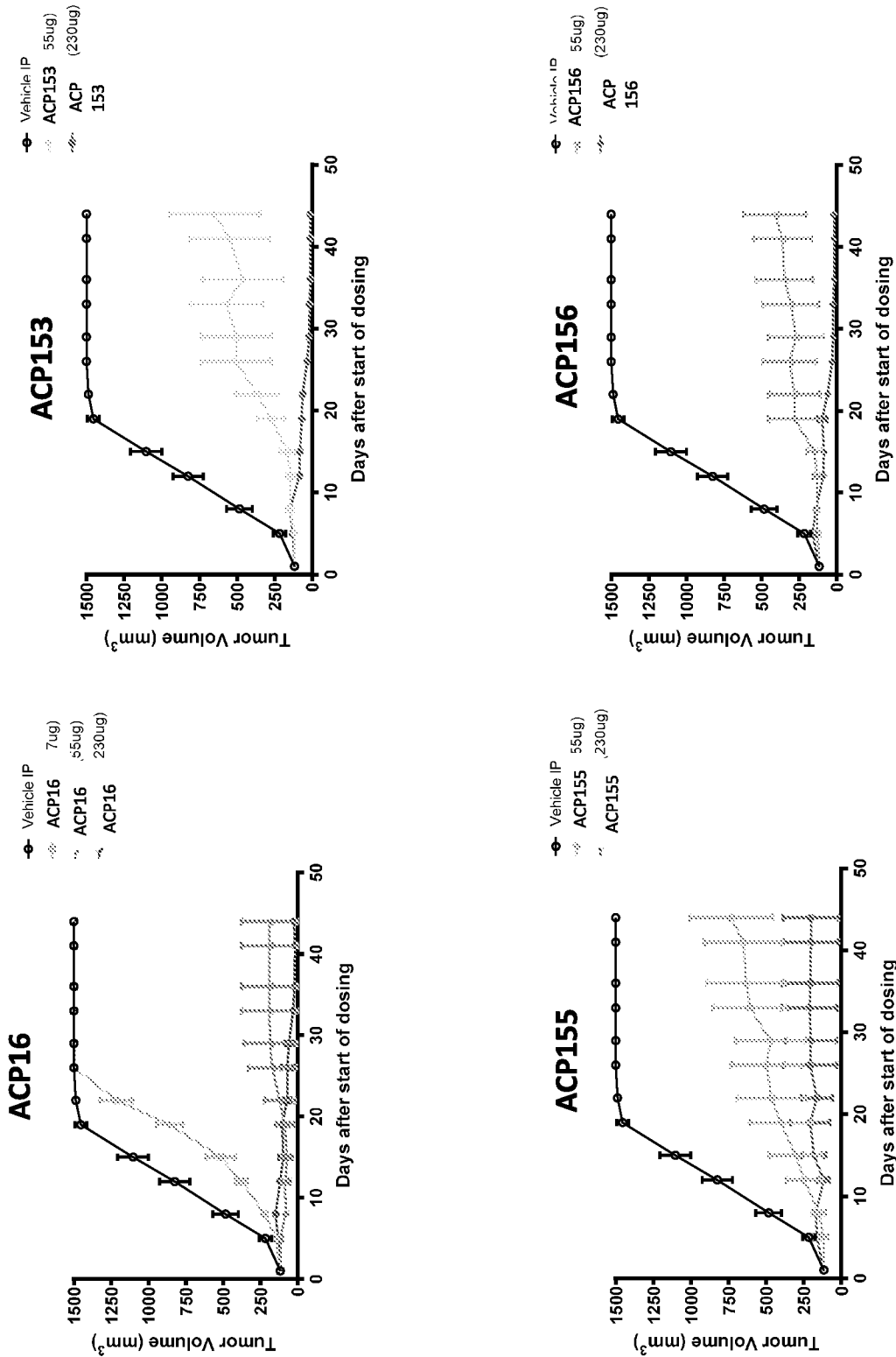


FIG. 49F

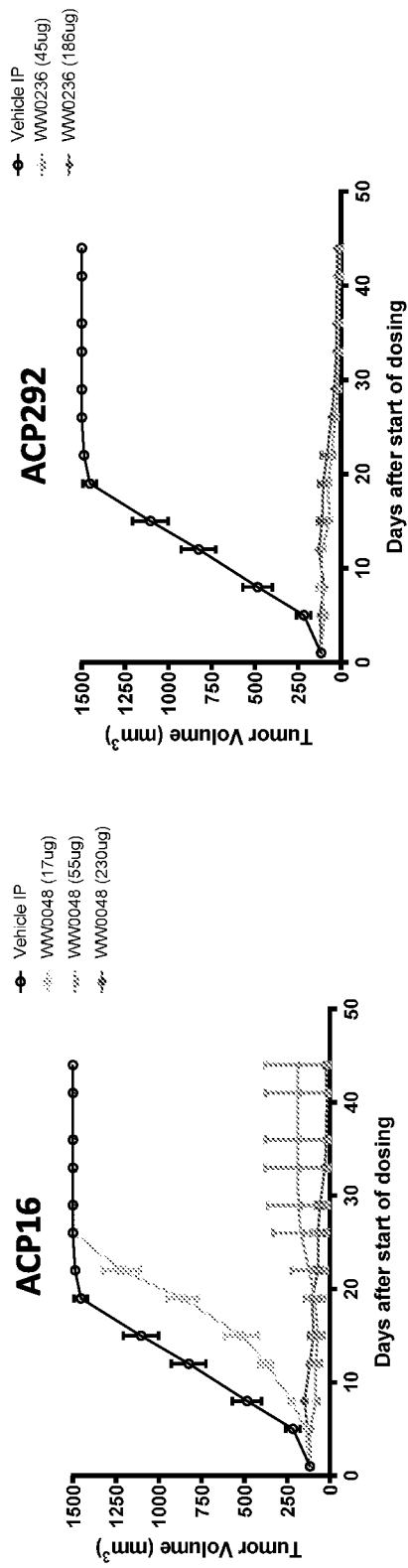


FIG. 49G

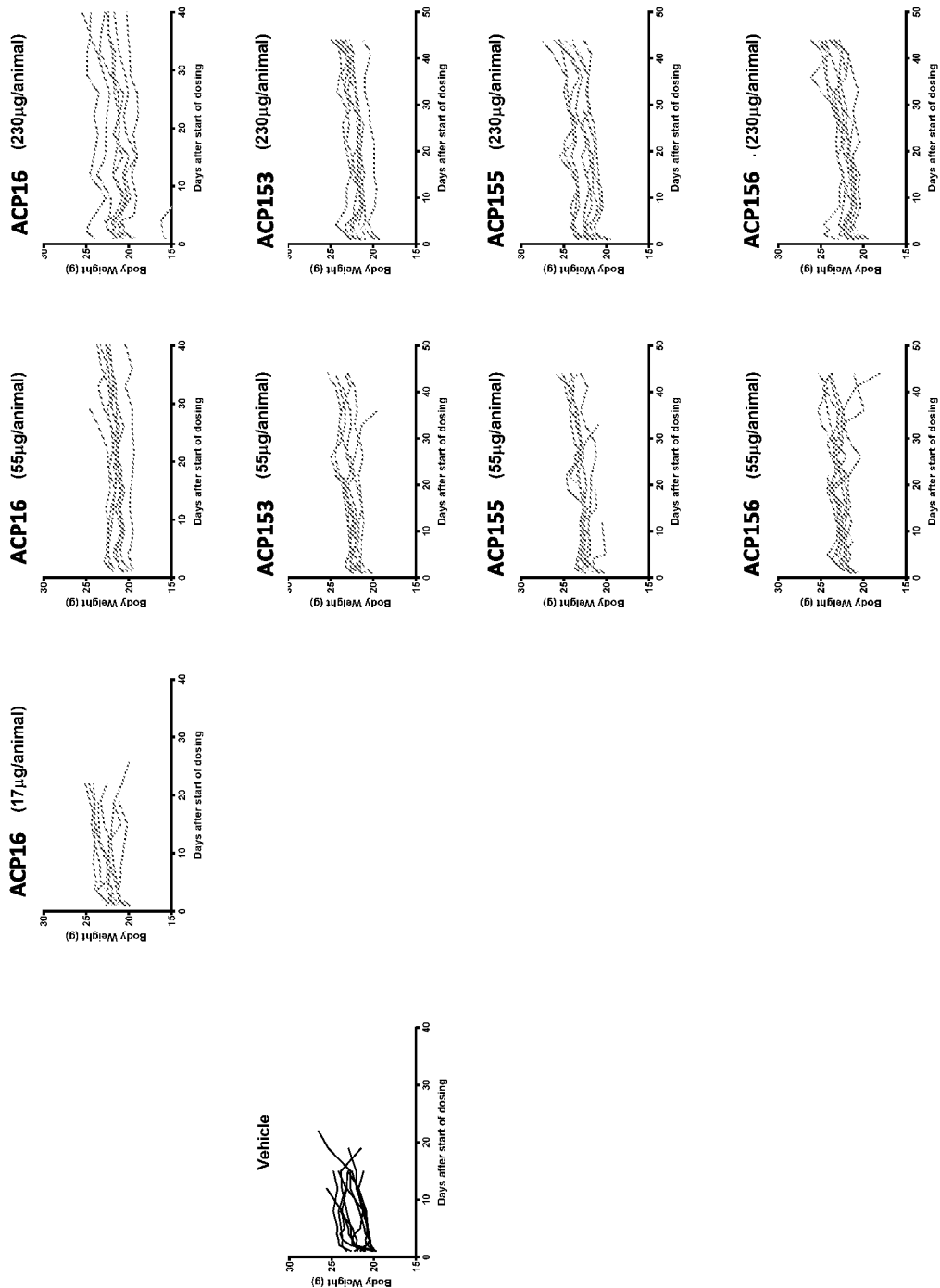


FIG. 49H

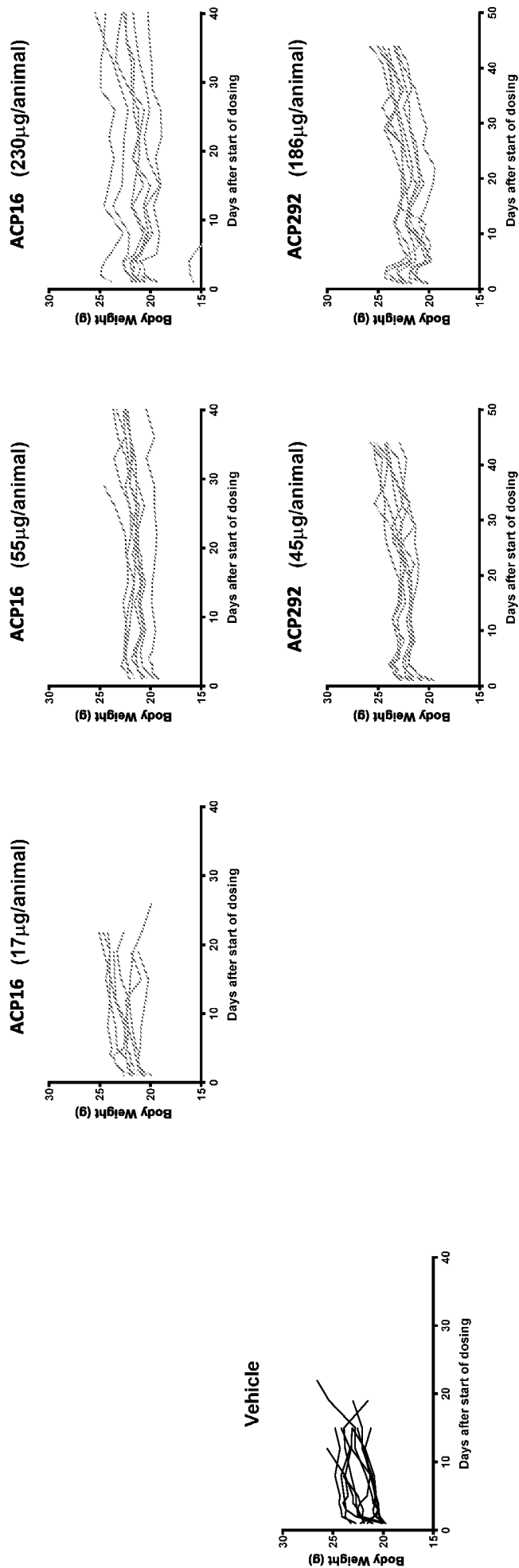


FIG. 49I

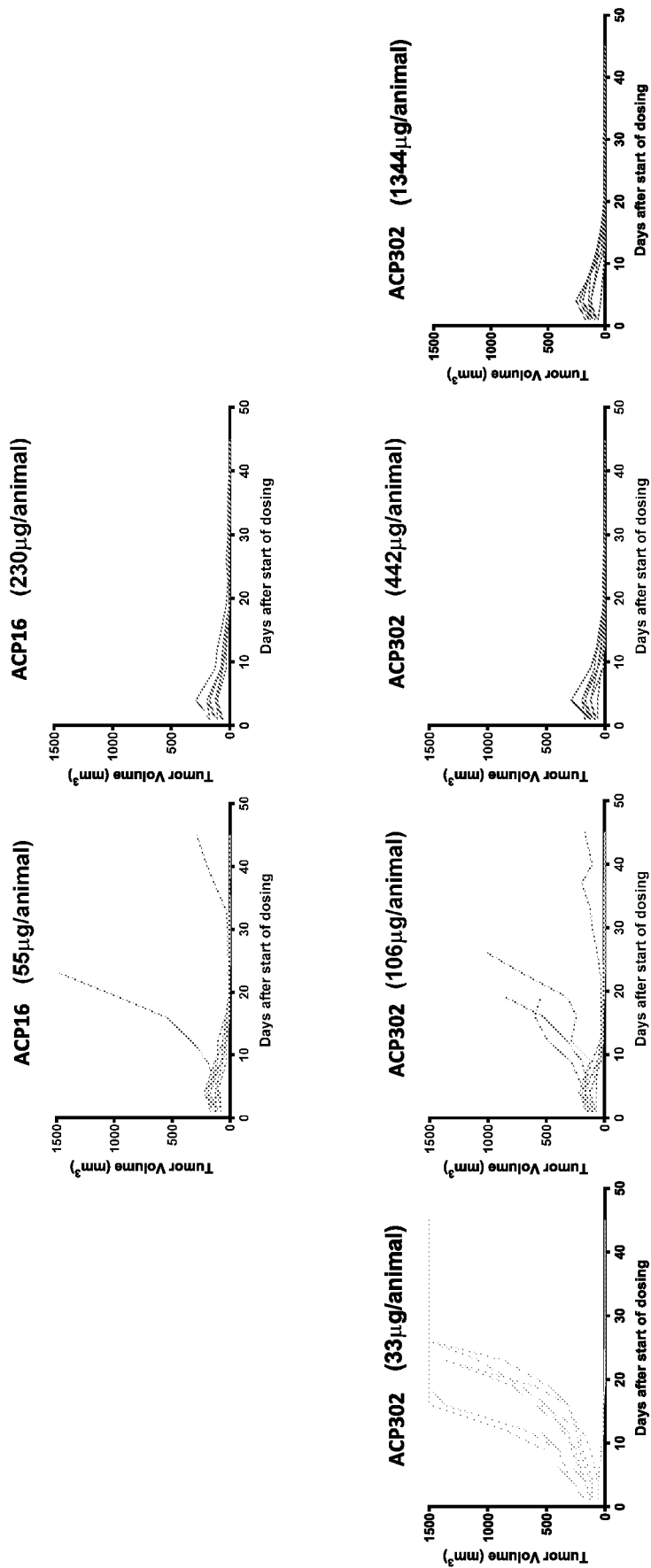


FIG. 50A

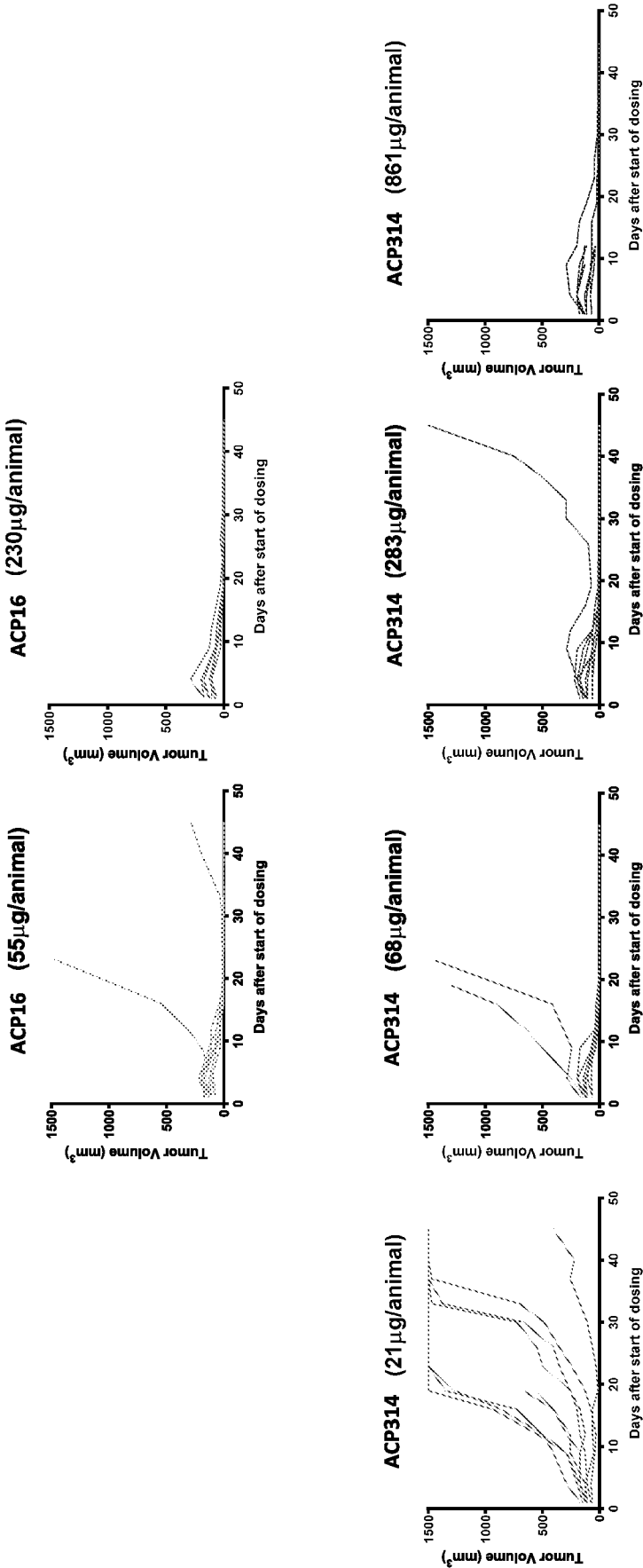


FIG. 50B

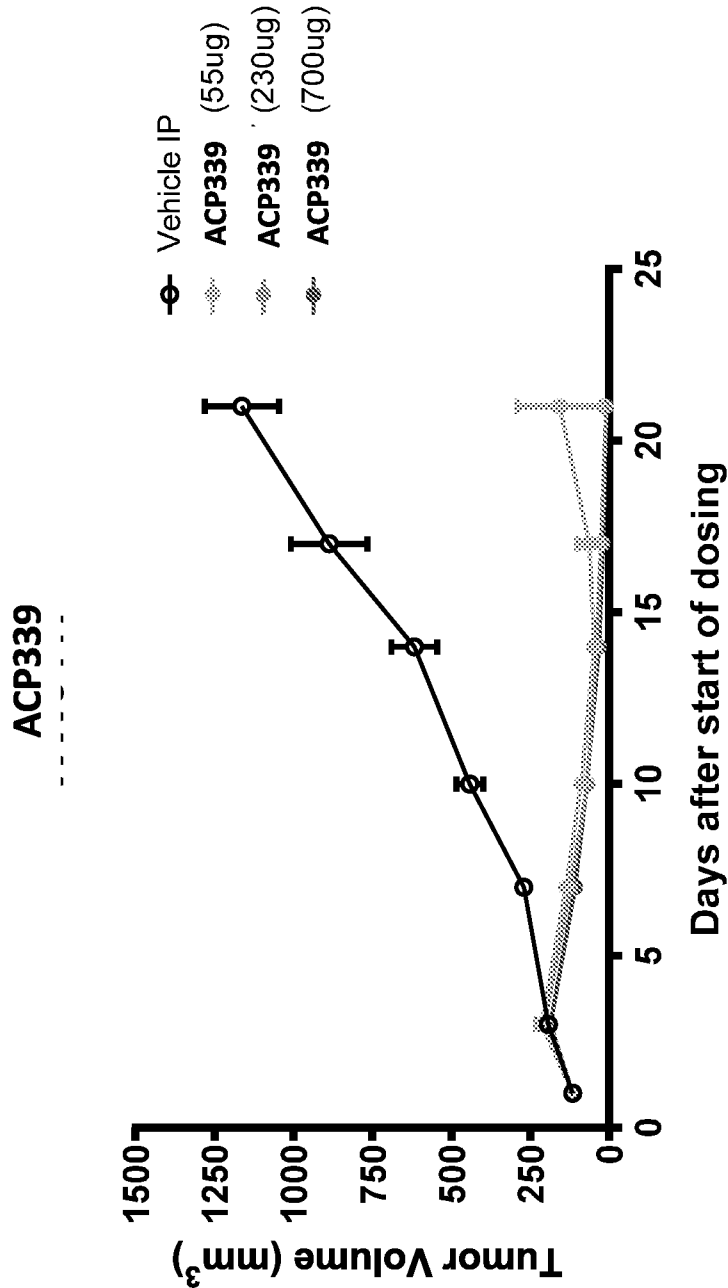
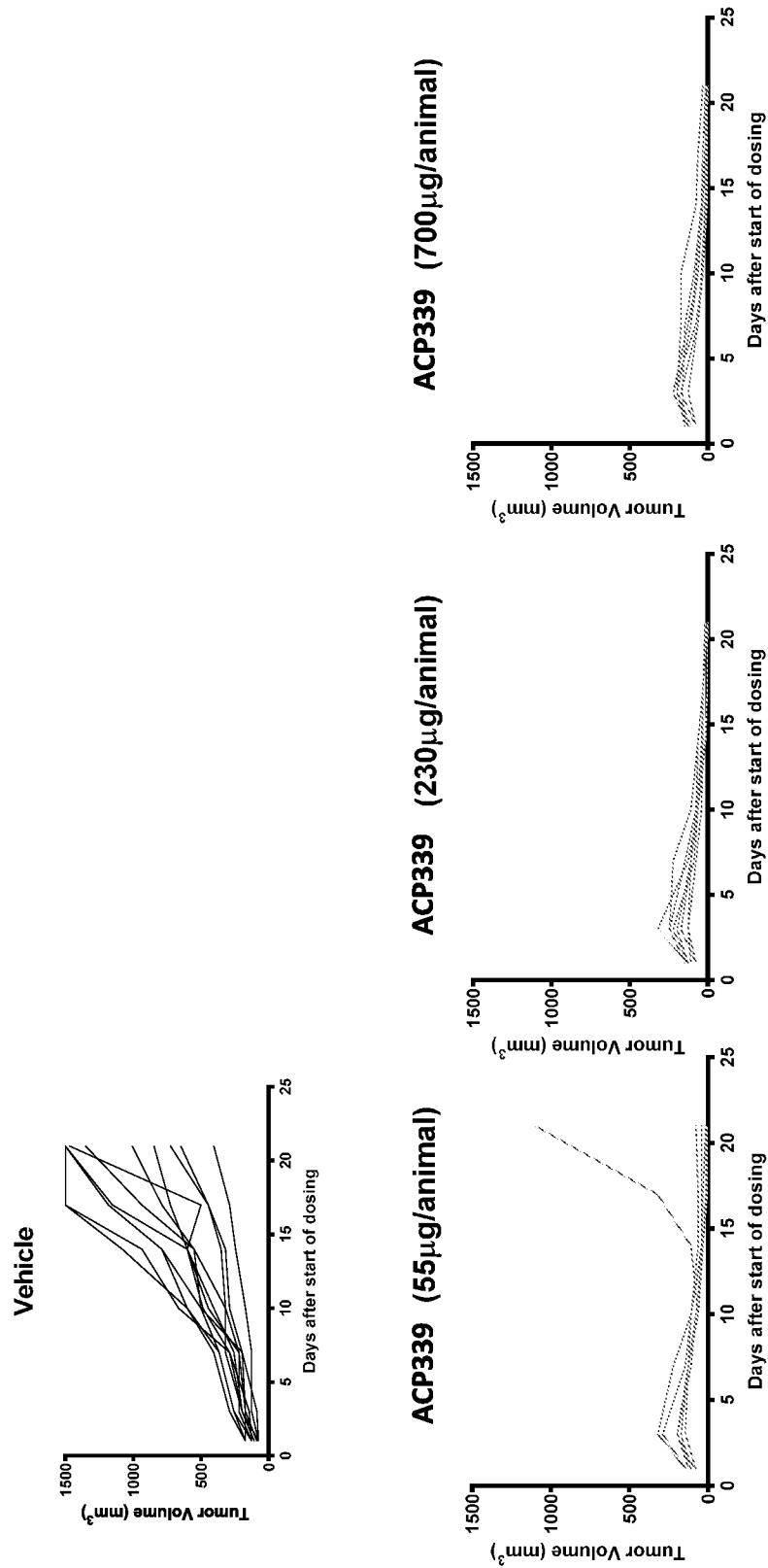


FIG. 51A

FIG. 51B



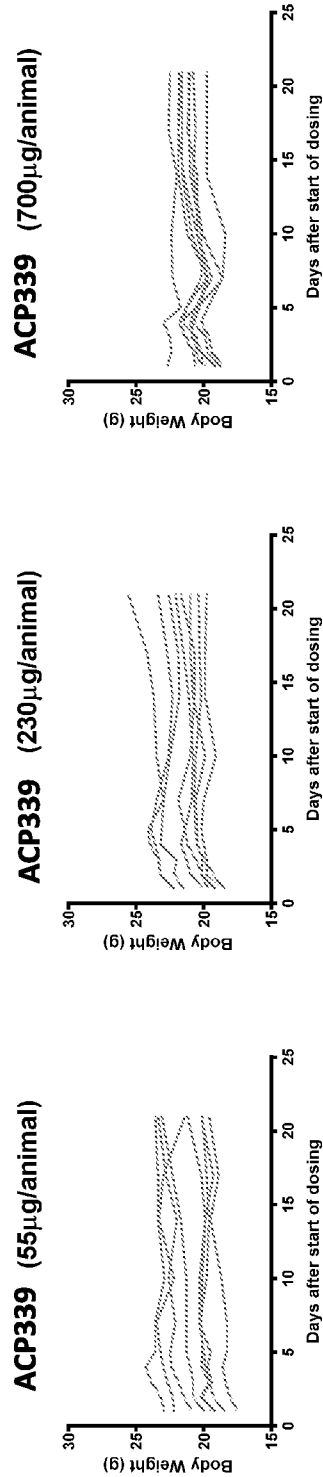
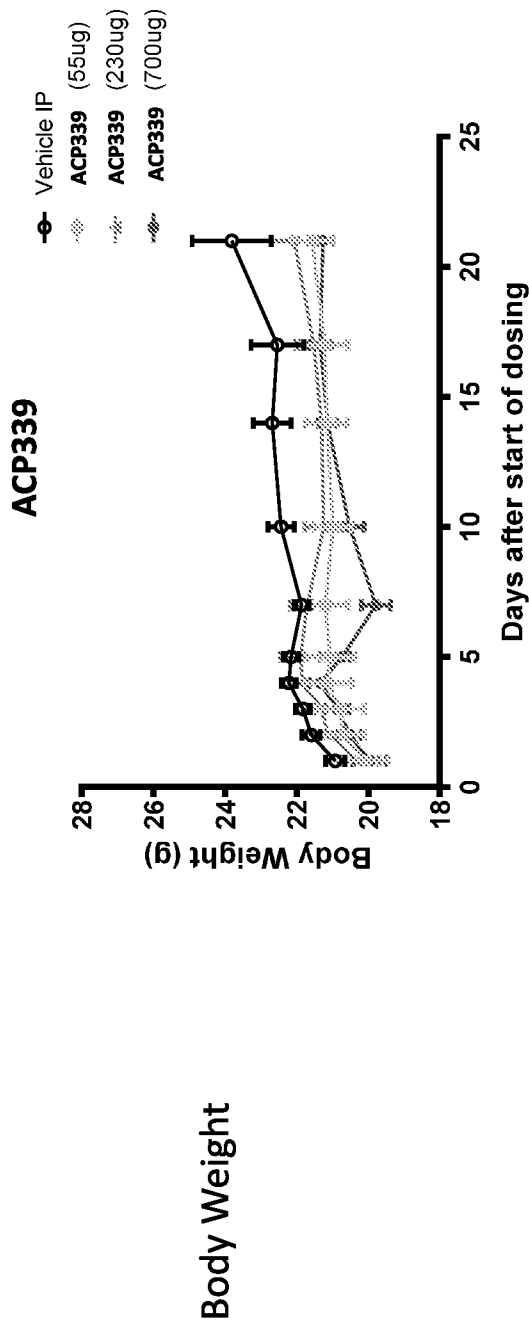


FIG. 51C

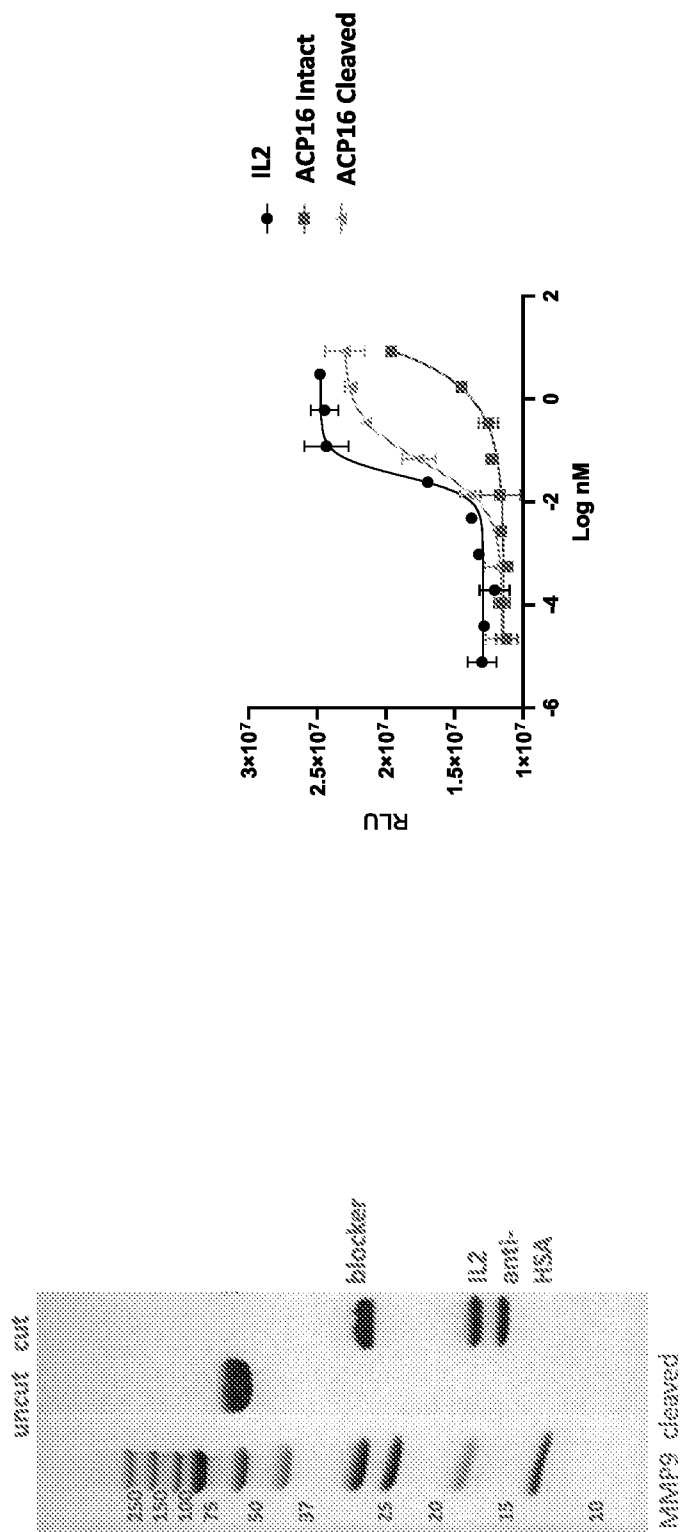


FIG. 52A

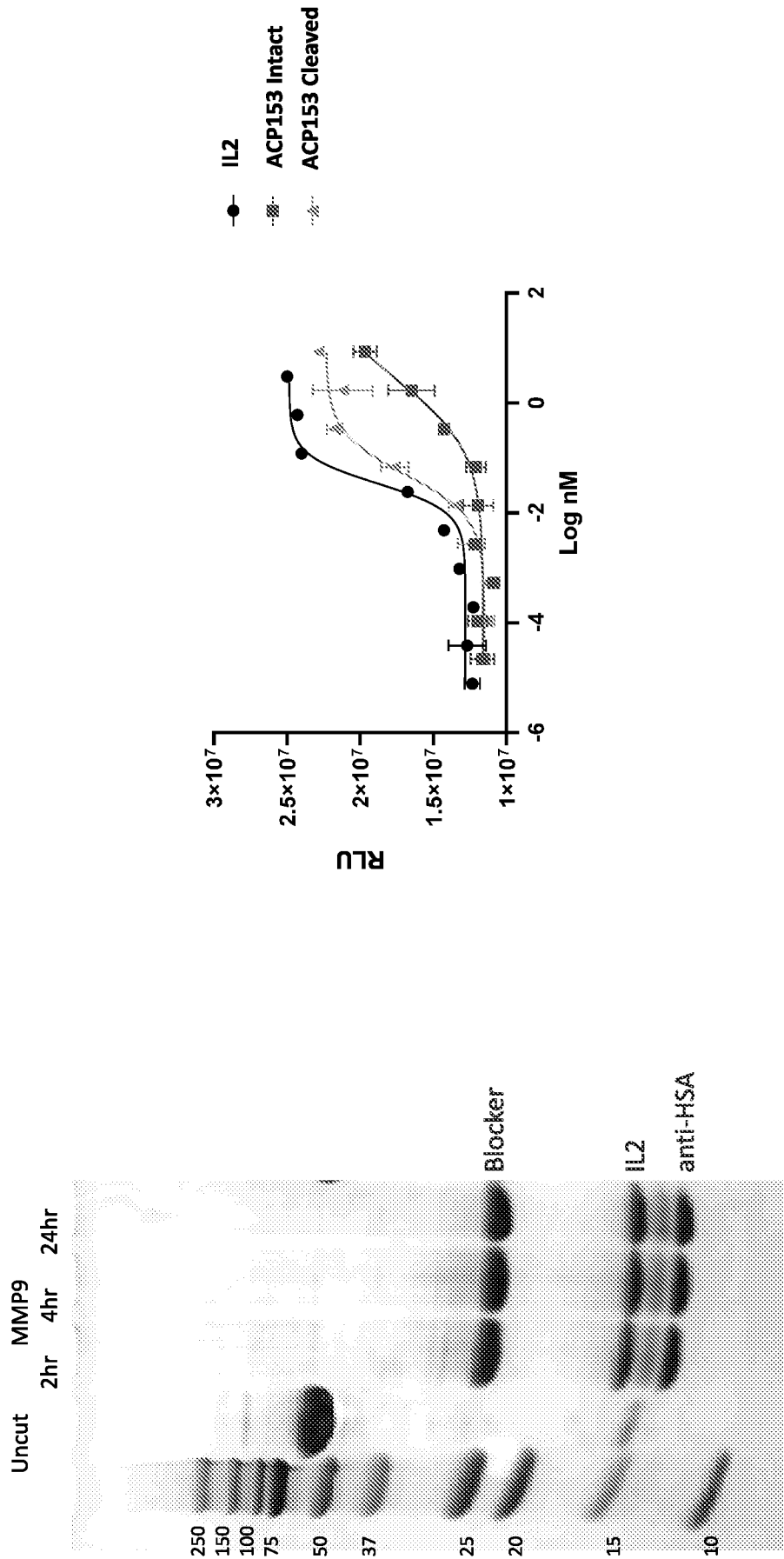


FIG. 52B

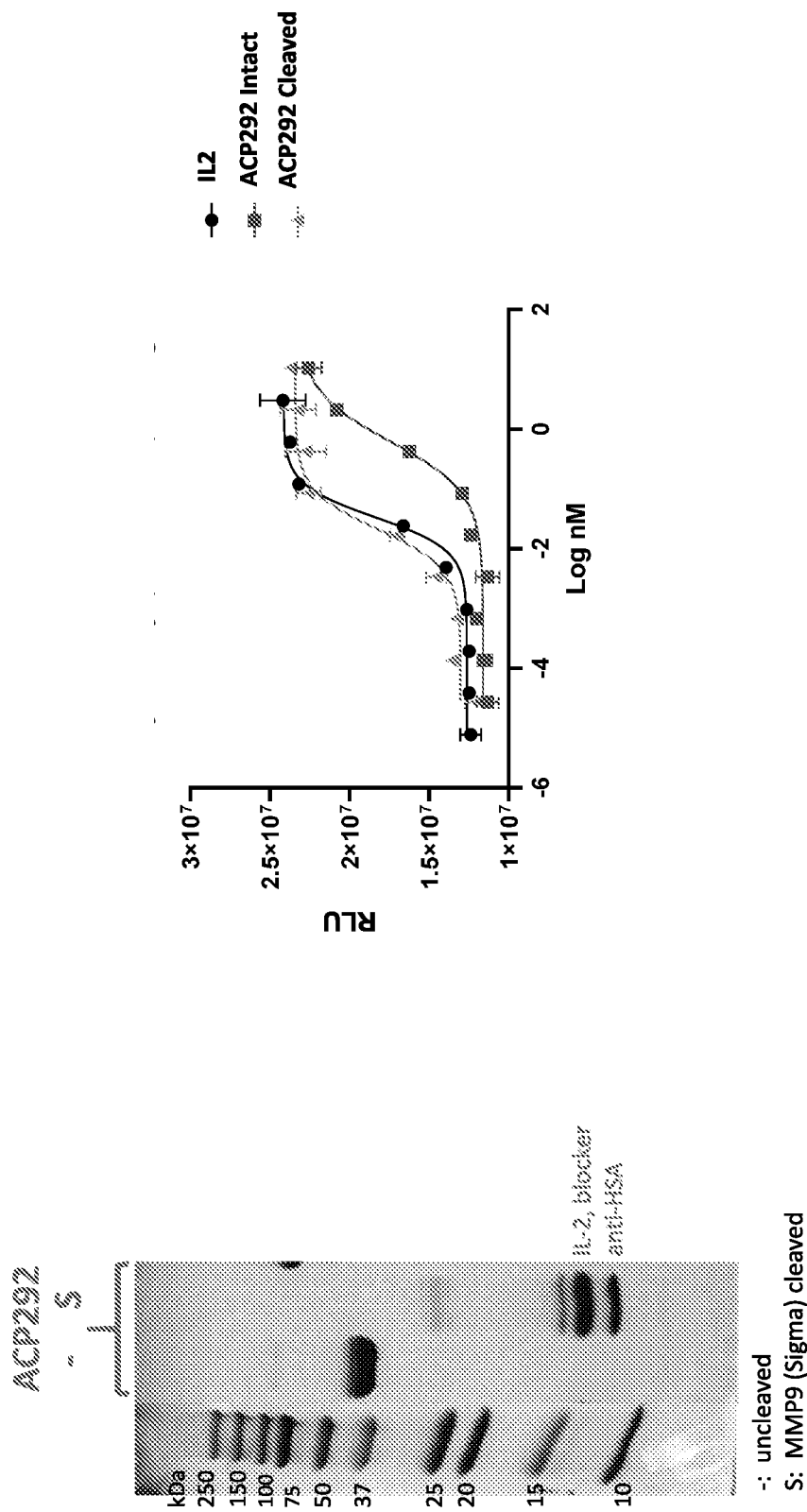


FIG. 52C

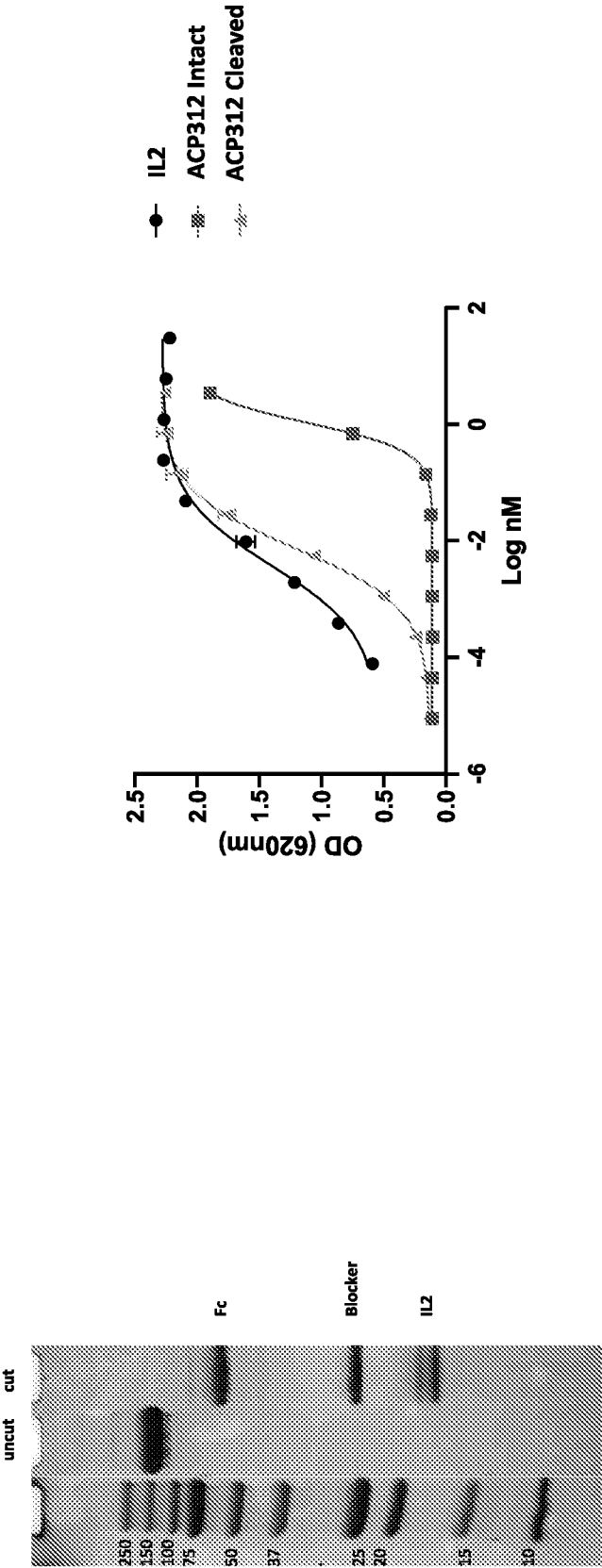


FIG. 52D

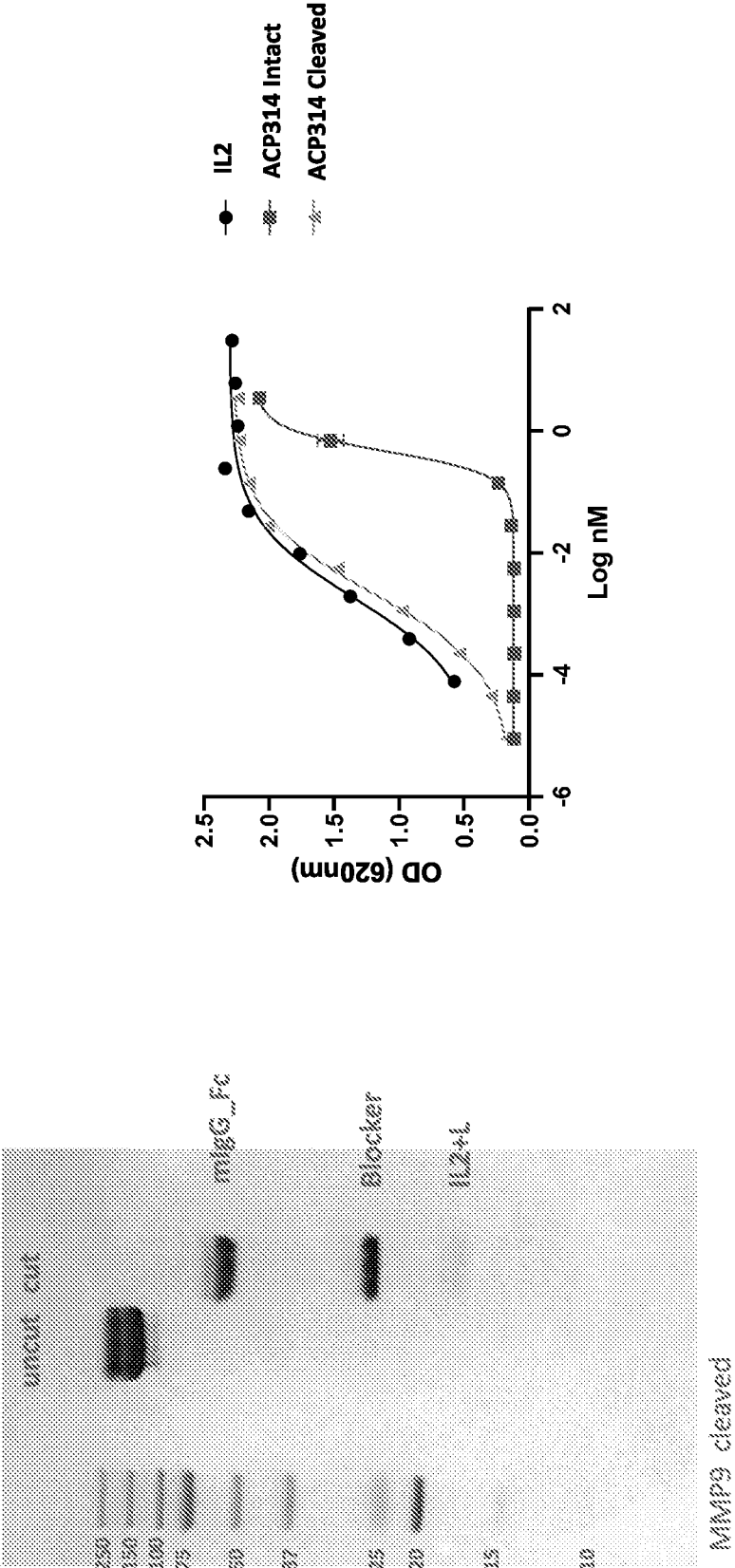


FIG. 52E

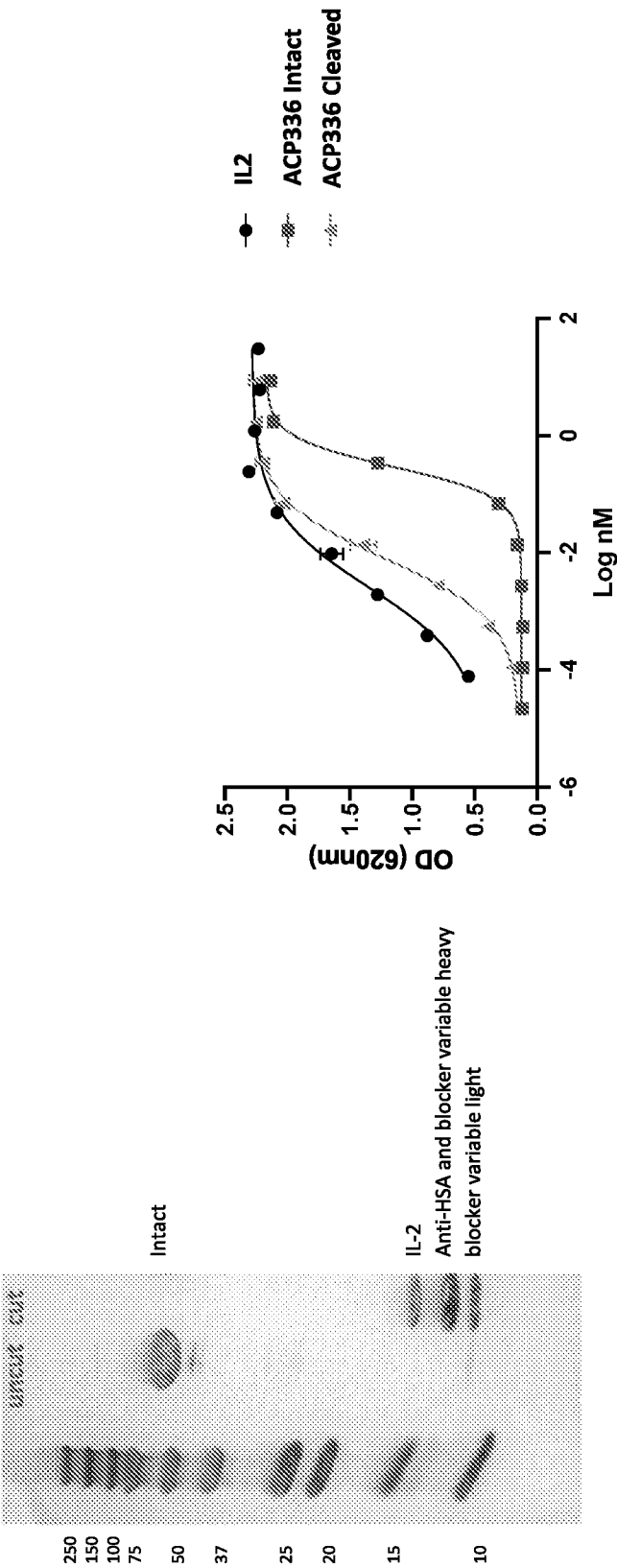


FIG. 52F

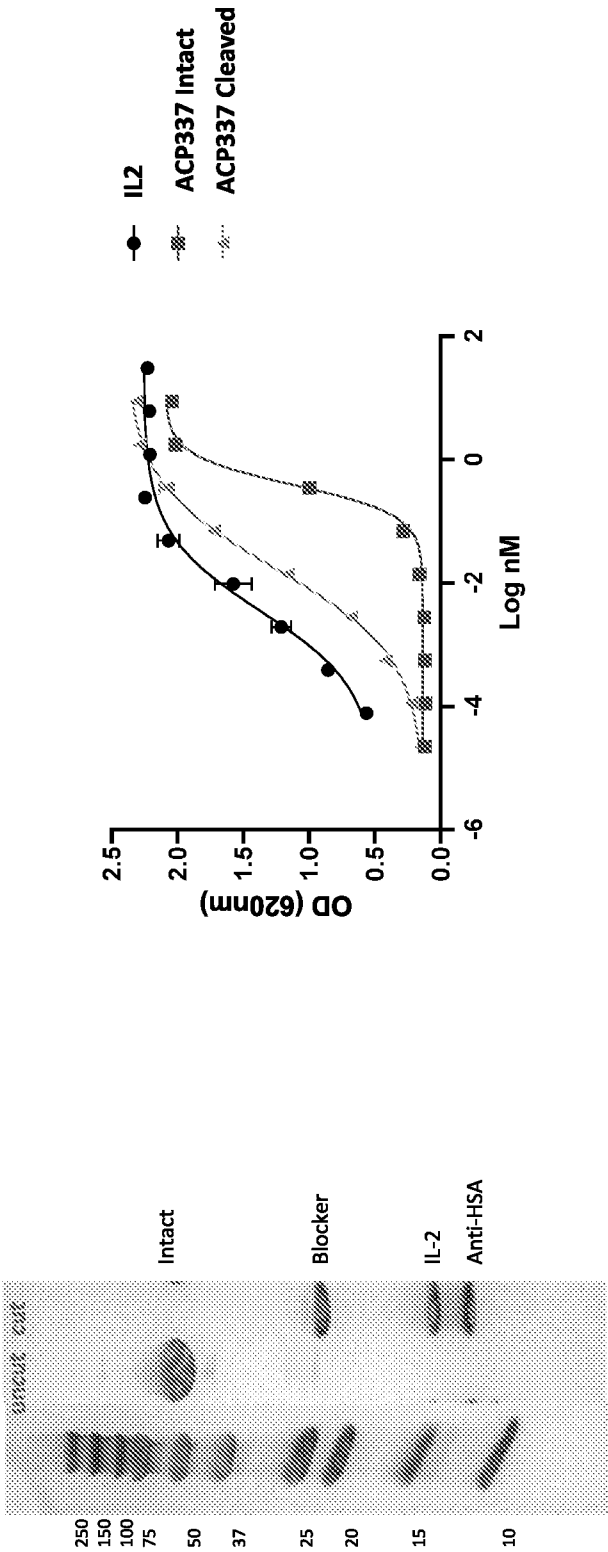


FIG. 52G

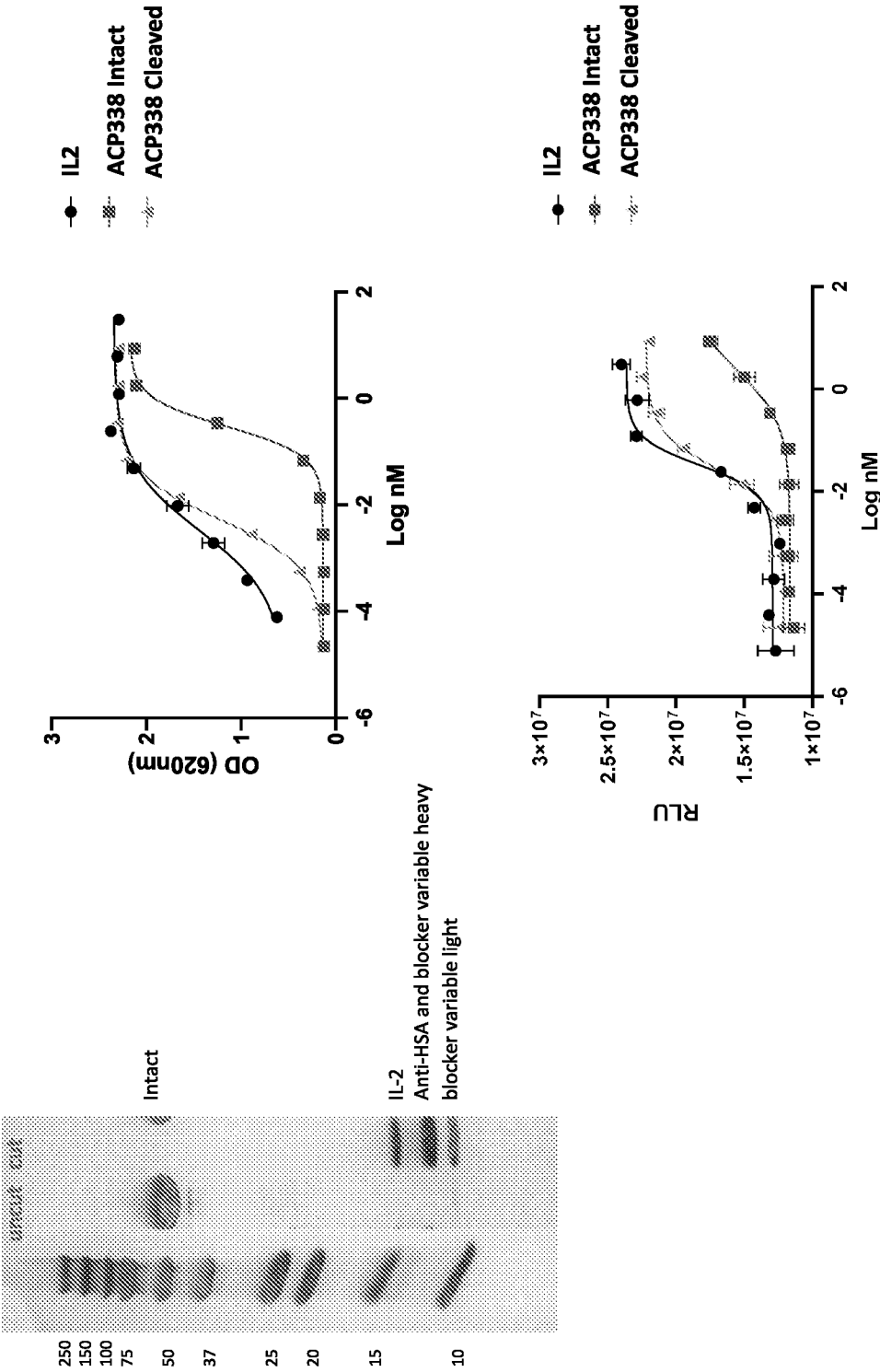


FIG. 52H

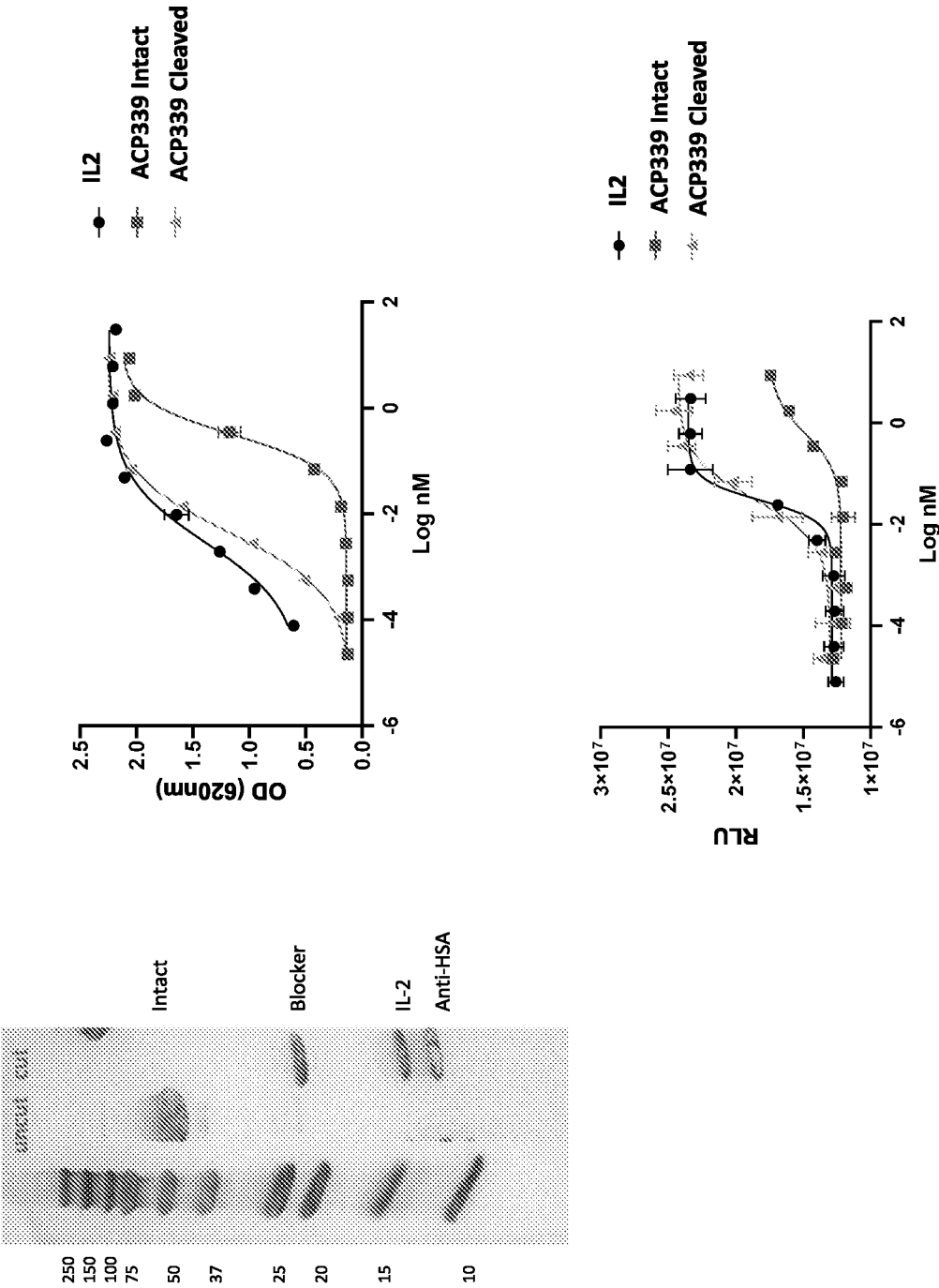


FIG. 52I

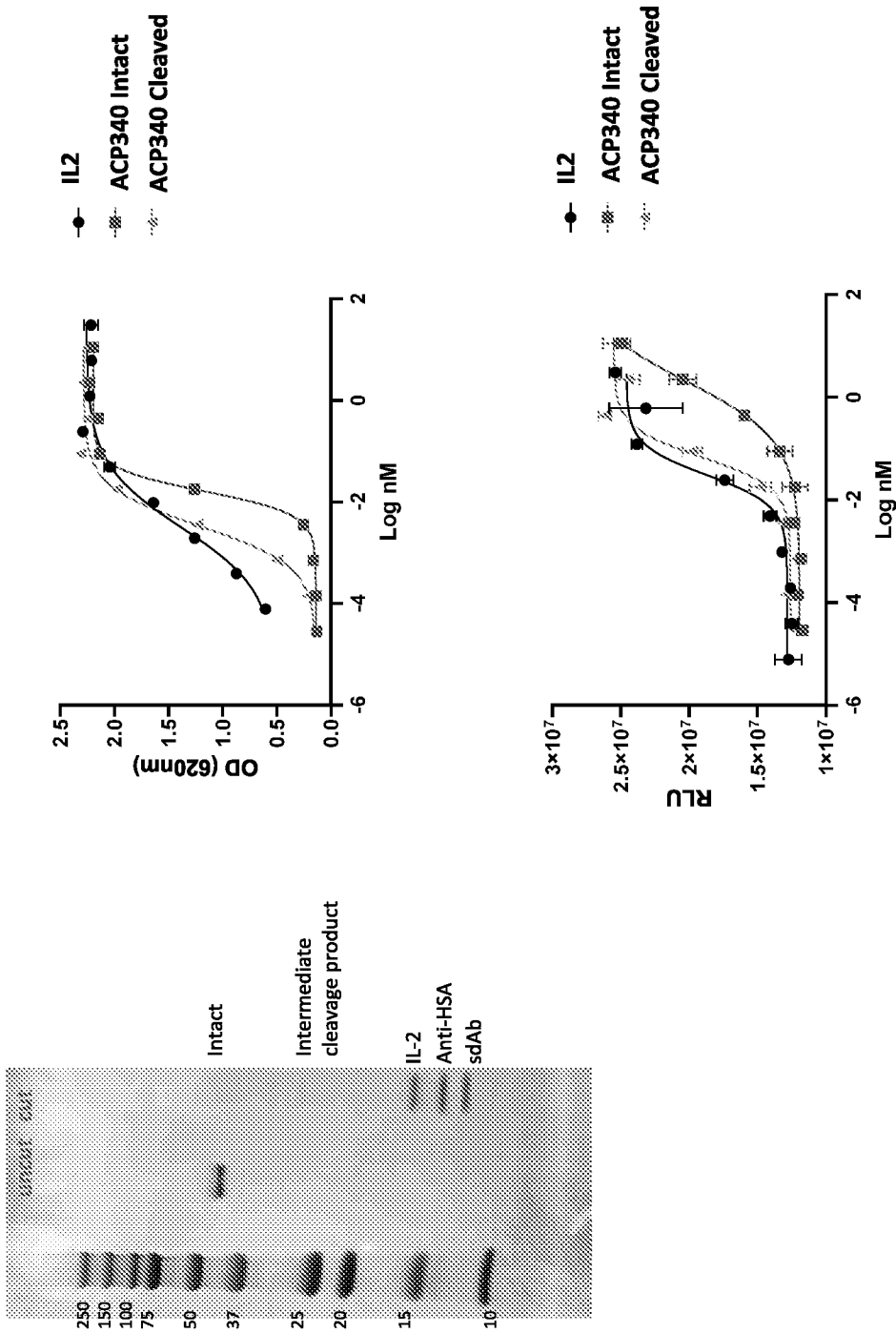


FIG. 52J

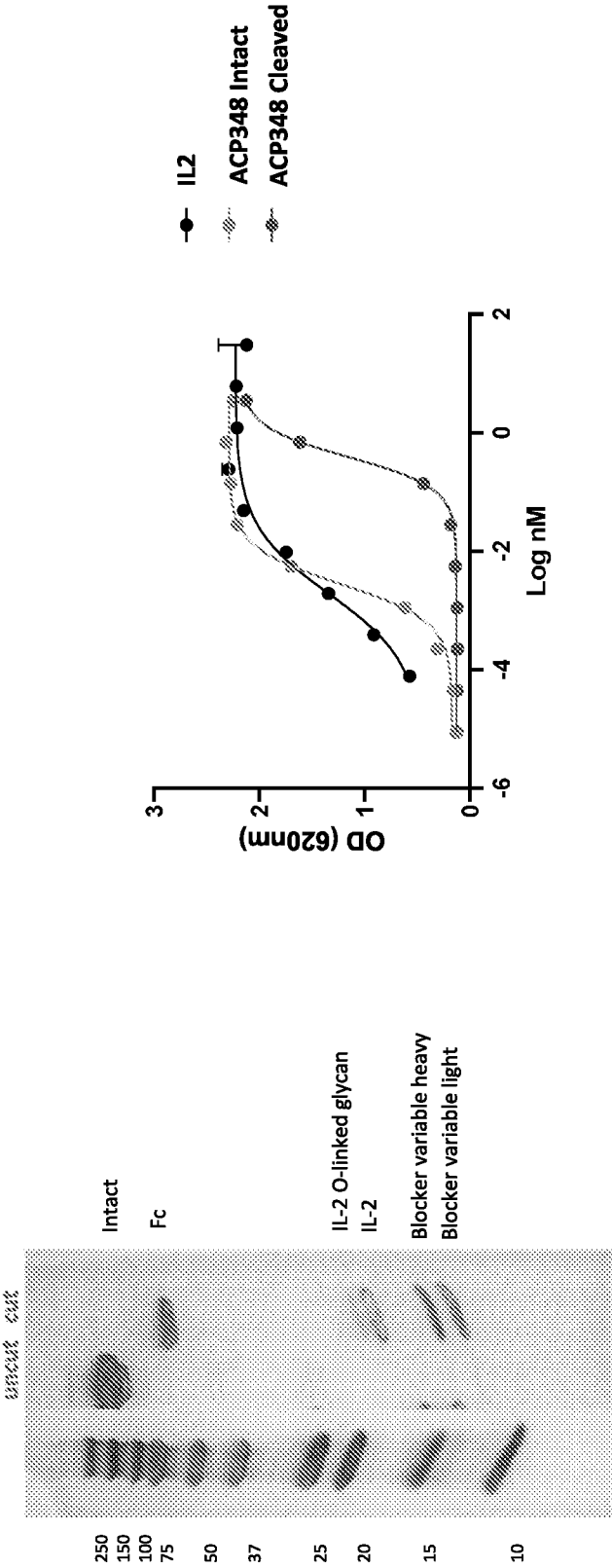


FIG. 52K

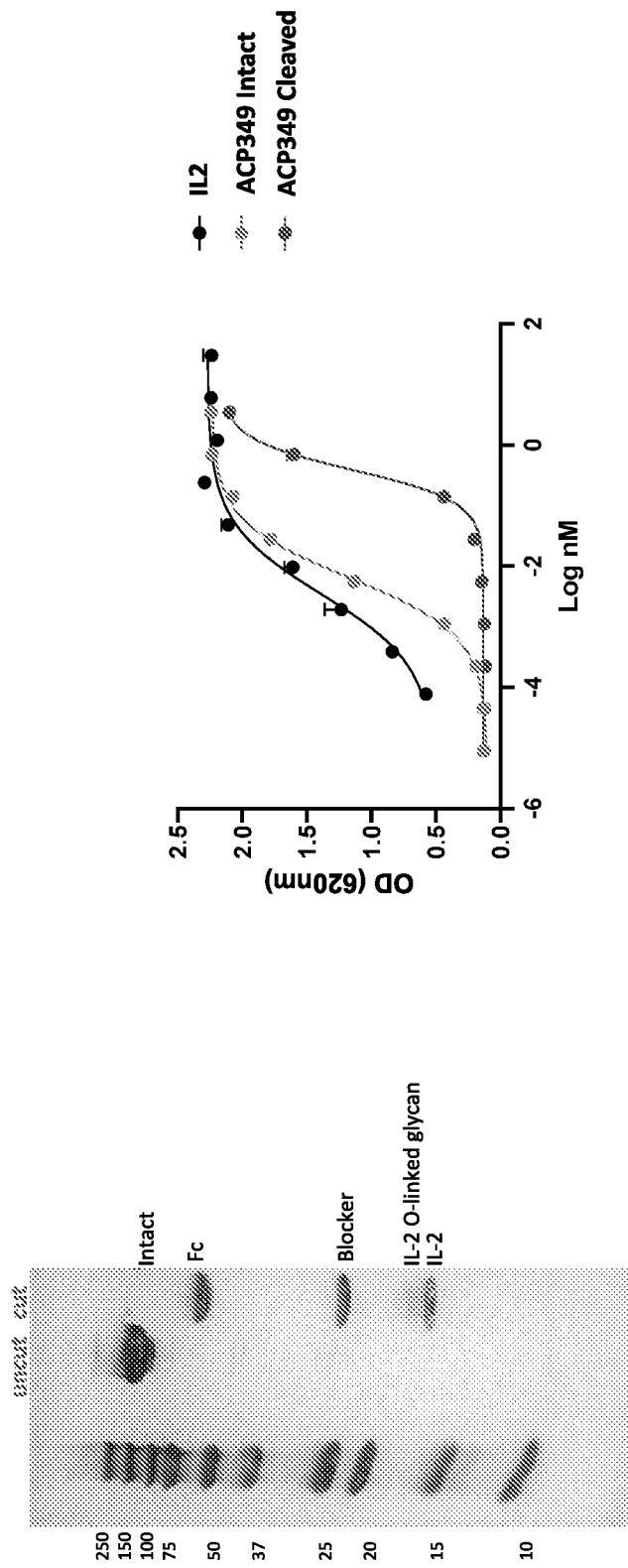


FIG. 52L

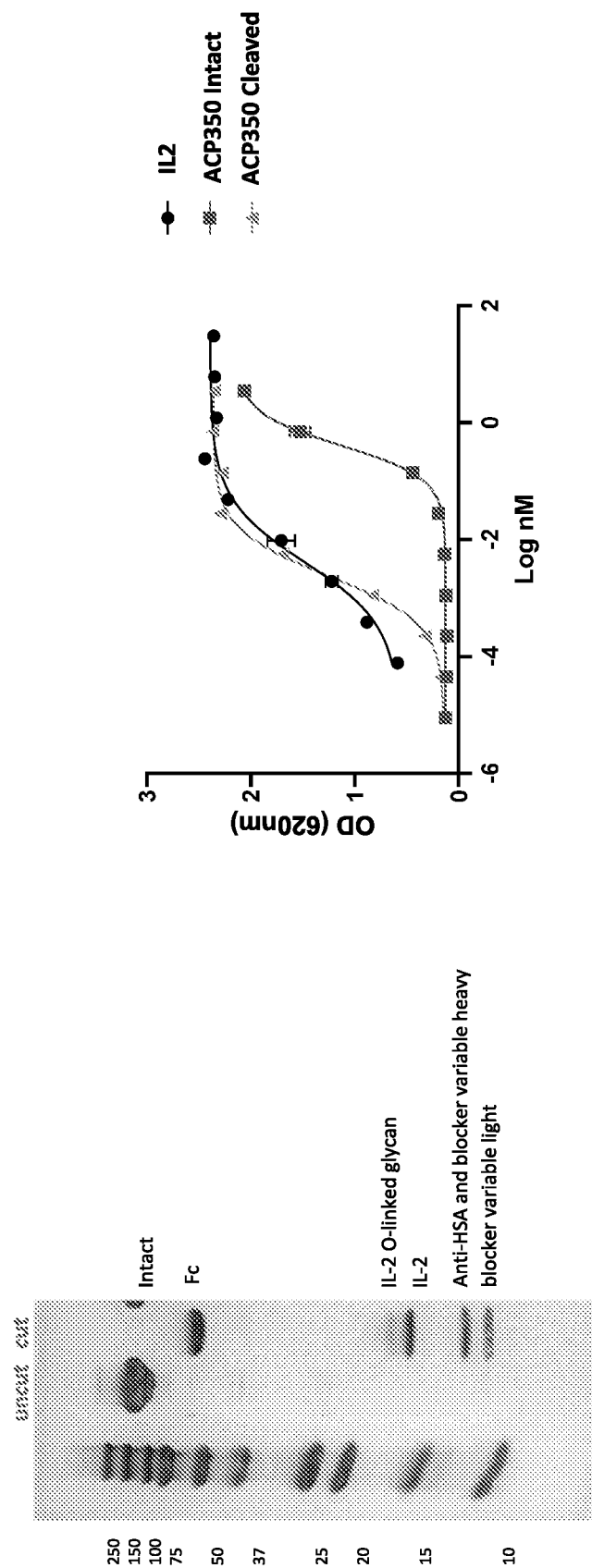


FIG. 52M

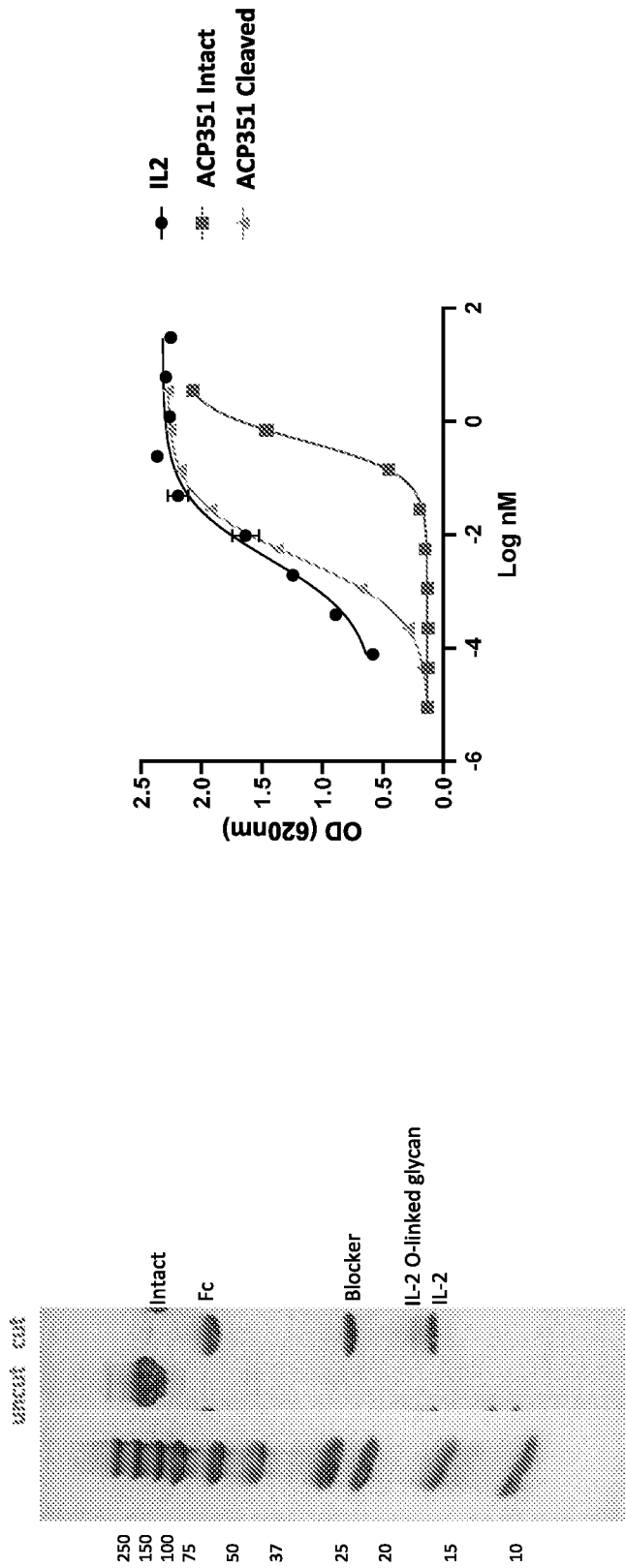


FIG. 52N

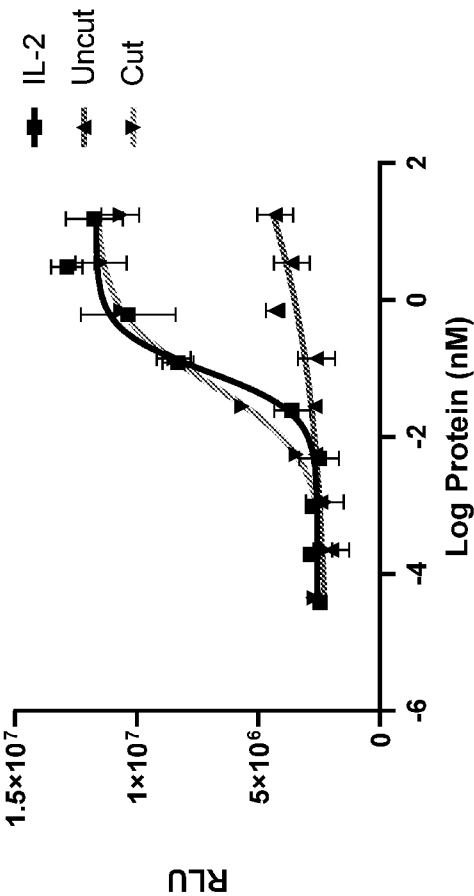


FIG. 53A

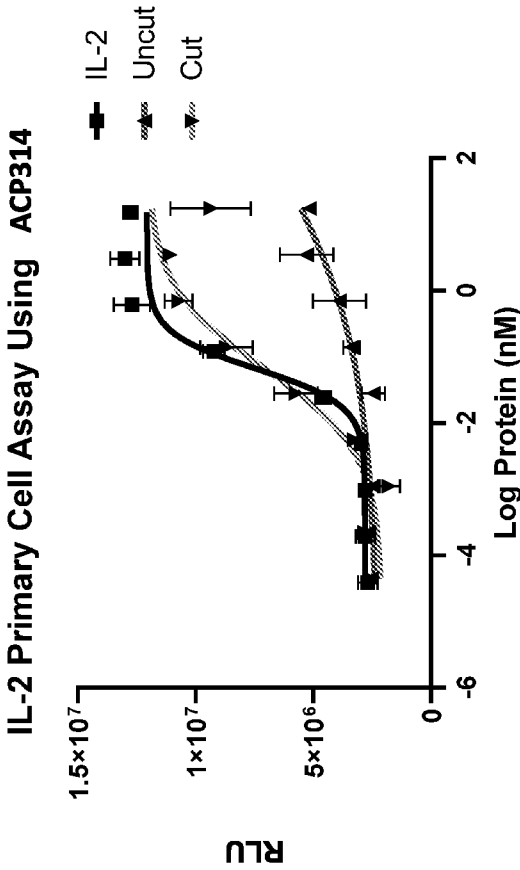


FIG. 53B

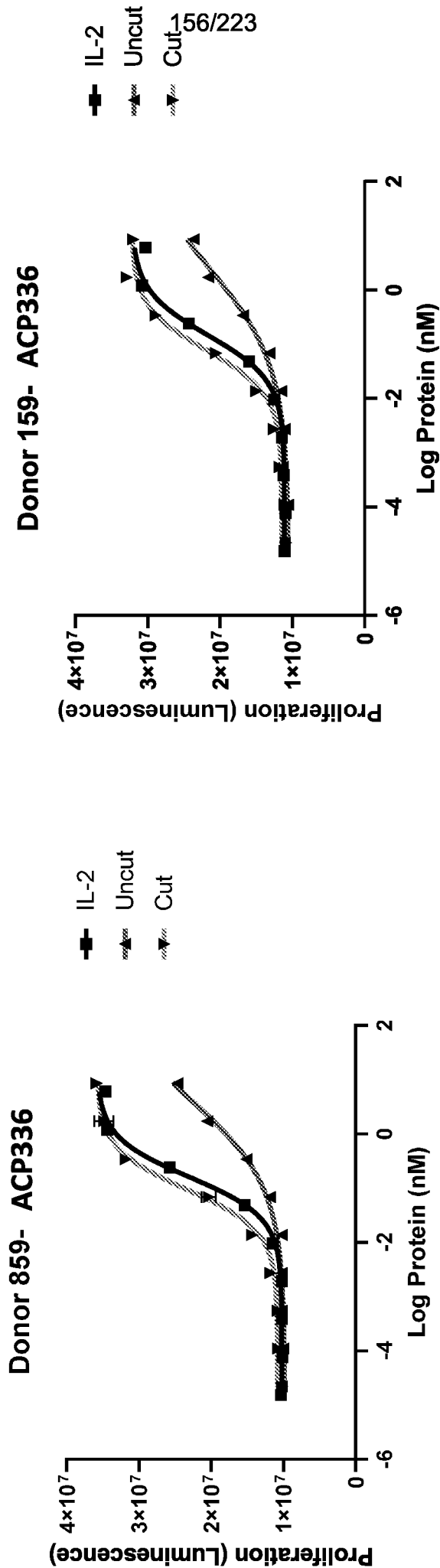


FIG. 54A

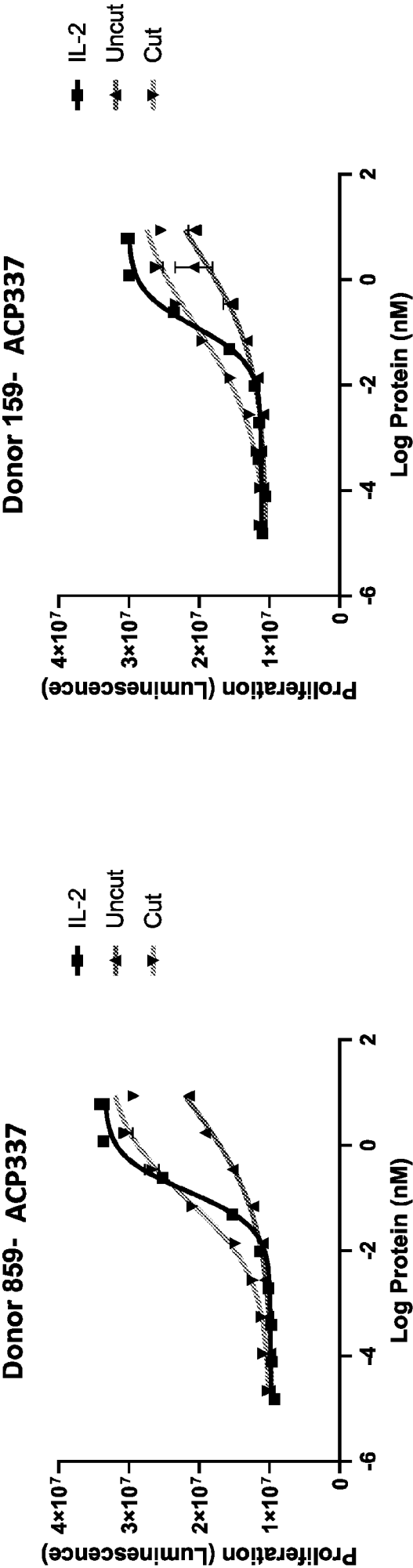


FIG. 54B

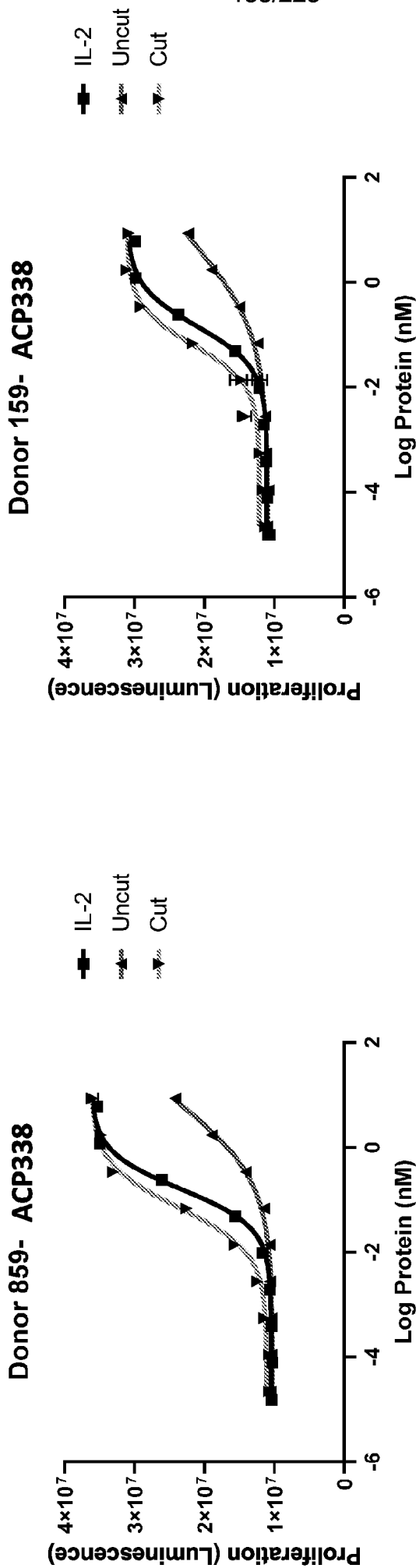


FIG. 54C

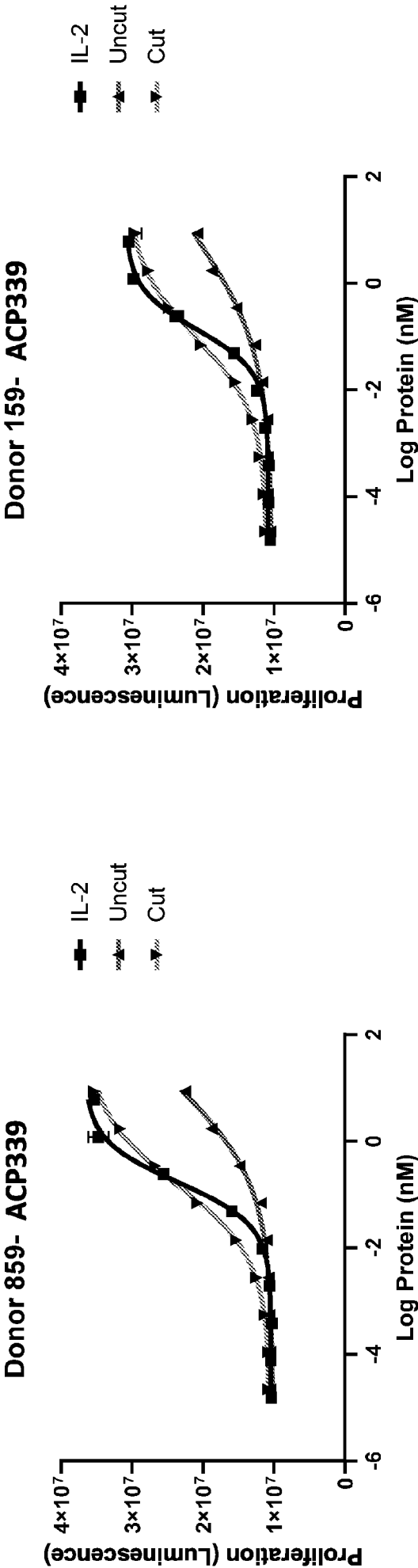


FIG. 54D

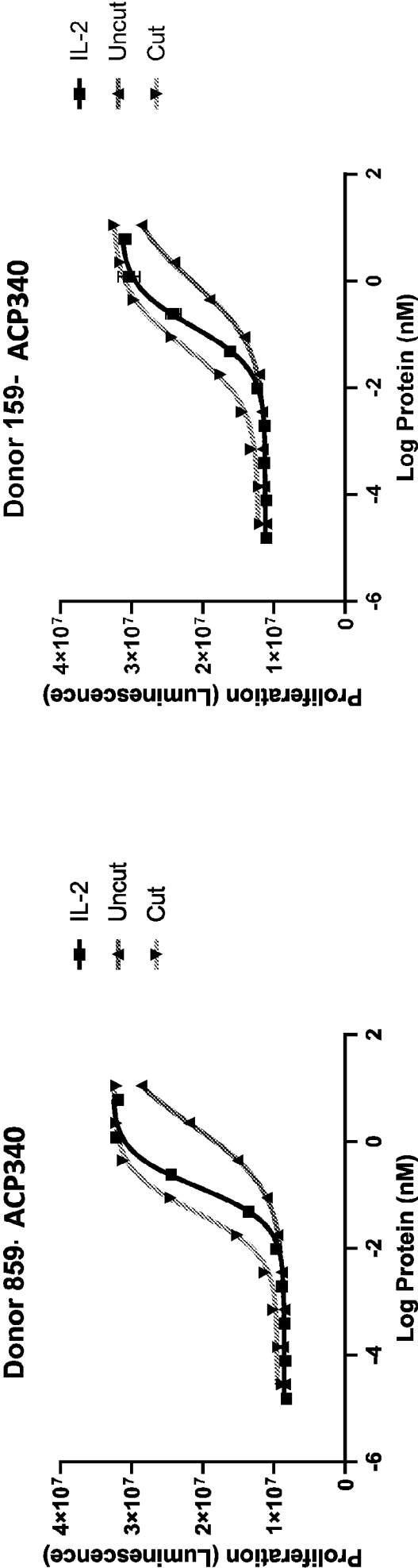


FIG. 54E

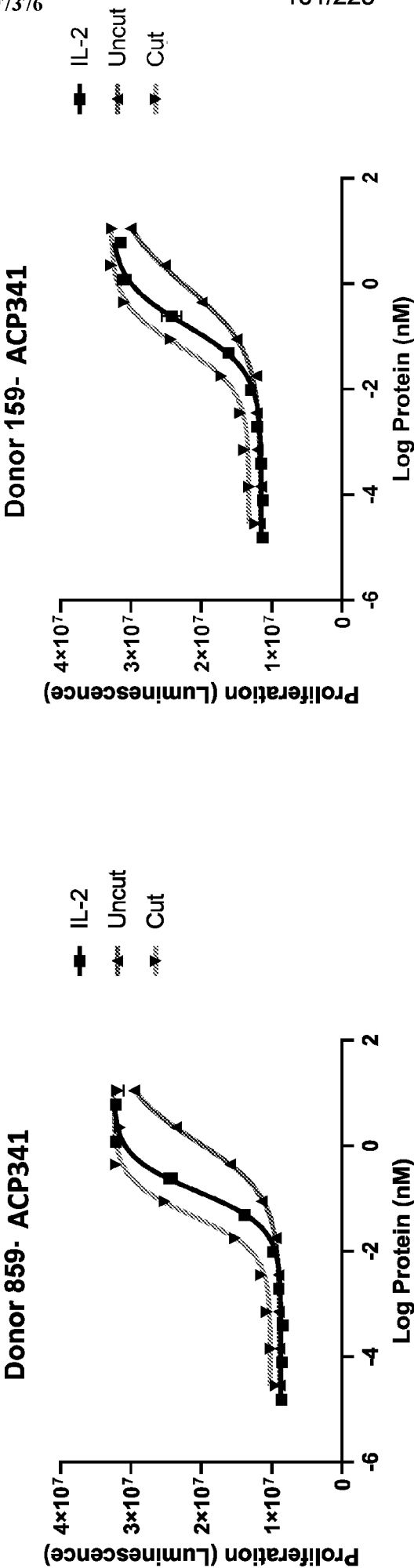


FIG. 54F

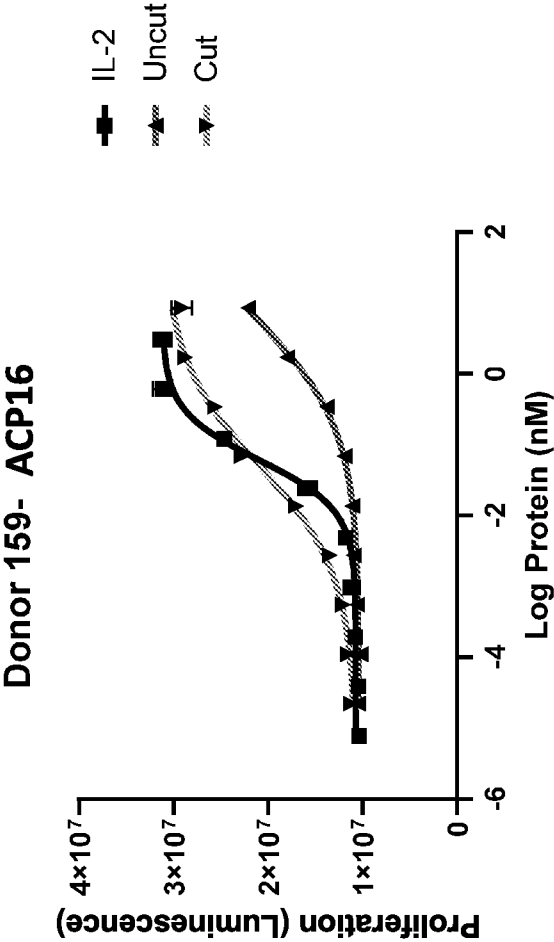


FIG. 54G

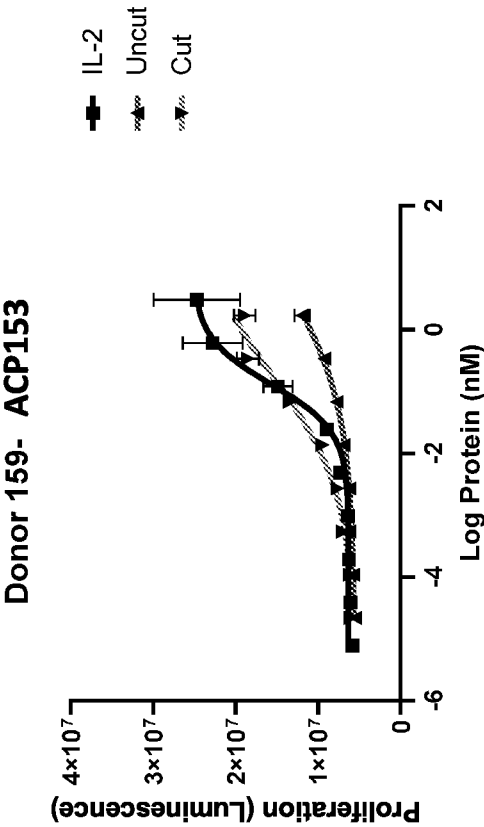


FIG. 54H

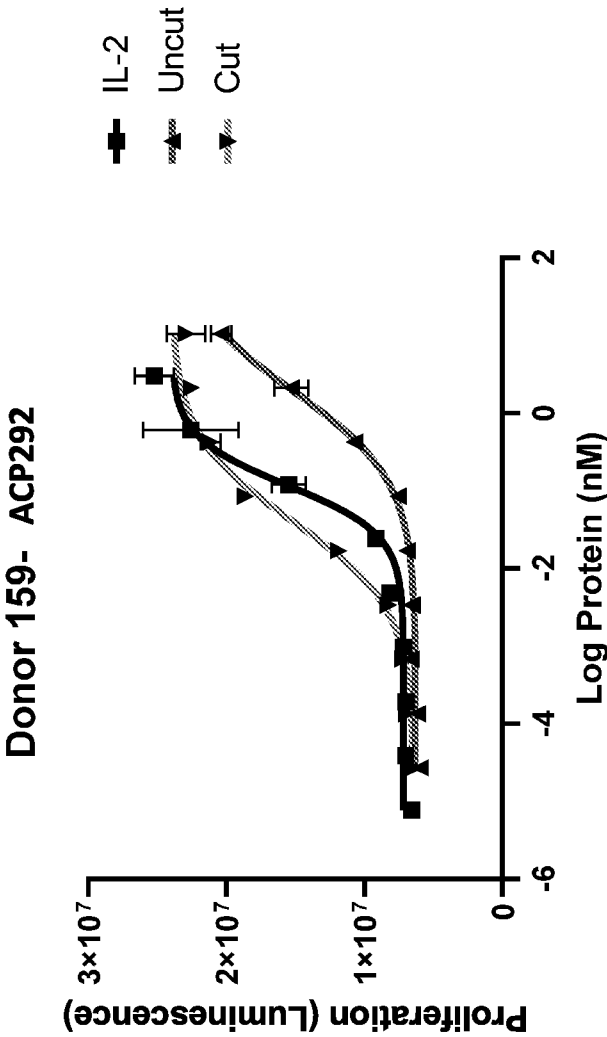


FIG. 54I

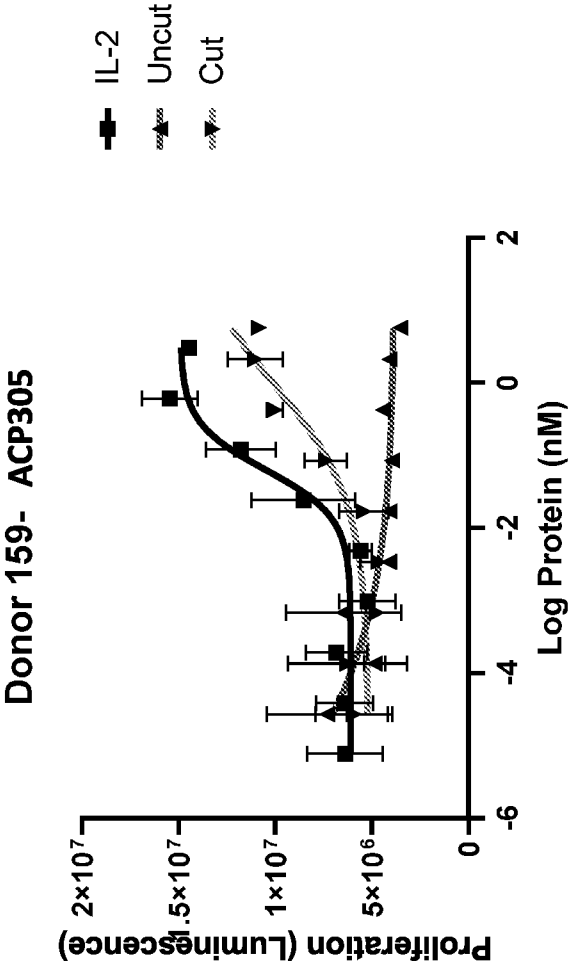


FIG. 54J

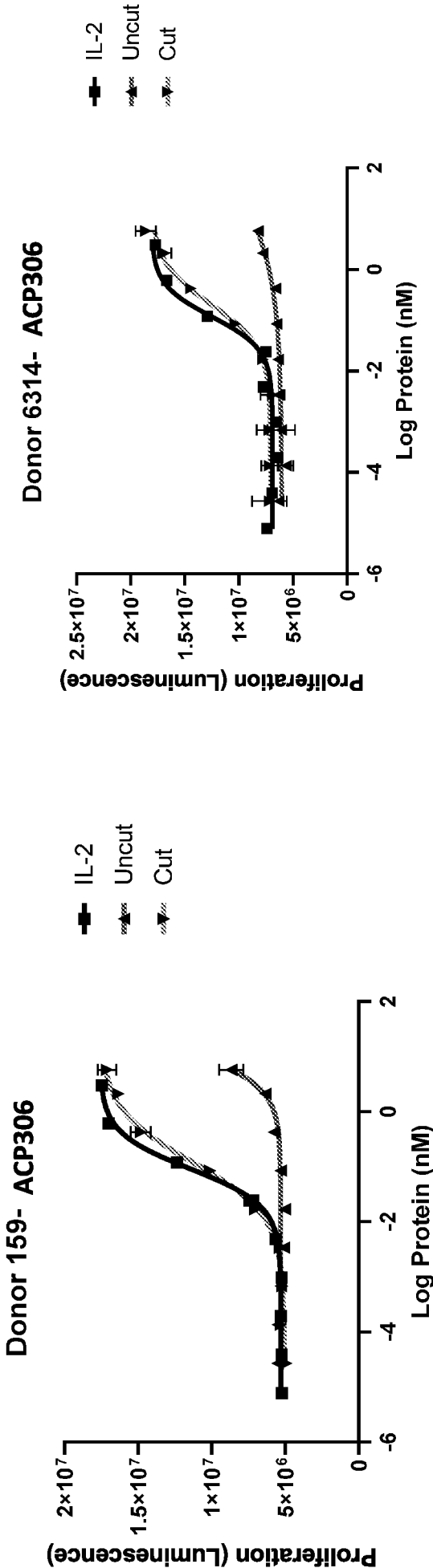


FIG. 54K

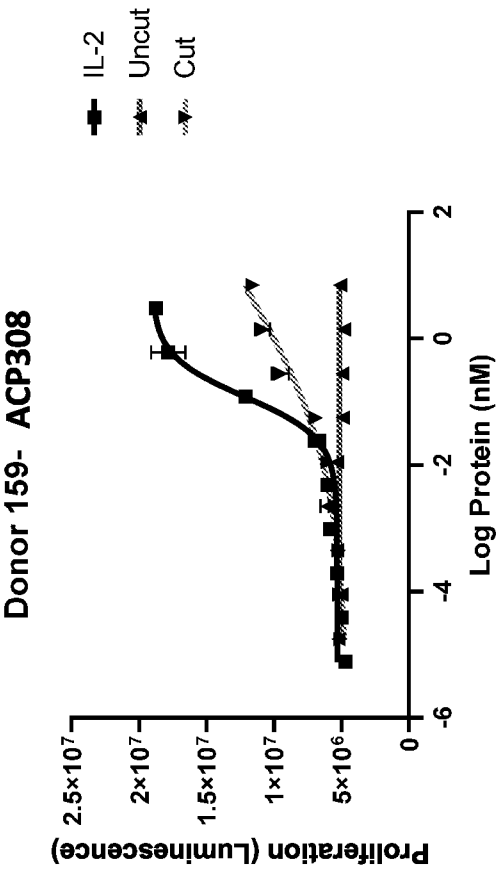


FIG. 54L

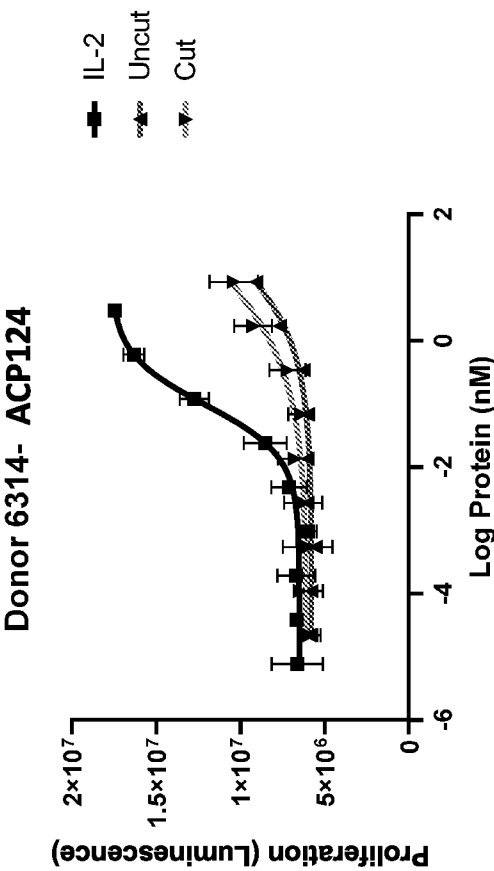


FIG. 54M

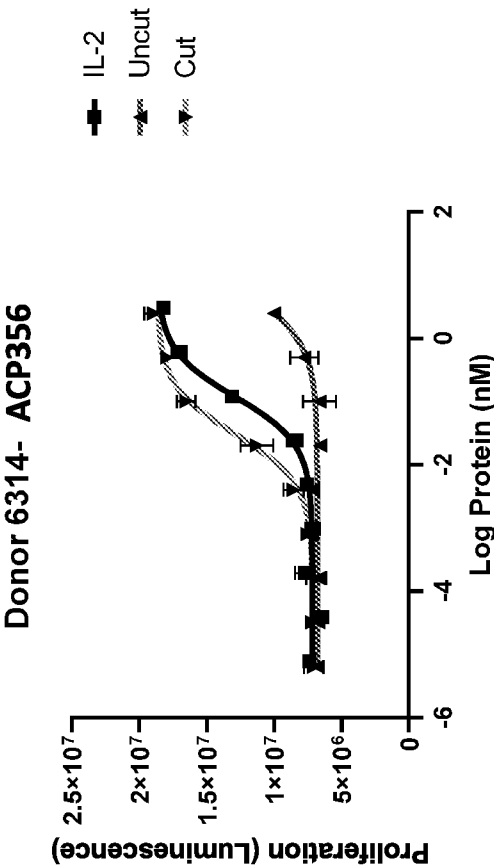


FIG. 54N

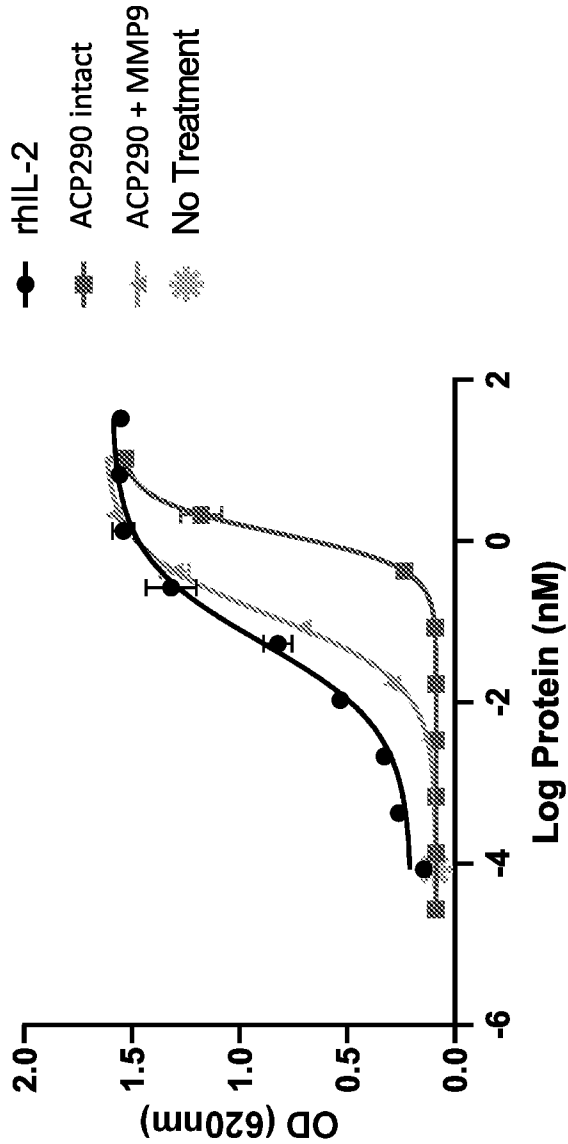
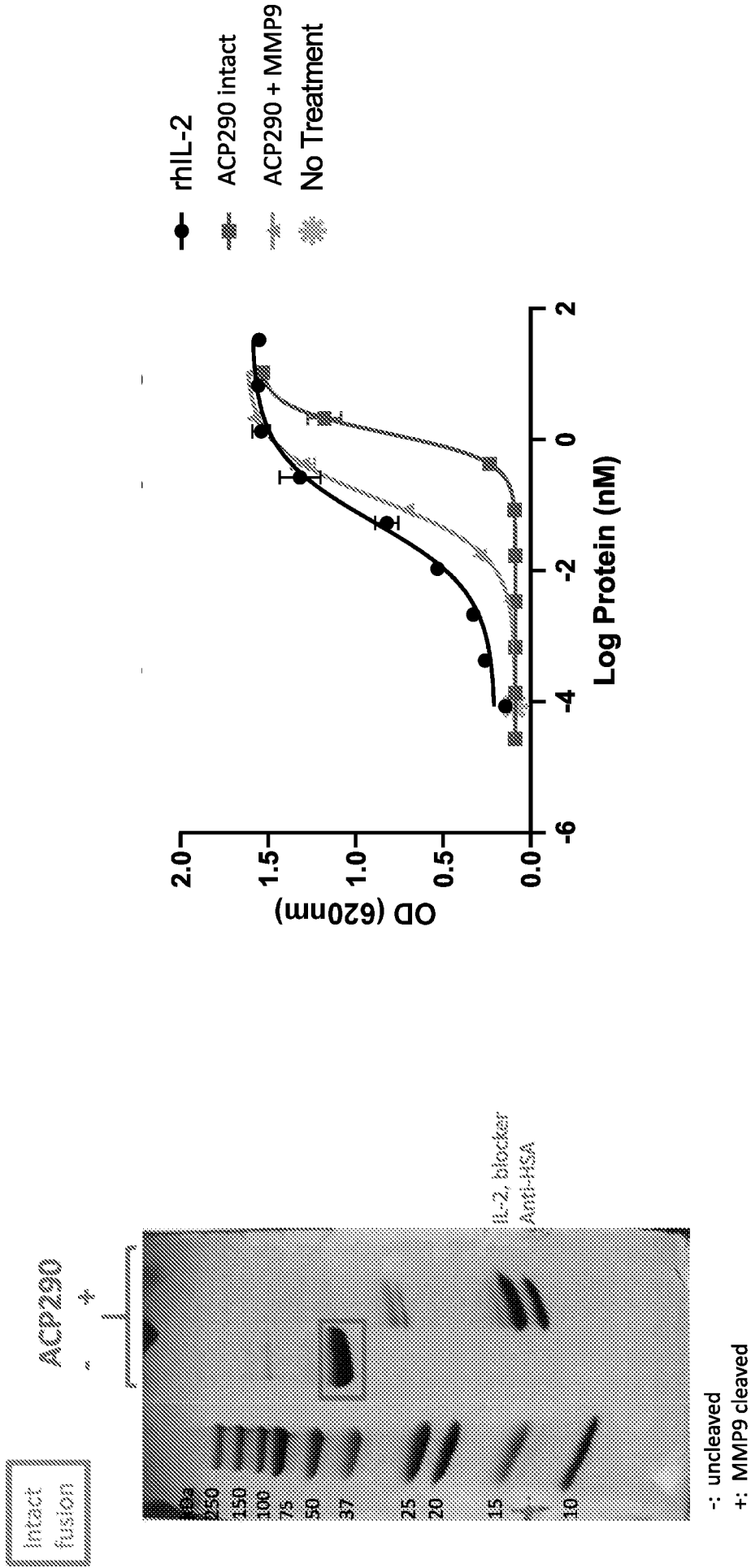


FIG. 55A

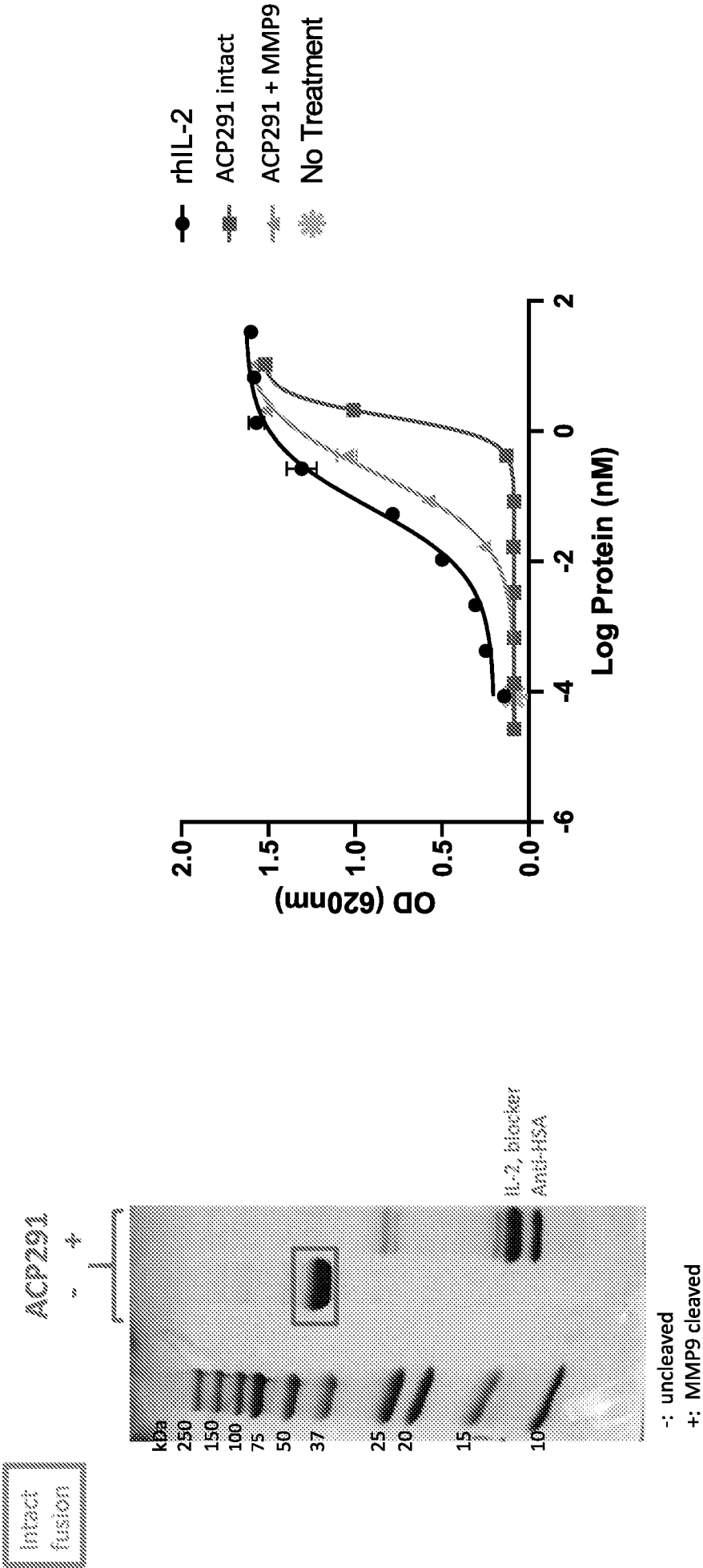


FIG. 55B

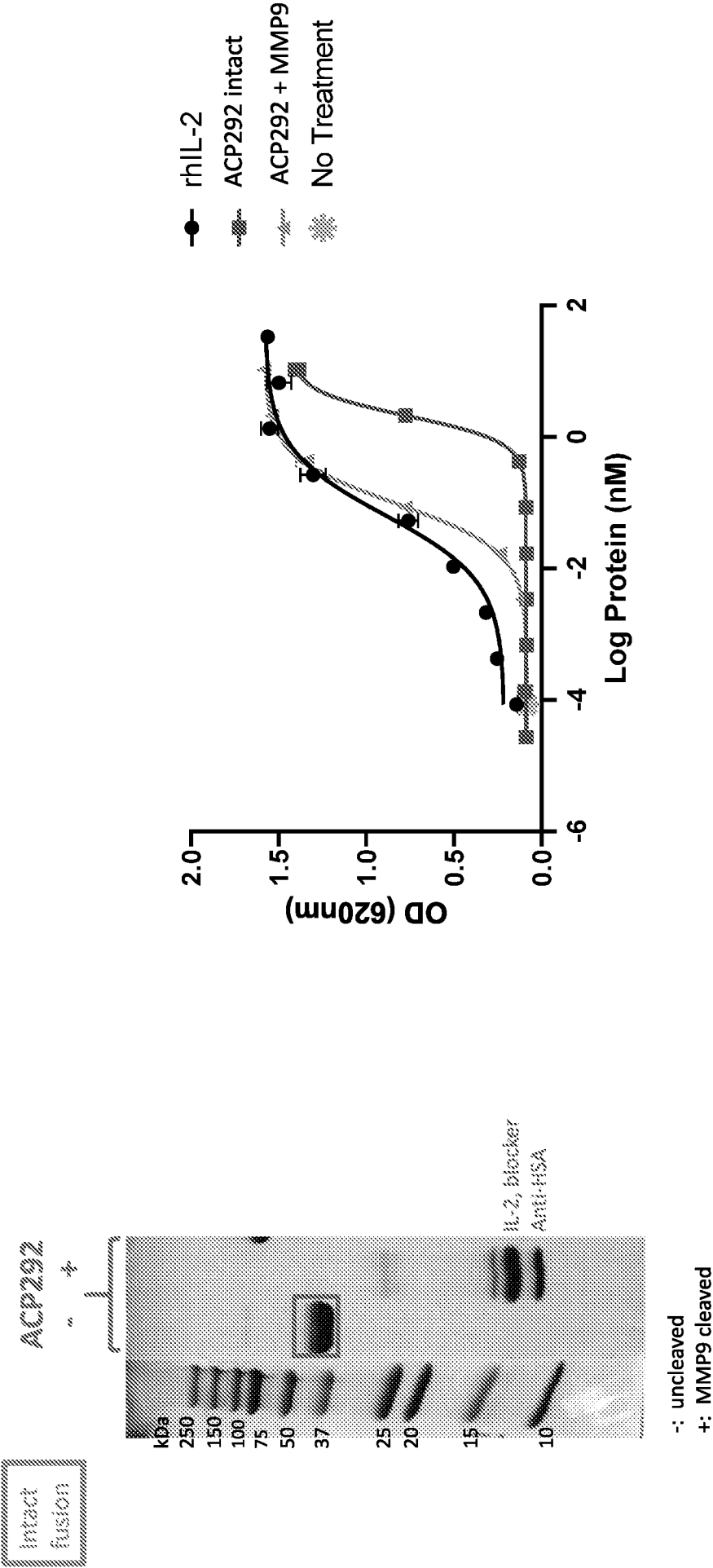


FIG. 55C

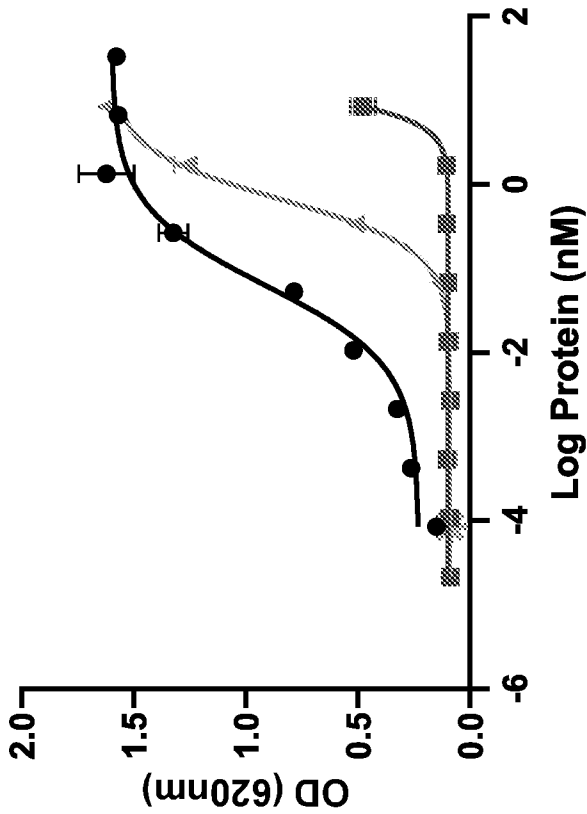
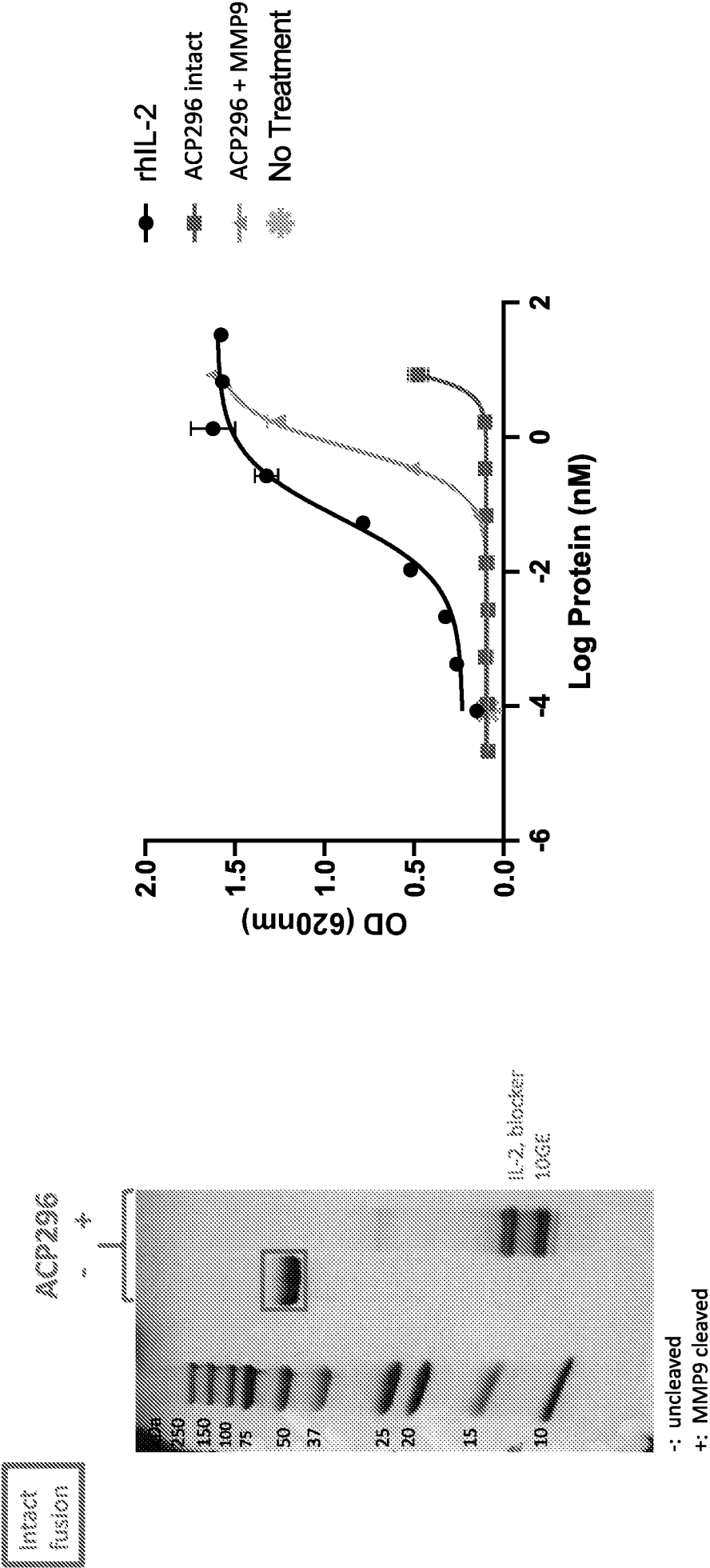
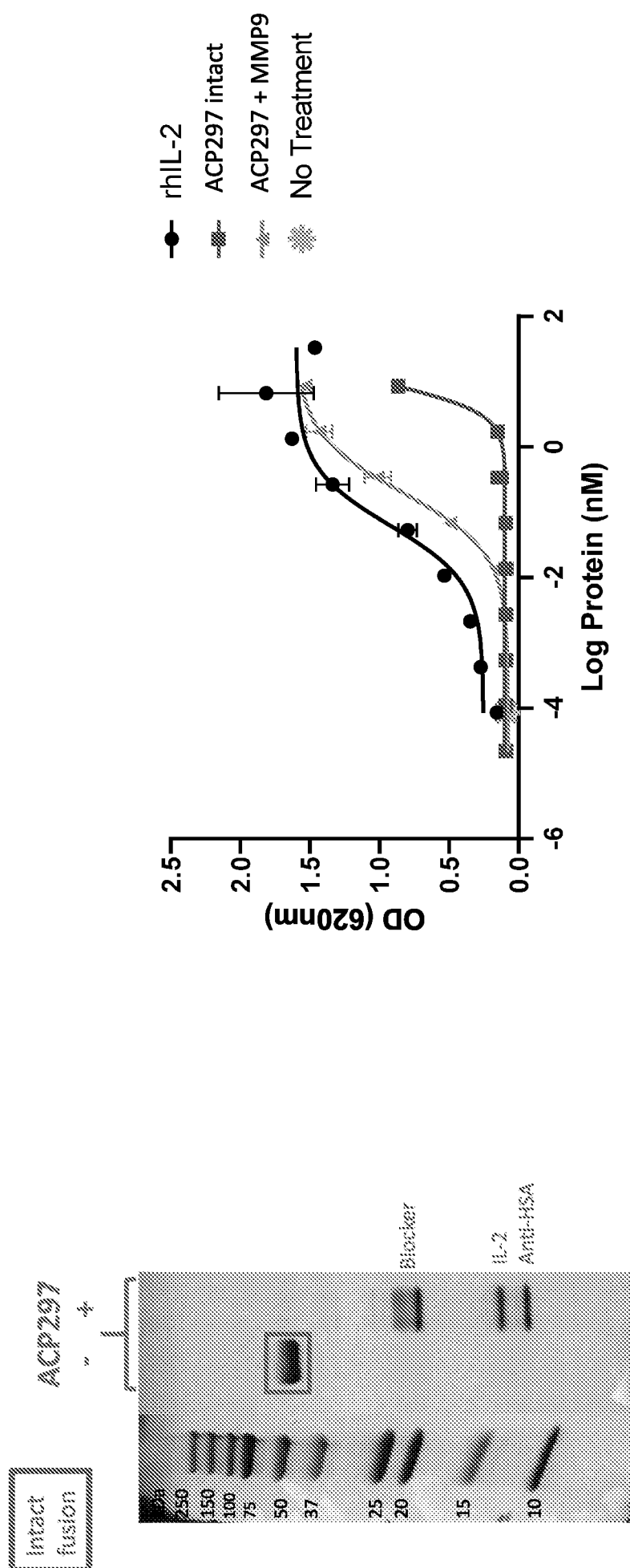
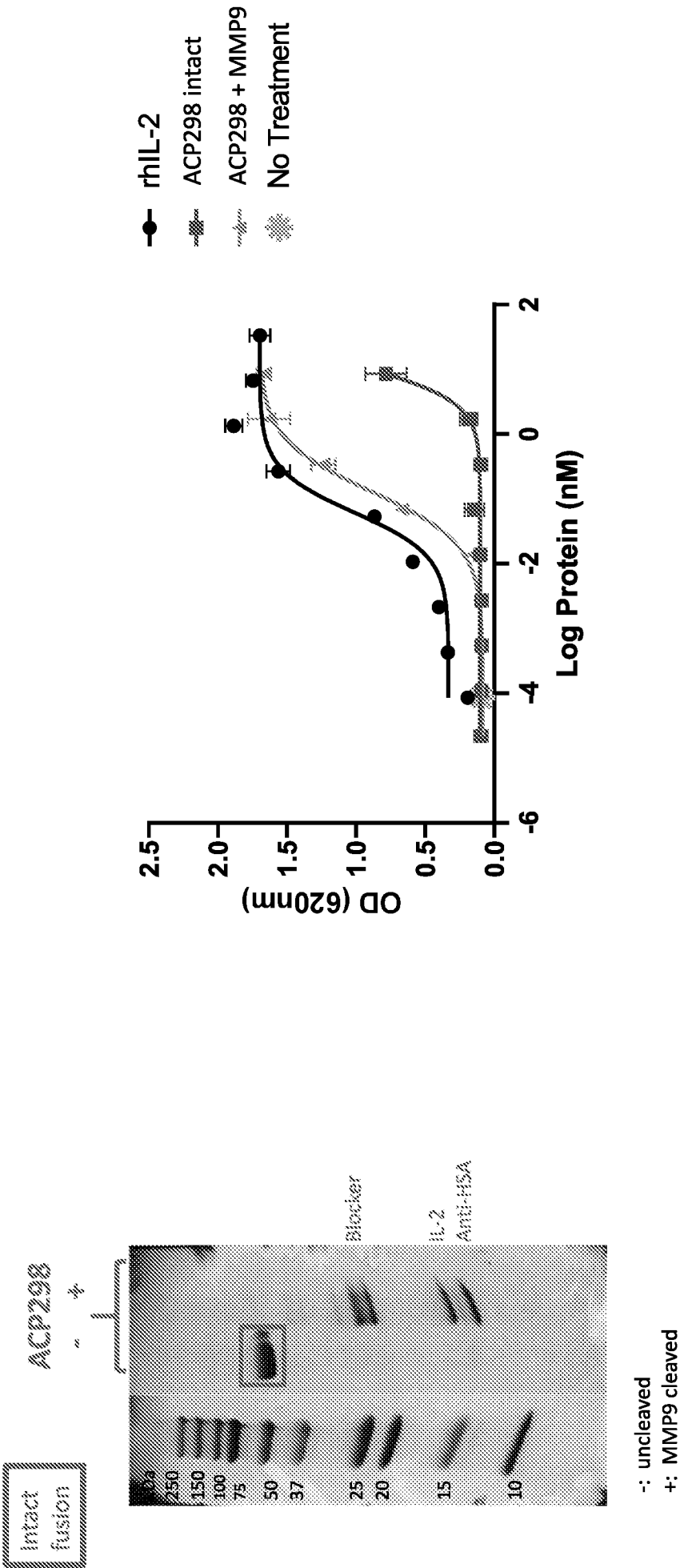


FIG. 55D



-: uncleaved
+: MMP9 cleaved

FIG. 55E



-: uncleaved
+: MMP9 cleaved

FIG. 55F

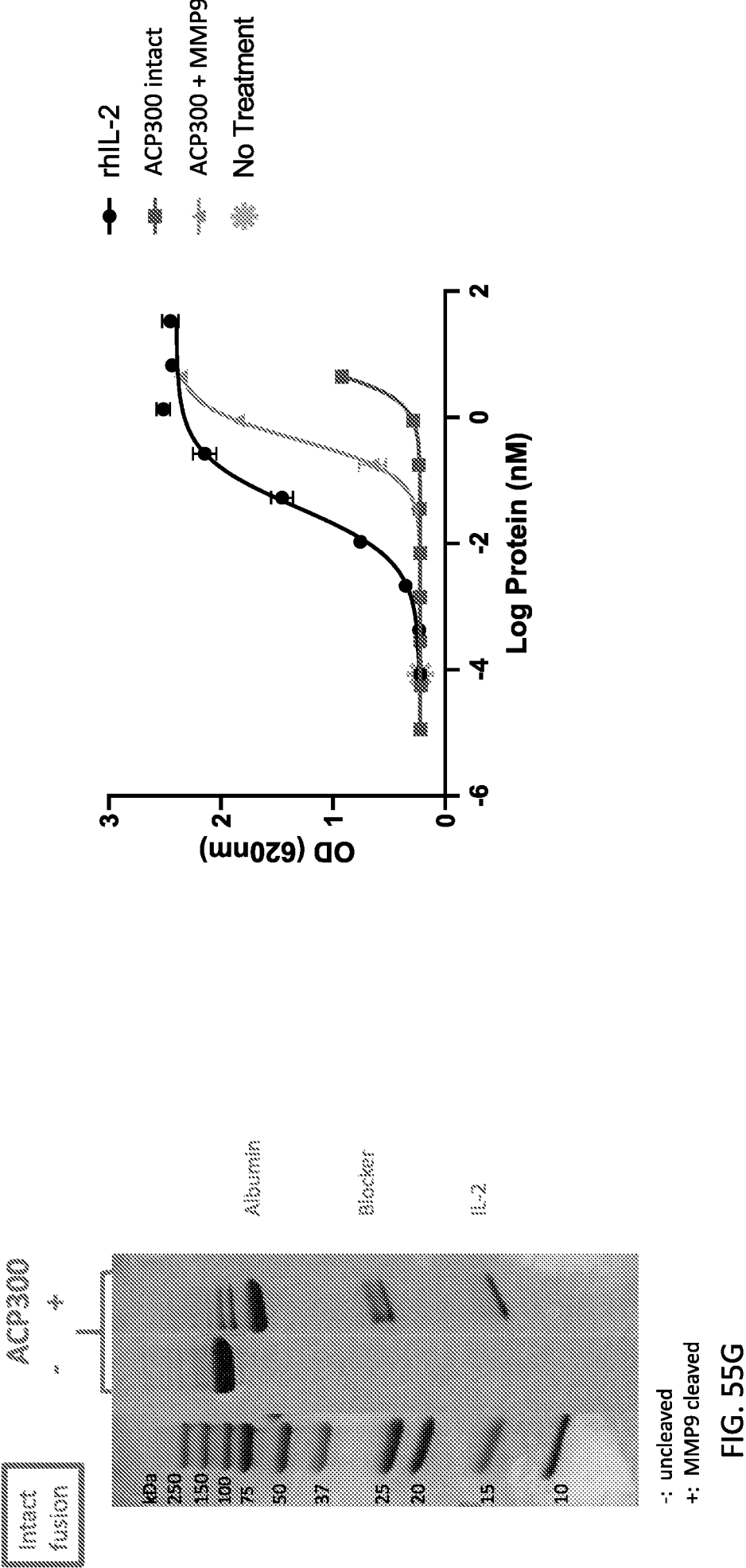


FIG. 55G

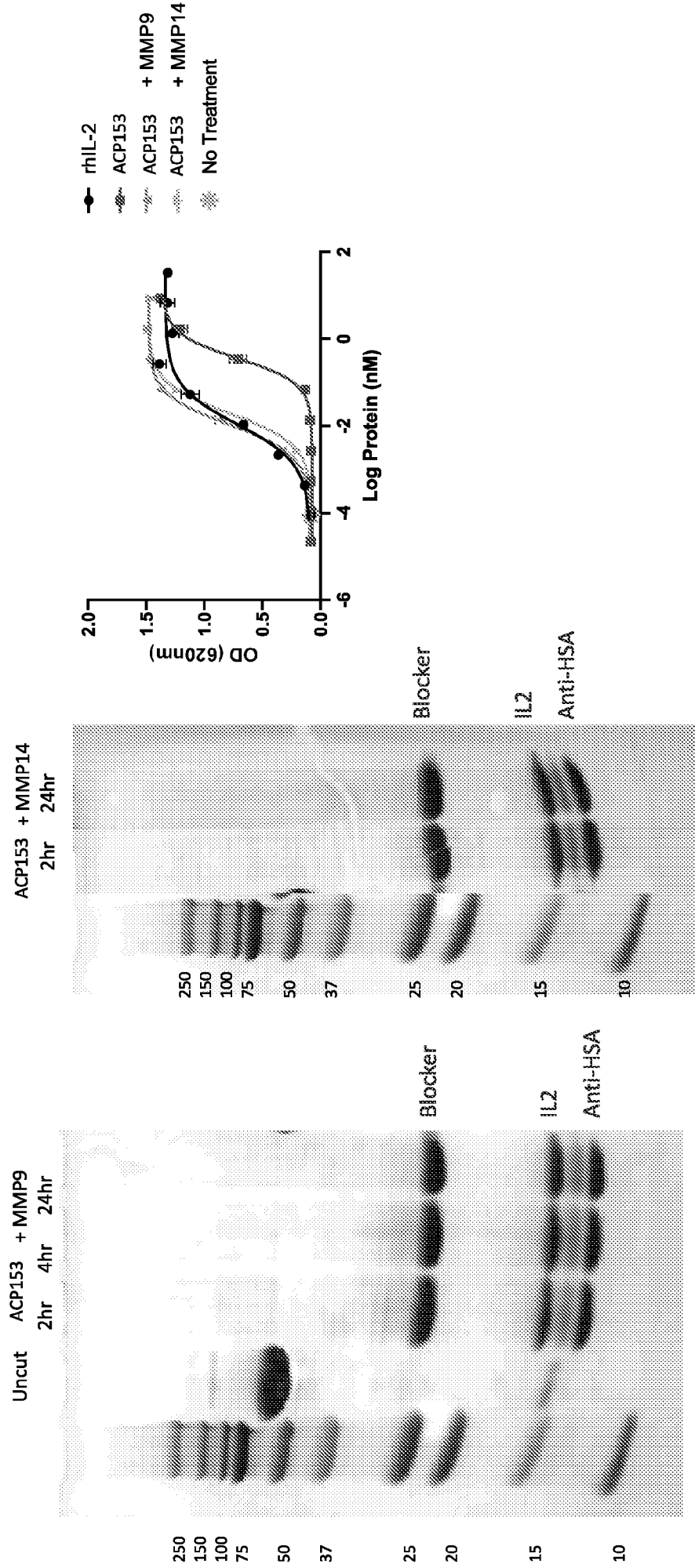


FIG. 55H

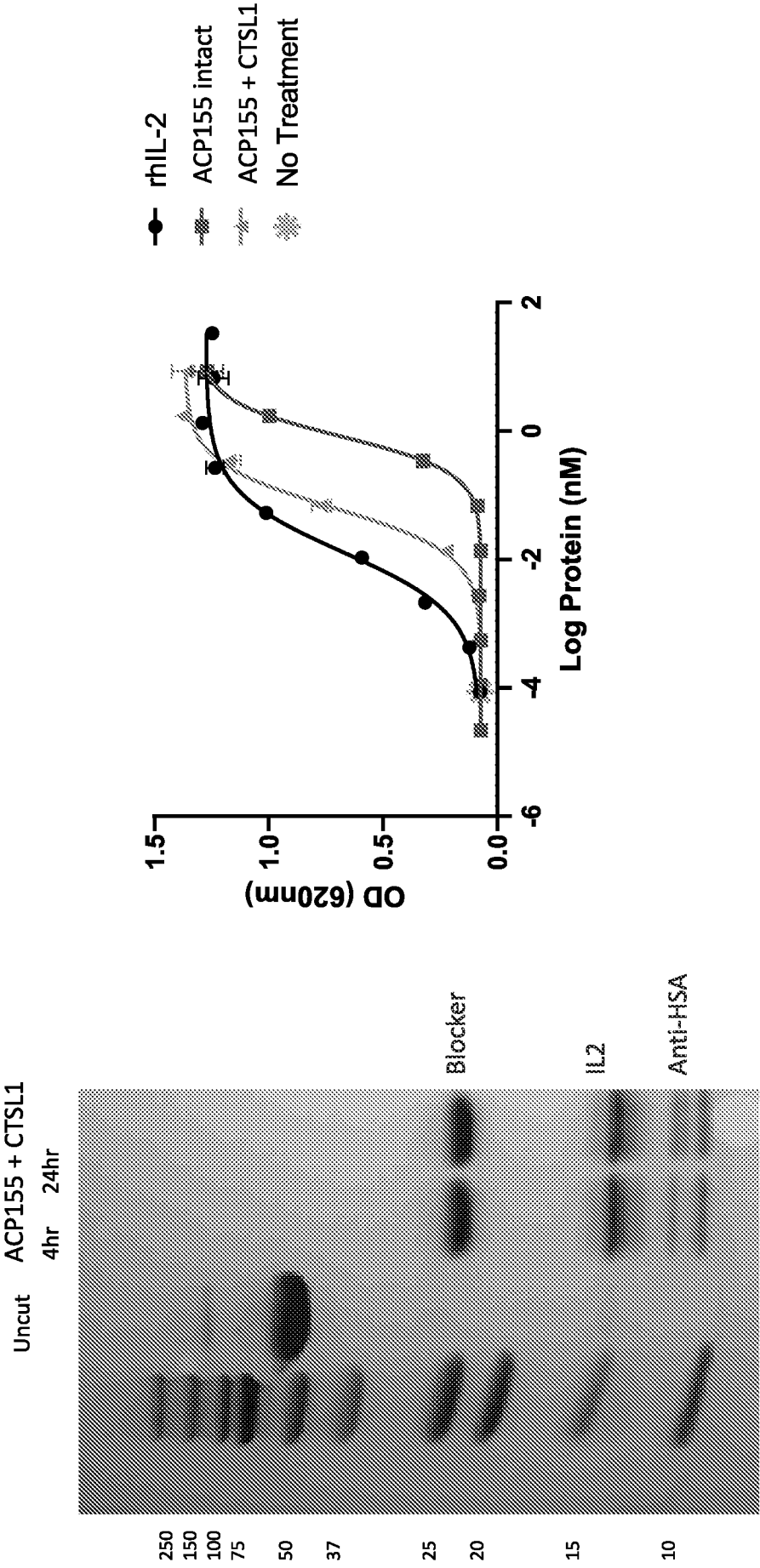


FIG. 55I

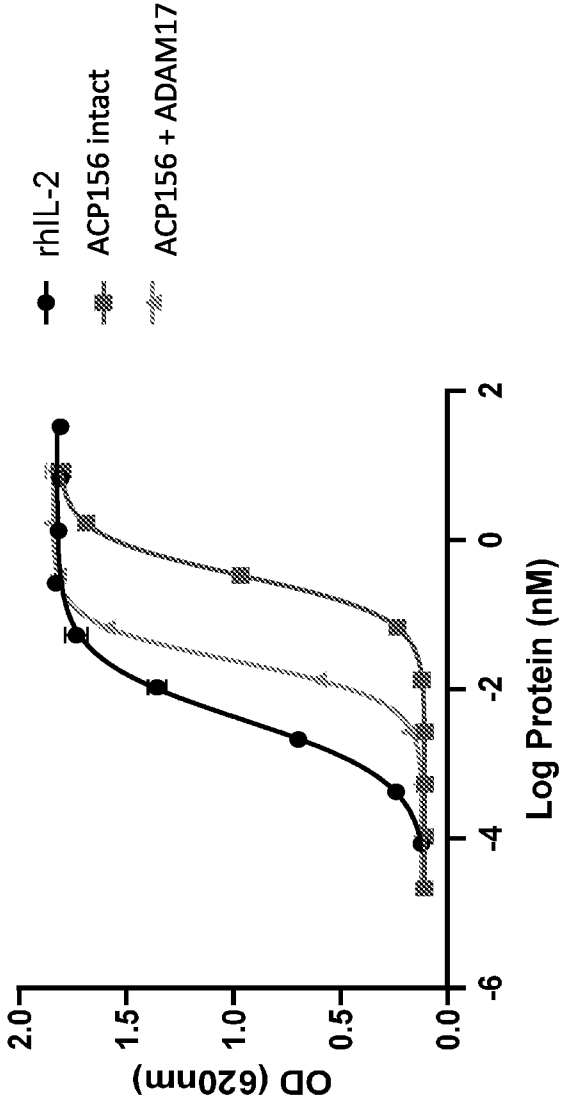
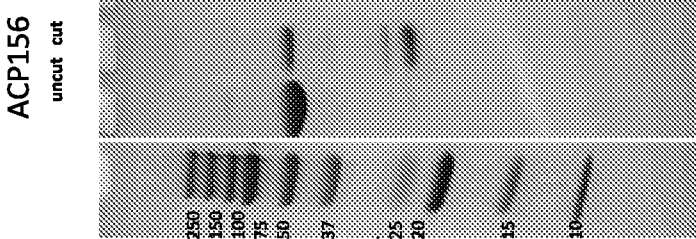


FIG. 55J

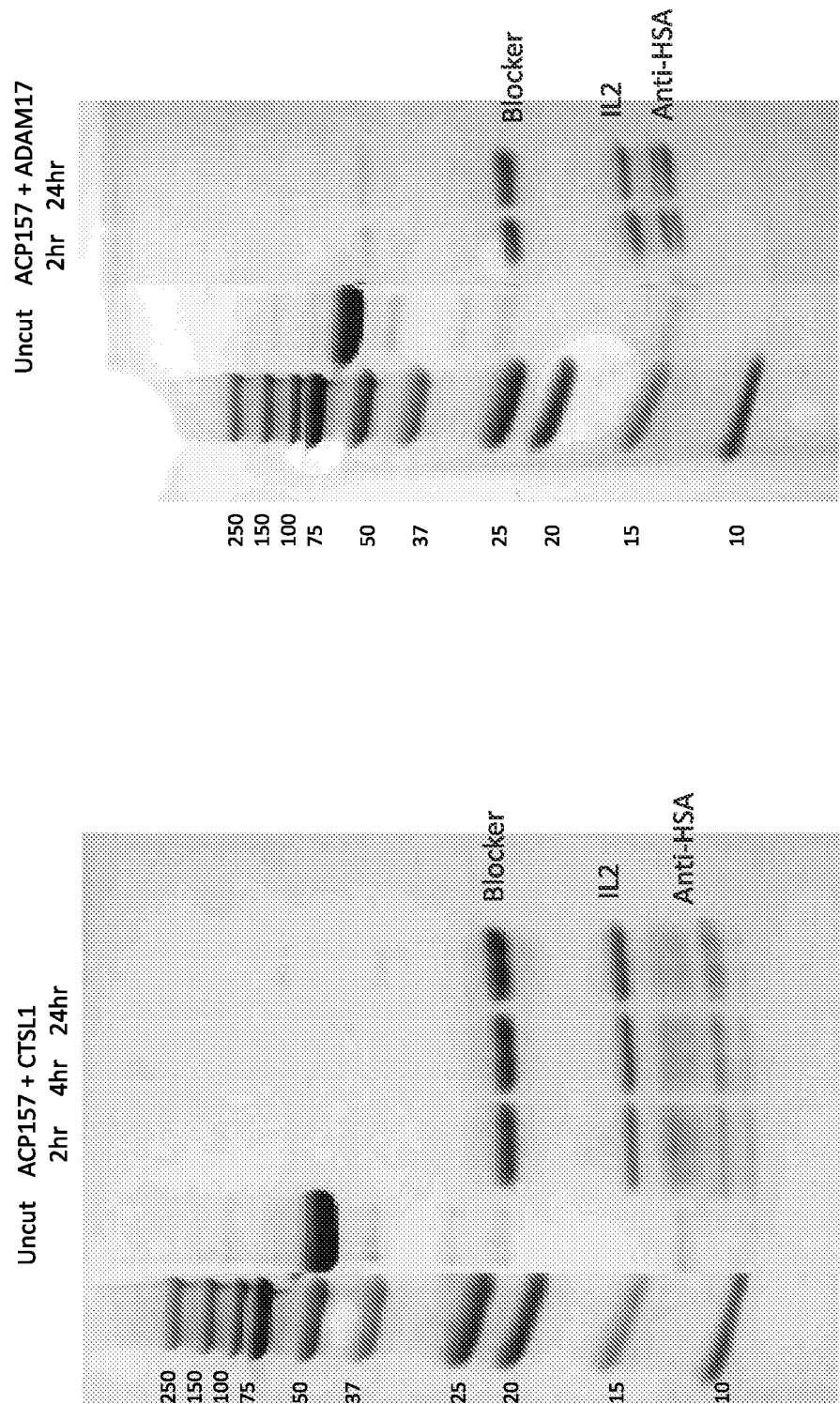


FIG. 55K

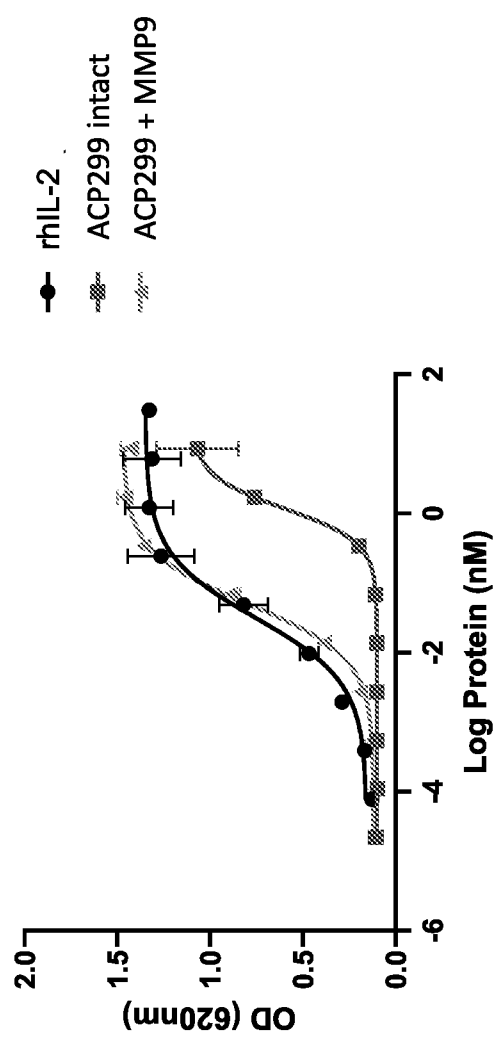


FIG. 55L

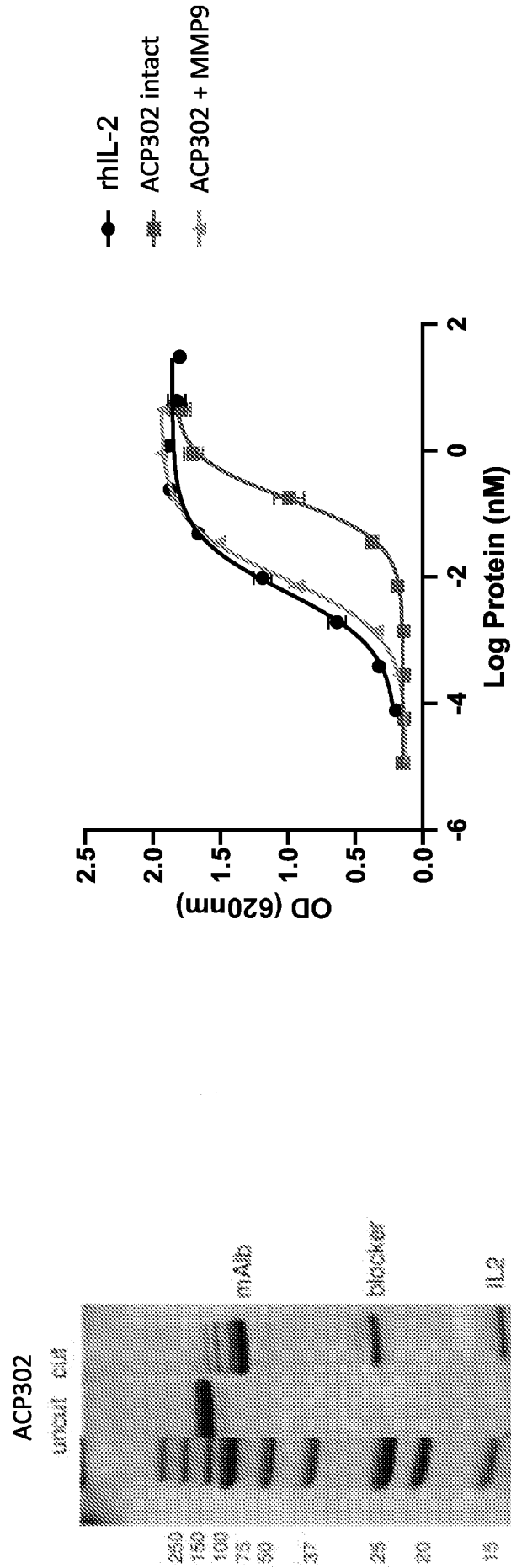
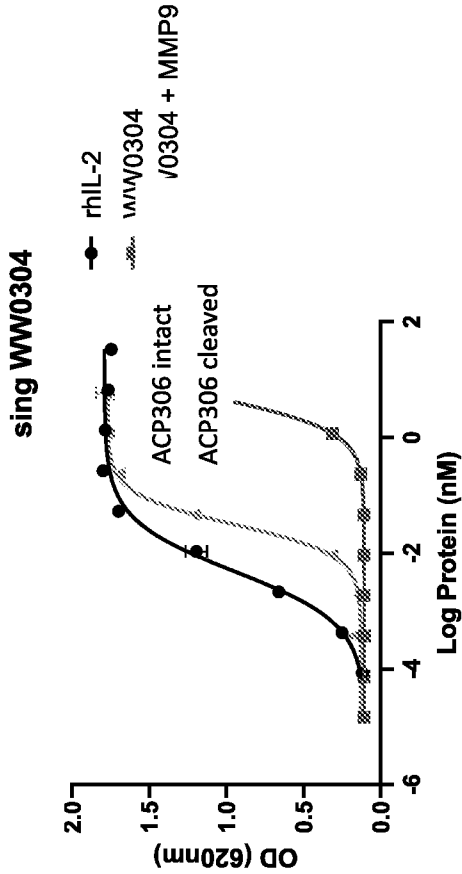
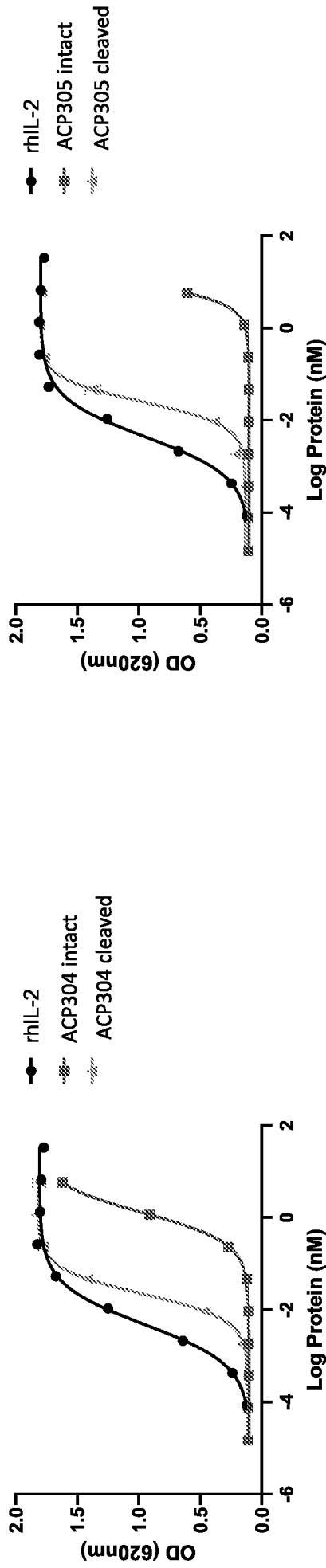


FIG. 55M

FIG. 55N



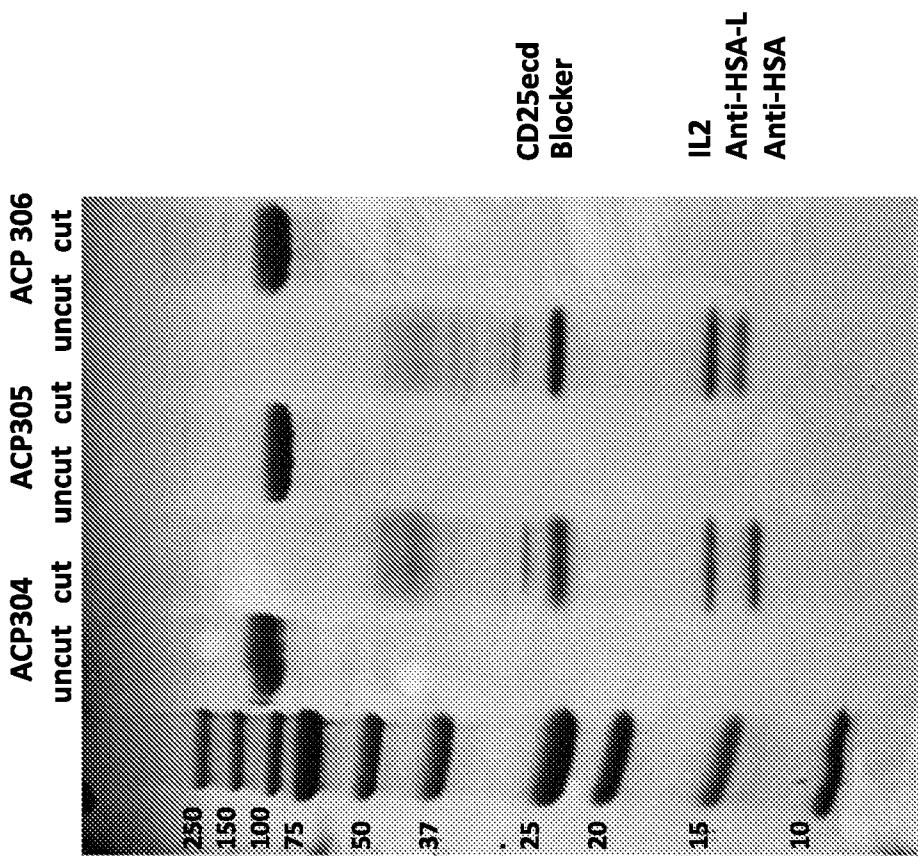


FIG. 56

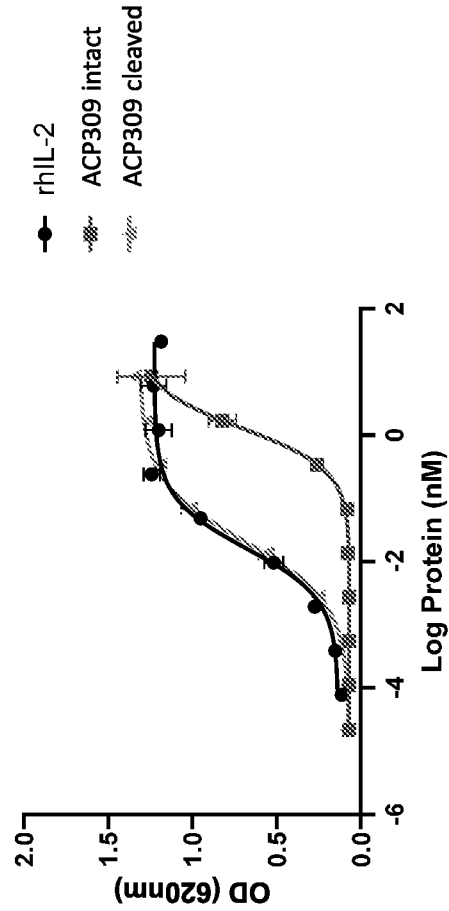


FIG. 57A

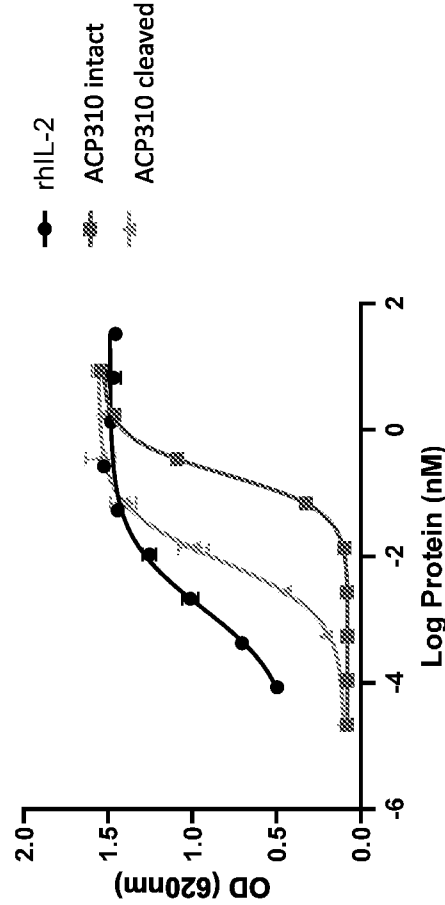


FIG. 57B

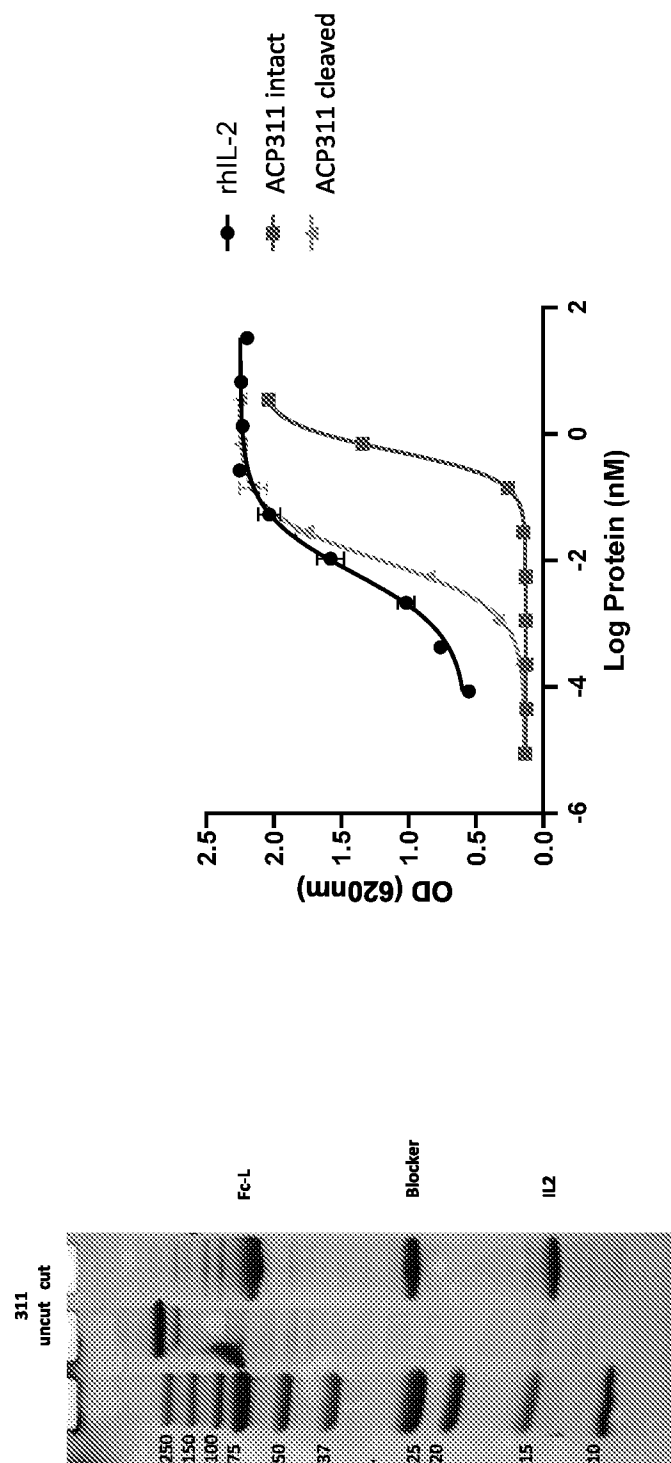


FIG. 57C

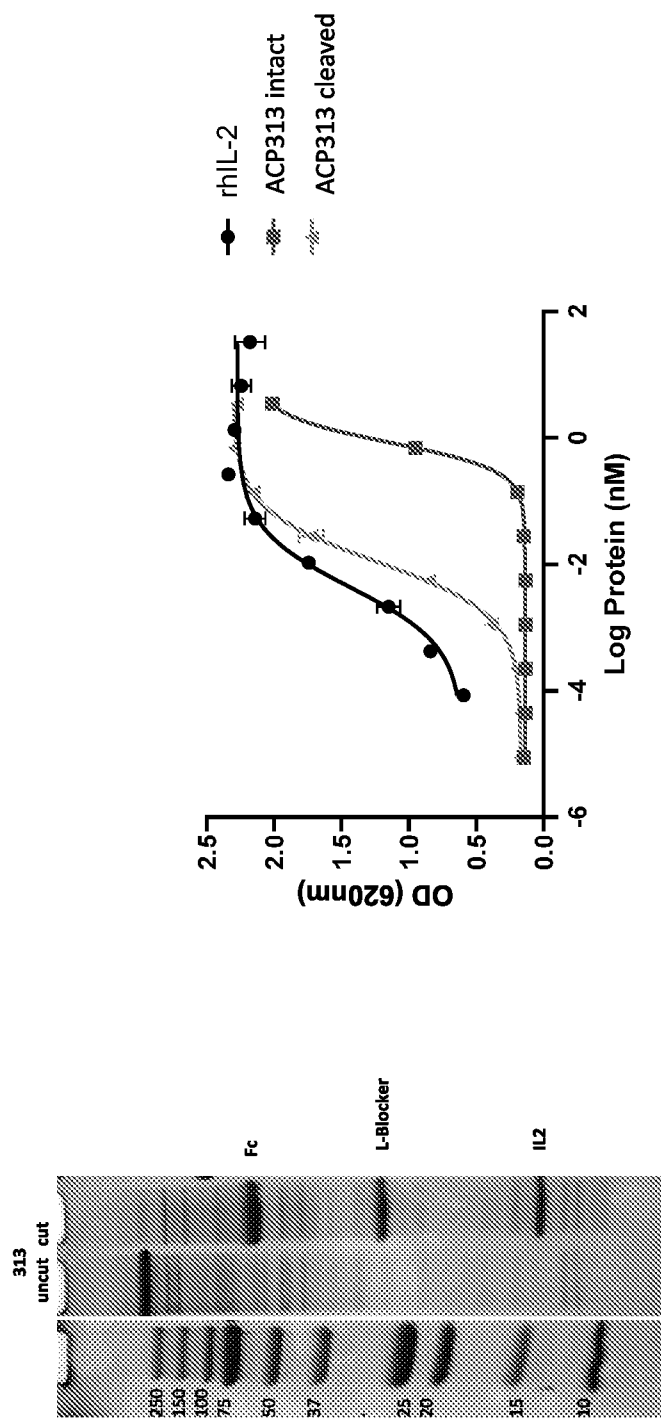


FIG. 57D

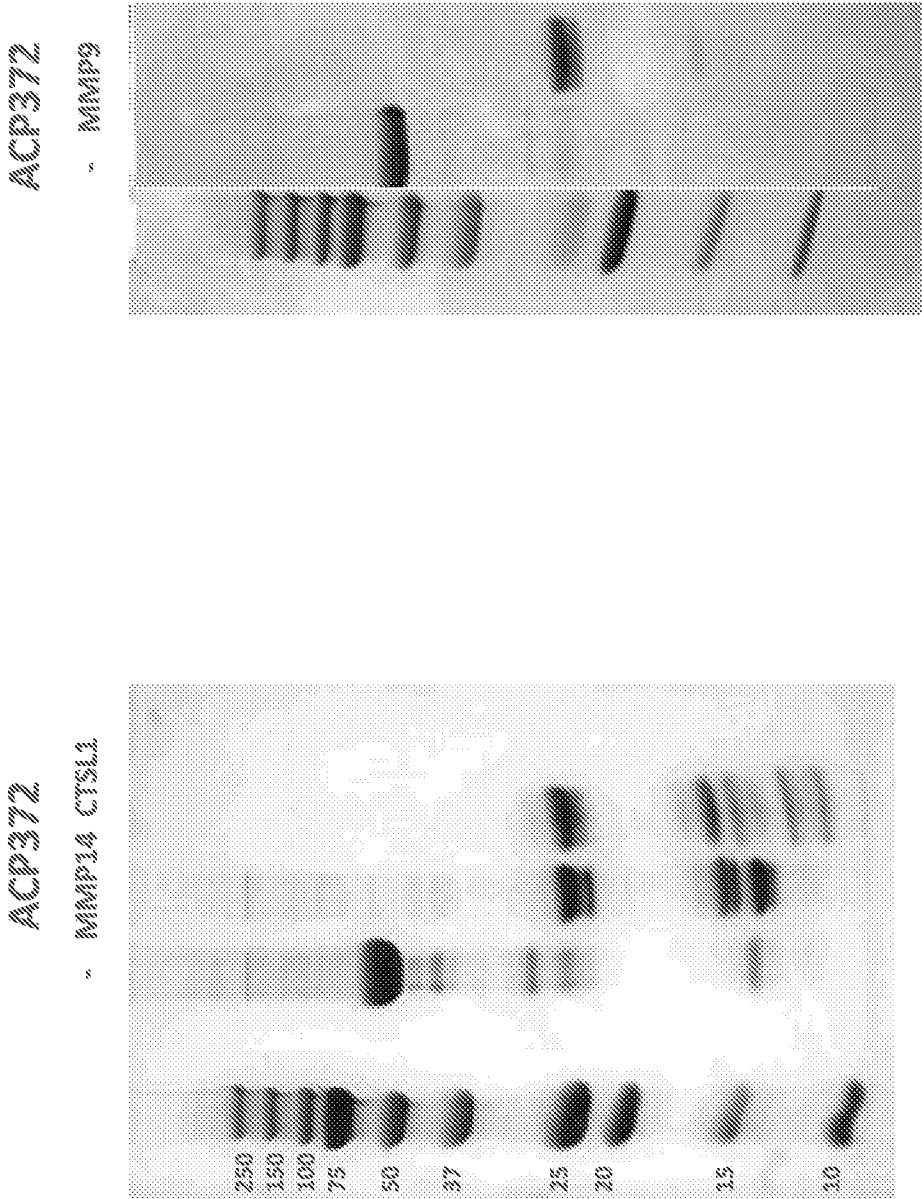


FIG. 58

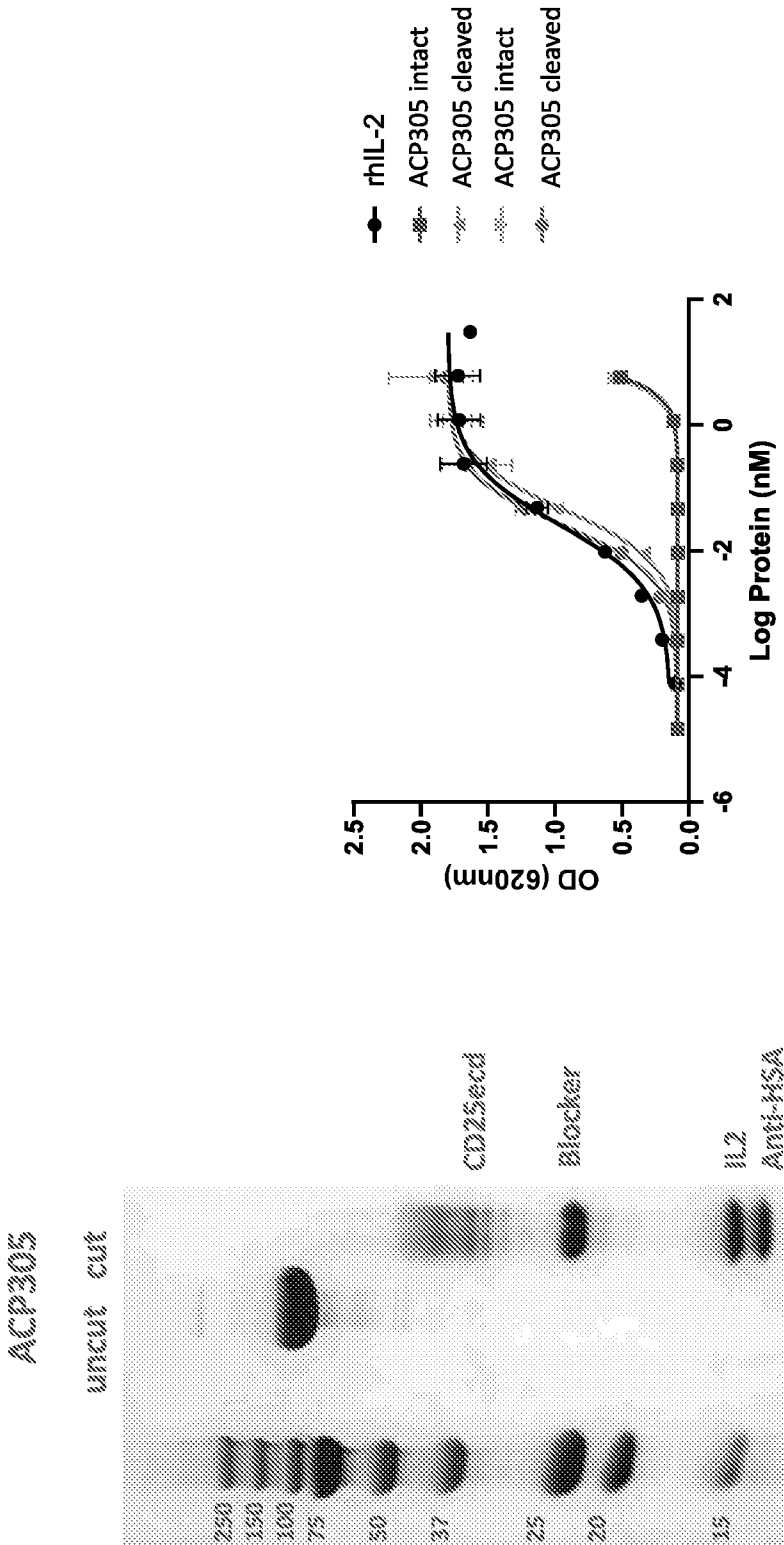


FIG. 59A

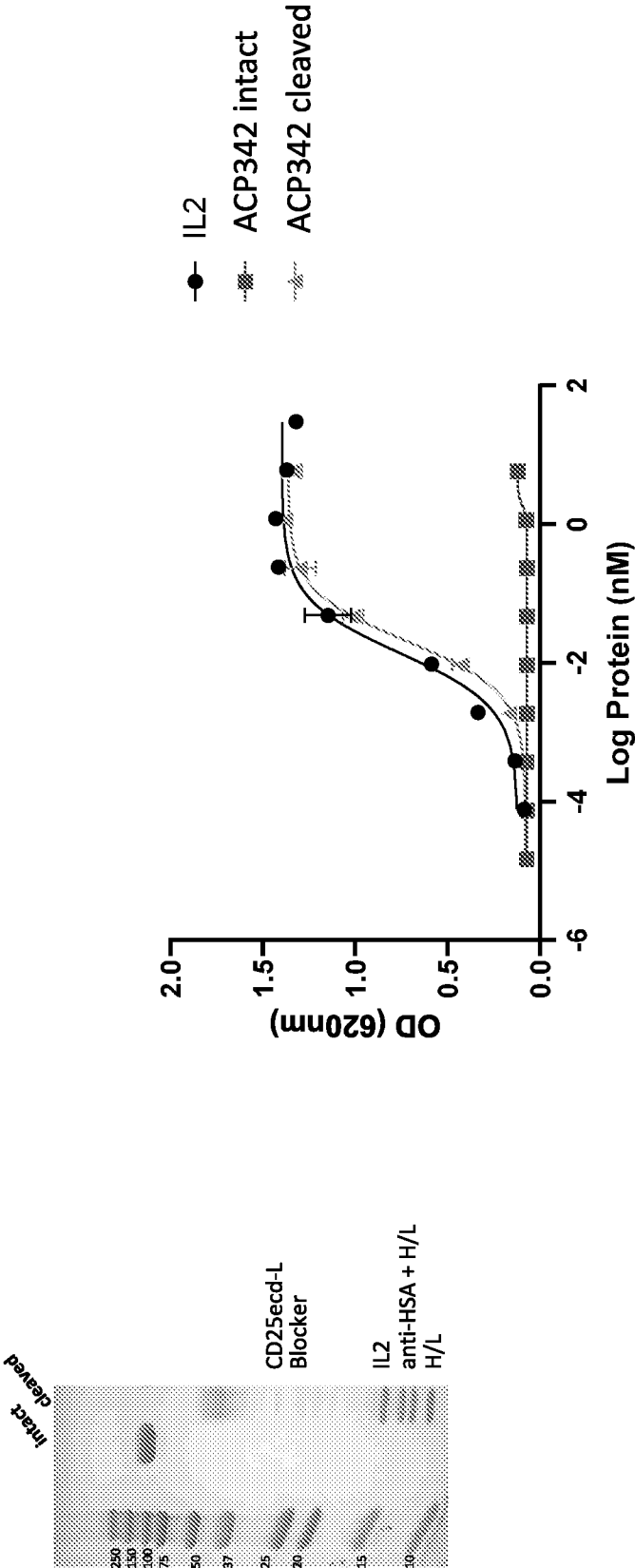


FIG. 59B

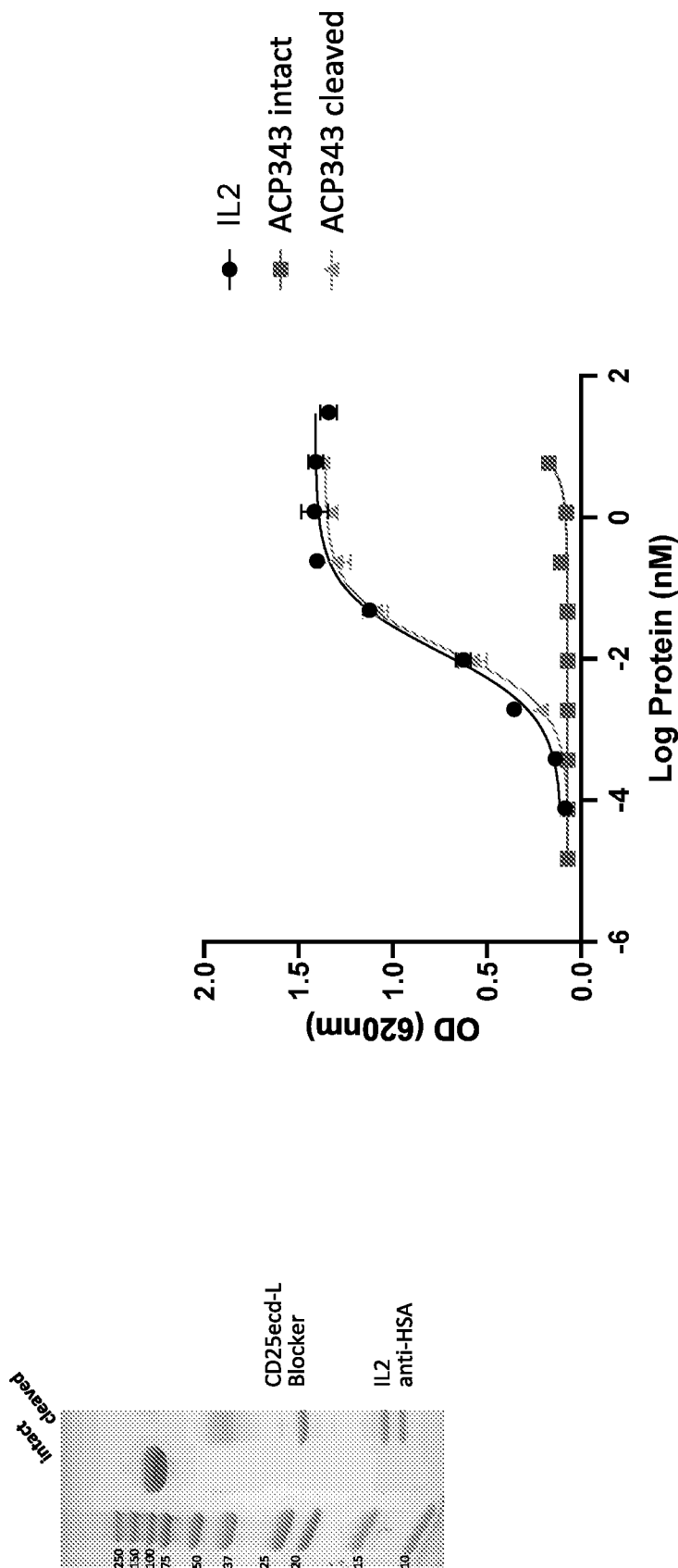


FIG. 59C

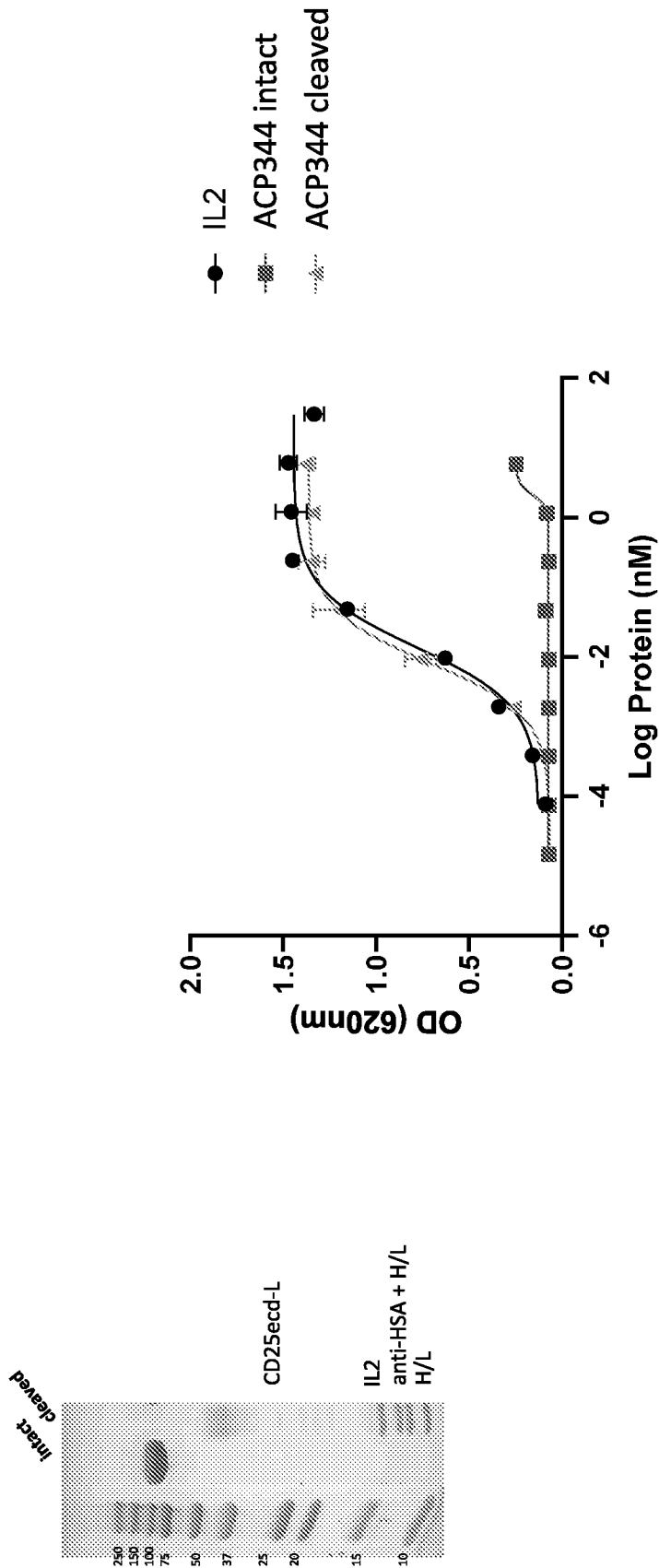


FIG. 59D

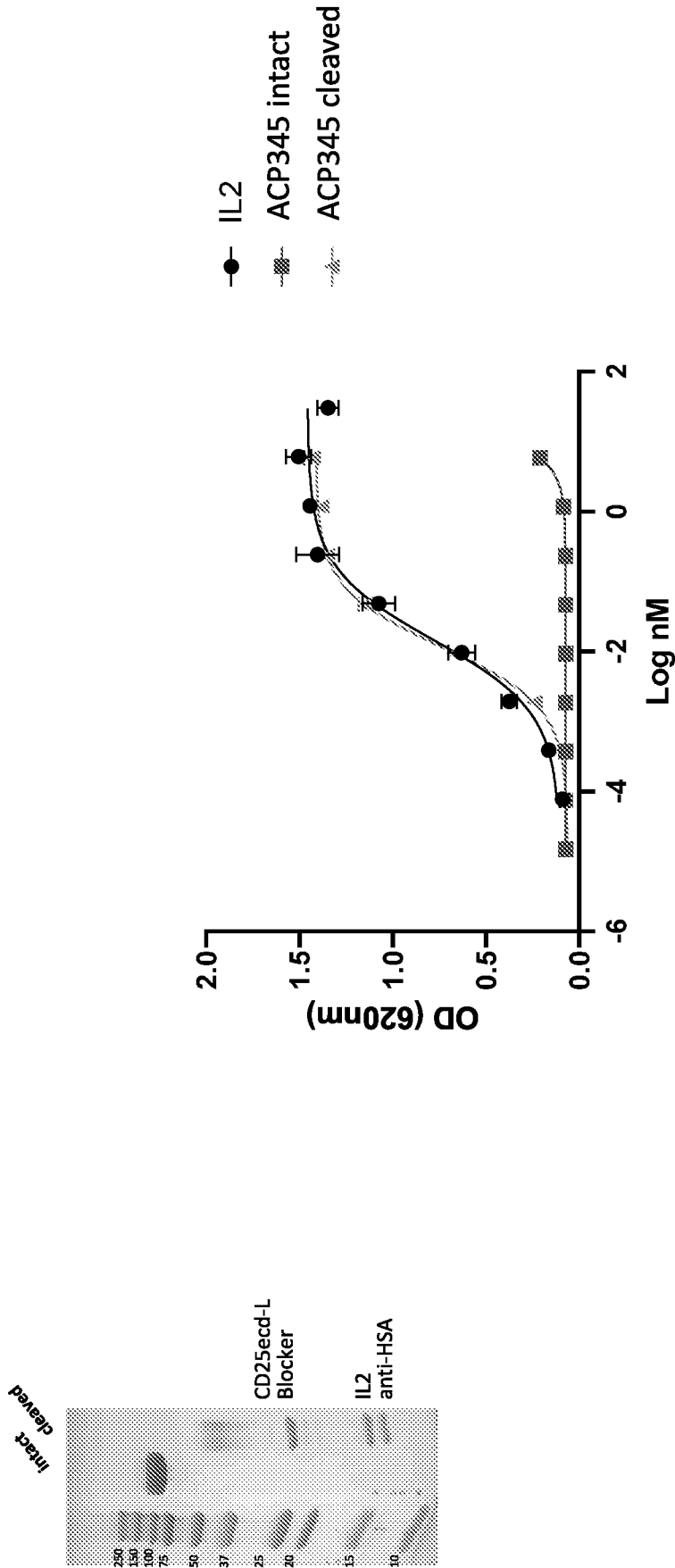


FIG. 59E

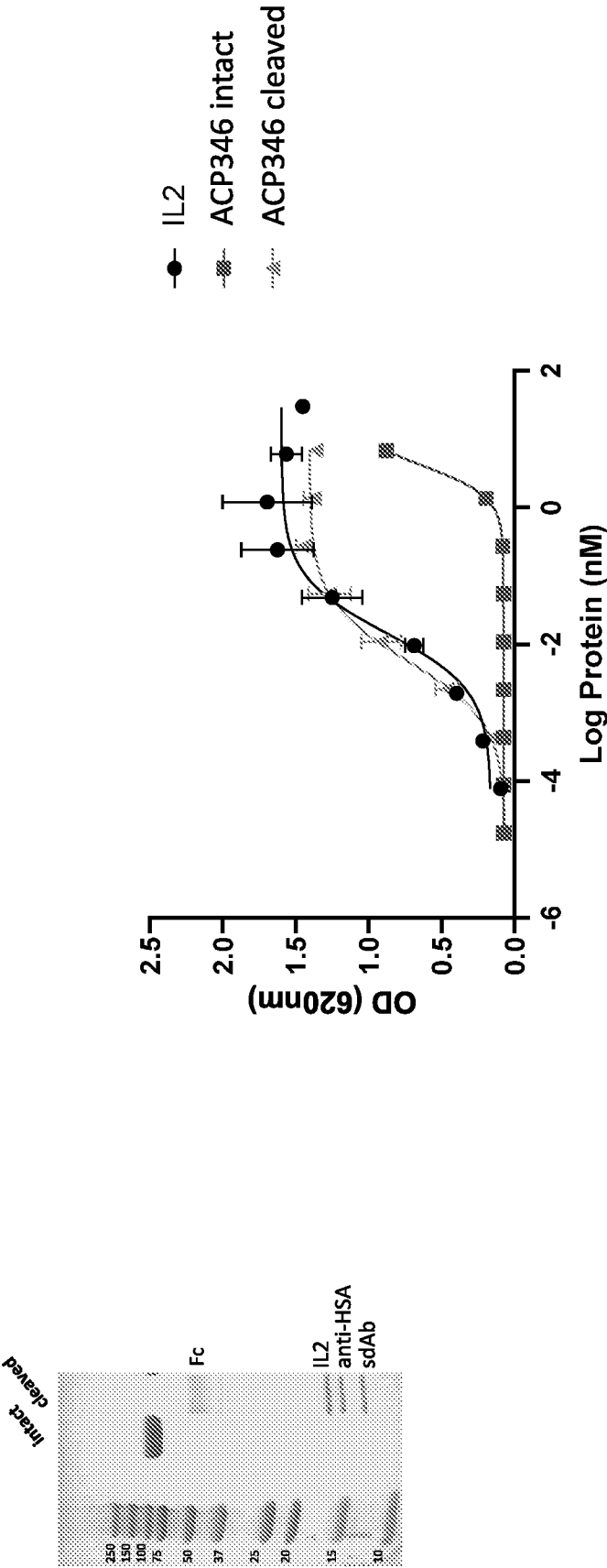


FIG. 59F

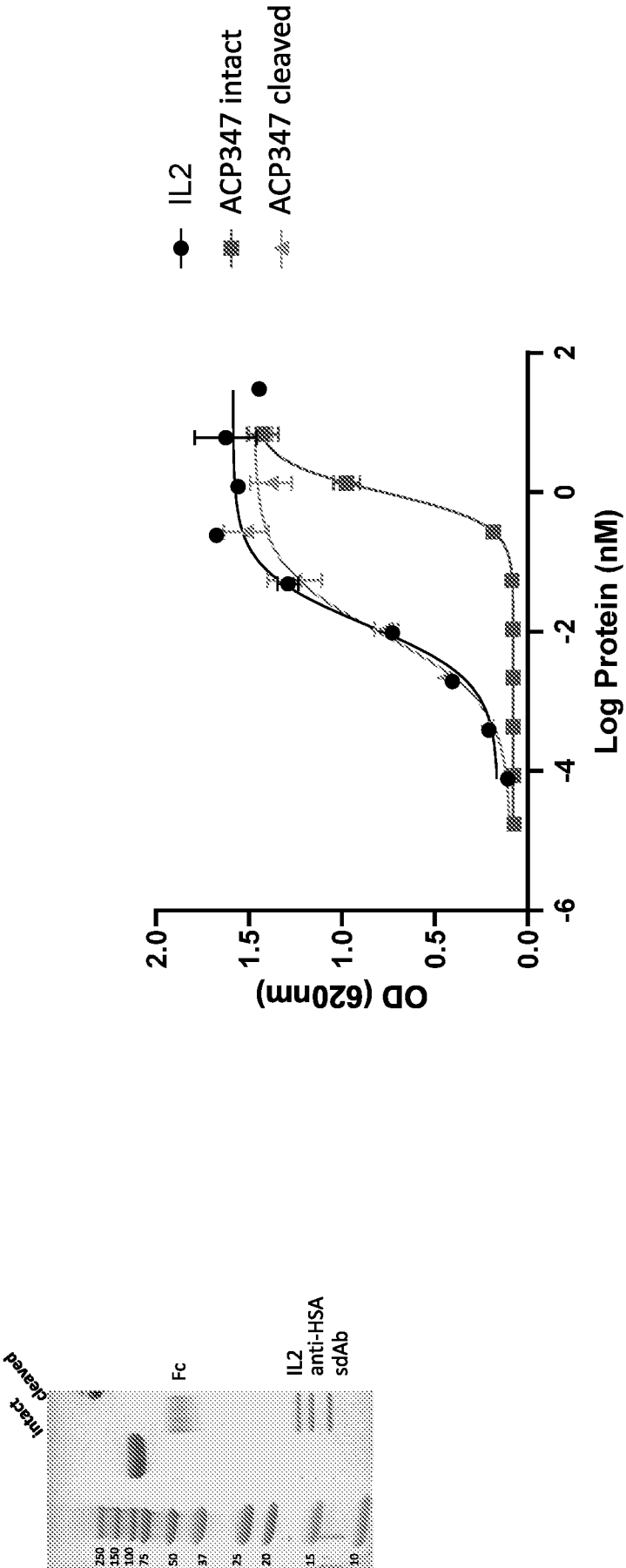


FIG. 59G

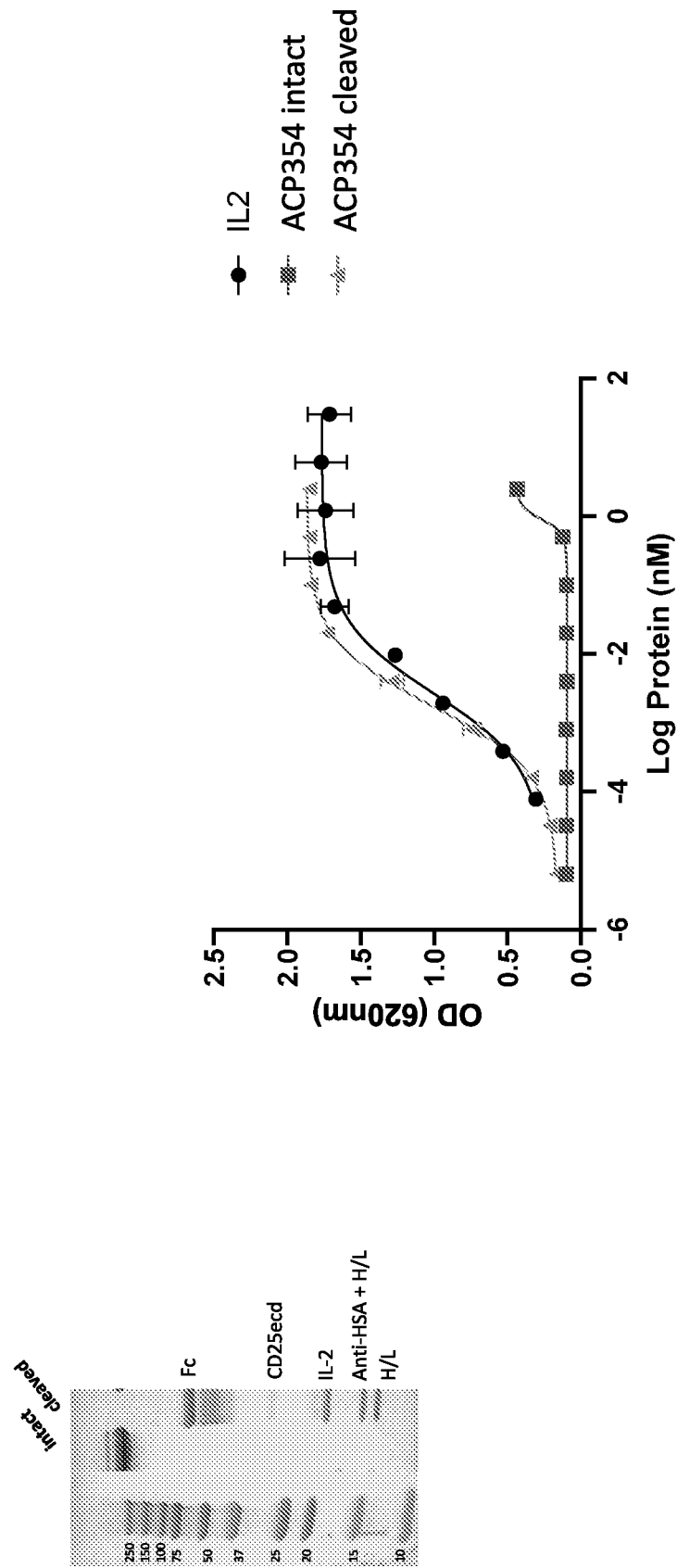


FIG. 59H

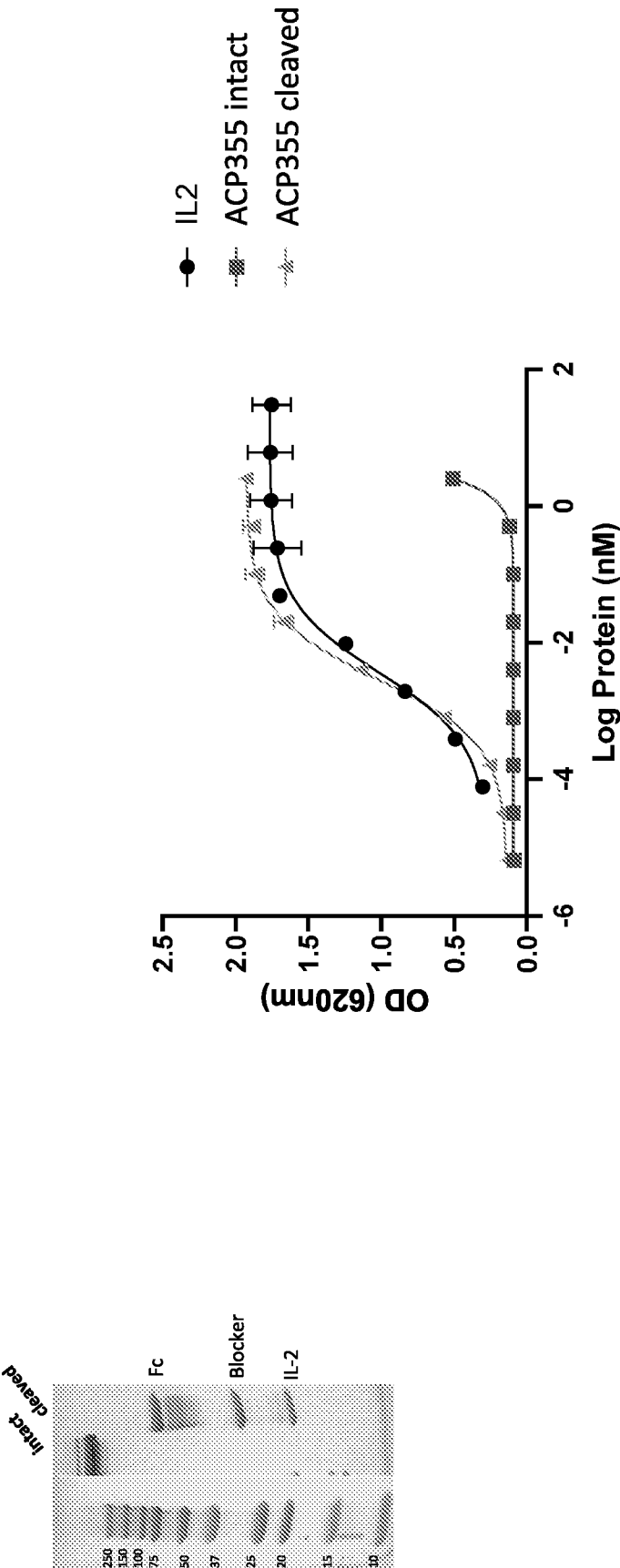


FIG. 59I

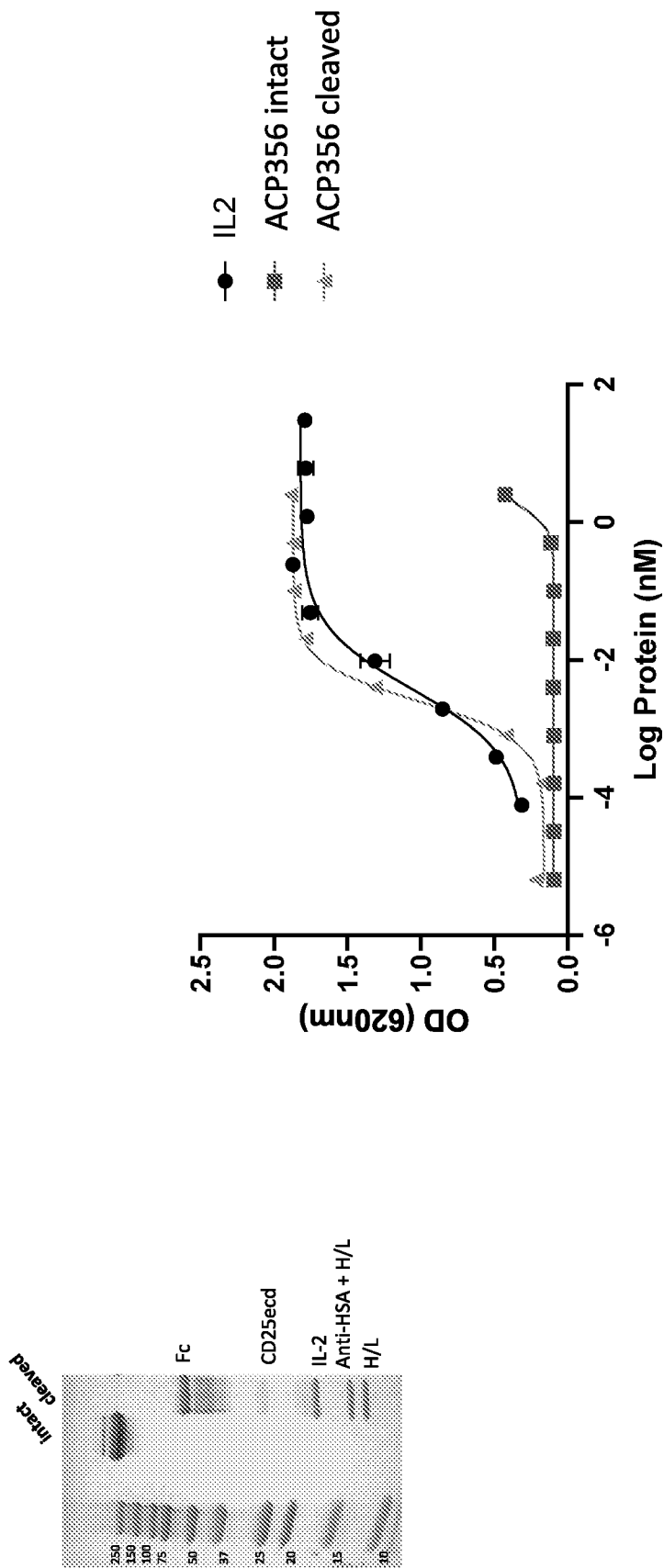


FIG.59J

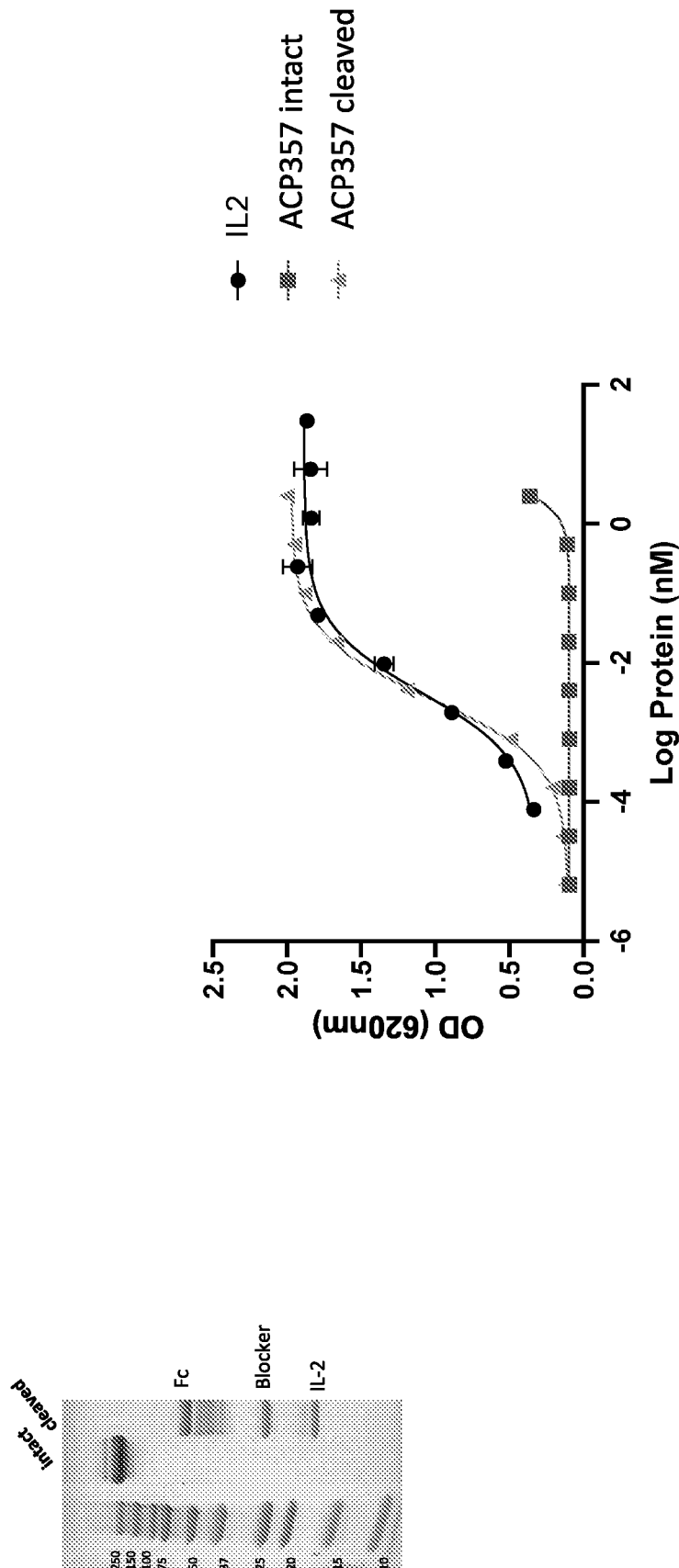


FIG. 59K

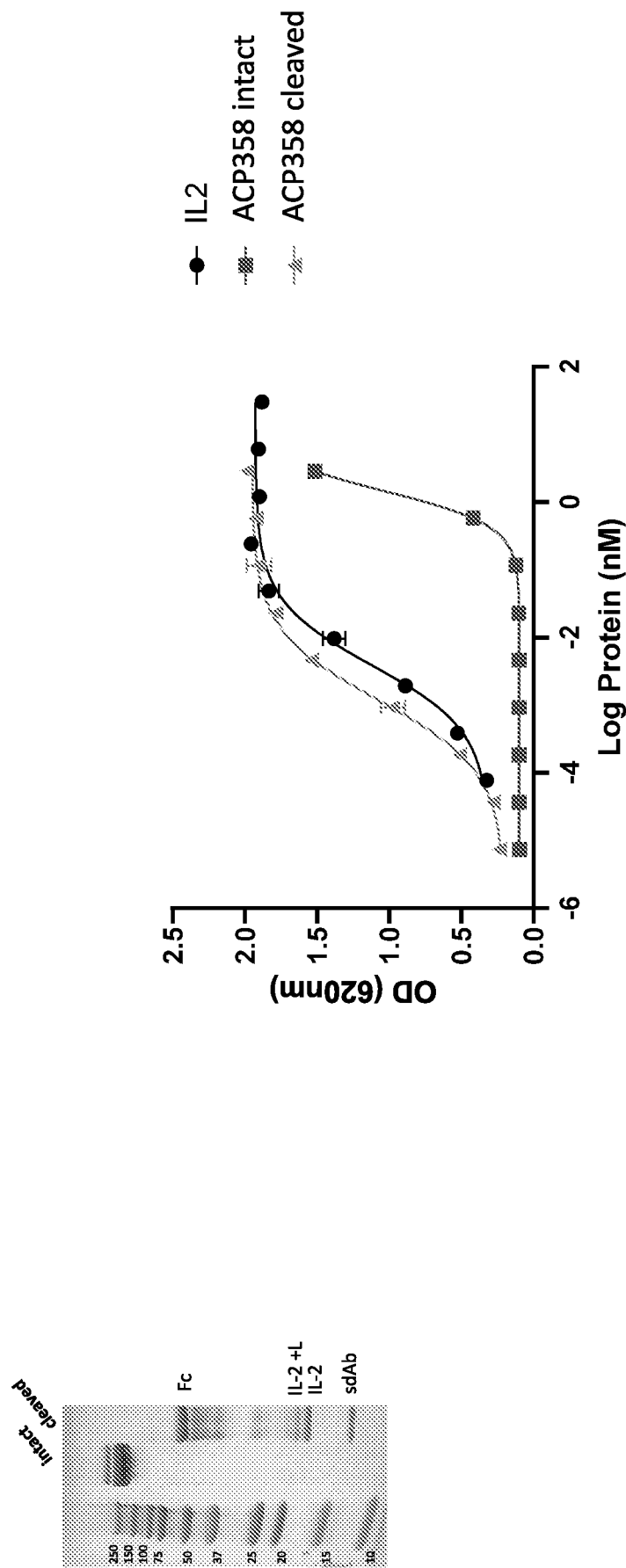


FIG. 59L

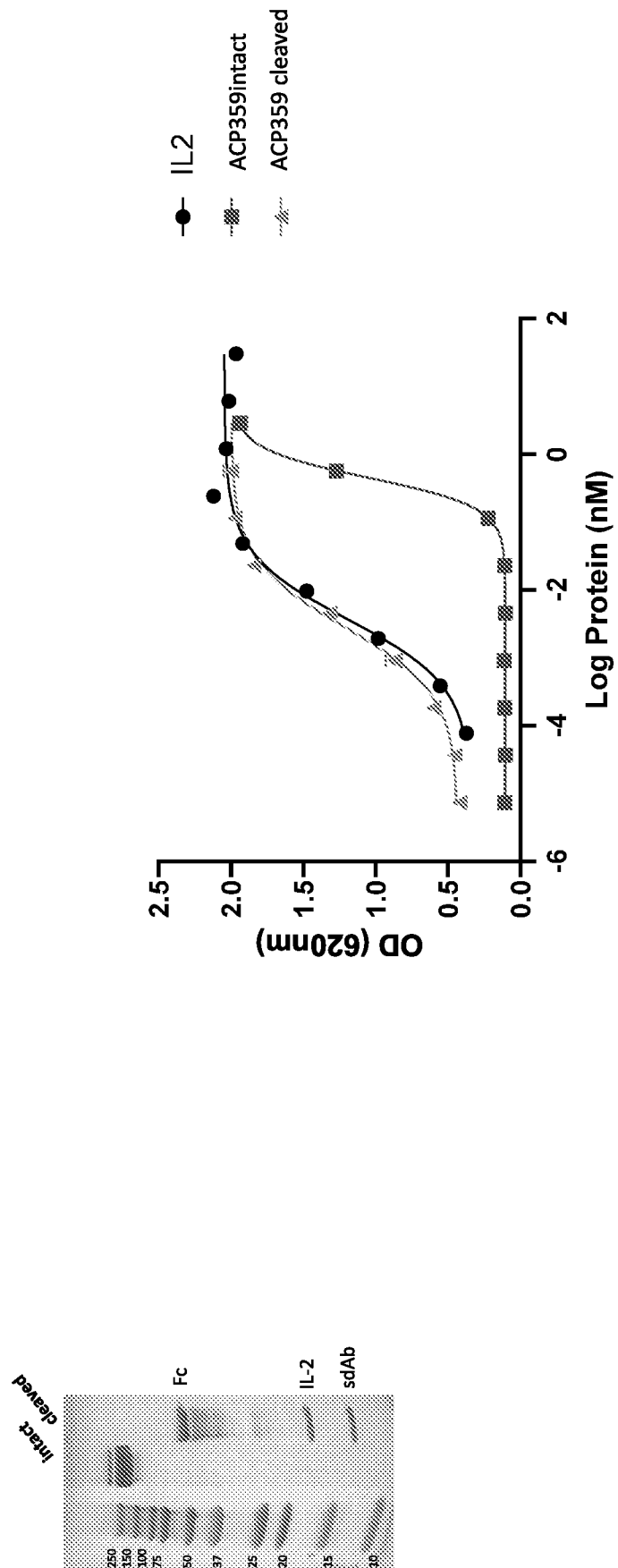


FIG. 59M

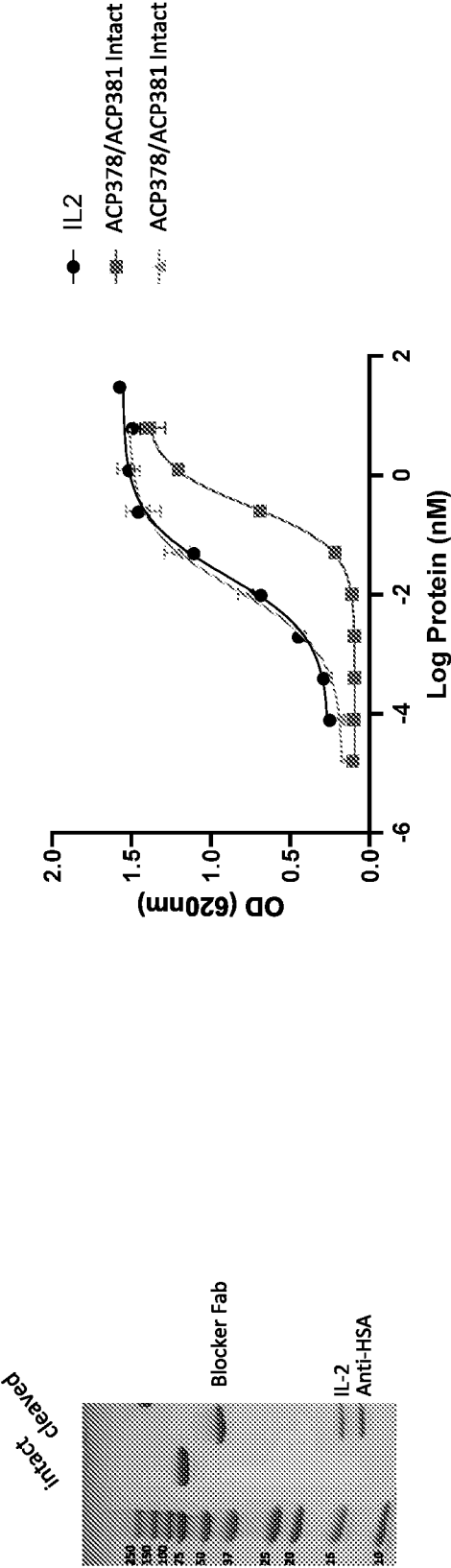


FIG. 59N

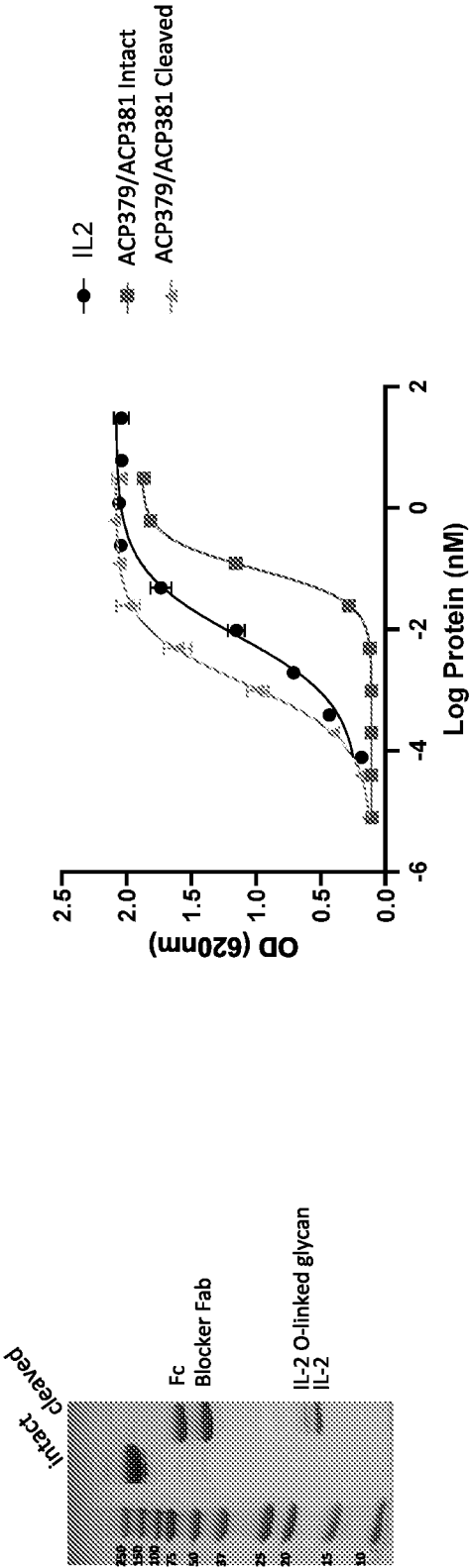


FIG. 590

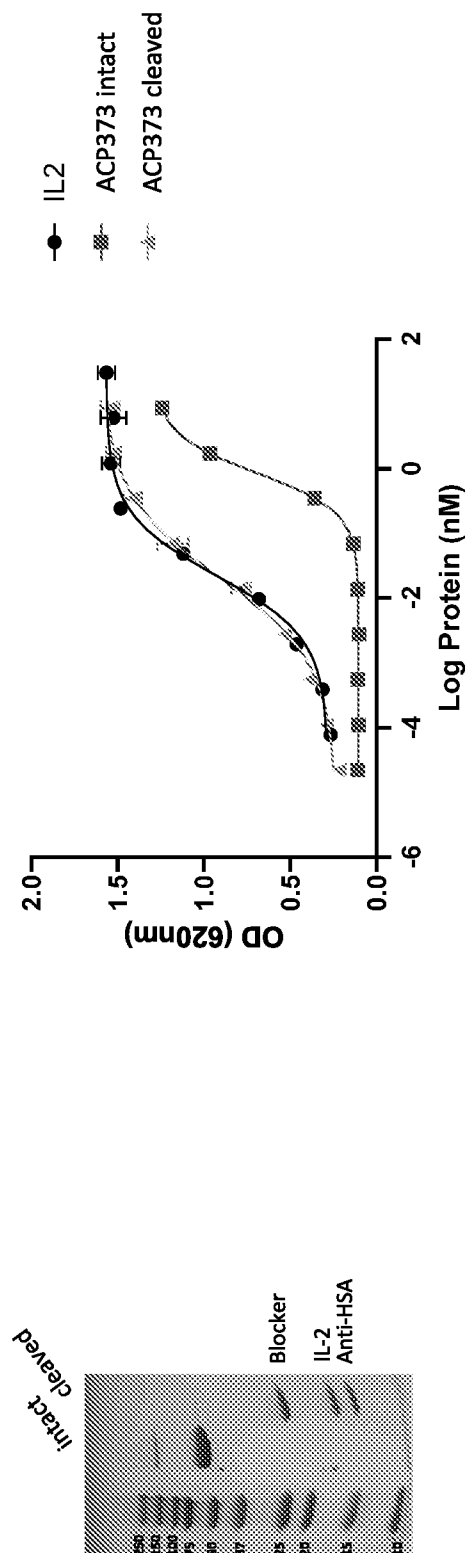


FIG. 59P

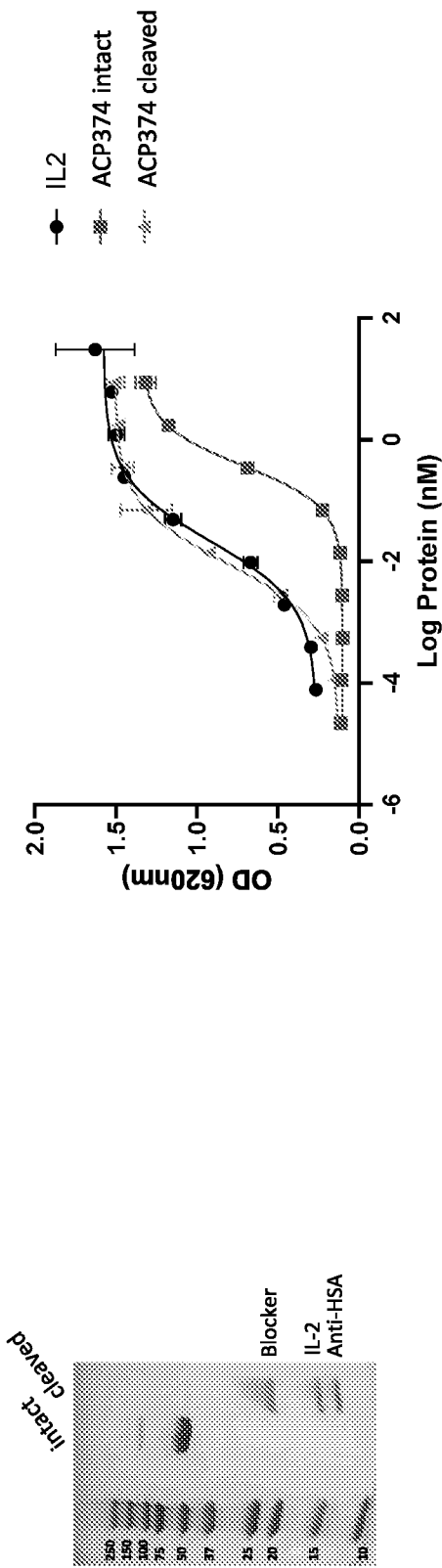


FIG. 59Q

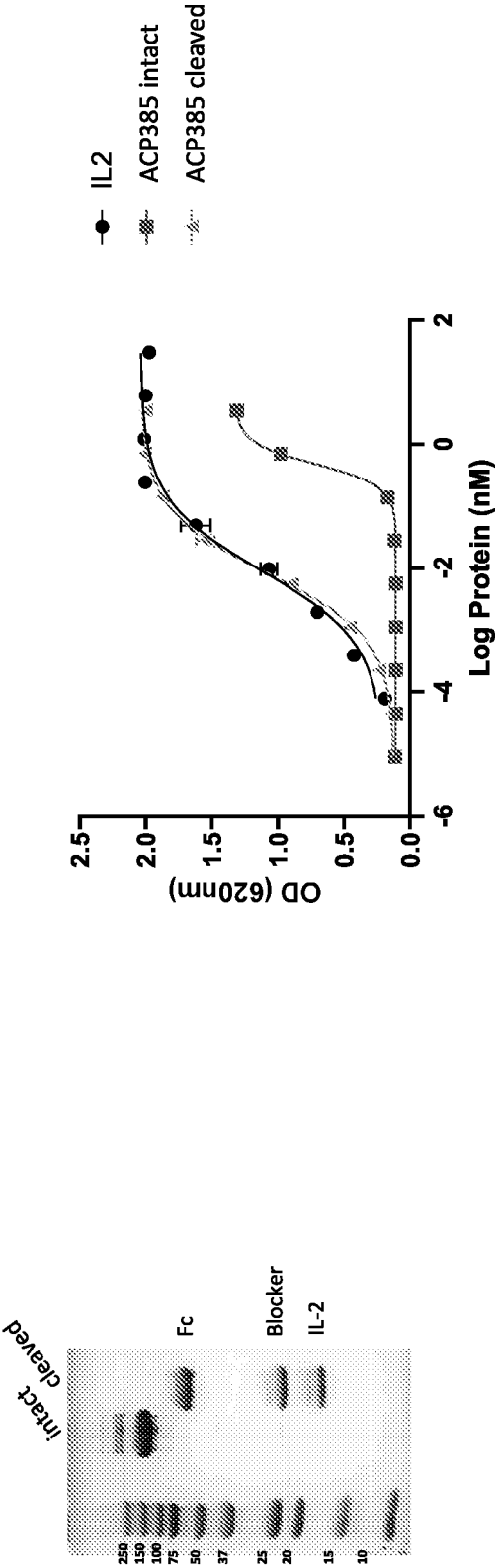


FIG. 59R

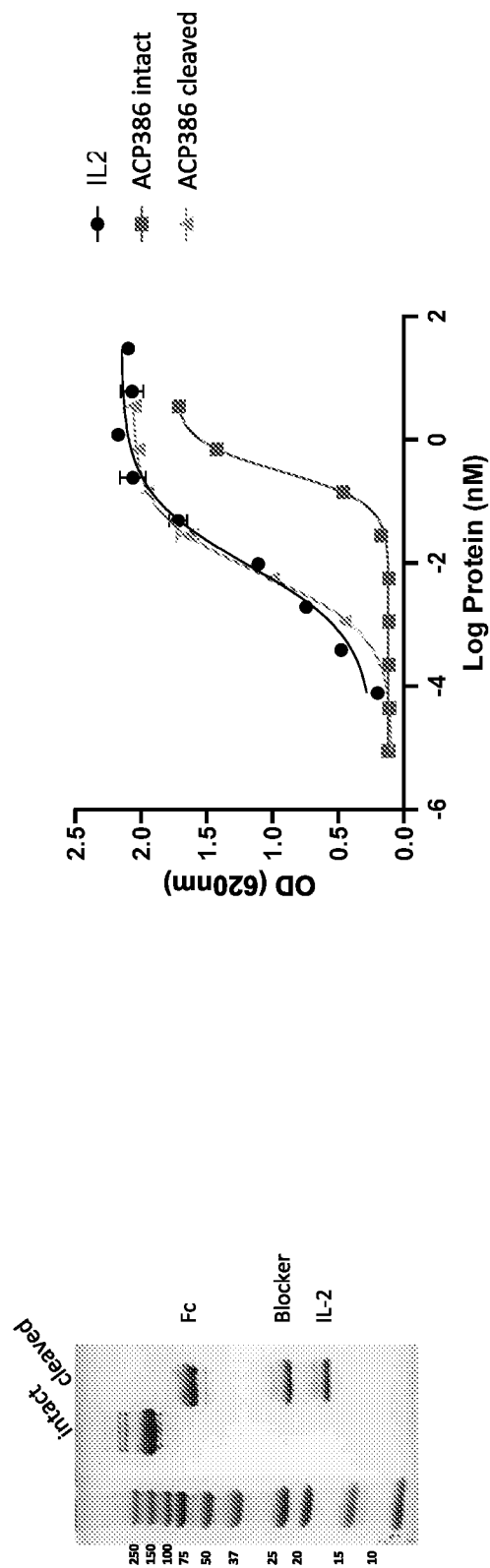


FIG. 59S

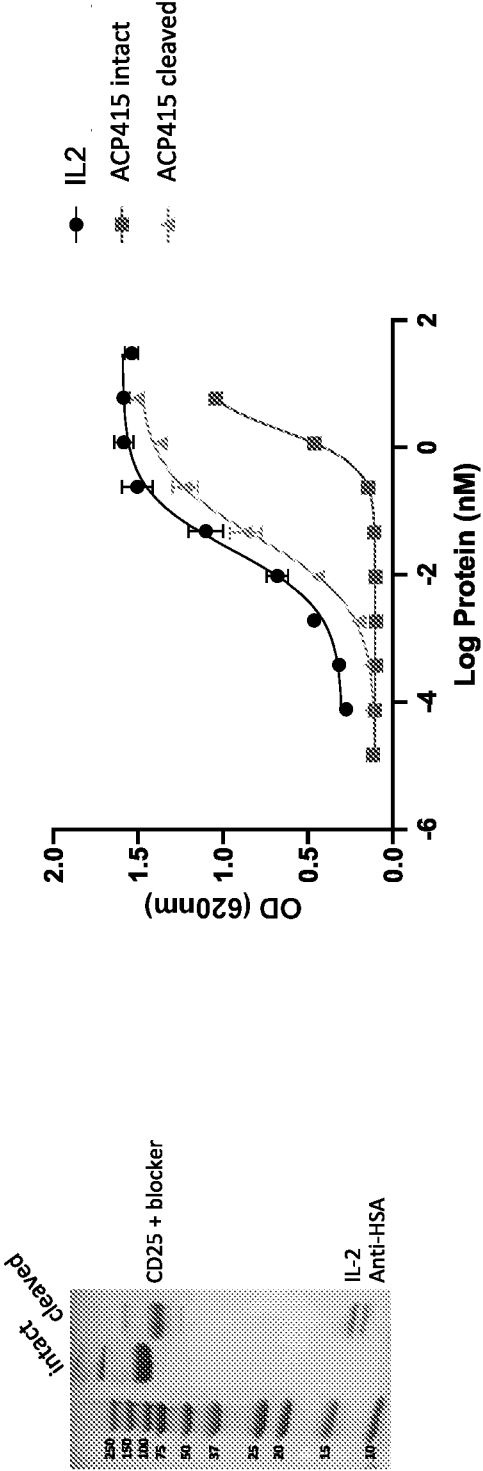


FIG. 59T

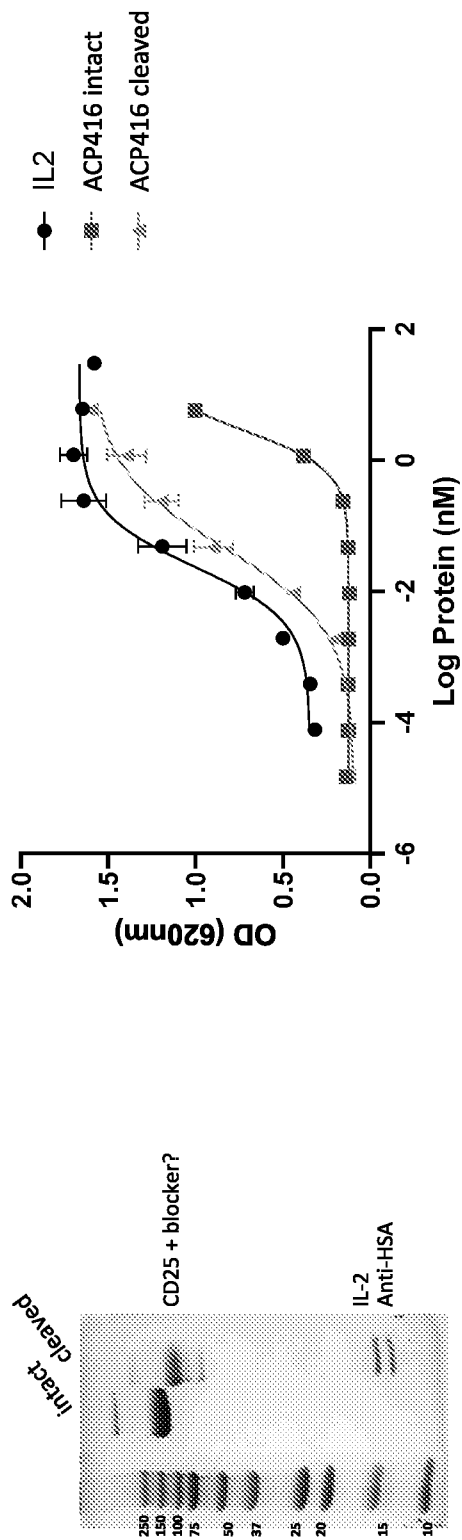


FIG. 59U

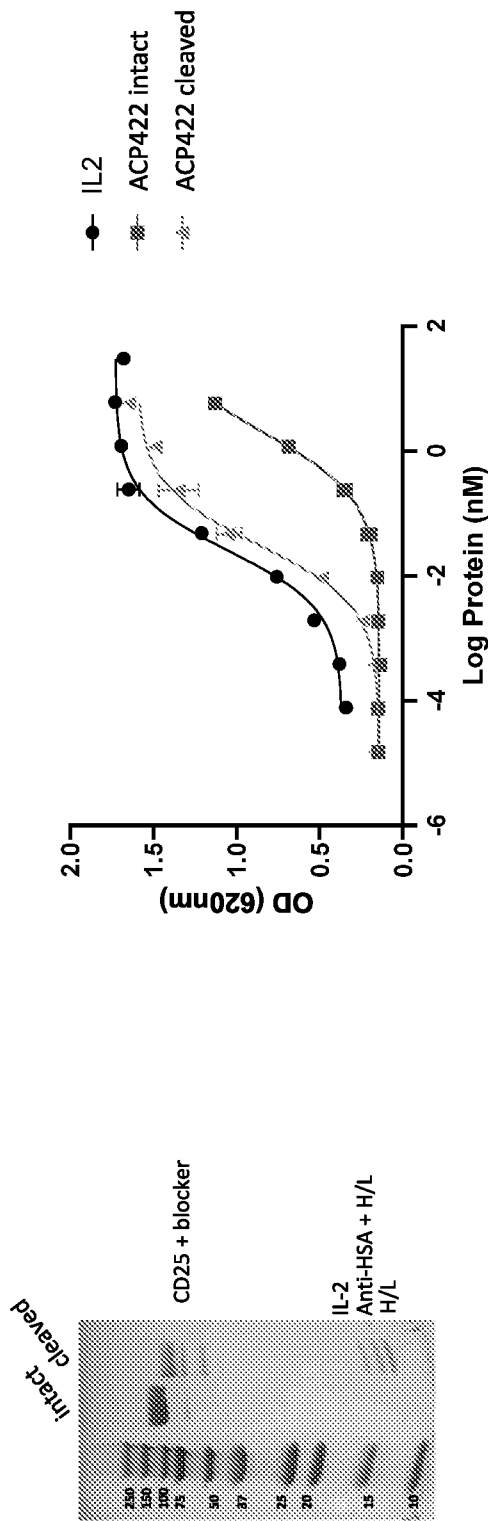


FIG. 59V

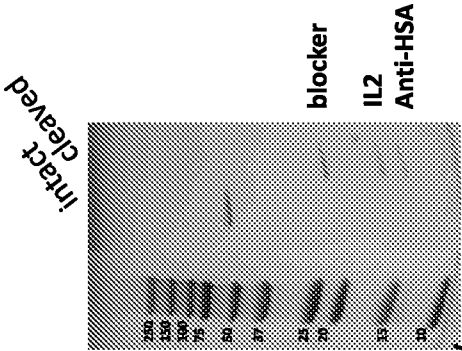
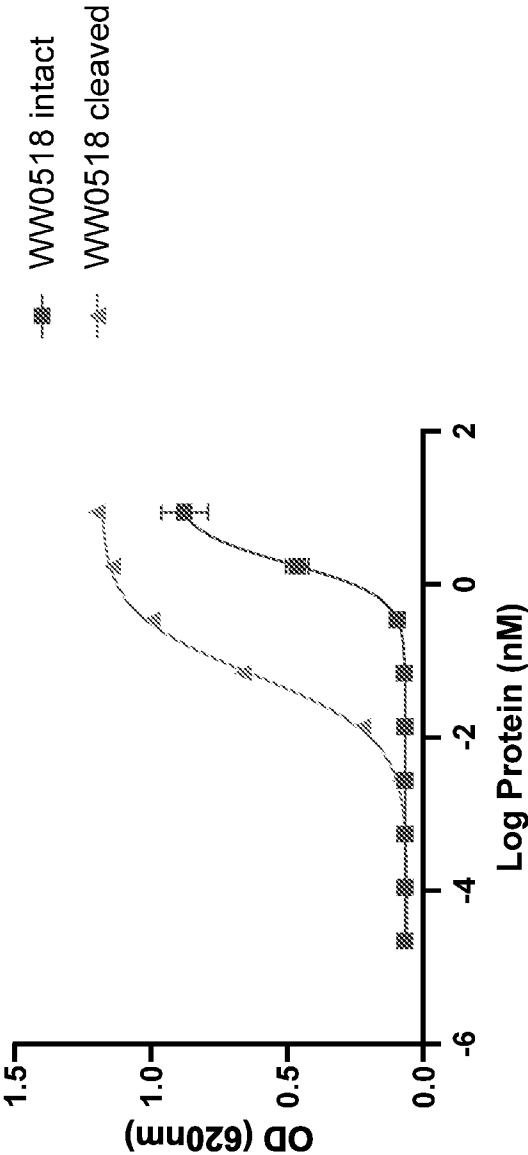


FIG. 59W

HekBlue IL2 Reporter Assay Using WW0520/524

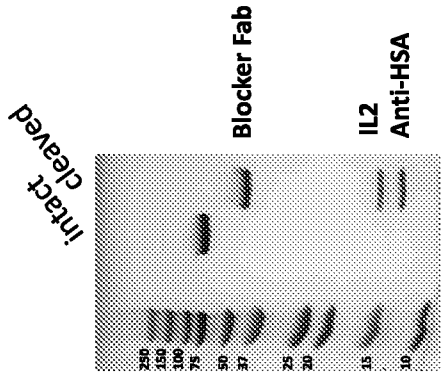
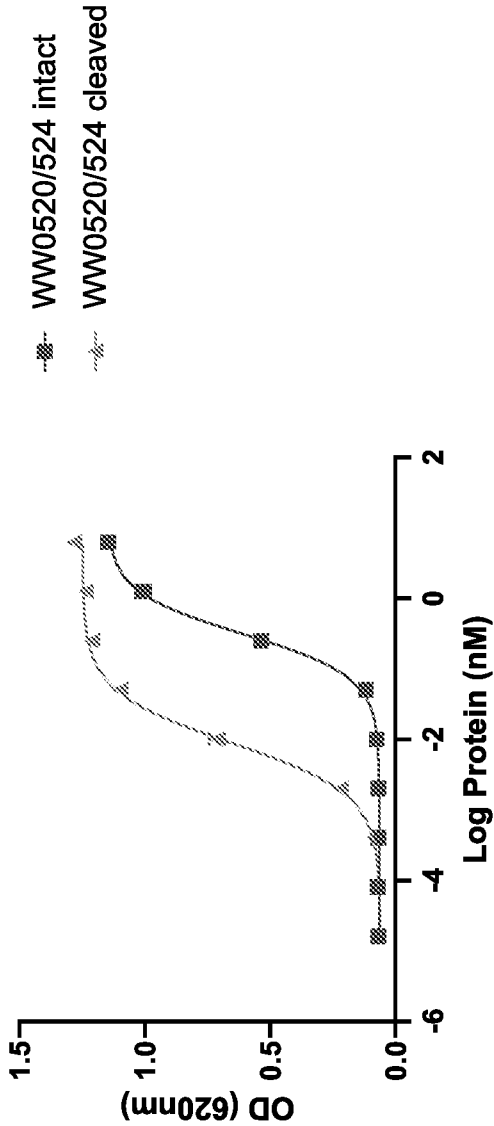


FIG. 59Z

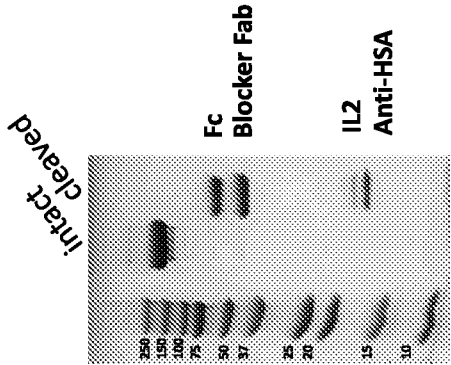
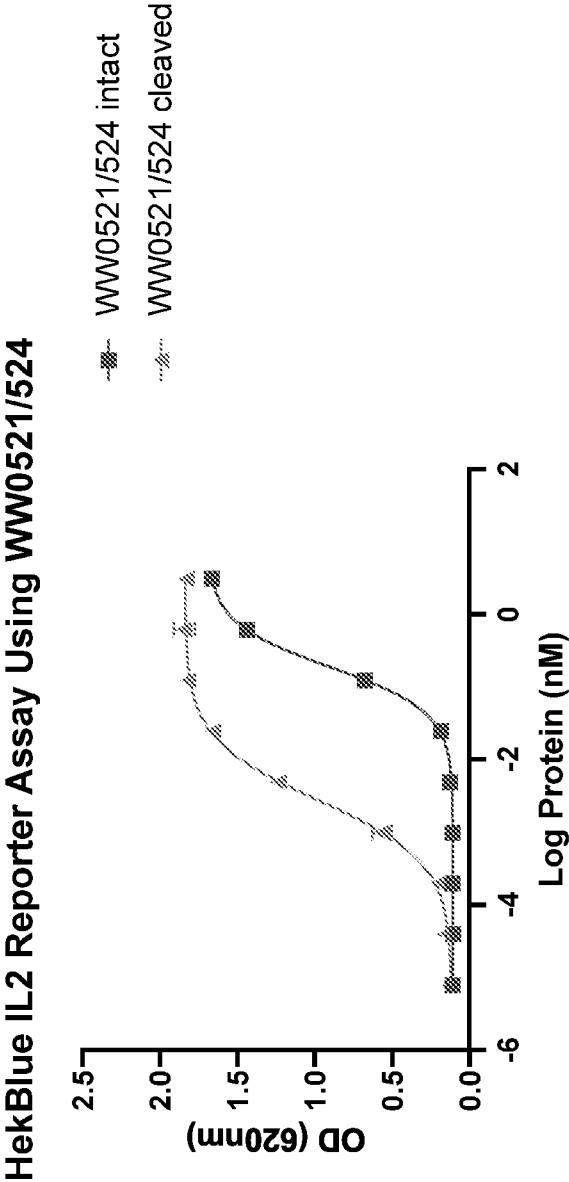


FIG. 59Y

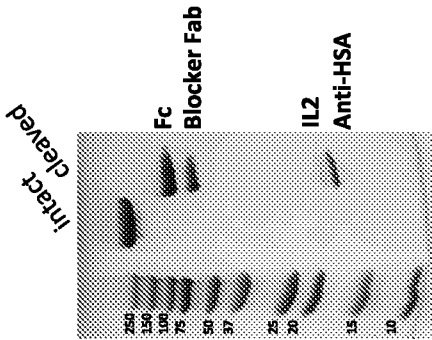
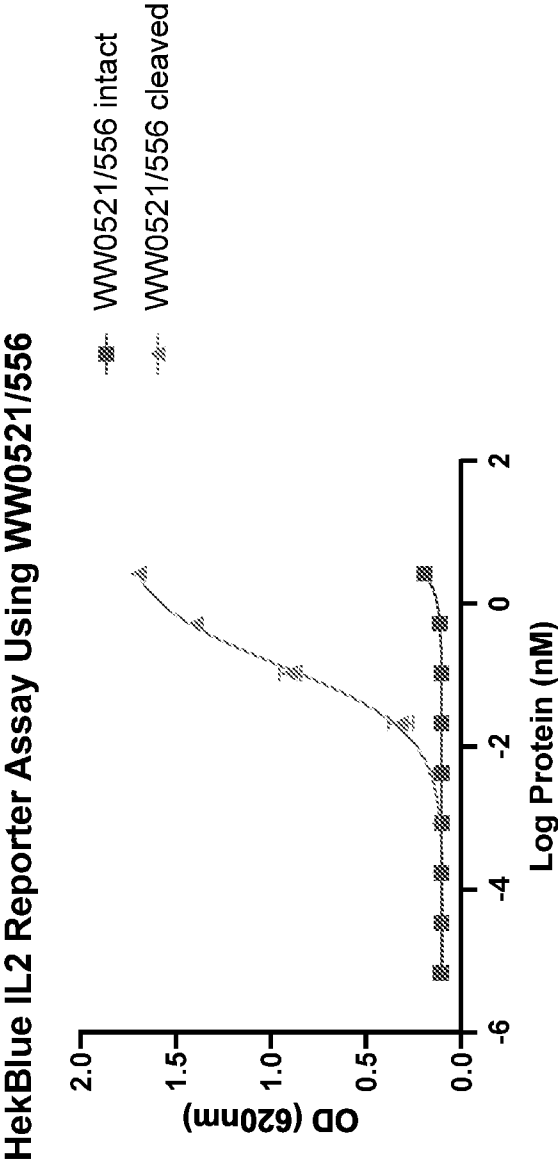


FIG. 59Z

FIG. 60A

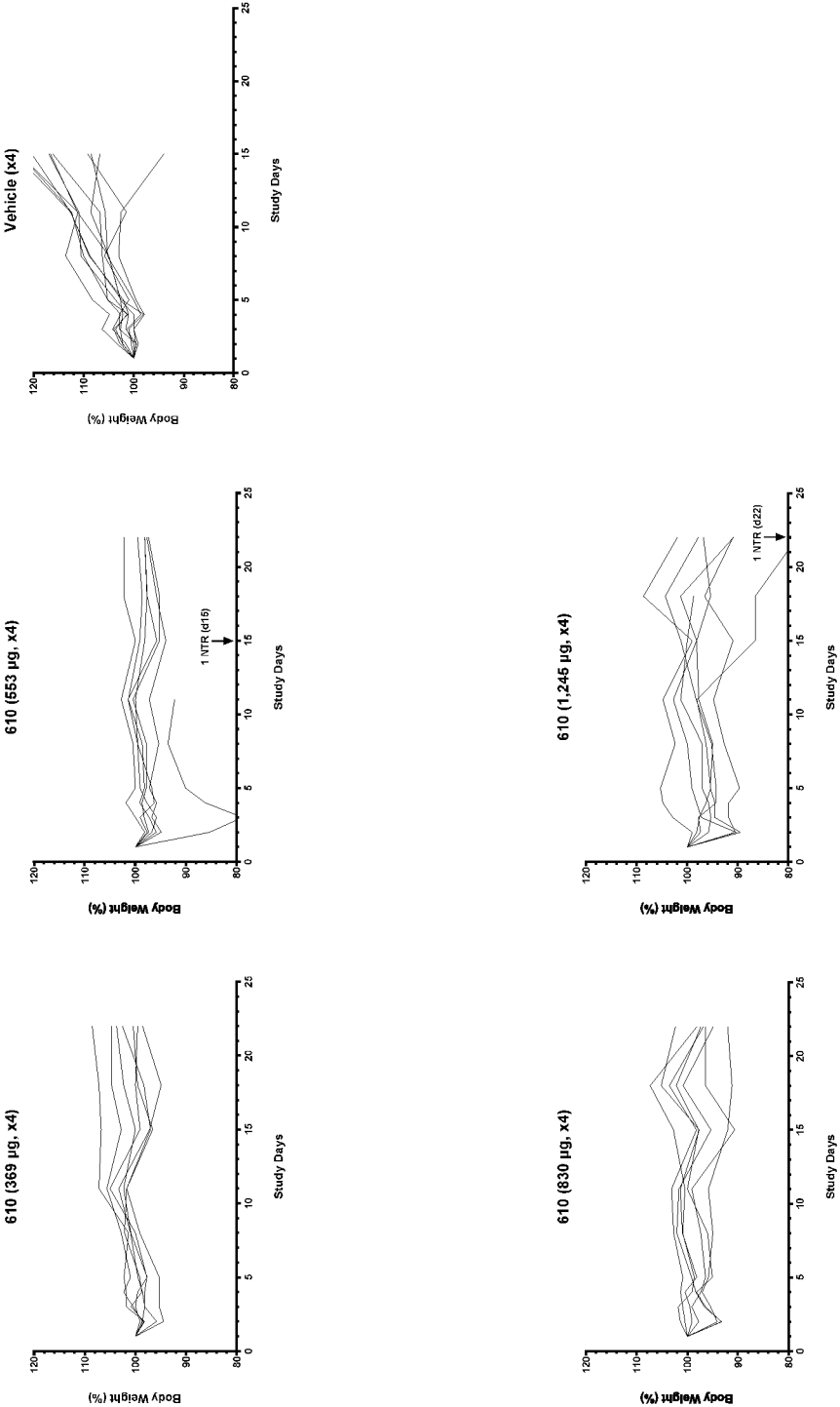


FIG. 60B

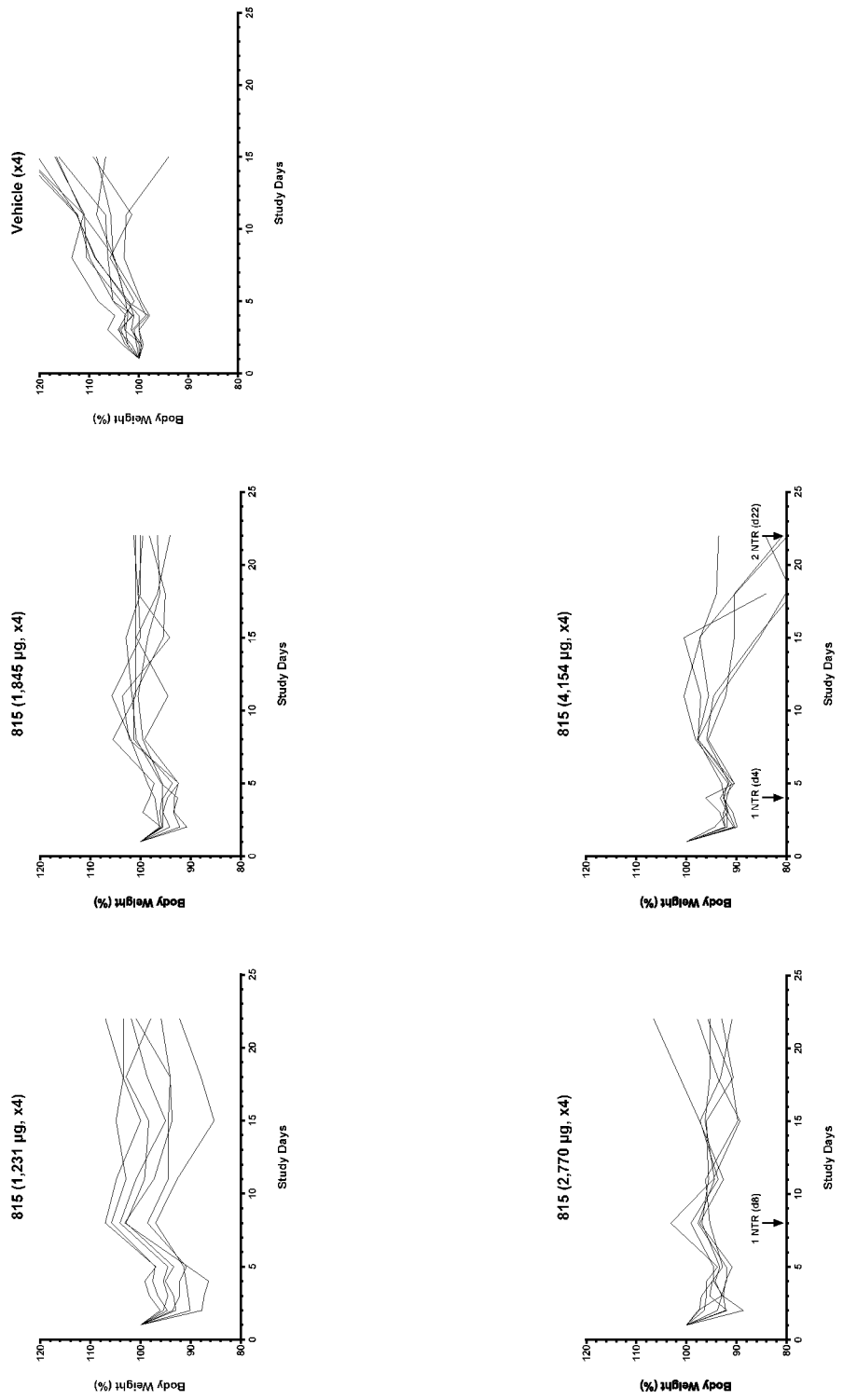


FIG. 60C

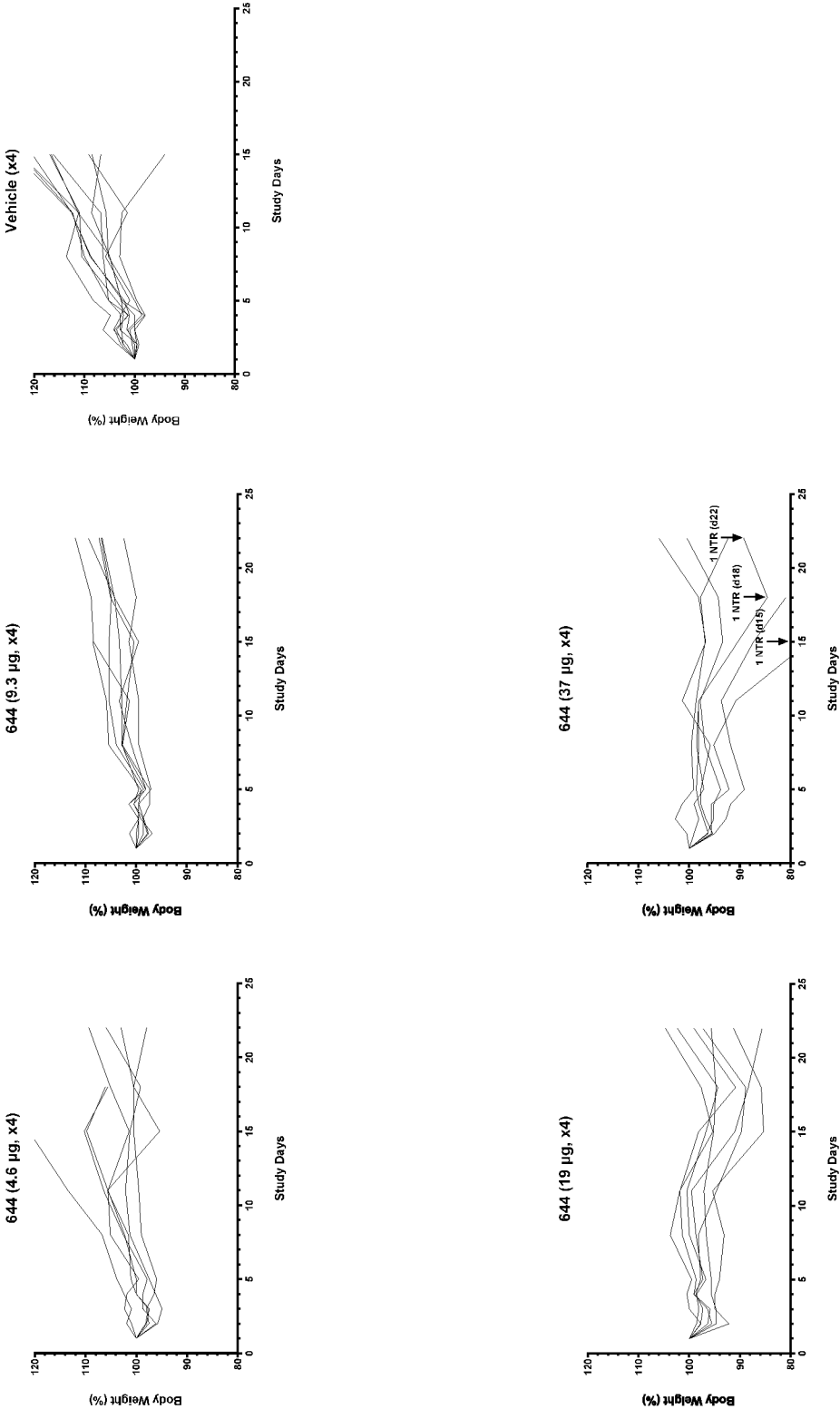


FIG. 60D

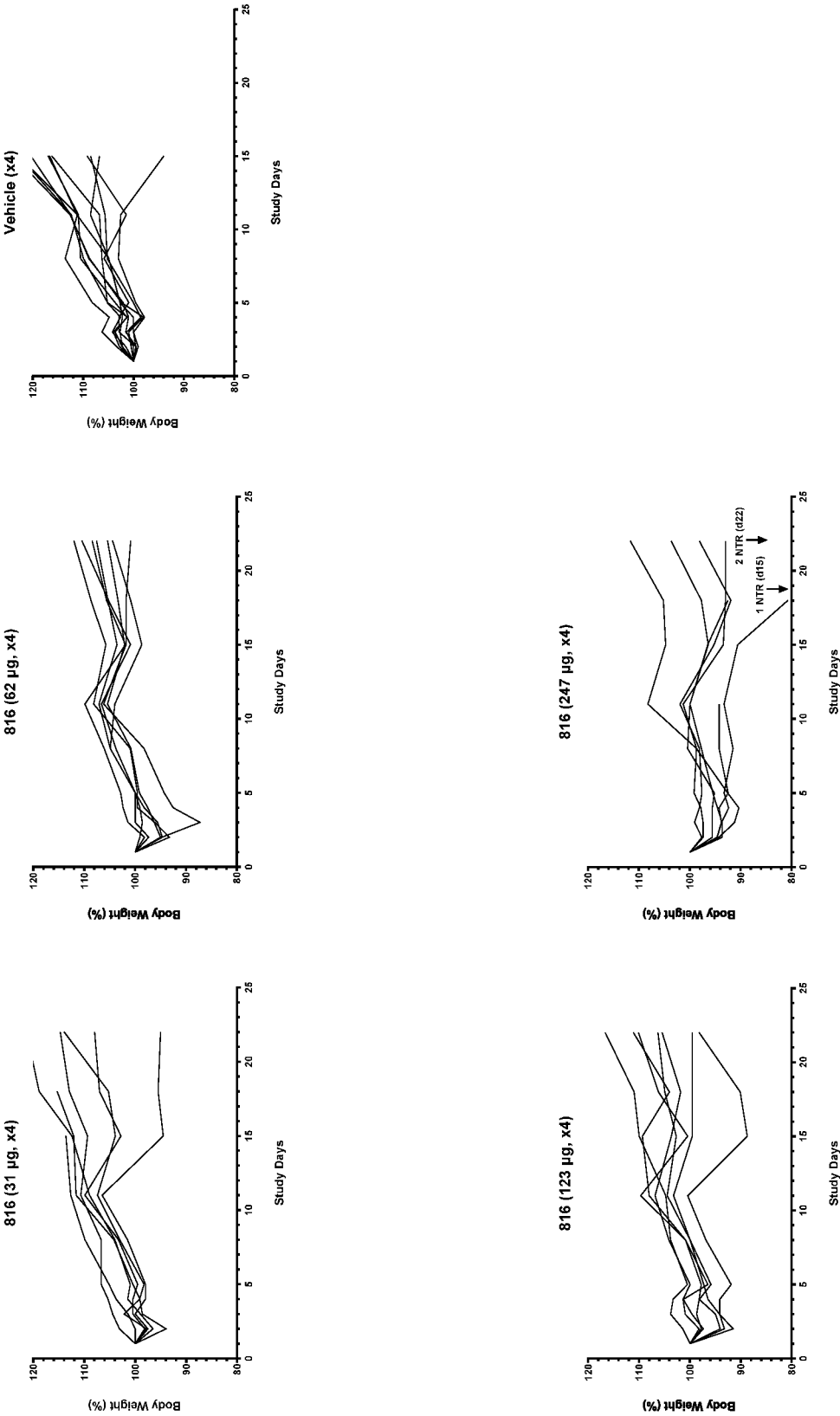


FIG. 60E

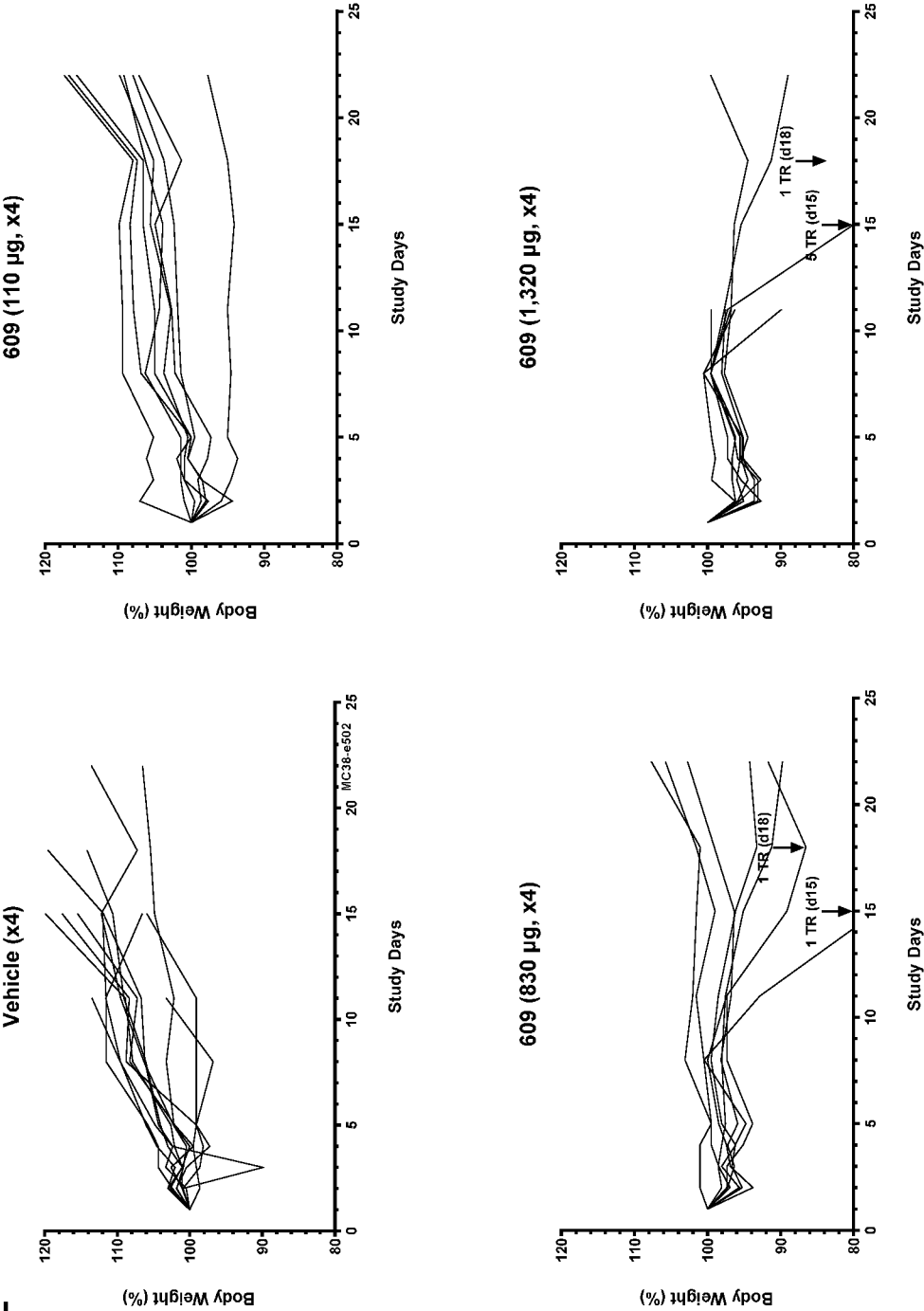


FIG. 60F

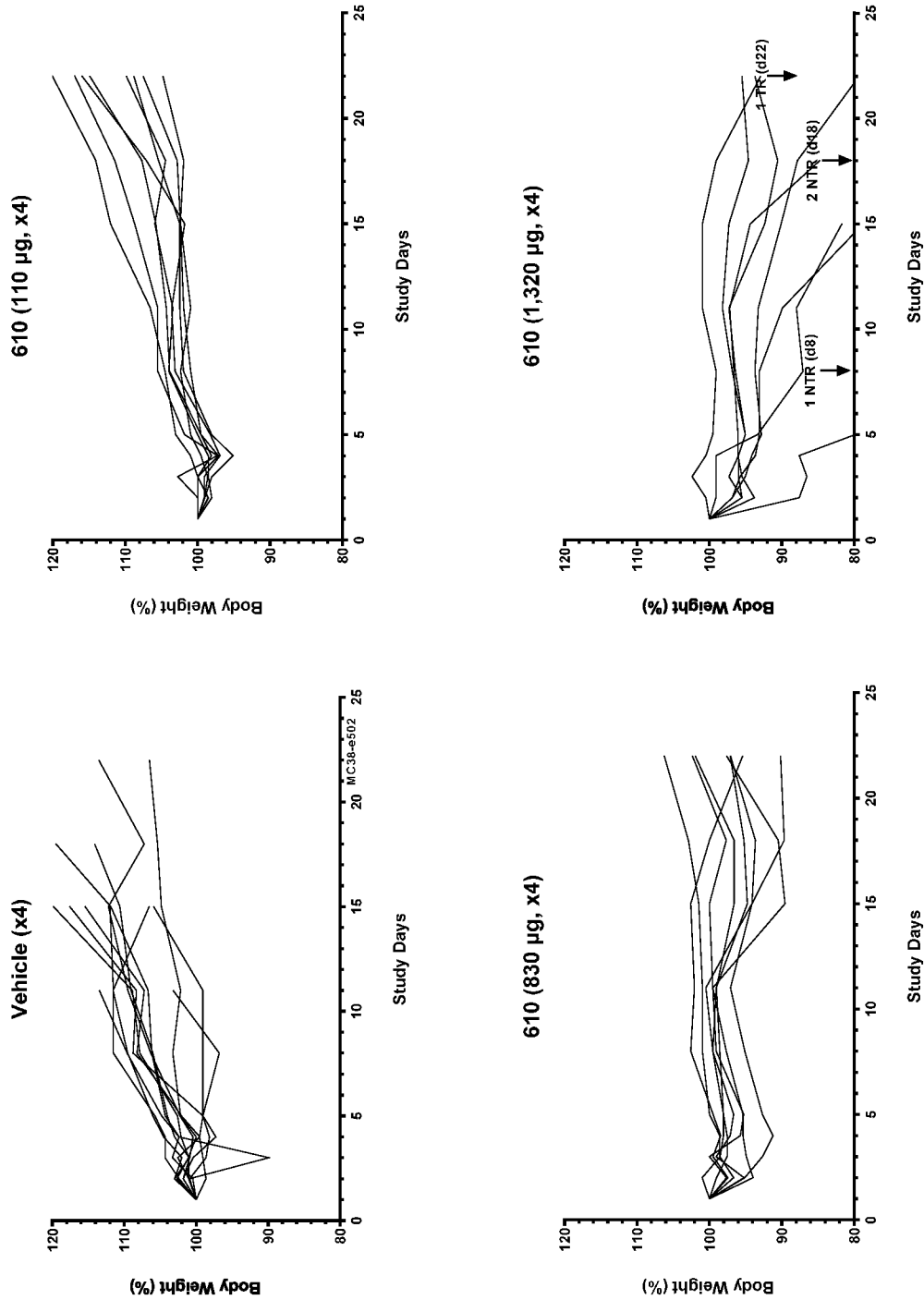


FIG. 60G

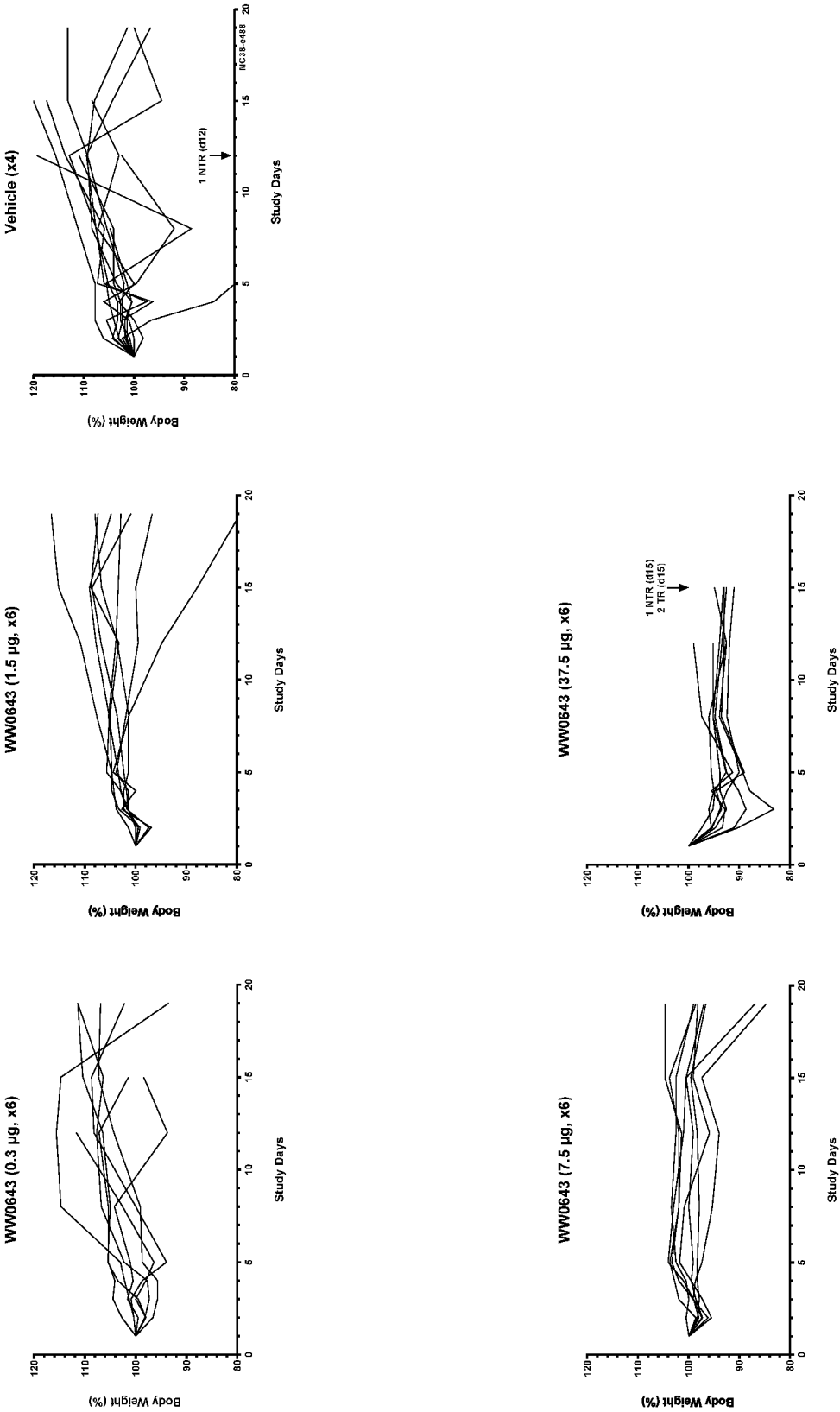
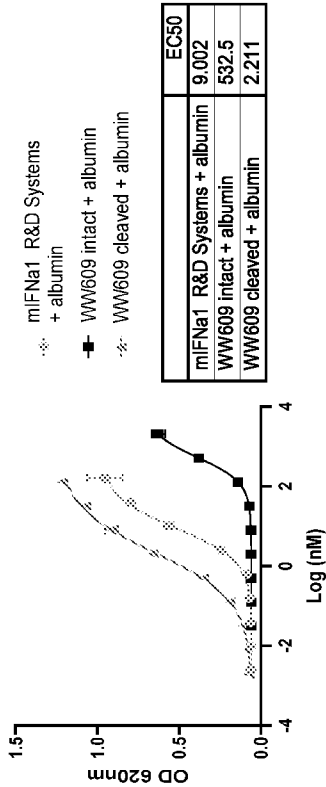


FIG. 61A

B16 Blue mIFNab Reporter assay with WW609 + Albumin



B16 Blue mIFNab Reporter assay with WW609 no Albumin

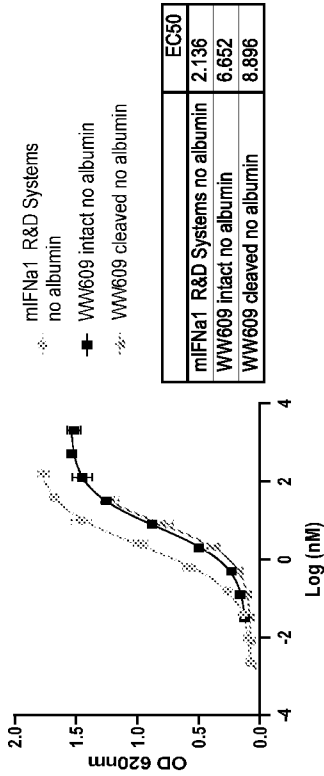
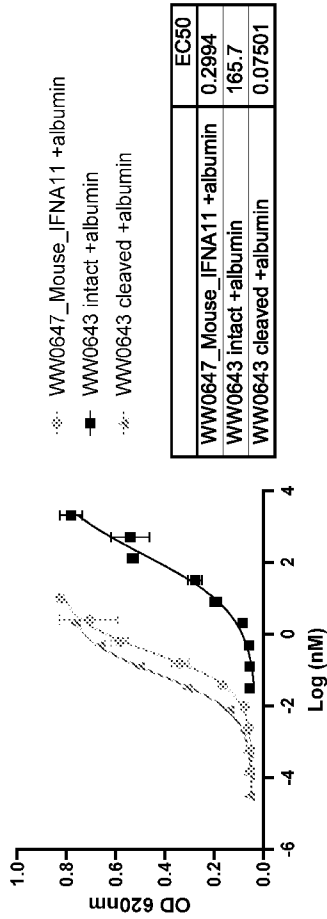


FIG. 61B

B16 Blue mIFNab Reporter assay with WW0643 + Albumin



B16 Blue mIFNab Reporter assay with WW0643 no albumin

