

Antibody Variable Heavy Chain (SEQ ID NO: 62)

QVQLQESGPSVKPQTLSICTYSGFSLNNNAVGVWRQAPGKVPESLVGCSSDGTCCYNSSALKSRIDTRTSKNQISLSVTTDDAAVYCTRGHYSYGVDLGTIDYWGPGILLTVSS

Kabat numbering and CDRs

123456789101112131415161718192021222324252627282930

QVQLQESGPSLVKPSQTLSLCTYSGFSLNNNAVGVWRQAPGKVPESLVGCSSDGTCCYNSSALKSRIDTRTSKNQISLSVTTDDAAVYCTRGHYSYGVDLGTIDYWGPGILLTVSS

HCDR1

31323334353637383940414243444546474849

NNAVGVWVRQAPEGKVPESESLV

HCDR2

50515253545556575859606162636465

GCSSTDGTCYVNSALKSS

Chothia numbering and CDRs

1234567891011121314151617181920212223242526272829303132333435363738394041424344454647484950515253545556575859606162636465

QVQLQESGPSLVKPSQTLSTLCTTVSGFSLNNNAVGVWVRQAPEGKVPESESLVGCSSDGTCCYVNSALKSS

HCDR1

262728293031323334353637383940414243444546474849

GFSLNNNAVGVWVRQAPEGKVPESESLV

HCDR2

50515253545556575859606162636465

GCSSTDGTCYVNSALKSS

Martin numbering and CDRs

1234567891011121314151617181920212223242526272829303132333435363738394041424344454647484950515253545556575859606162636465

QVQLQESGPSLVKPSQTLSTLCTTVSGFSLNNNAVGVWVRQAPEGKVPESESLV

HCDR1

262728293031323334353637383940414243444546474849

GFSLNNNAVGVWVRQAPEGKVPESESLV

HCDR2

50515253545556575859606162636465

GCSSTDGTCYVNSALKSS

Antibody Variable Light Chain (SEQ ID NO: 433)

QAVLTQPSVSGSLGQRVSI TCSGSSSNVGGNSVGVWQHLPGLGKTIYDINSRPSGVPDRSGSRGNTATINSLOAEDGYYCVTGSDTHDILVSGSTRILVIG

Kabat numbering and CDRs

|       |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |     |     |       |    |    |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
|-------|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|-----|-----|-------|----|----|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| LCDR1 |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |     |     | LCDR2 |    |    |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| 1     | 2  | 3  | 4  | 5  | 6  | 7  | 8  | 9  | 11 | 12 | 13 | 14 | 15 | 16 | 17 | 18 | 19 | 20 | 21 | 22 | 23 | 24 | 25 | 26 | 27 | 27a | 27b | 27c   | 28 |    |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| Q     | A  | V  | L  | T  | Q  | P  | S  | S  | V  | S  | G  | S  | L  | G  | Q  | R  | V  | S  | I  | T  | C  | S  | G  | S  | S  | N   | V   | G     | G  | G  |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| 35    | 36 | 37 | 38 | 39 | 40 | 41 | 42 | 43 | 44 | 45 | 46 | 47 | 48 | 49 | 50 | 51 | 52 | 53 | 54 | 55 | 56 | 57 | 58 | 59 | 60 | 61  | 62  | 63    | 64 | 65 |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| G     | S  | S  | S  | N  | V  | G  | G  | G  | N  | S  | V  | G  | W  | Y  | Q  | H  | L  | P  | G  | S  | G  | L  | K  | T  | I  | I   | Y   | D     | T  | N  |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| 57    | 58 | 59 | 60 | 61 | 62 | 63 | 64 | 65 | 66 | 67 | 68 | 69 | 70 | 71 | 72 | 73 | 74 | 75 | 76 | 77 | 78 | 79 | 80 | 81 | 82 | 83  | 84  | 85    | 86 | 87 |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| S     | G  | V  | P  | D  | R  | F  | S  | G  | S  | R  | P  | S  | G  | V  | P  | D  | R  | F  | S  | G  | S  | R  | P  | S  | G  | V   | P   | D     | R  | F  | S |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |

Chothia numbering and CDRs

|       |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |       |     |     |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
|-------|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|-------|-----|-----|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| LCDR1 |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    | LCDR2 |     |     |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| 1     | 2  | 3  | 4  | 5  | 6  | 7  | 8  | 9  | 11 | 12 | 13 | 14 | 15 | 16 | 17 | 18 | 19 | 20 | 21 | 22 | 23 | 24 | 25 | 26 | 27 | 28 | 29 | 30    | 30a | 30c |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| Q     | A  | V  | L  | T  | Q  | P  | S  | S  | V  | S  | G  | S  | L  | G  | Q  | R  | V  | S  | I  | T  | C  | S  | G  | S  | S  | N  | V  | G     | G   | N   |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| 35    | 36 | 37 | 38 | 39 | 40 | 41 | 42 | 43 | 44 | 45 | 46 | 47 | 48 | 49 | 50 | 51 | 52 | 53 | 54 | 55 | 56 | 57 | 58 | 59 | 60 | 61 | 62 | 63    | 64  | 65  |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| G     | S  | S  | S  | N  | V  | G  | G  | G  | N  | S  | V  | G  | W  | Y  | Q  | H  | L  | P  | G  | S  | G  | L  | K  | T  | I  | I  | Y  | D     | T   | N   |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| 57    | 58 | 59 | 60 | 61 | 62 | 63 | 64 | 65 | 66 | 67 | 68 | 69 | 70 | 71 | 72 | 73 | 74 | 75 | 76 | 77 | 78 | 79 | 80 | 81 | 82 | 83 | 84 | 85    | 86  | 87  |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| S     | G  | V  | P  | D  | R  | F  | S  | G  | S  | R  | P  | S  | G  | V  | P  | D  | R  | F  | S  | G  | S  | R  | P  | S  | G  | V  | P  | D     | R   | F   | S |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |

Martin numbering and CDRs

|       |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |       |     |     |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
|-------|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|----|-------|-----|-----|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| LCDR1 |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    |    | LCDR2 |     |     |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| 1     | 2  | 3  | 4  | 5  | 6  | 7  | 8  | 9  | 11 | 12 | 13 | 14 | 15 | 16 | 17 | 18 | 19 | 20 | 21 | 22 | 23 | 24 | 25 | 26 | 27 | 28 | 29 | 30    | 30a | 30c |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| Q     | A  | V  | L  | T  | Q  | P  | S  | S  | V  | S  | G  | S  | L  | G  | Q  | R  | V  | S  | I  | T  | C  | S  | G  | S  | S  | N  | V  | G     | G   | N   |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| 35    | 36 | 37 | 38 | 39 | 40 | 41 | 42 | 43 | 44 | 45 | 46 | 47 | 48 | 49 | 50 | 51 | 52 | 53 | 54 | 55 | 56 | 57 | 58 | 59 | 60 | 61 | 62 | 63    | 64  | 65  |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| G     | S  | S  | S  | N  | V  | G  | G  | G  | N  | S  | V  | G  | W  | Y  | Q  | H  | L  | P  | G  | S  | G  | L  | K  | T  | I  | I  | Y  | D     | T   | N   |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| 57    | 58 | 59 | 60 | 61 | 62 | 63 | 64 | 65 | 66 | 67 | 68 | 69 | 70 | 71 | 72 | 73 | 74 | 75 | 76 | 77 | 78 | 79 | 80 | 81 | 82 | 83 | 84 | 85    | 86  | 87  |   |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| S     | G  | V  | P  | D  | R  | F  | S  | G  | S  | R  | P  | S  | G  | V  | P  | D  | R  | F  | S  | G  | S  | R  | P  | S  | G  | V  | P  | D     | R   | F   | S |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |

Fig. 1 (cont)

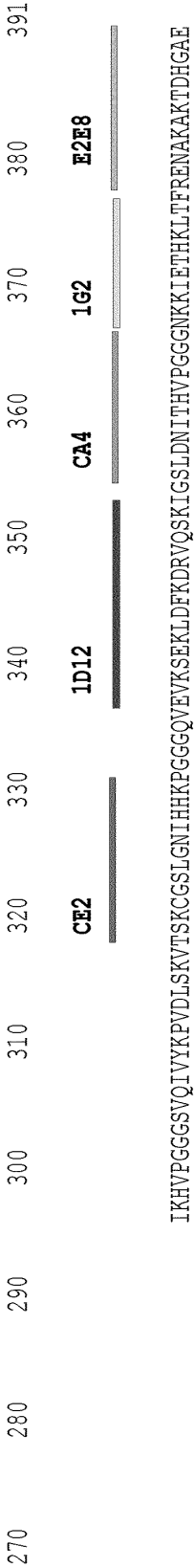


Fig. 2

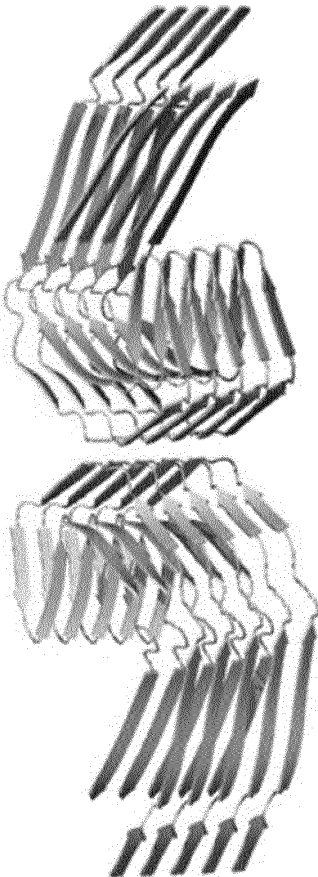


Fig. 3

319-331 CE2

337-355 1D12

355-367 CA4

367-379 1G2

379-391 E2E8

<sup>297</sup>IKHVPGGGSQIVY KPDLSKVTSKGSLGNIHH KPGGGQVEVK SEKLDKDRV QSKIGSLDNI THVPGGNNKK IETHKLTFRE NAKAKTDHGAE<sup>391</sup>

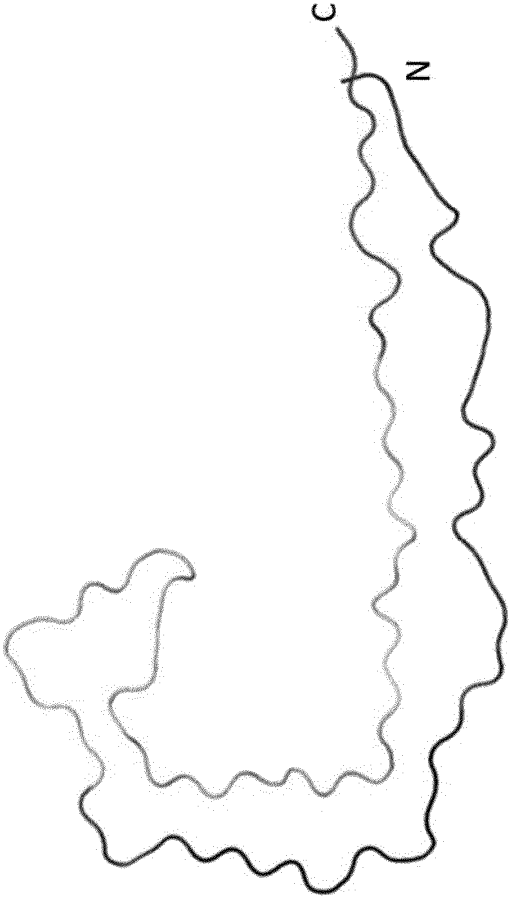


Fig. 4

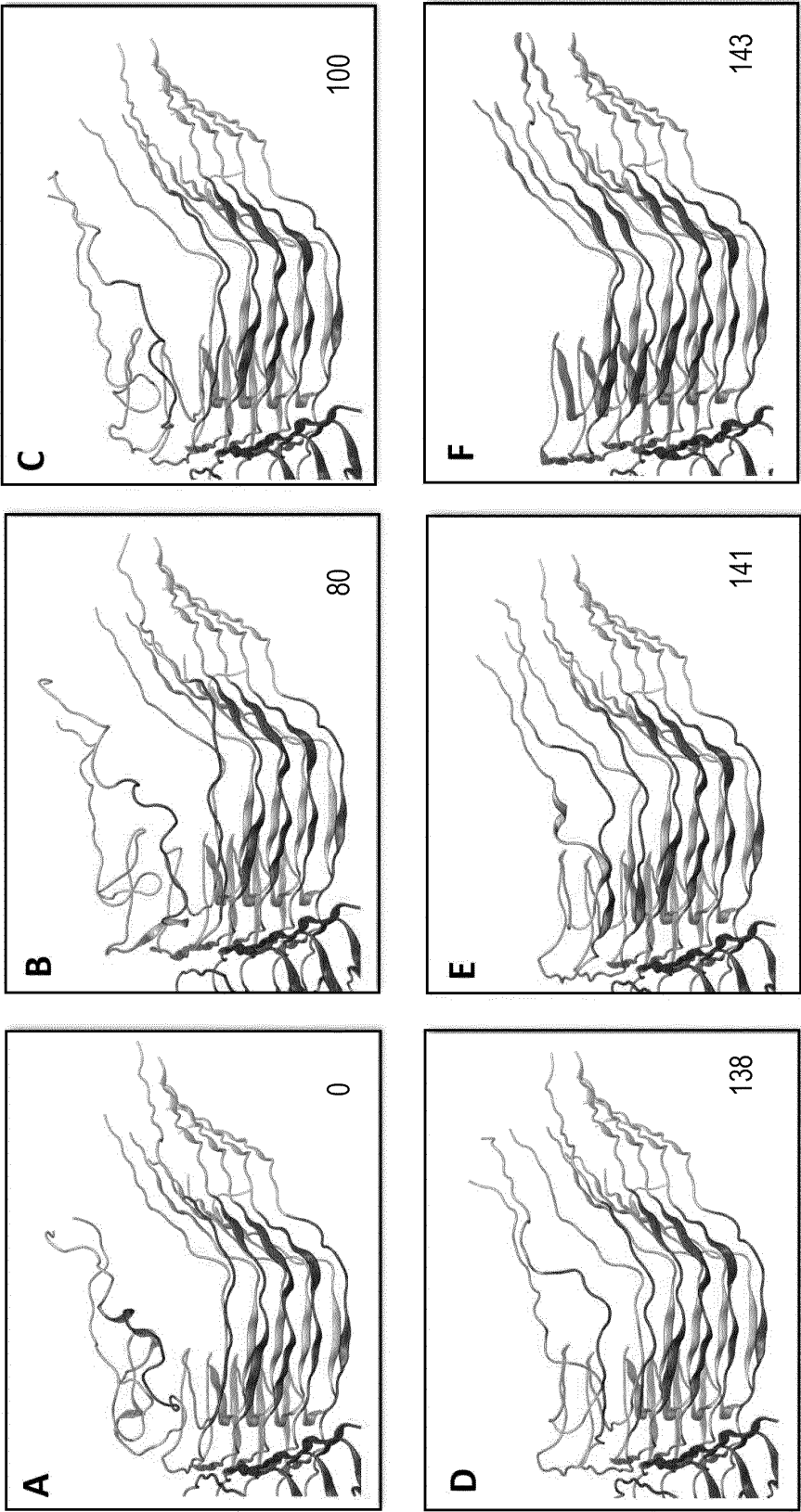


Fig. 5

|     |     |     |     |     |            |     |             |     |            |     |            |             |
|-----|-----|-----|-----|-----|------------|-----|-------------|-----|------------|-----|------------|-------------|
| 270 | 280 | 290 | 300 | 310 | 320        | 330 | 340         | 350 | 360        | 370 | 380        | 391         |
|     |     |     |     |     | <b>CE2</b> |     | <b>1D12</b> |     | <b>CA4</b> |     | <b>1G2</b> | <b>E2E8</b> |
|     |     |     |     |     |            |     |             |     |            |     |            |             |
| (1) |     |     |     |     |            |     |             |     |            |     |            |             |
|     |     |     |     |     |            |     |             |     |            |     |            |             |
| (2) |     |     |     |     |            |     |             |     |            |     |            |             |
|     |     |     |     |     |            |     |             |     |            |     |            |             |
| (3) |     |     |     |     |            |     |             |     |            |     |            |             |

Fig. 6

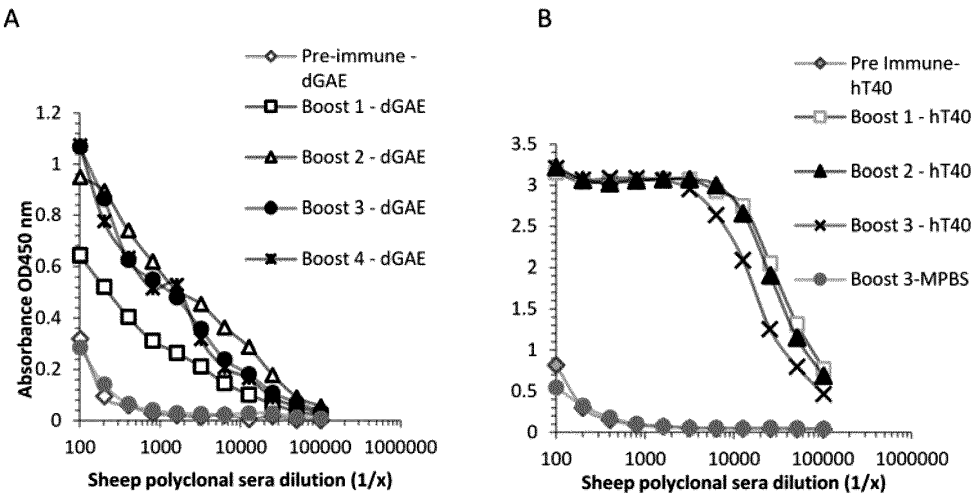


Fig. 7

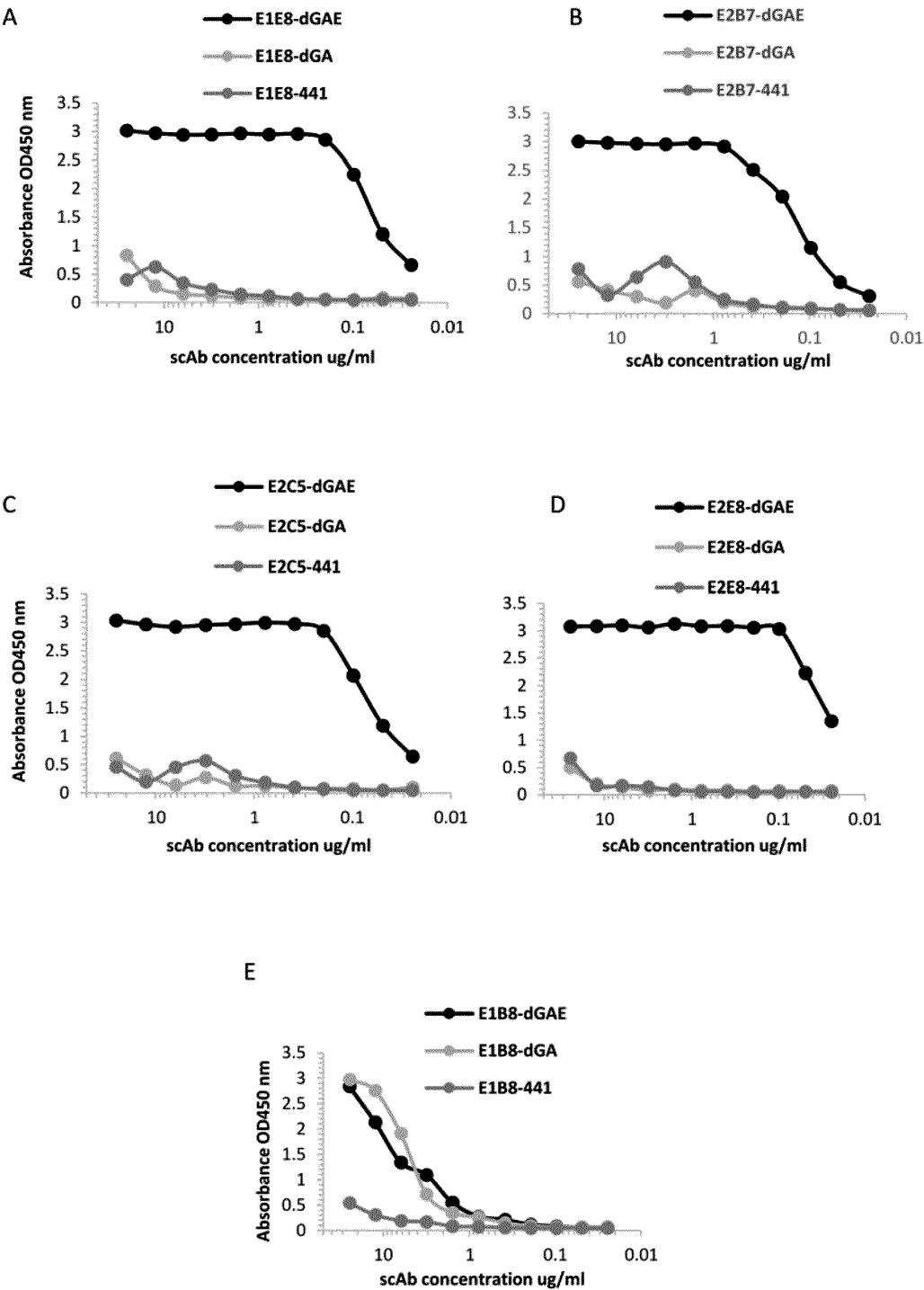


Fig. 8

9/64

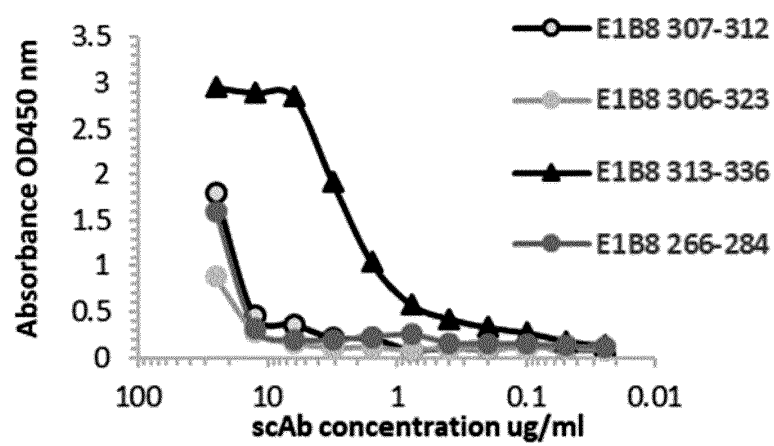
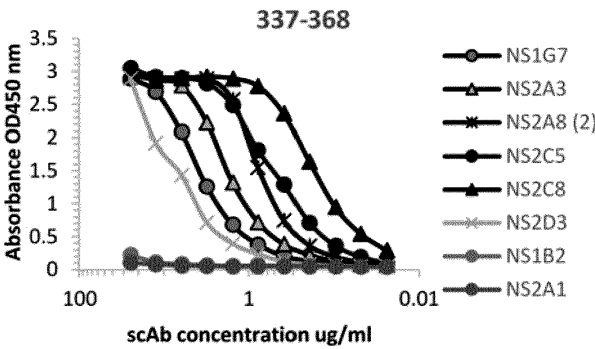
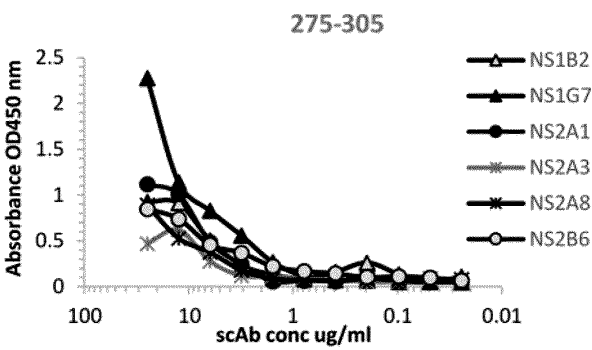


Fig. 9

A



B



C

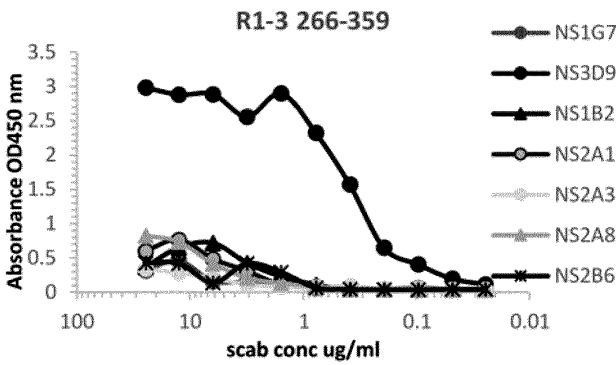
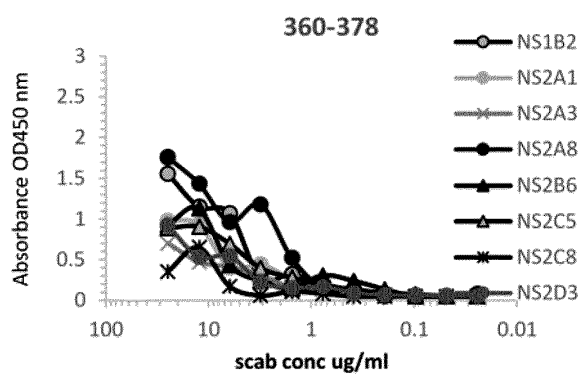


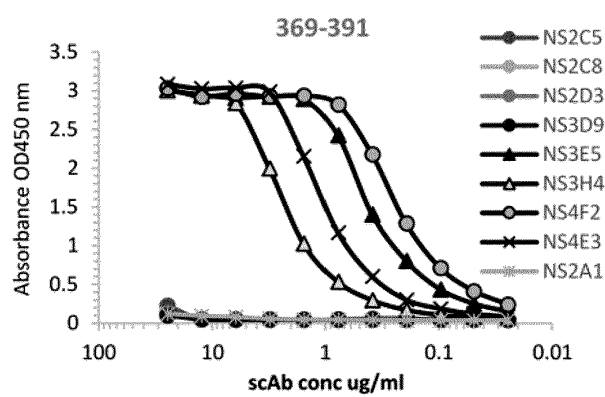
Fig. 10

11/64

D



E



F

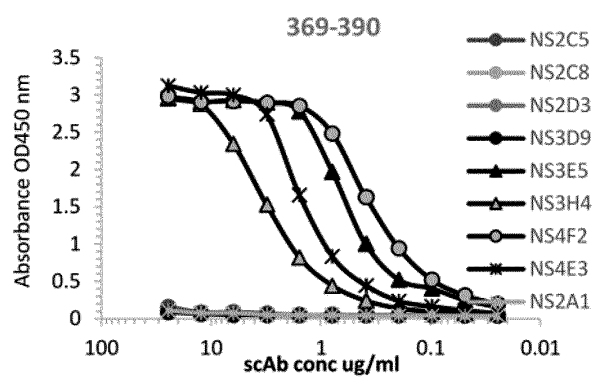


Fig. 10 (cont)

12/64

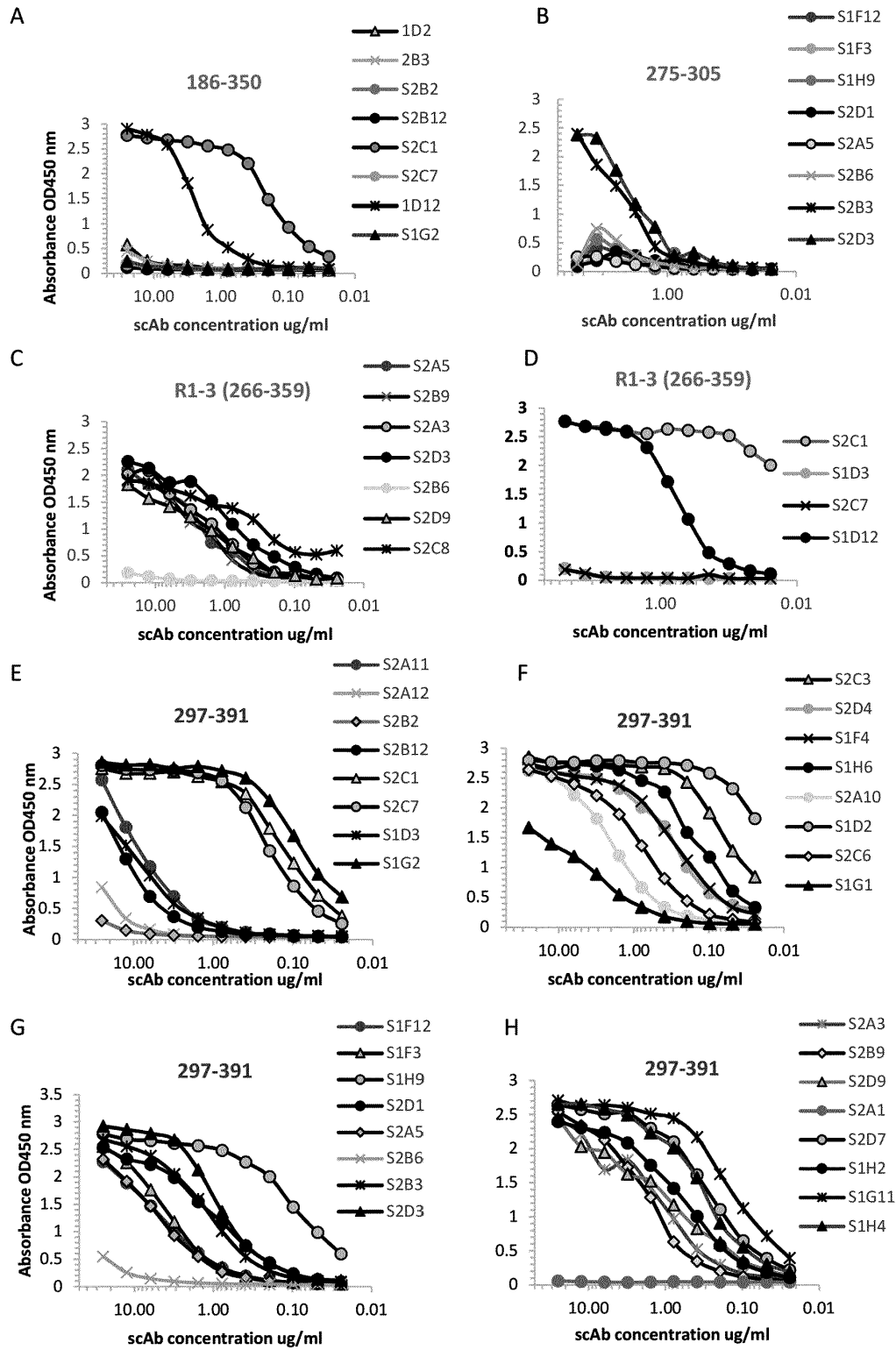


Fig. 11

13/64

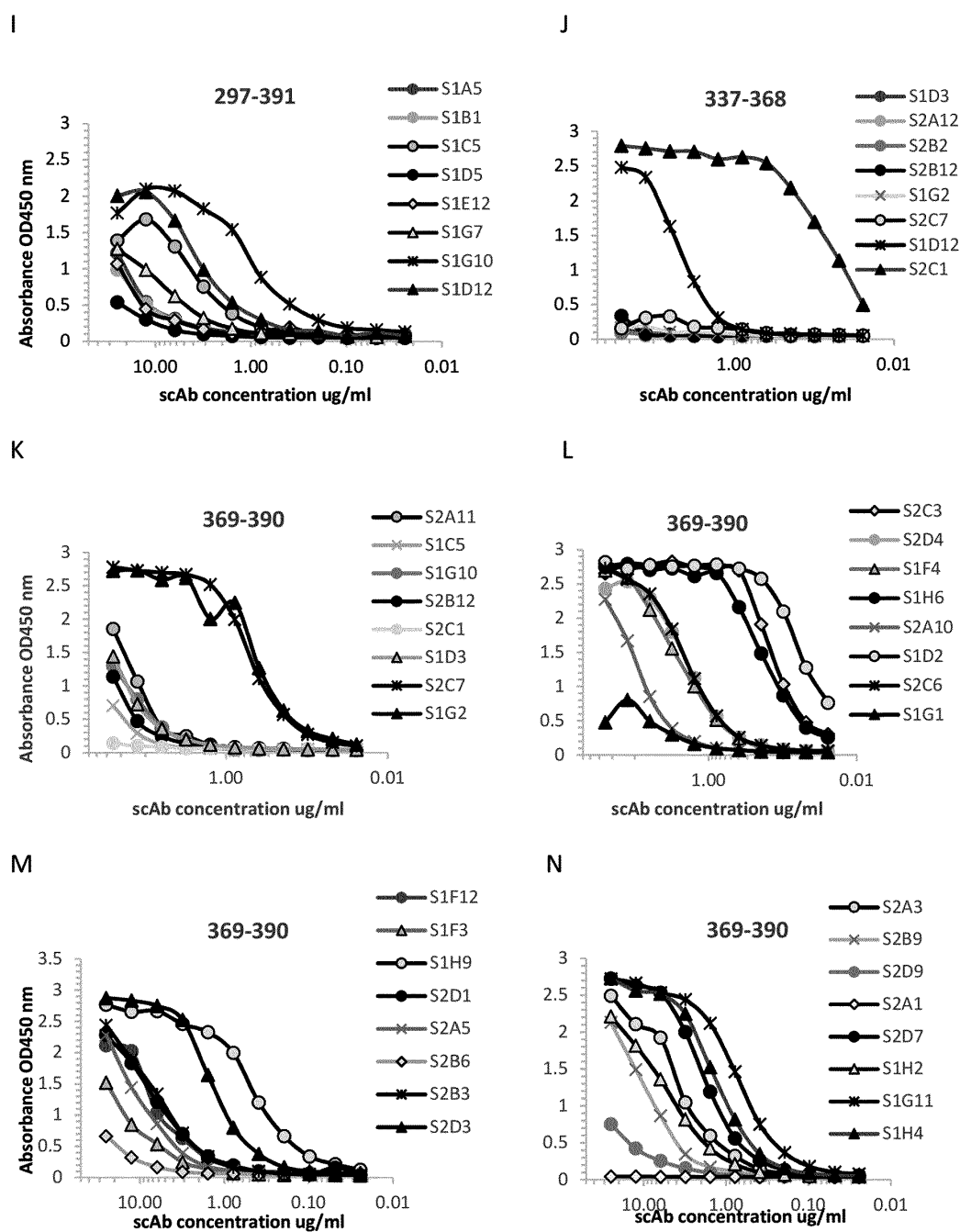
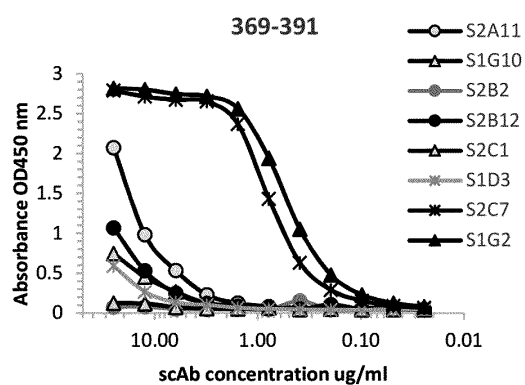


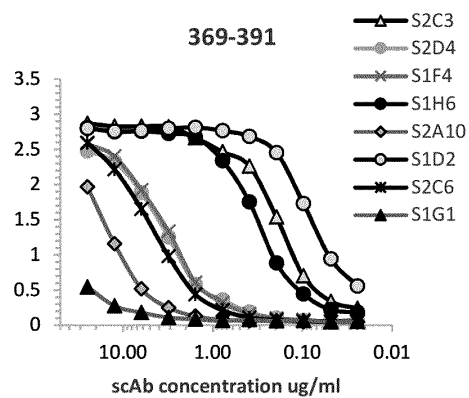
Fig. 11 (cont)

14/64

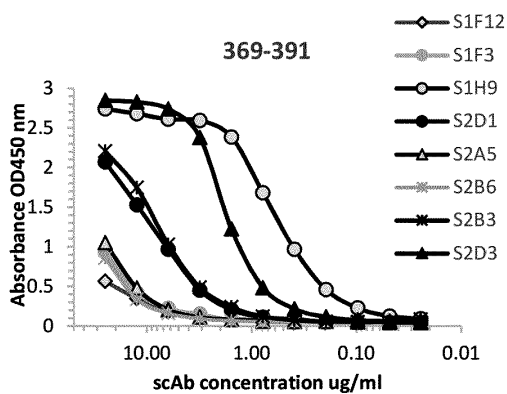
O



P



Q



R

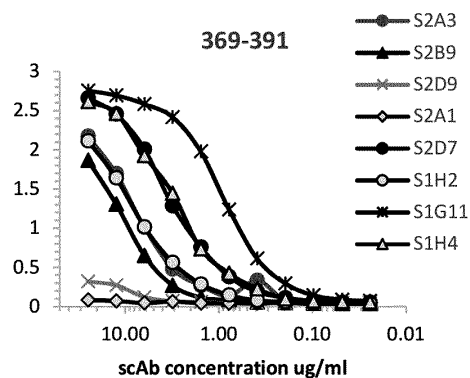


Fig. 11 (cont)

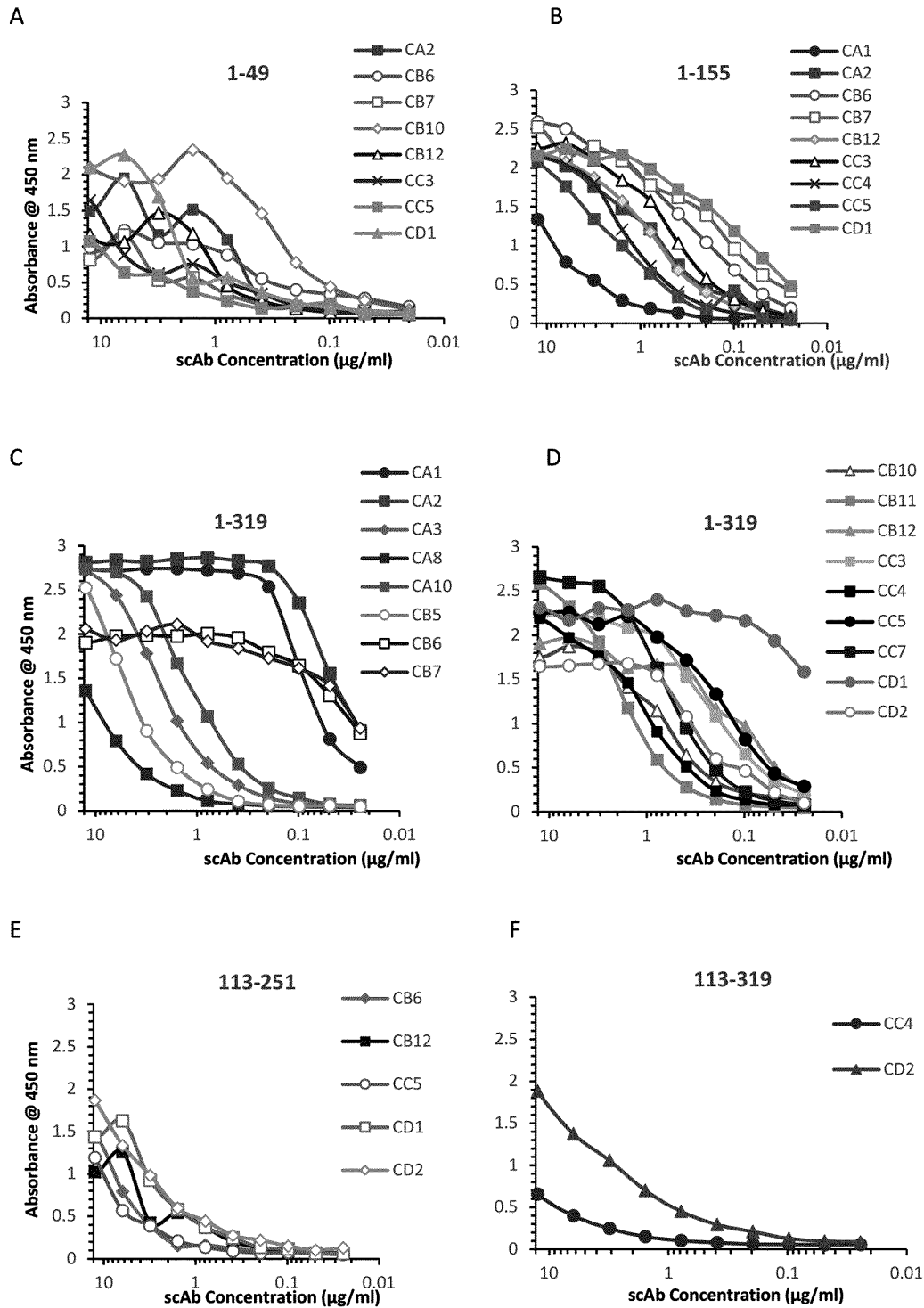


Fig. 12

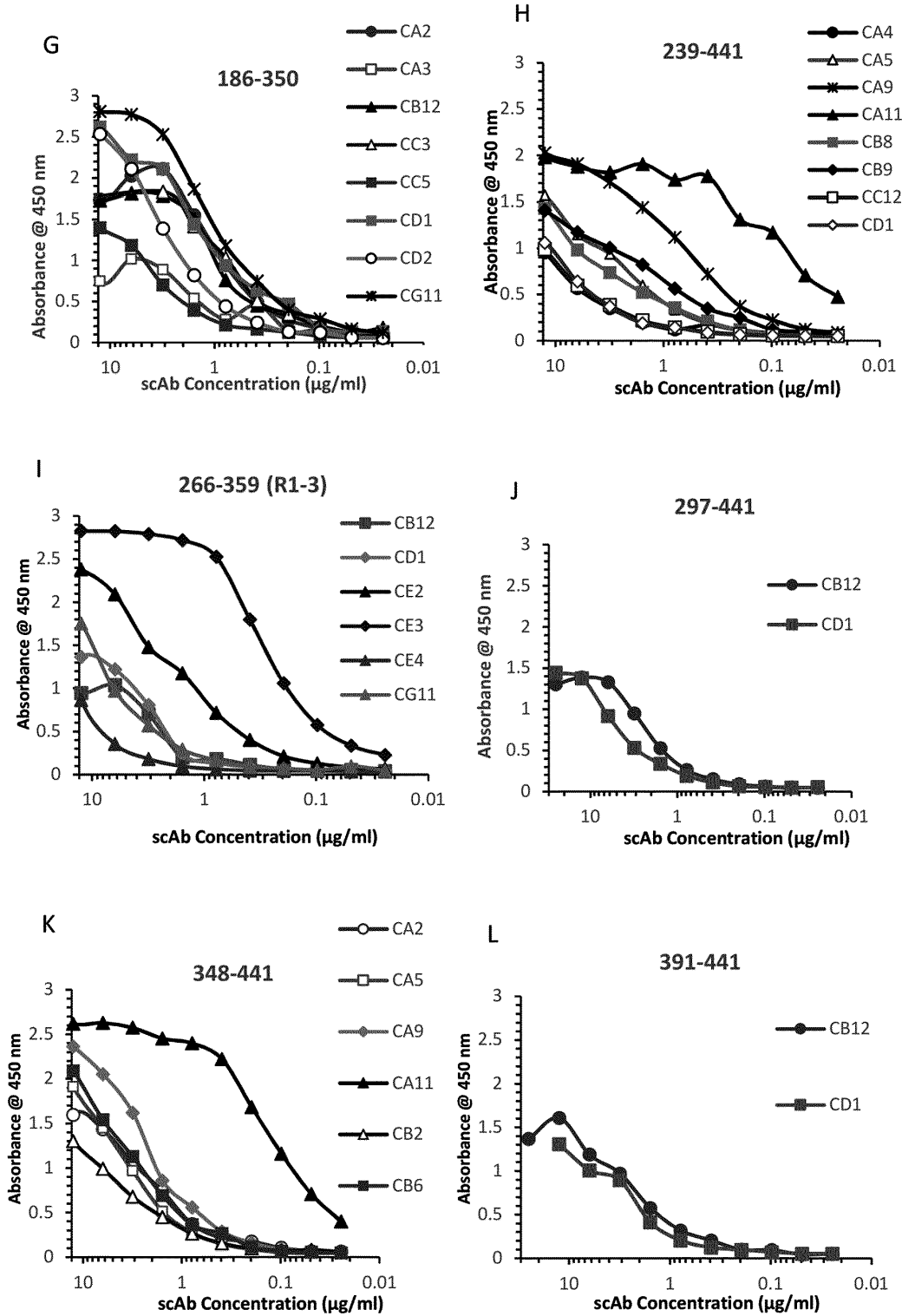


Fig. 12 (cont)

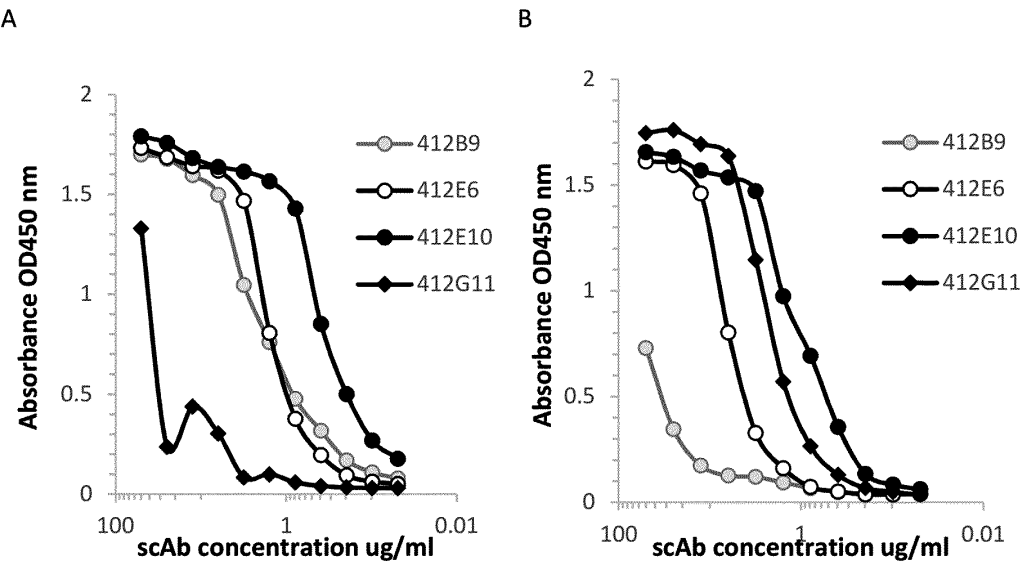


Fig. 13

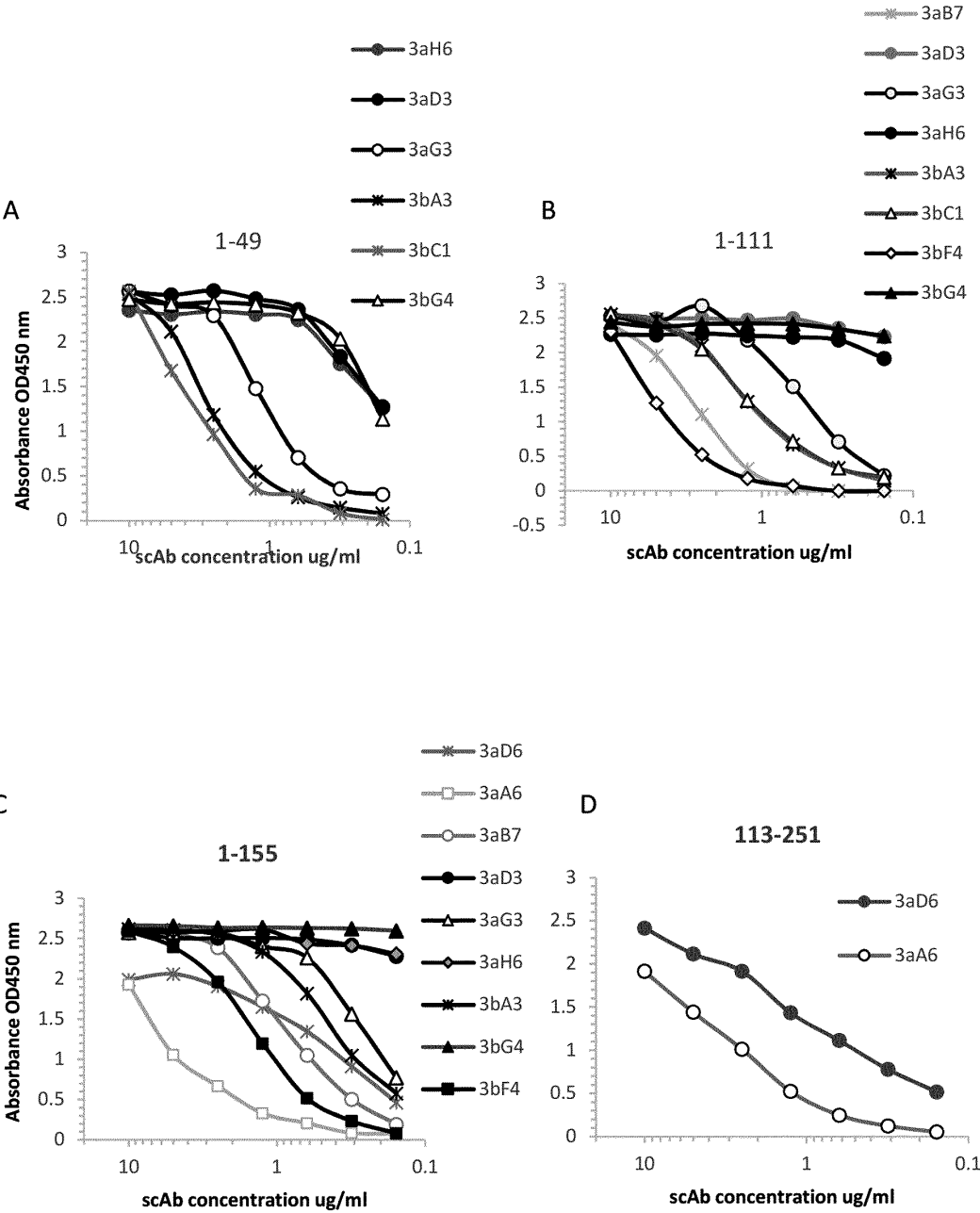
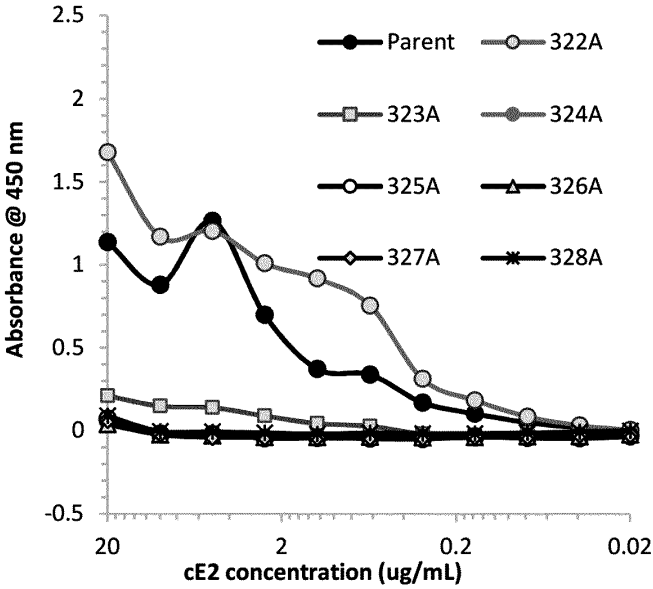


Fig. 14

A



B

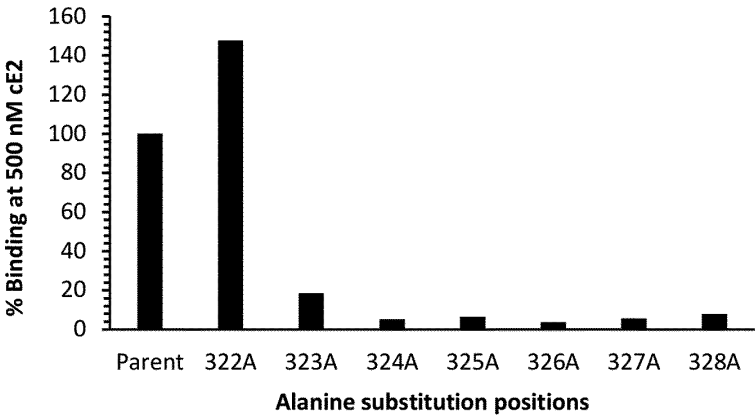


Fig. 15

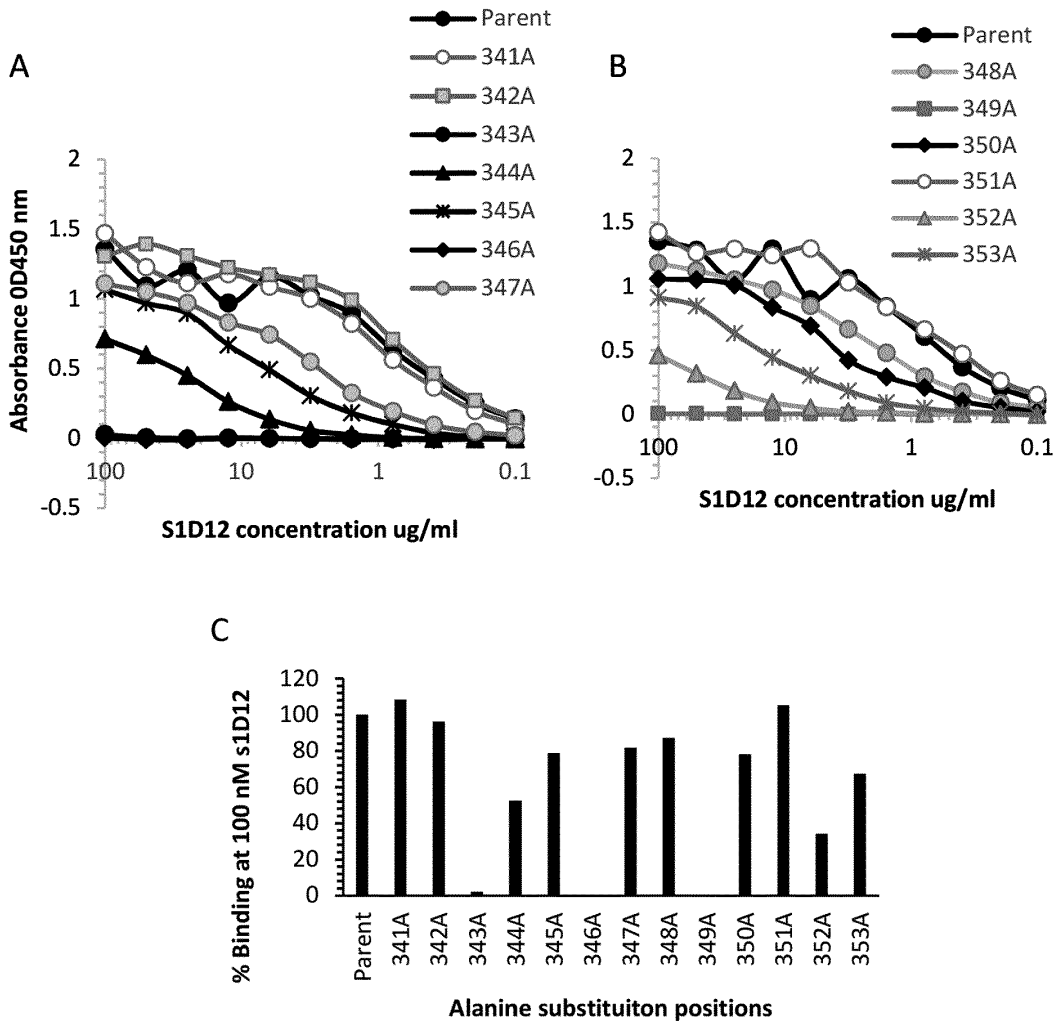


Fig. 16

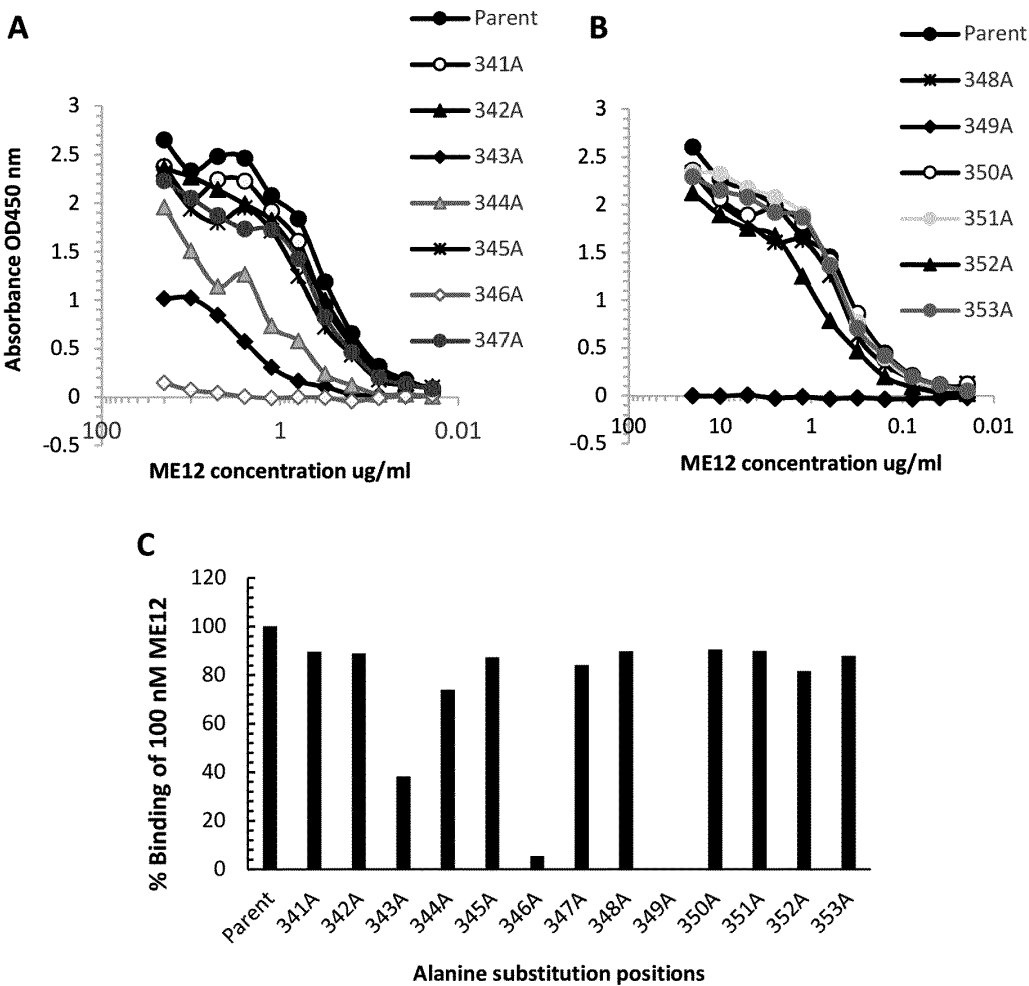
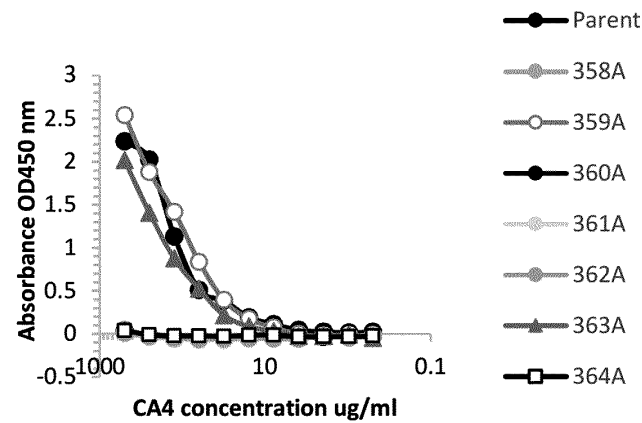


Fig. 17

22/64

A



B

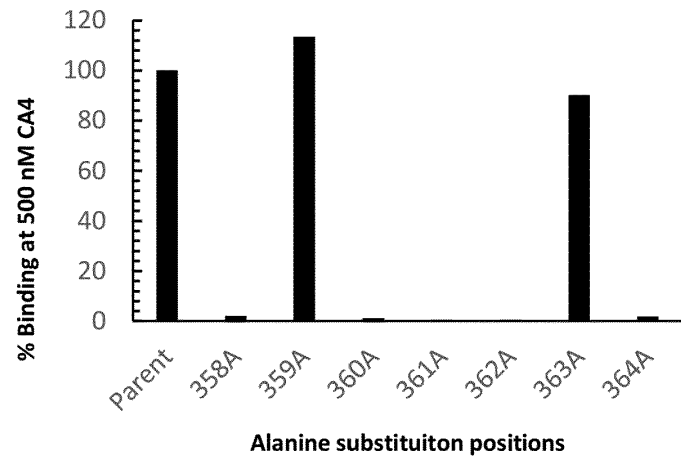


Fig. 18

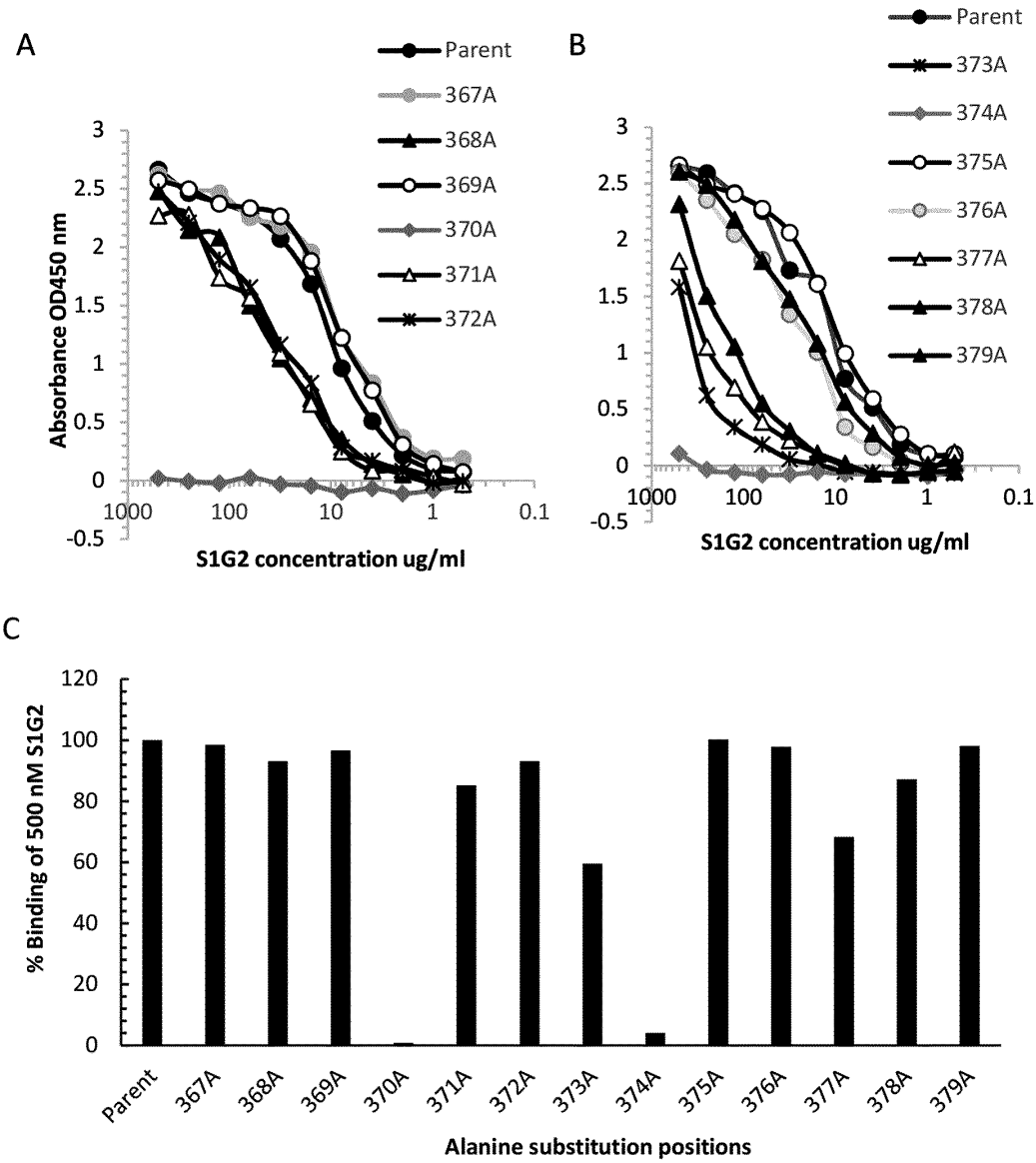
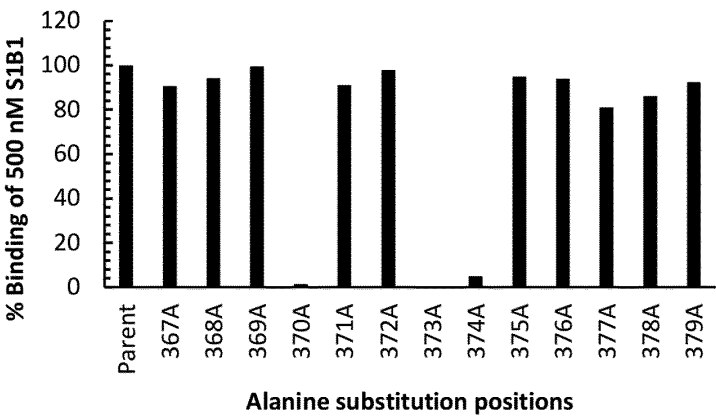


Fig. 19

**A**



**B**

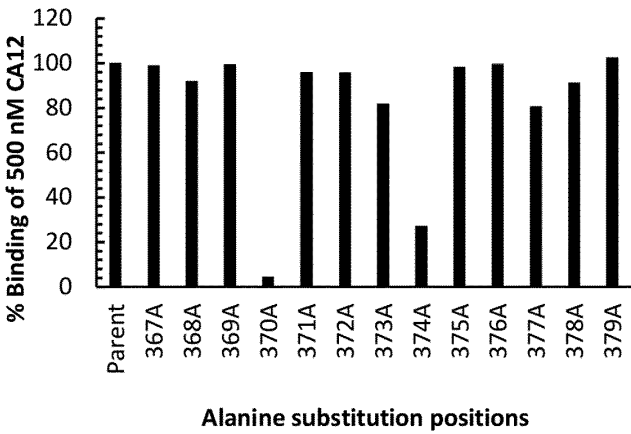
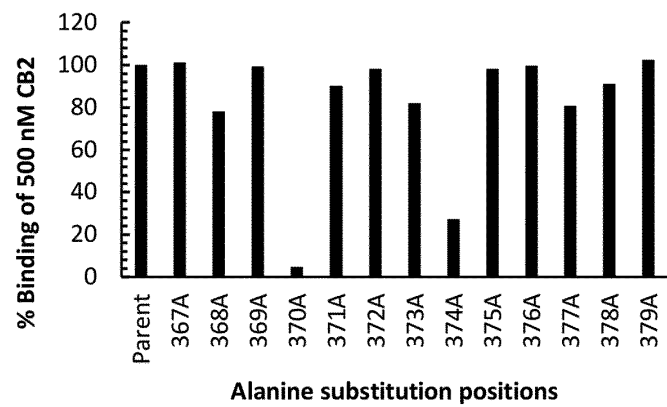


Fig. 20

C



D

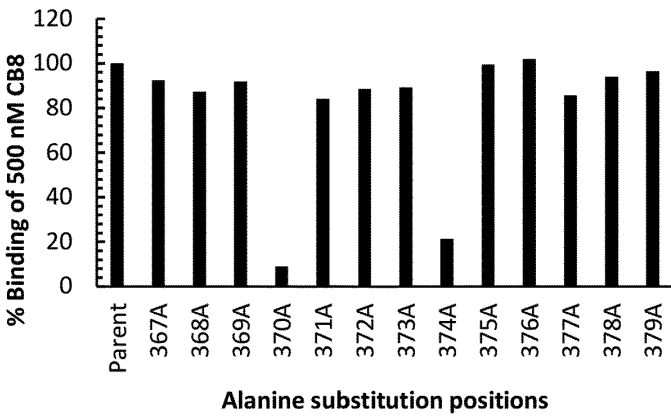


Fig. 20 (cont)

26/64

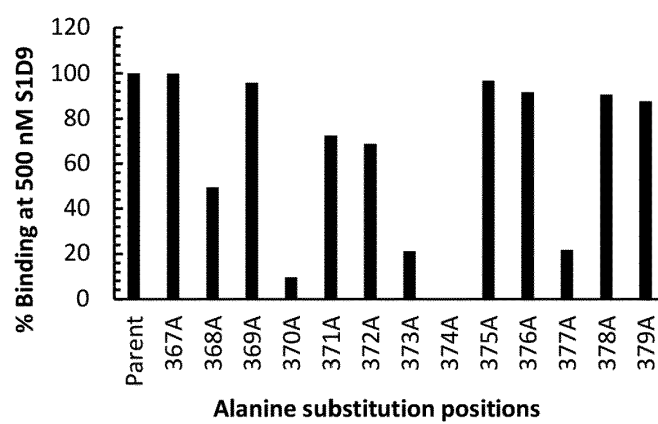
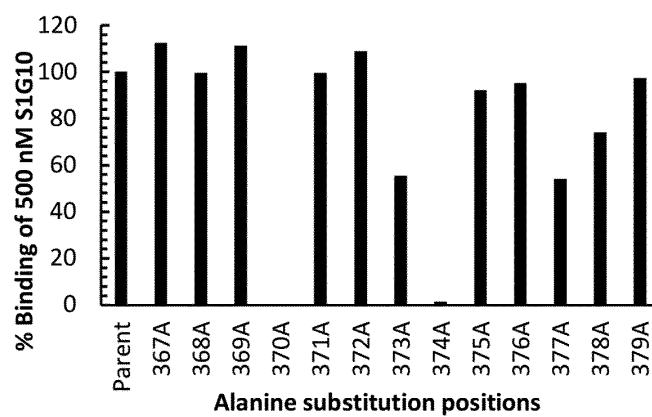
**E****F**

Fig. 20 (cont)

27/64

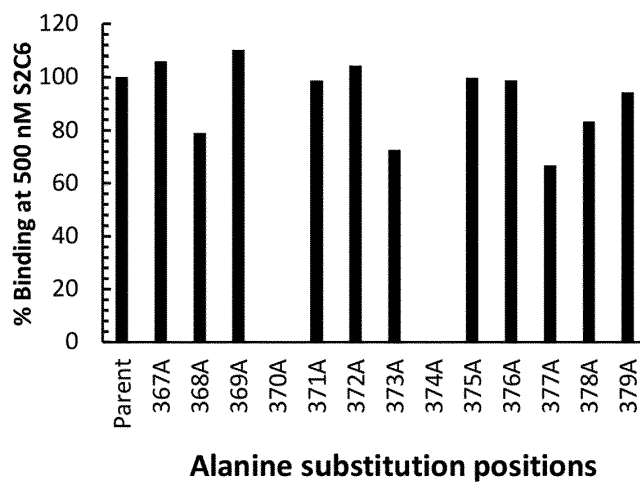
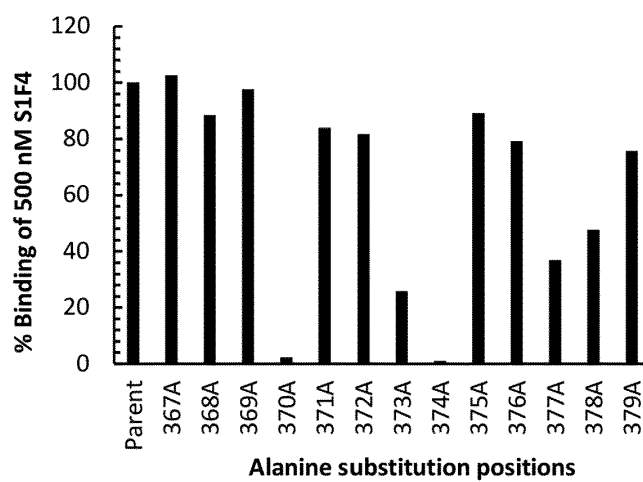
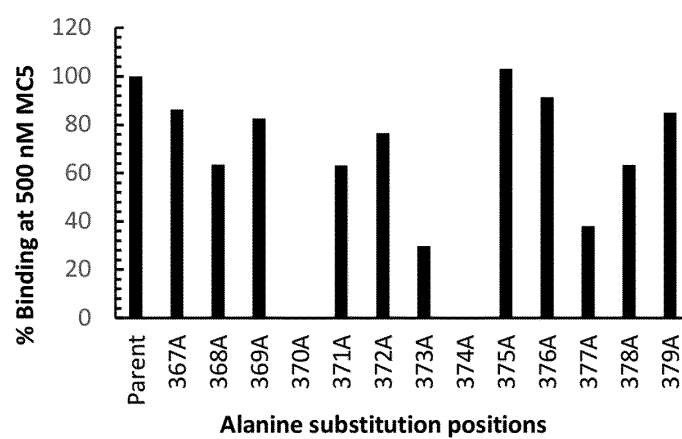
**G****H**

Fig. 20 (cont)

28/64

I



J

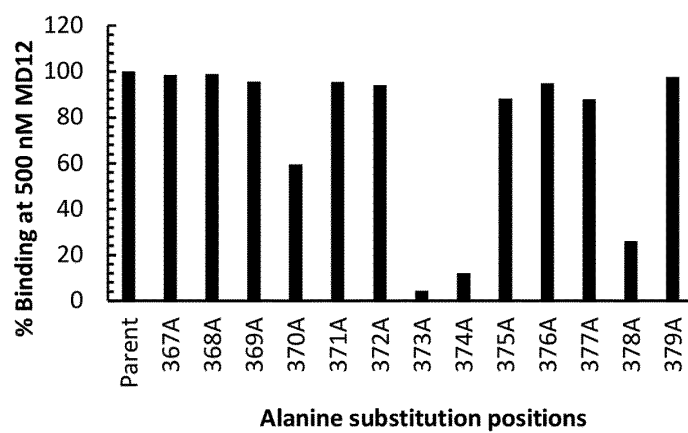


Fig. 20 (cont)

29/64

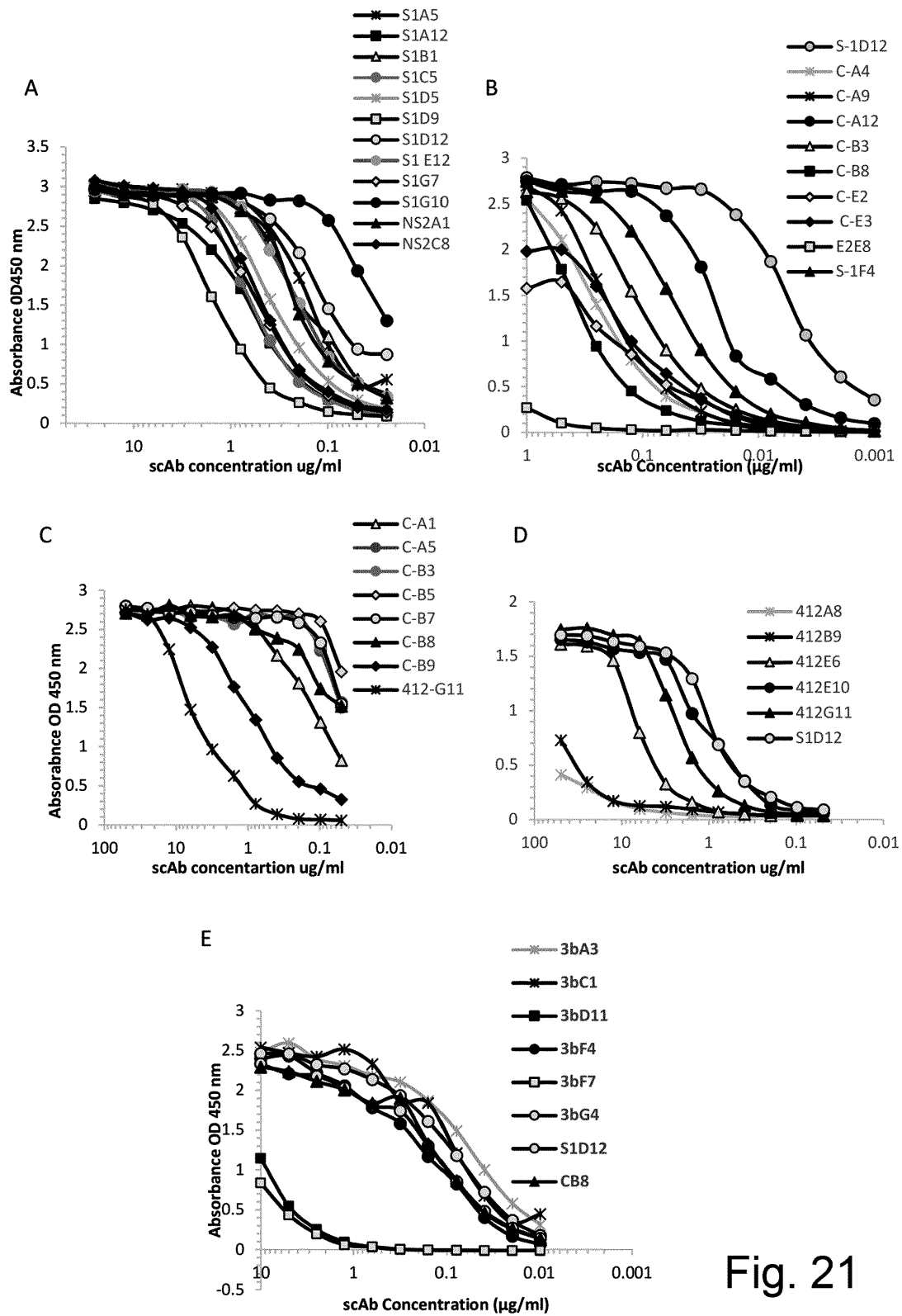


Fig. 21

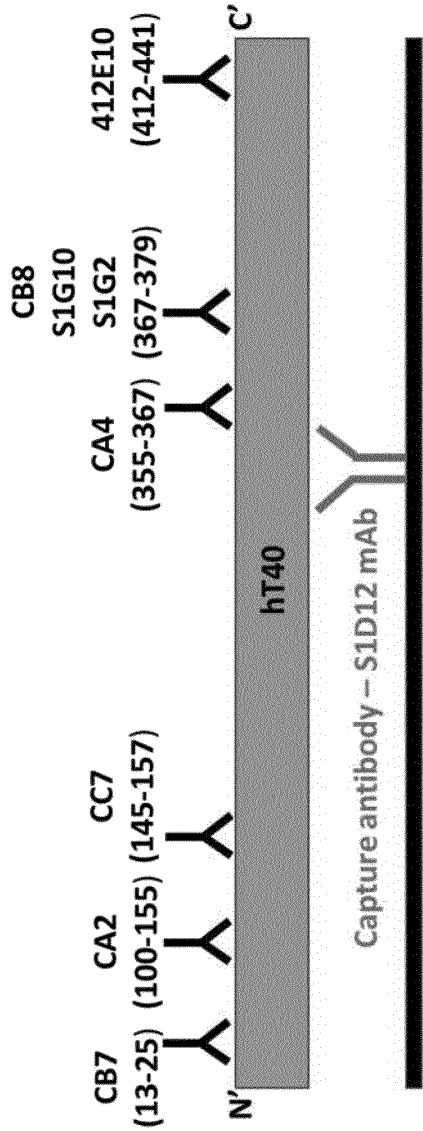


Fig. 22

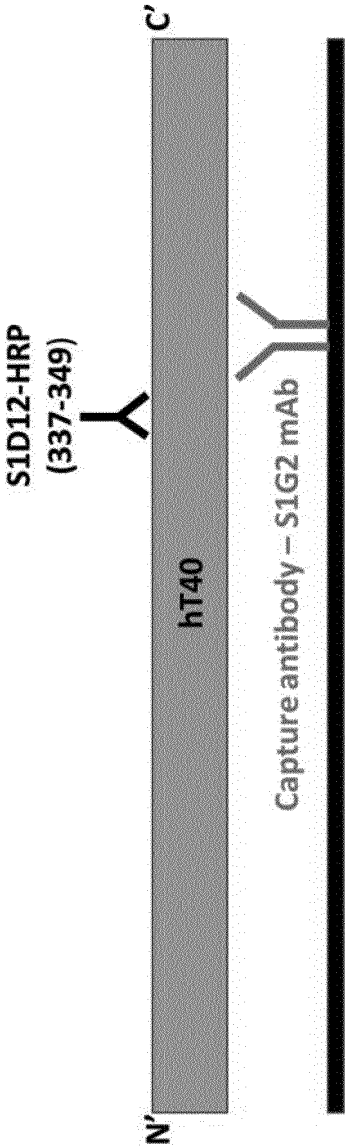


Fig. 23

32/64

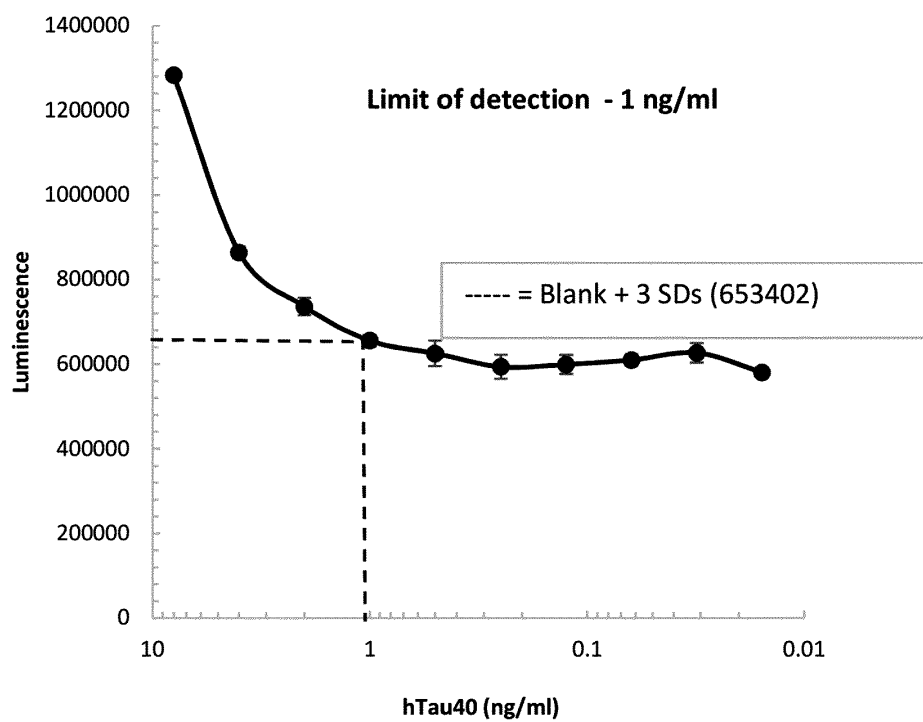


Fig. 24

33/64

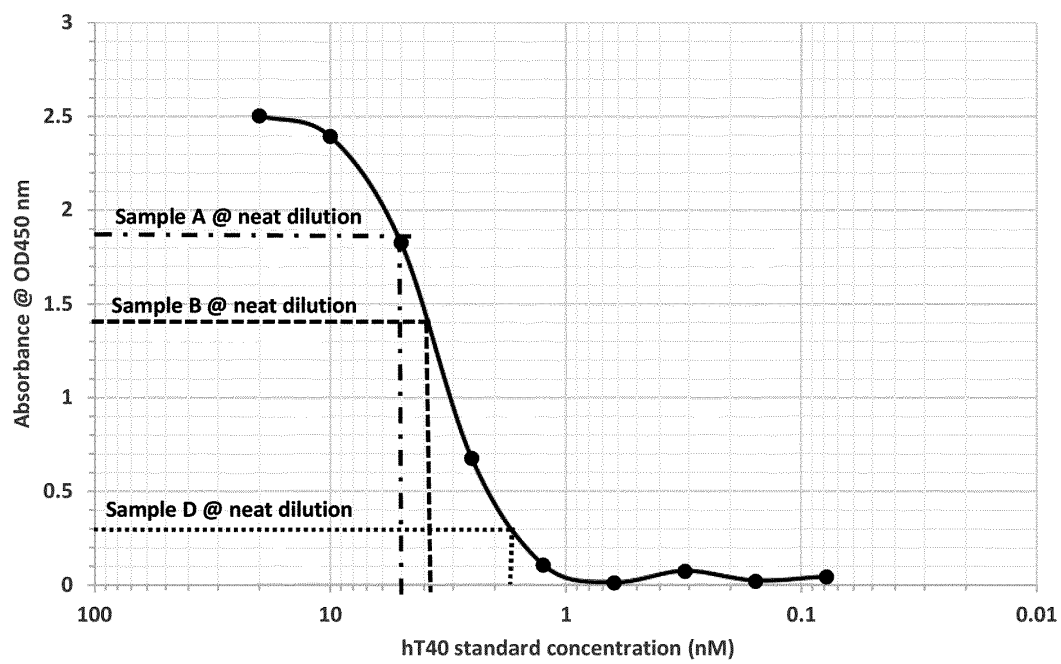


Fig. 25

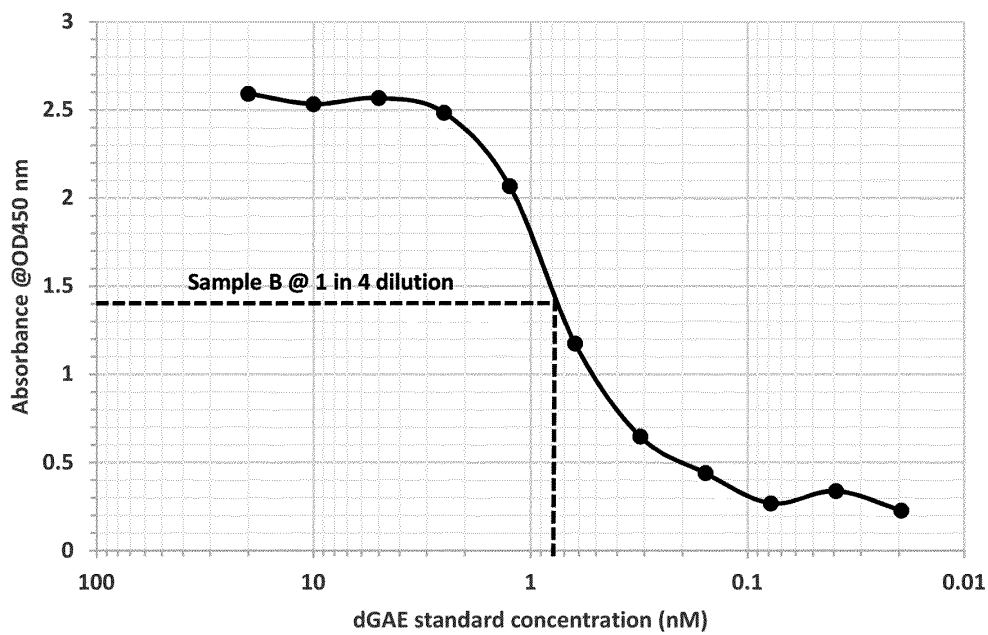


Fig. 26

35/64

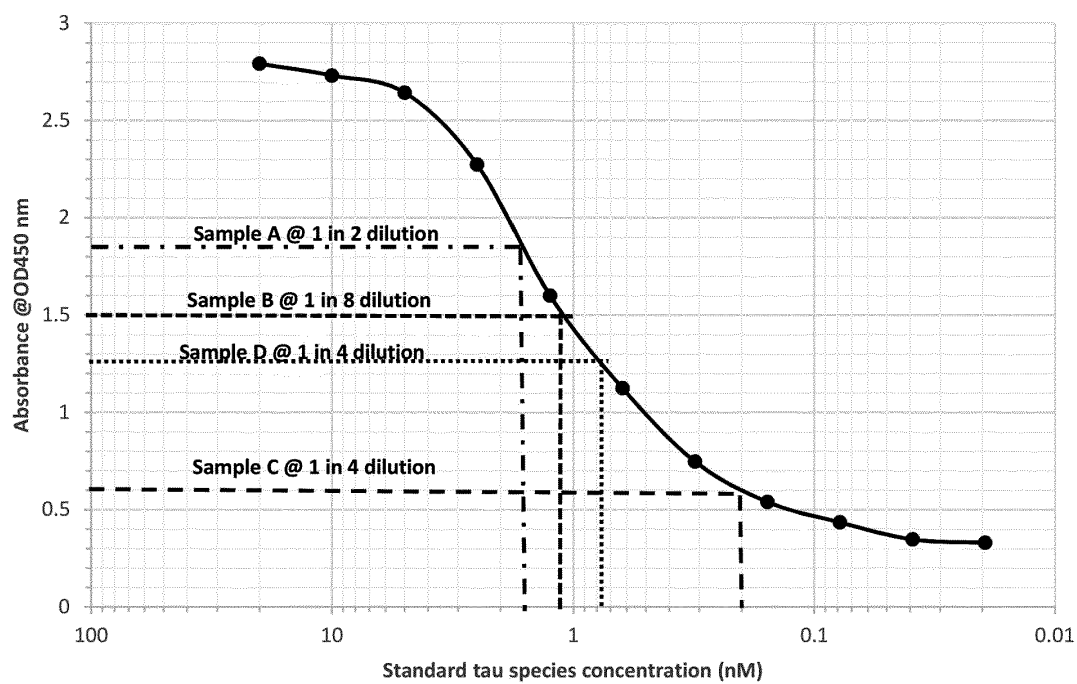


Fig. 27

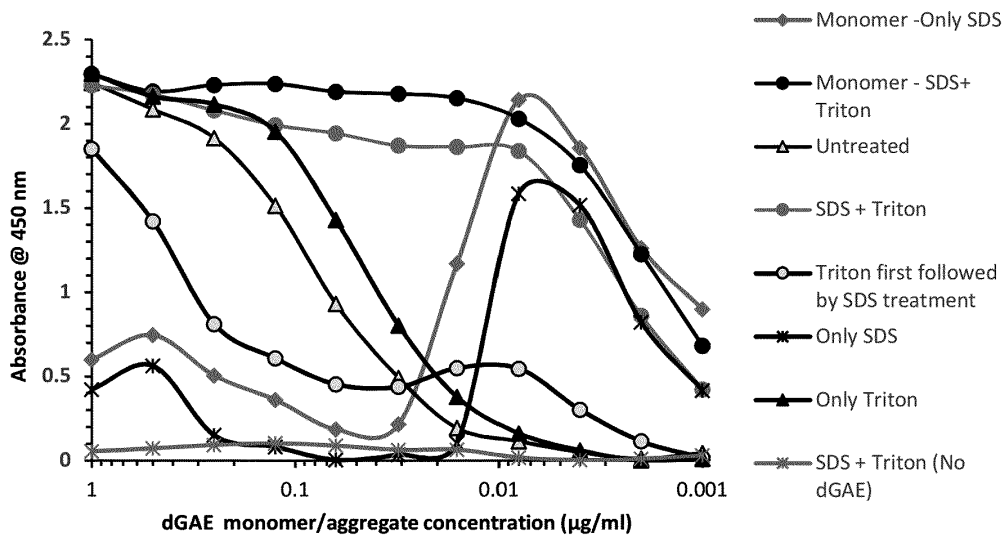


Fig. 28

37/64

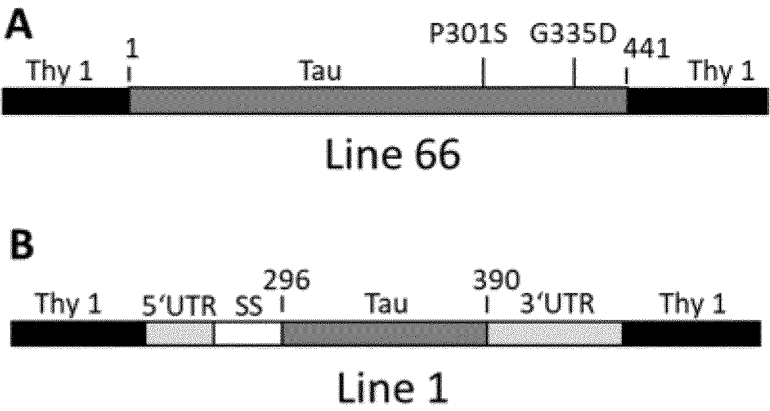


Fig. 29

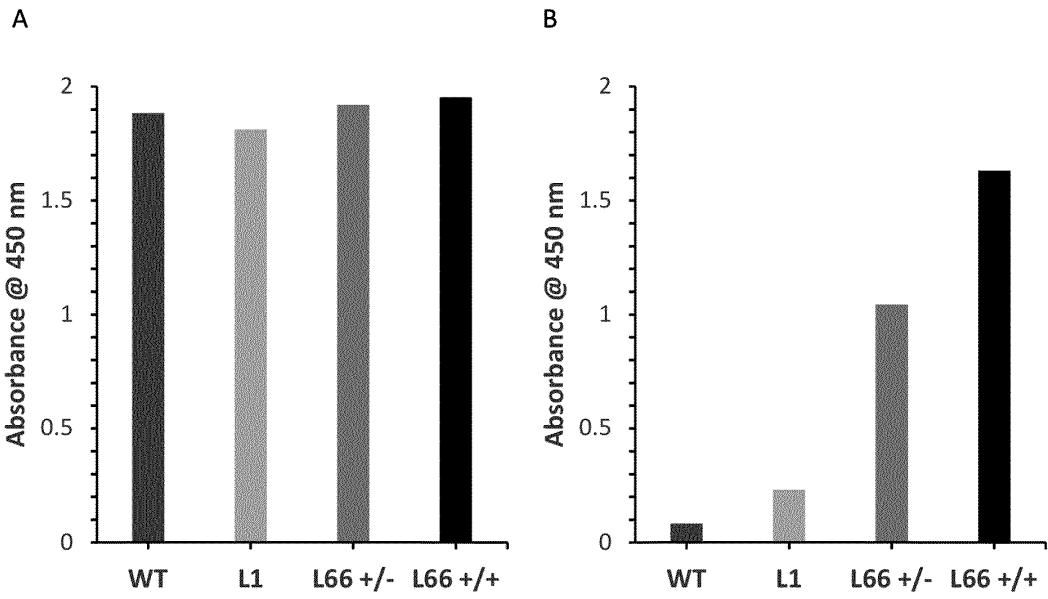


Fig. 30

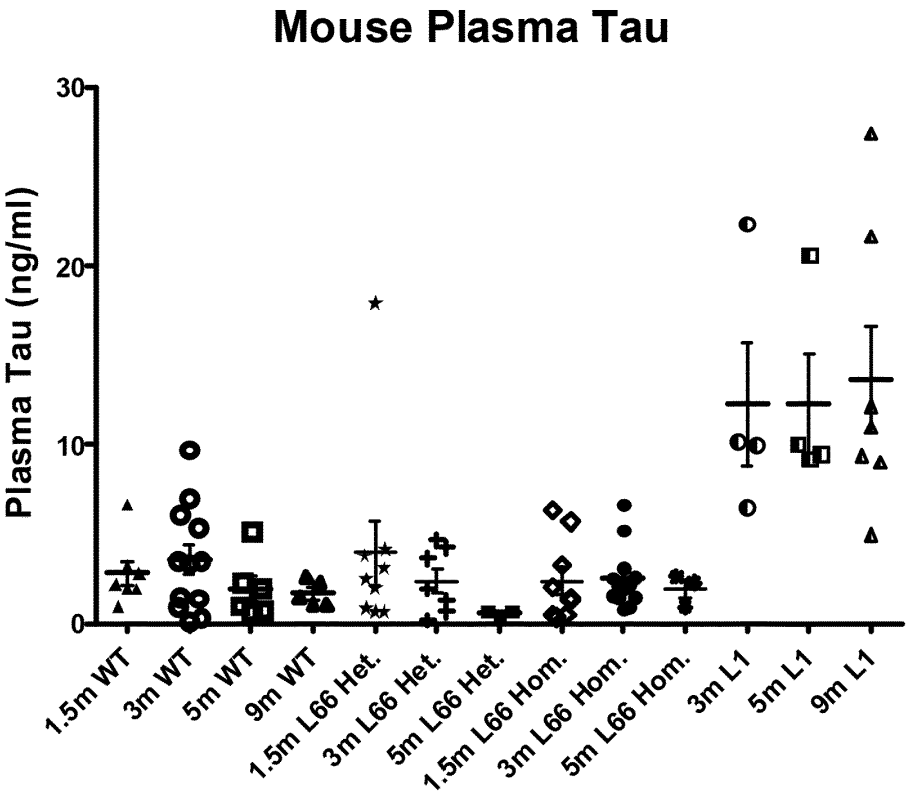


Fig. 31

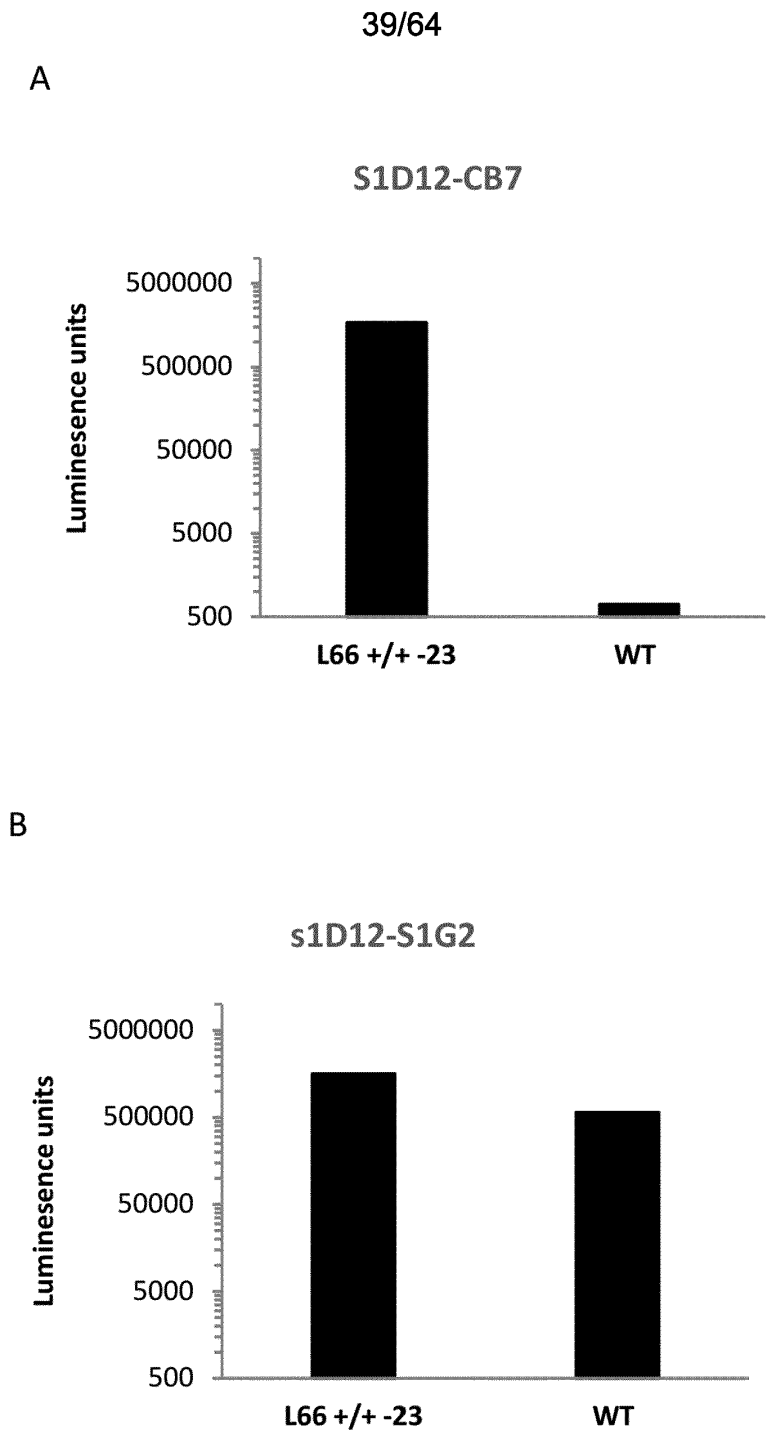


Fig. 32

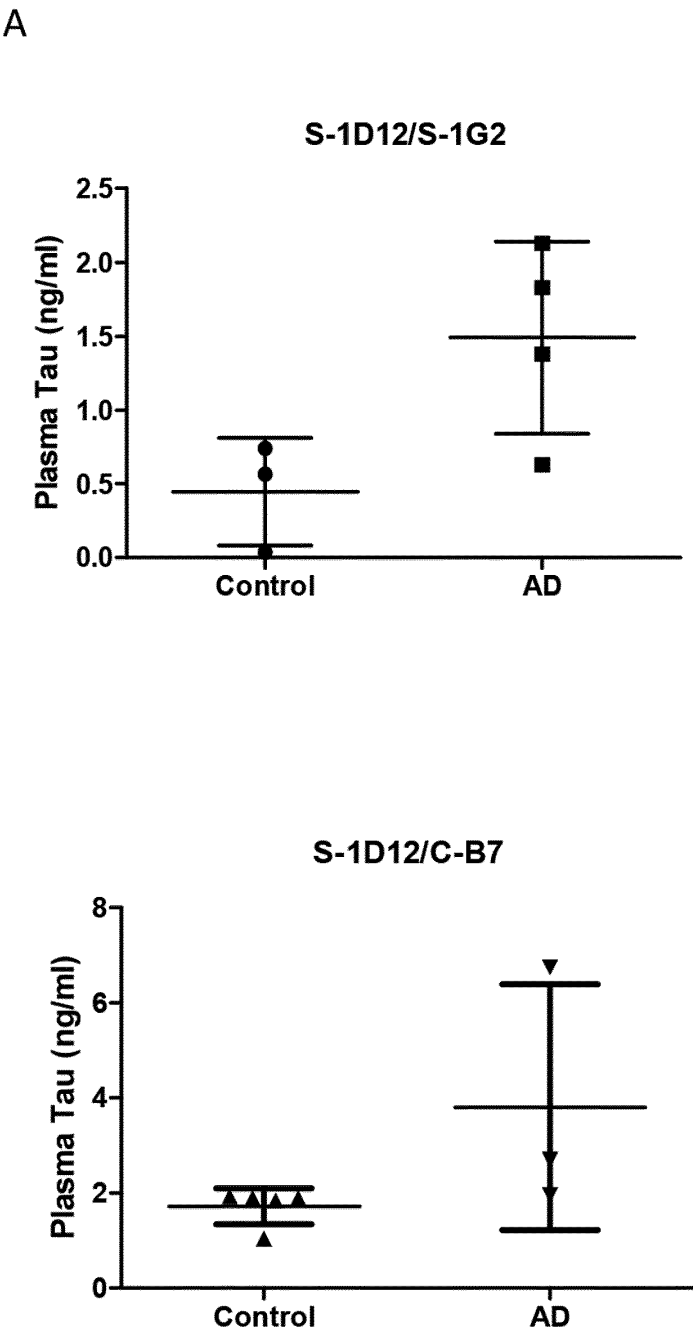


Fig. 33

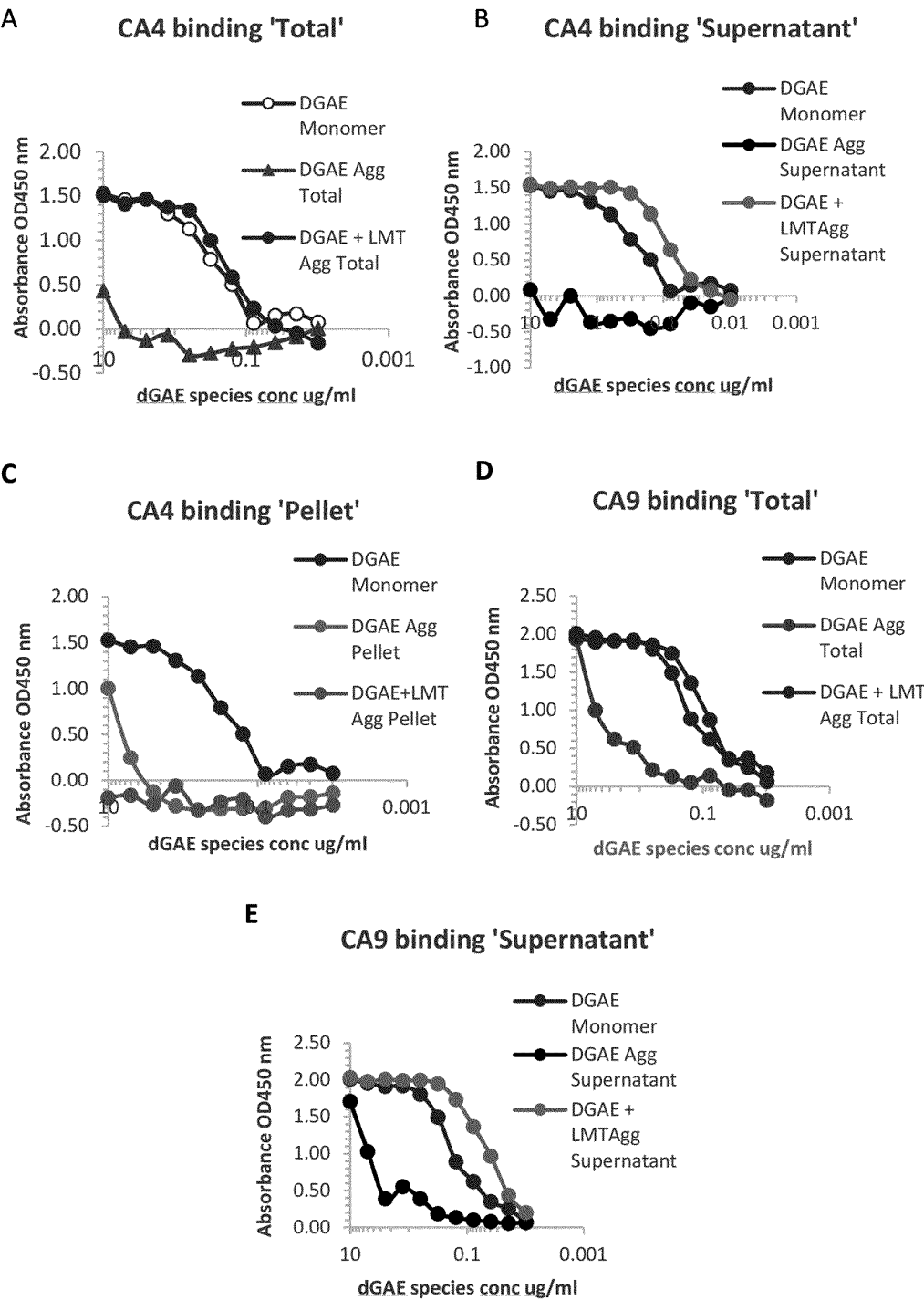


Fig. 34

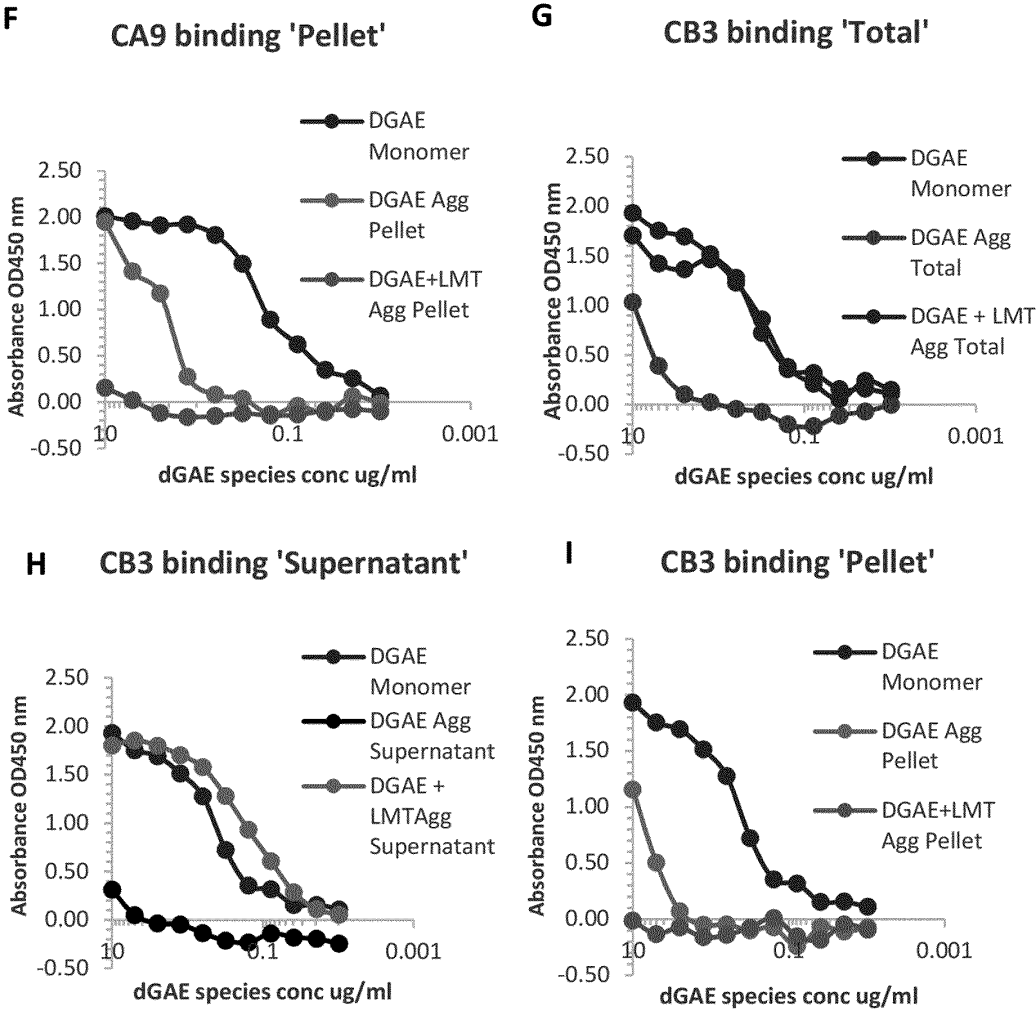


Fig. 34 (cont)

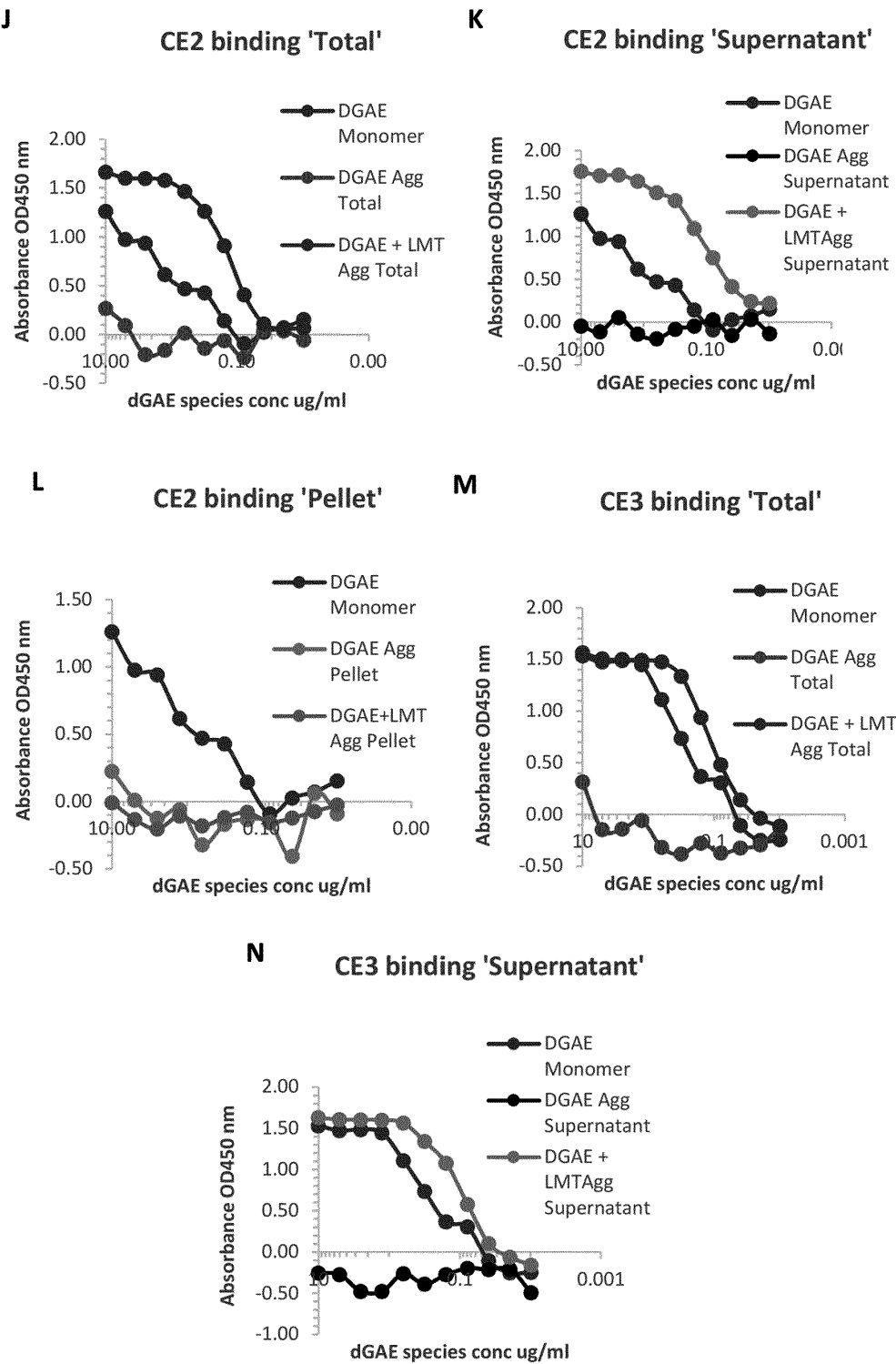


Fig. 34 (cont)

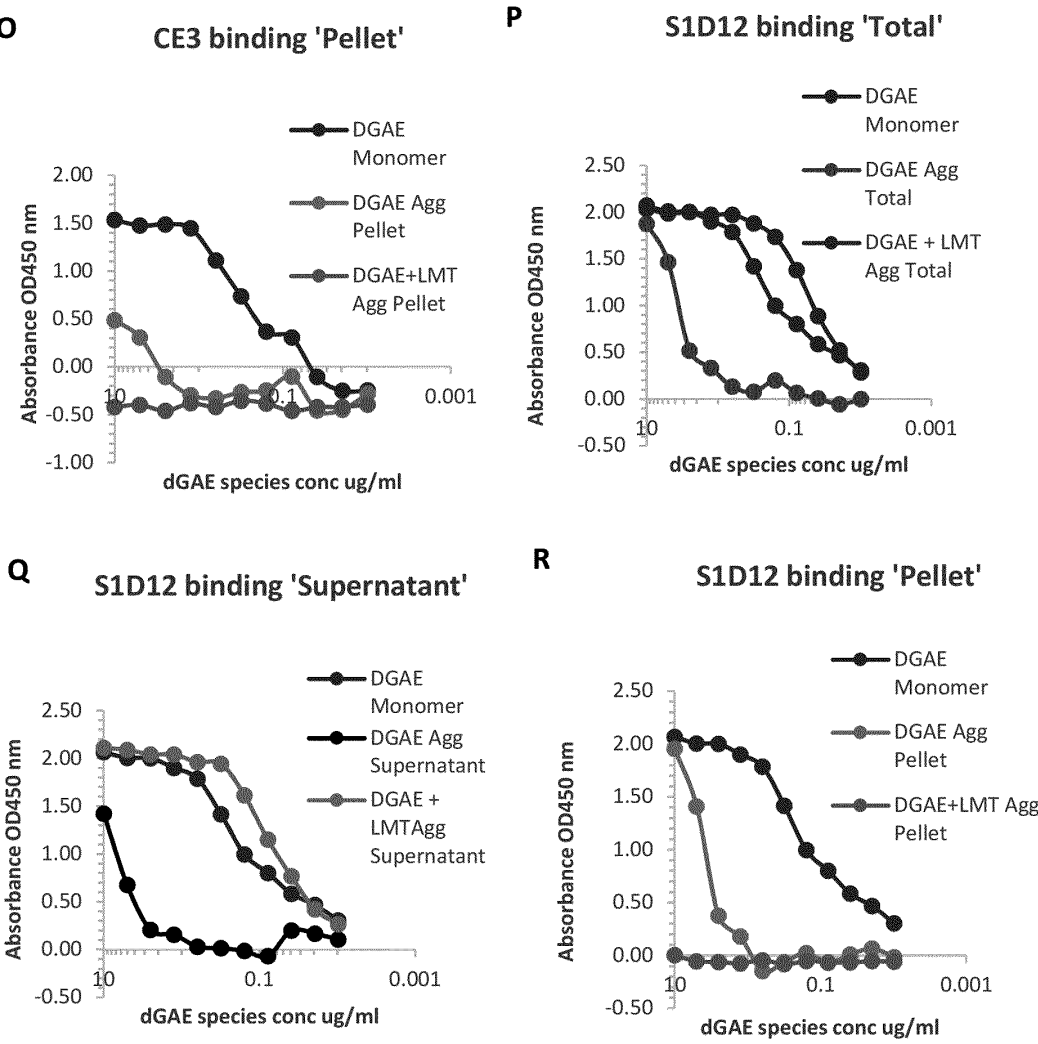


Fig. 34 (cont)

45/64

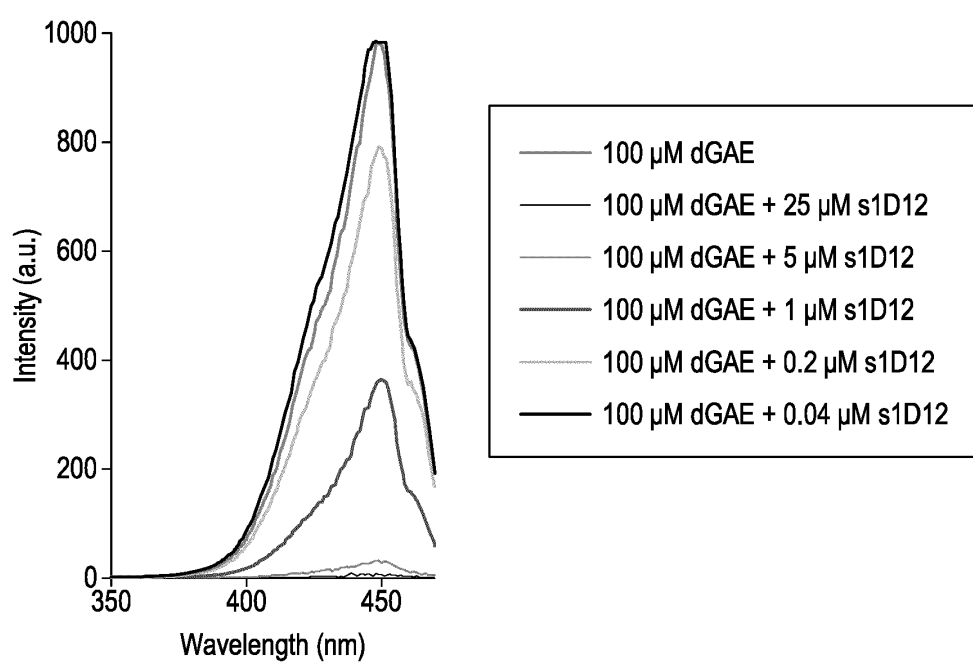


Fig. 35

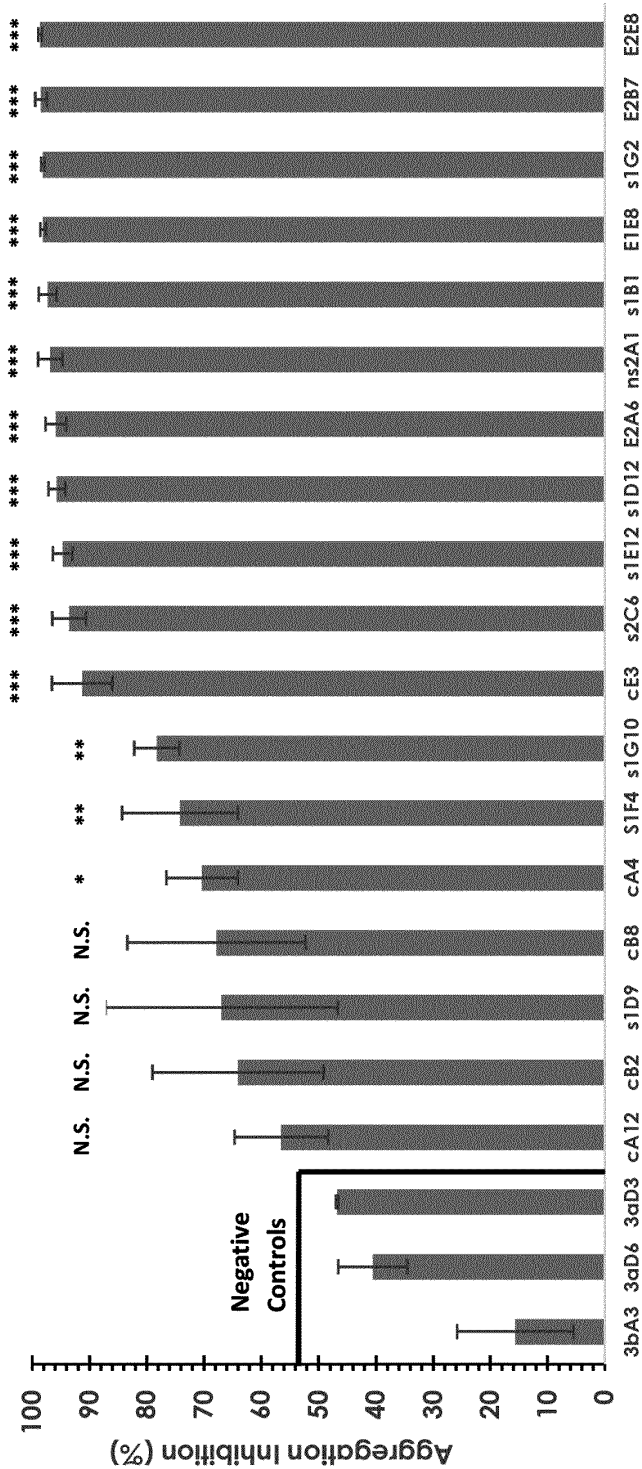


Fig. 36

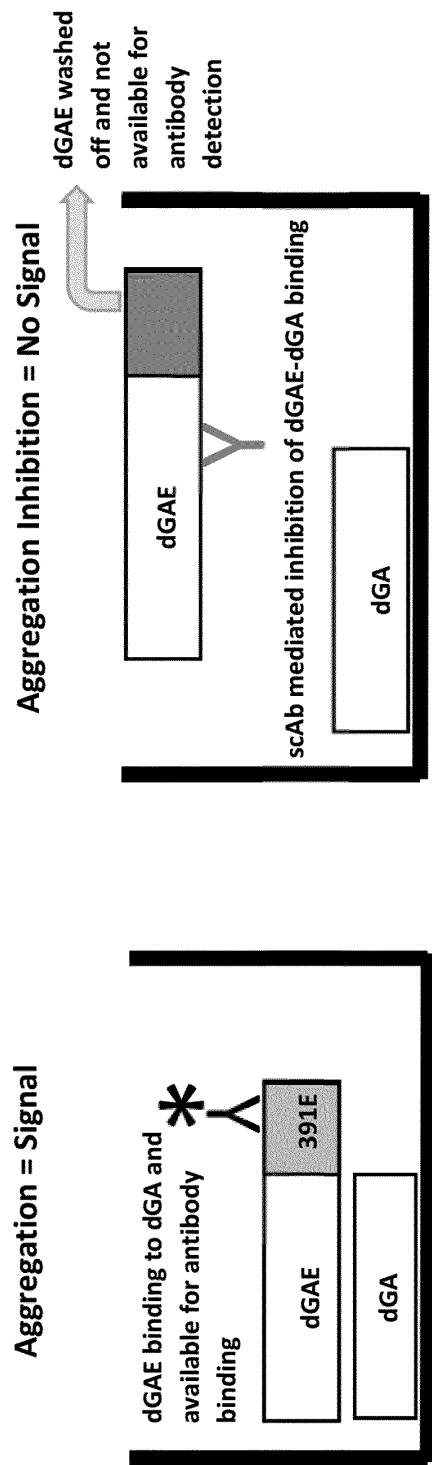


Fig. 37

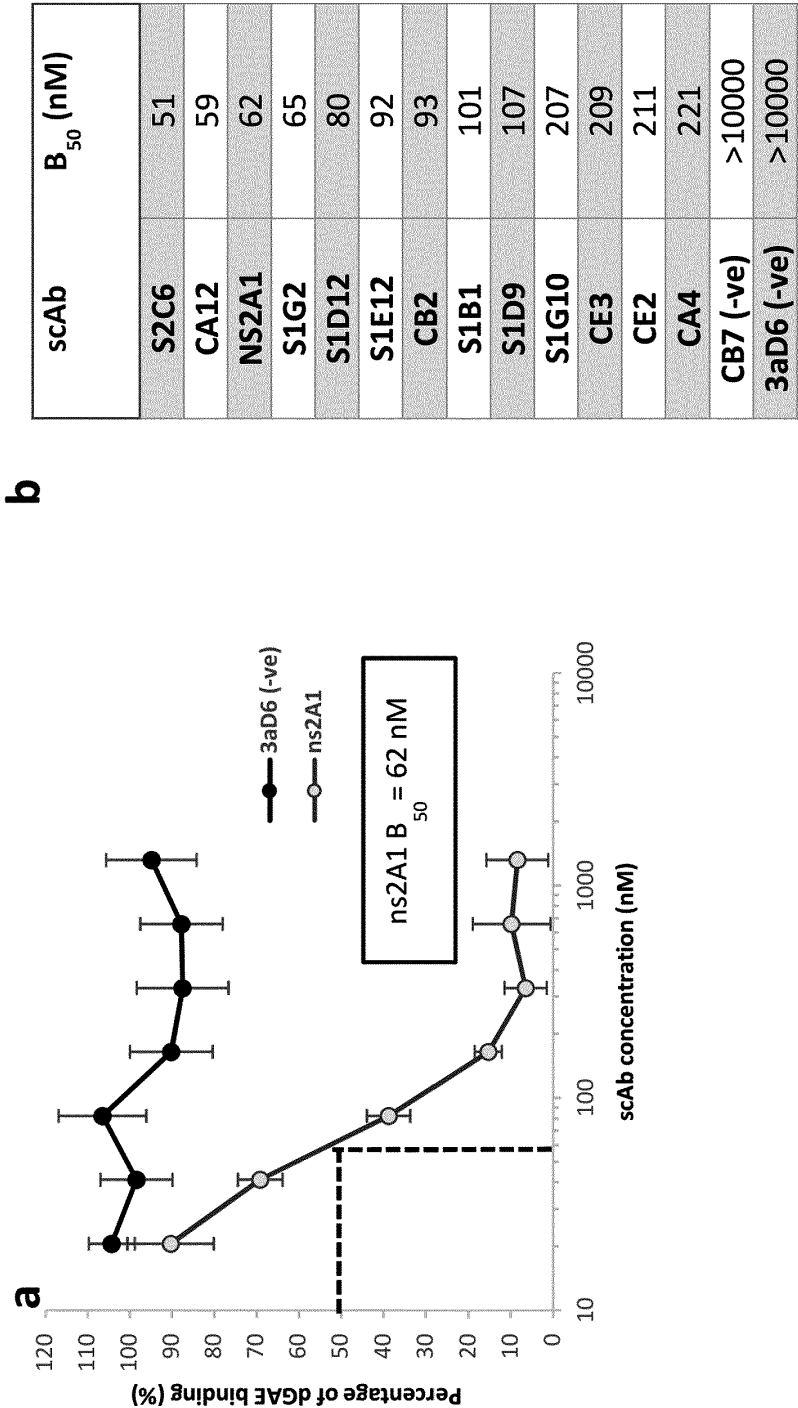


Fig. 38

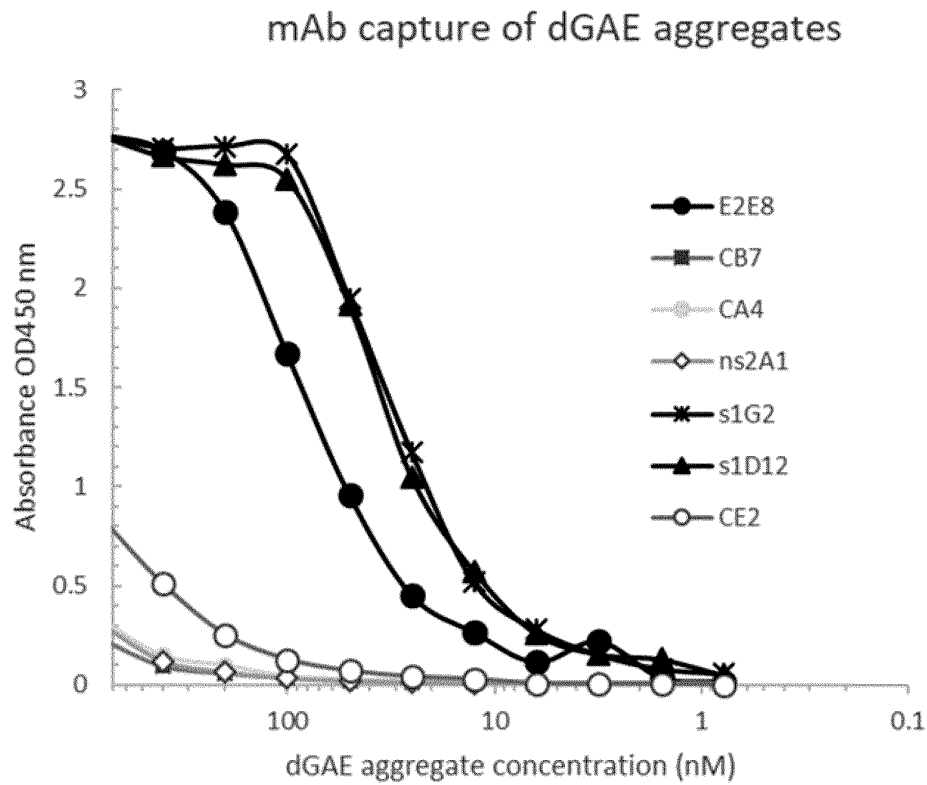


Fig. 39

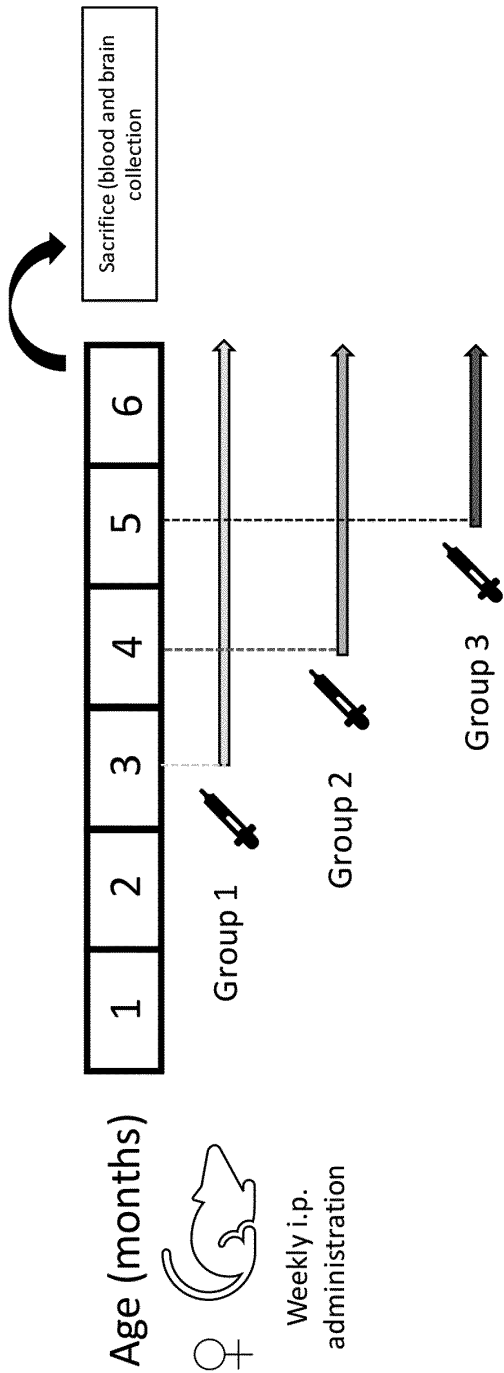


Fig. 40

51/64

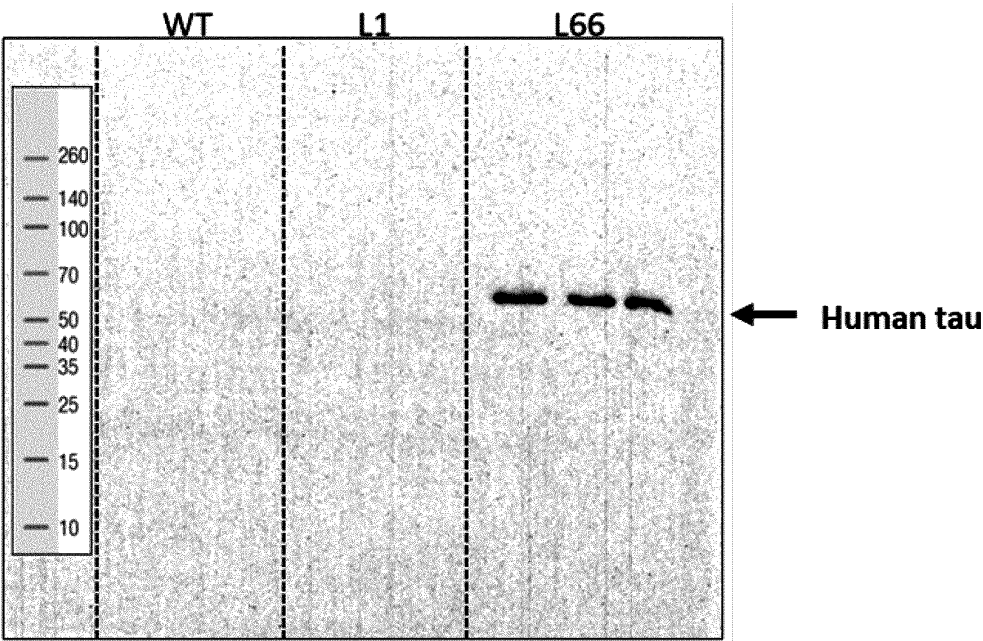


Fig. 41

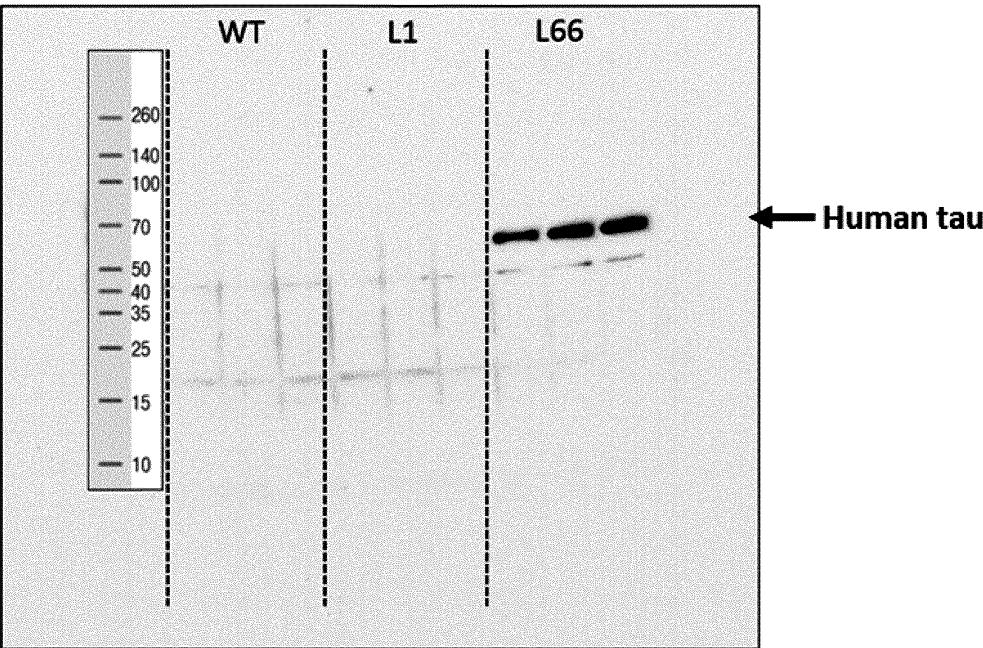


Fig. 42

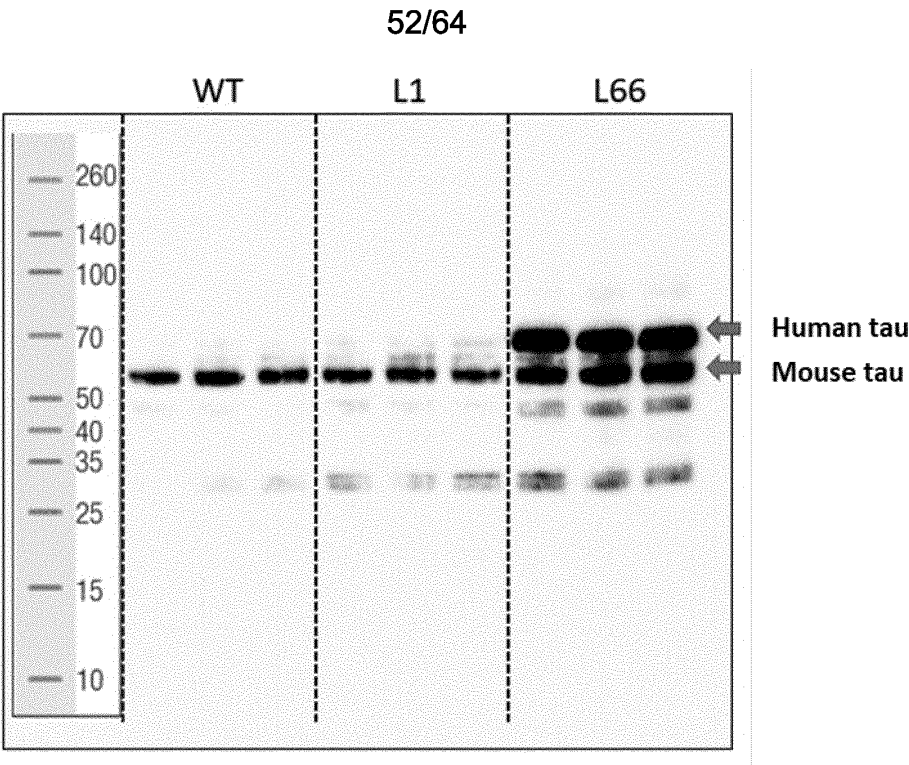


Fig. 43

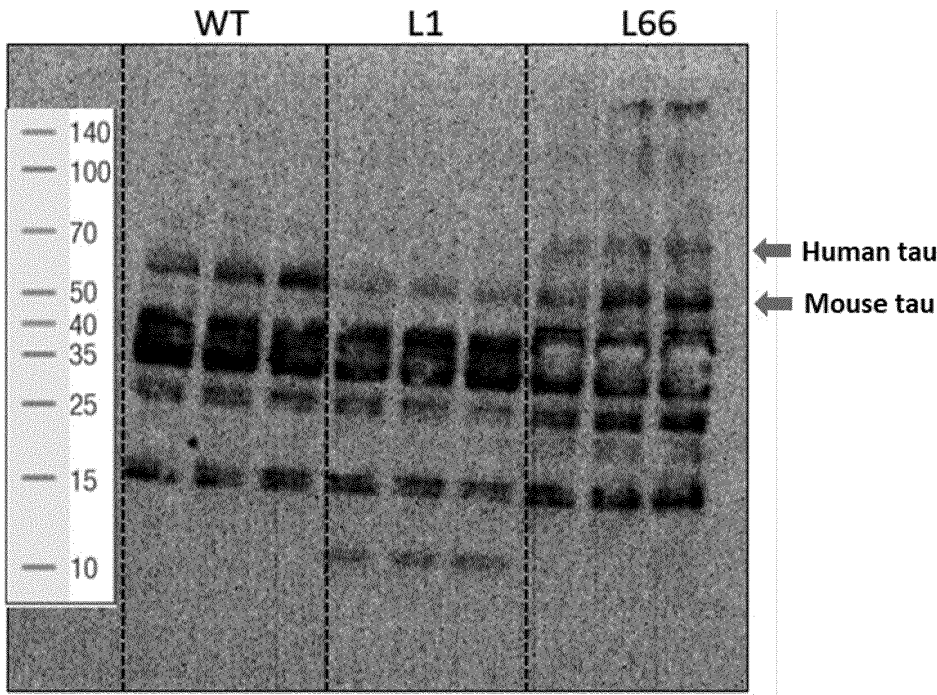


Fig. 44

Sequence alignment of human and mouse tau

```

CB7 (13-25)
P10636-8|TAU_HUMAN MAEPRQEFVEMDHAGTYGLGDRKQGGYTMHQDQEGDTDAGLKESPLQPTEDGSEEPG 60
P10637-2|TAU_MOUSE MADPRQEFDTMEDHAGDY-----TLLQDQEGDMHGLKESPPQPPADDGAEPEG 49
**::*****:..***** * **::***** * ***** * ***** * ***** * *****

P10636-8|TAU_HUMAN SETSDAKSTPTAEDVTAPLVDEGAPGKQAAQPHTEIPEGTTAEAGIGDTPSLEDEAAG 120
P10637-2|TAU_MOUSE SETSDAKSTPTAEDVTAPLVDERAPDKQAAQPHTEIPEGITAEAGIGDTPNQEDQAAG 109
*****

P10636-8|TAU_HUMAN HVTQARMVSKSDGTGSDDKKAKGADGK--TKIATPRGAAPPQKGQANATRIPAKTPPA 178
P10637-2|TAU_MOUSE HVTQARVASKD--RTGNDEKKAKGADGKTGAKIATPRGAASPAQKGTSNATRIPAKTTPS 167
*****:..** **::***** **::***** * ***** * ***** * *****

P10636-8|TAU_HUMAN PKTPPSSGEPKSGDRSGYSSPGSPGTPGSRSRTPSLPTPTPREPKKVAVRTPPKSPSS 238
P10637-2|TAU_MOUSE PKTPPGSGEPKSGERSGYSSPGSPGTPGSRSRTPSLPTPTPREPKKVAVRTPPKSPSA 227
*****.*****:*****:*****:*****:*****:*****:*****:*****:*****:

P10636-8|TAU_HUMAN AKSRLQTAPEVMPDLKNVSKIGSTENLKHQPGGKQVQIINKKLDLSNVQSKCGSKONIK 298
P10637-2|TAU_MOUSE SKSRLQTAPEVMPDLKNVRSKIGSTENLKHQPGGKQVQIINKKLDLSNVQSKCGSKONIK 287
:*****:*****:*****:*****:*****:*****:*****:*****:*****:

P10636-8|TAU_HUMAN HVPGGGSVQIVYKPVDSLKVTSKCGSLGNIHHKPGGGQVEVKSEKIDFQORVQSKIGSLD 358
P10637-2|TAU_MOUSE HVPGGGSVQIVYKPVDSLKVTSKCGSLGNIHHKPGGGQVEVKSEKIDFQORVQSKIGSLD 347
*****
S-1G2 (367-379)
P10636-8|TAU_HUMAN NITHVPGGGNKKIETHKLTFRENAKAKTDHGAEIVYKSPVSGDTSPRHLSNVSTGSID 418
P10637-2|TAU_MOUSE NITHVPGGGNKKIETHKLTFRENAKAKTDHGAEIVYKSPVSGDTSPRHLSNVSTGSID 407
*****

P10636-8|TAU_HUMAN MVDSPQLATLADSVSASLAKQGL 441
P10637-2|TAU_MOUSE MVDSPQLATLADSVSASLAKQGL 430

```

Fig. 45

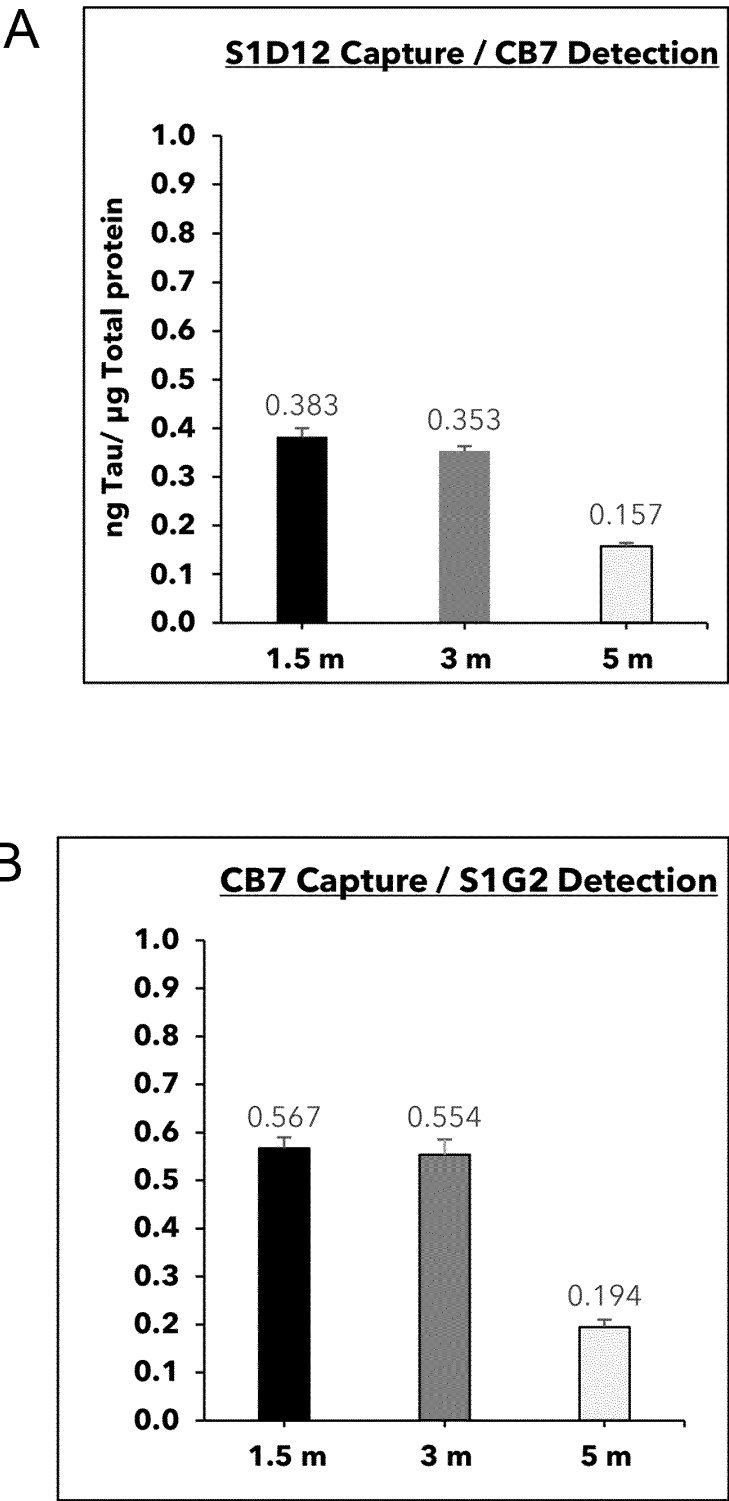
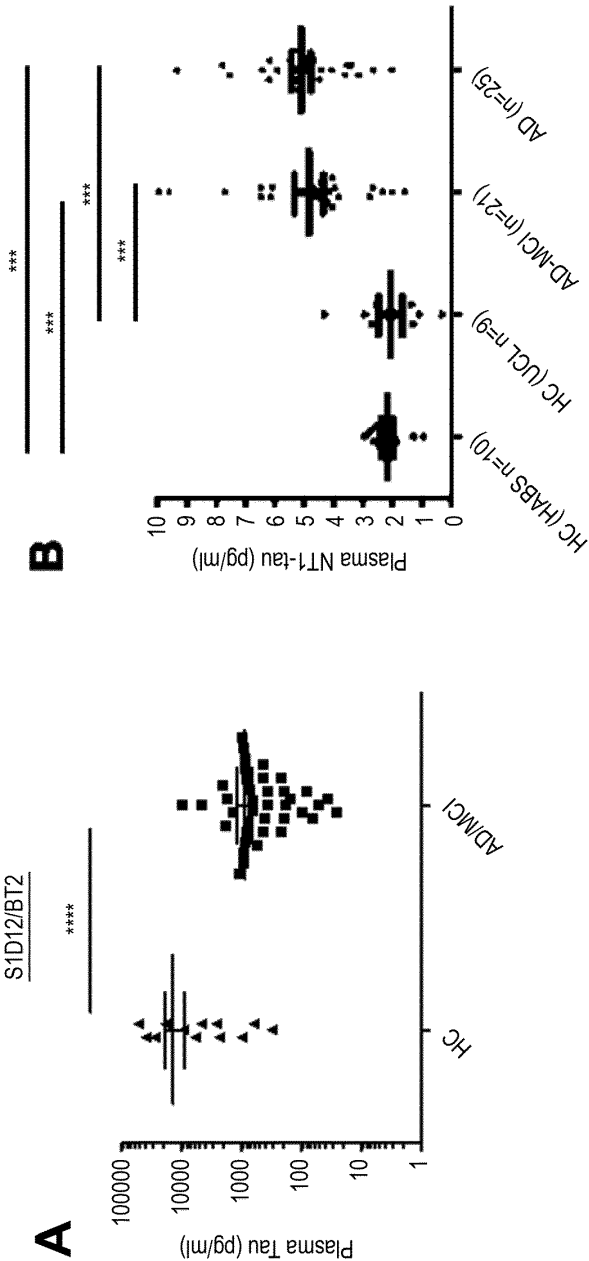
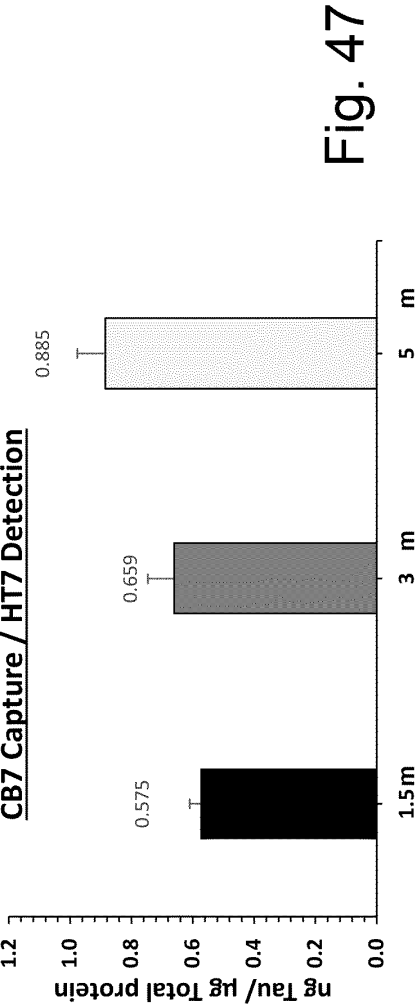


Fig. 46



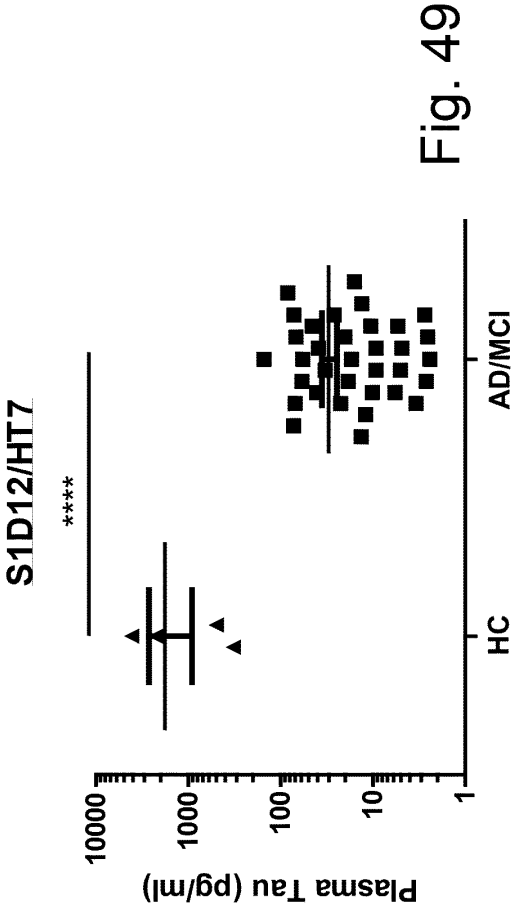


Fig. 49

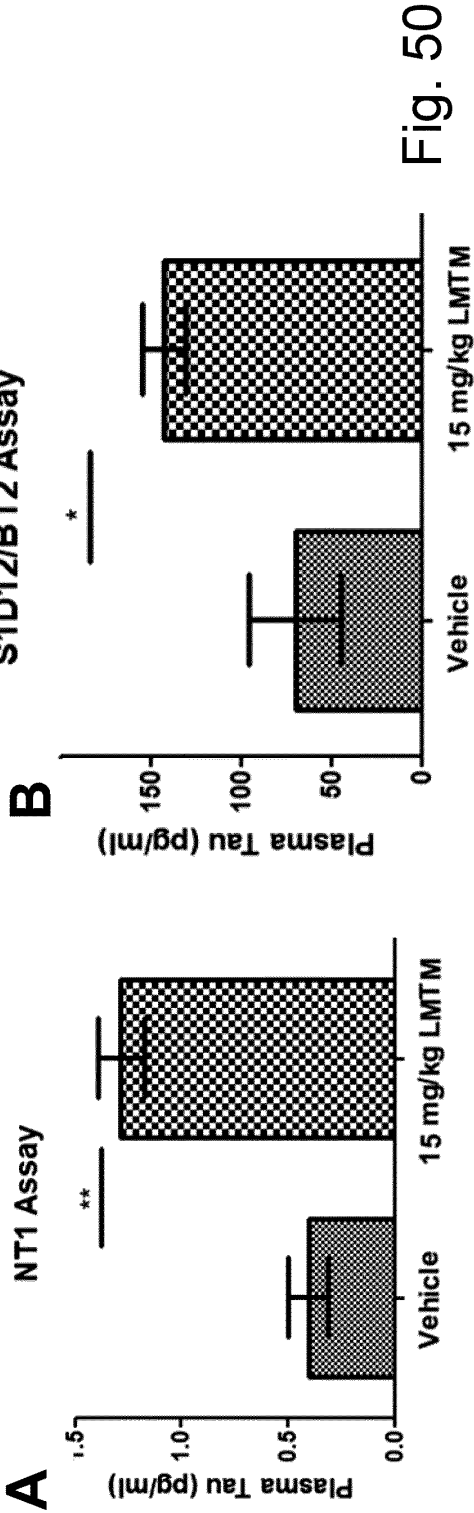


Fig. 50

57/64

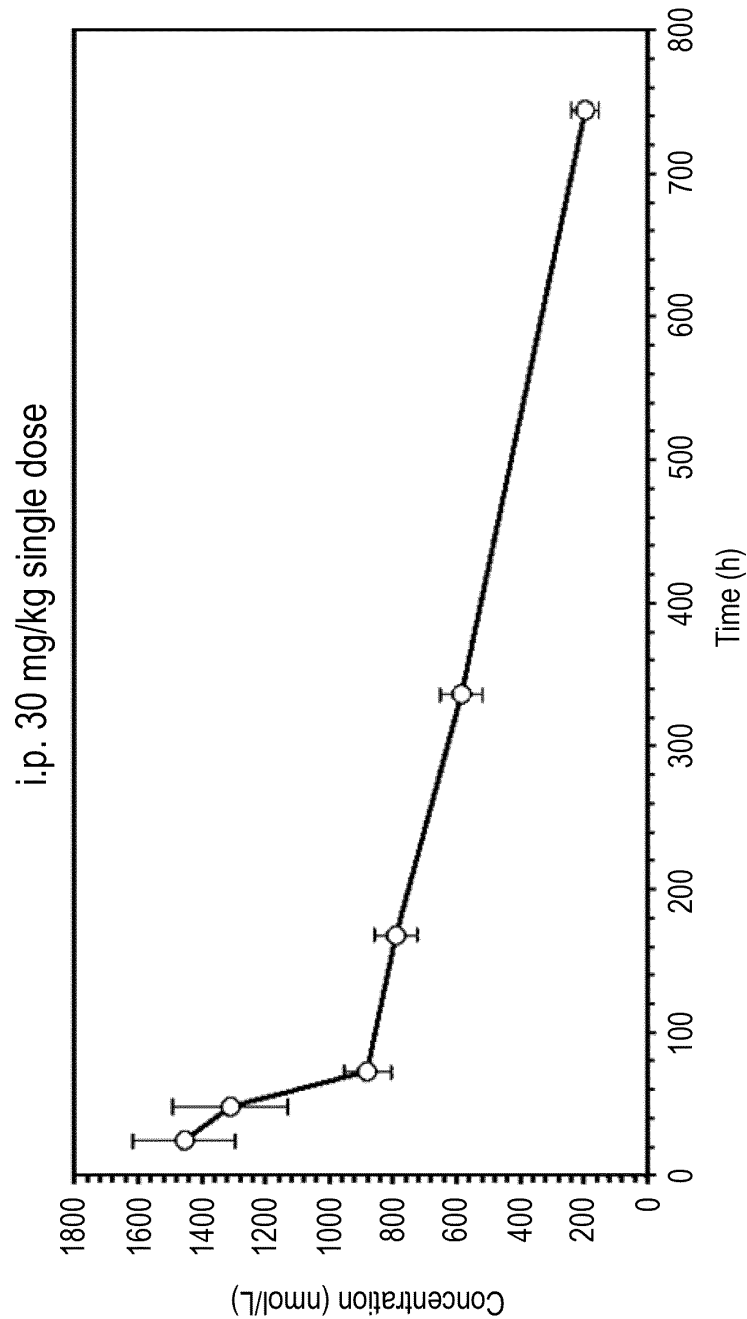


Fig. 51

58/64

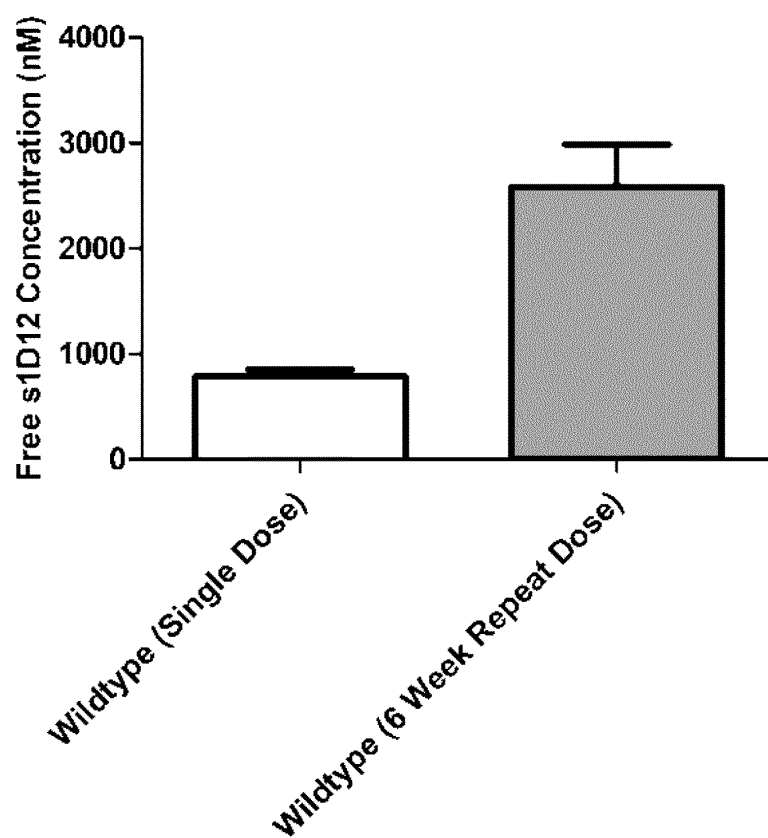


Fig. 52

59/64

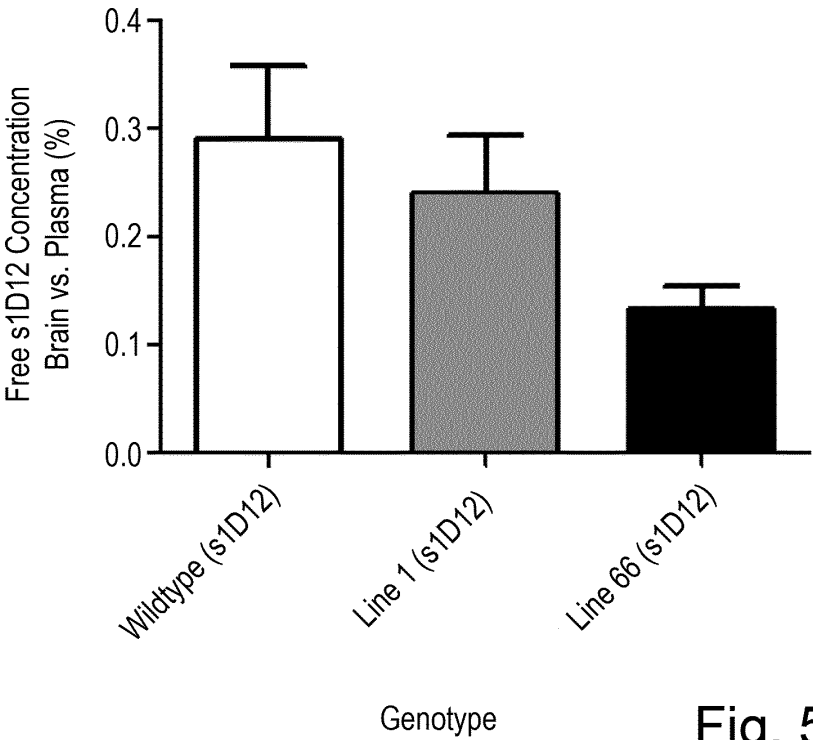


Fig. 53

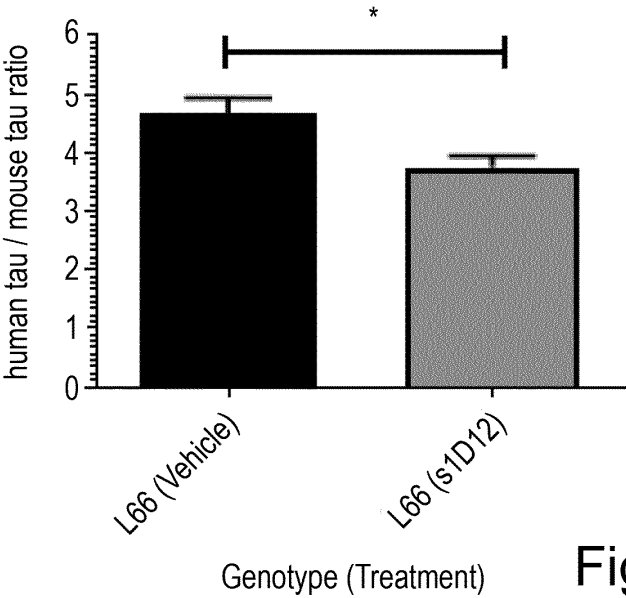


Fig. 54

60/64

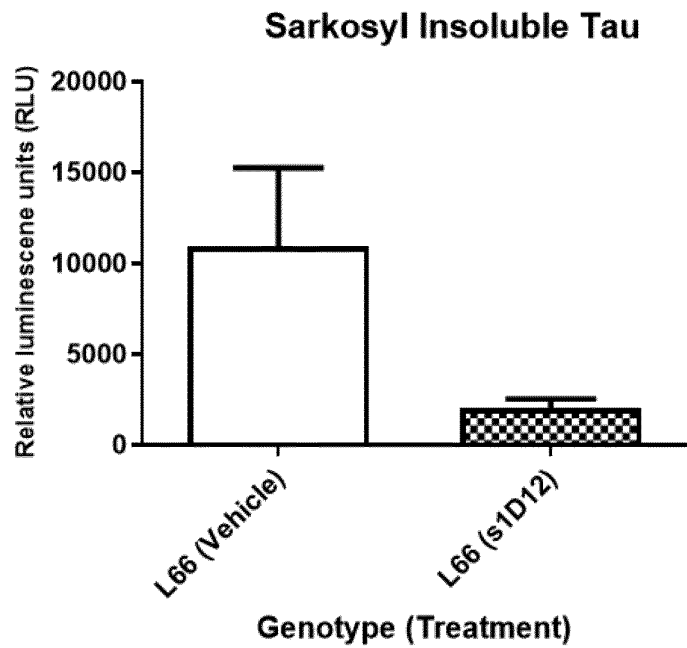


Fig. 55

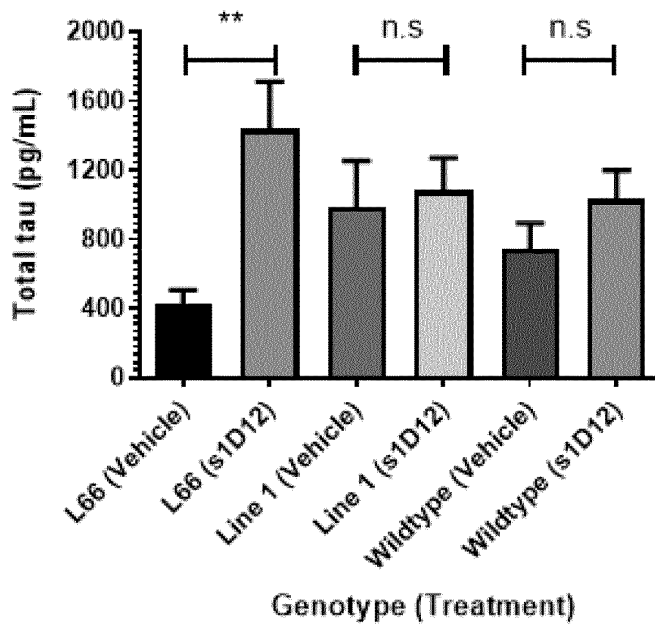


Fig. 56

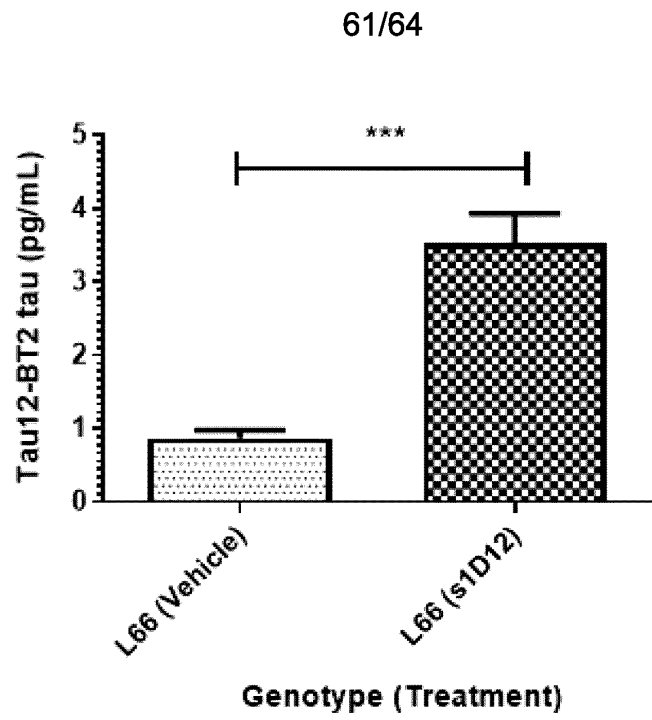


Fig. 57

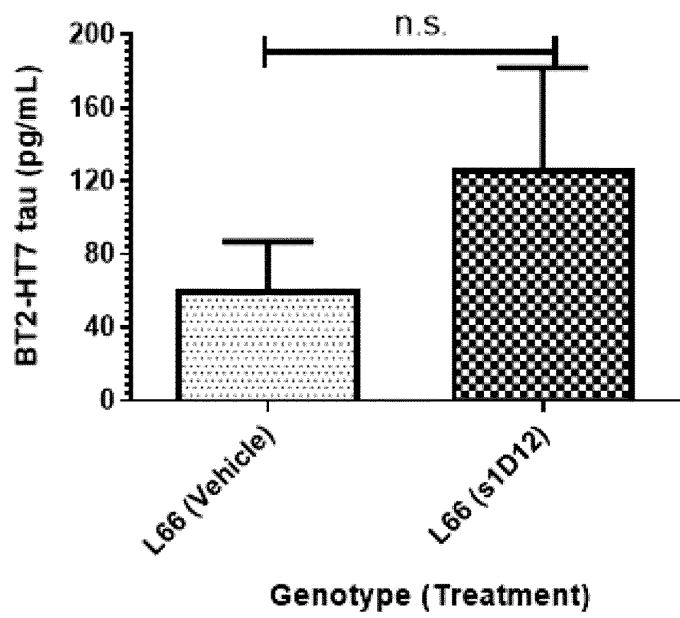


Fig. 58

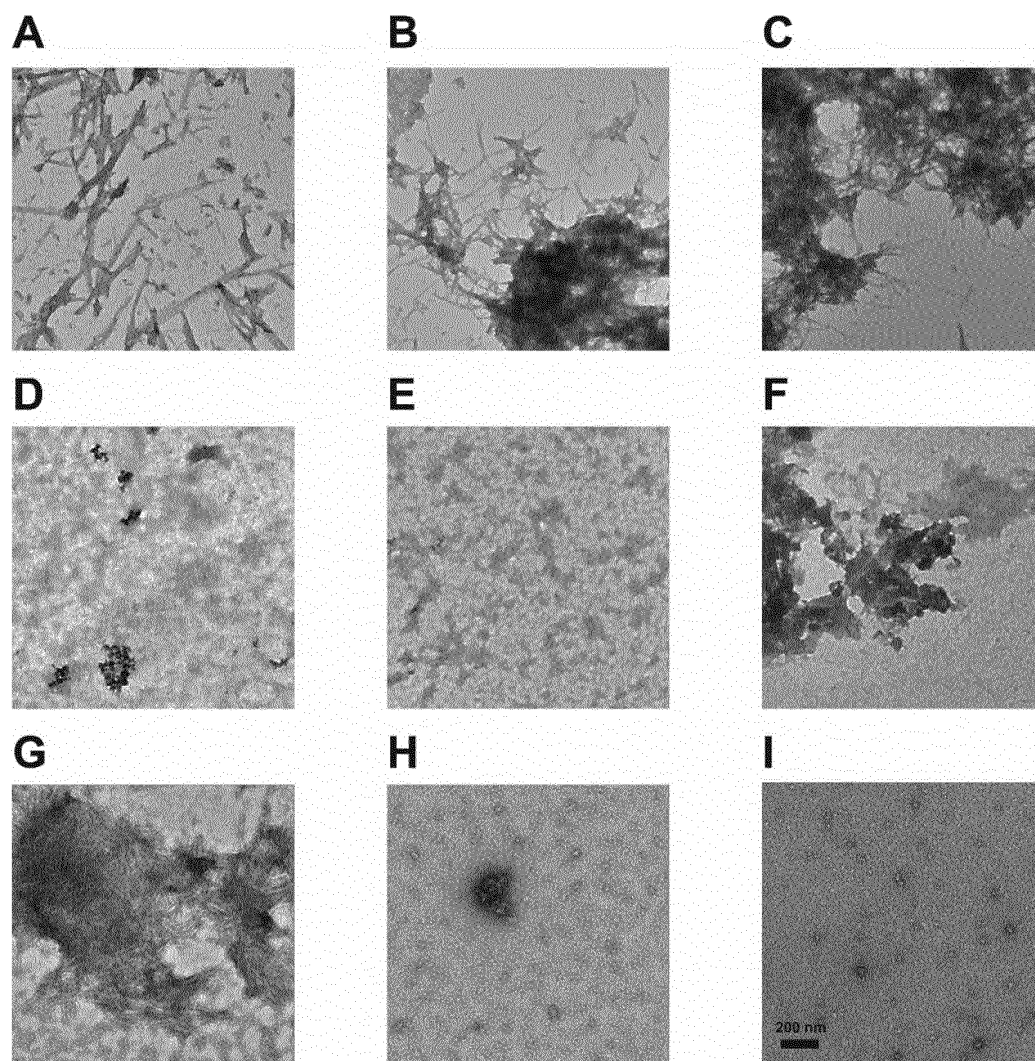


Fig. 59

63/64

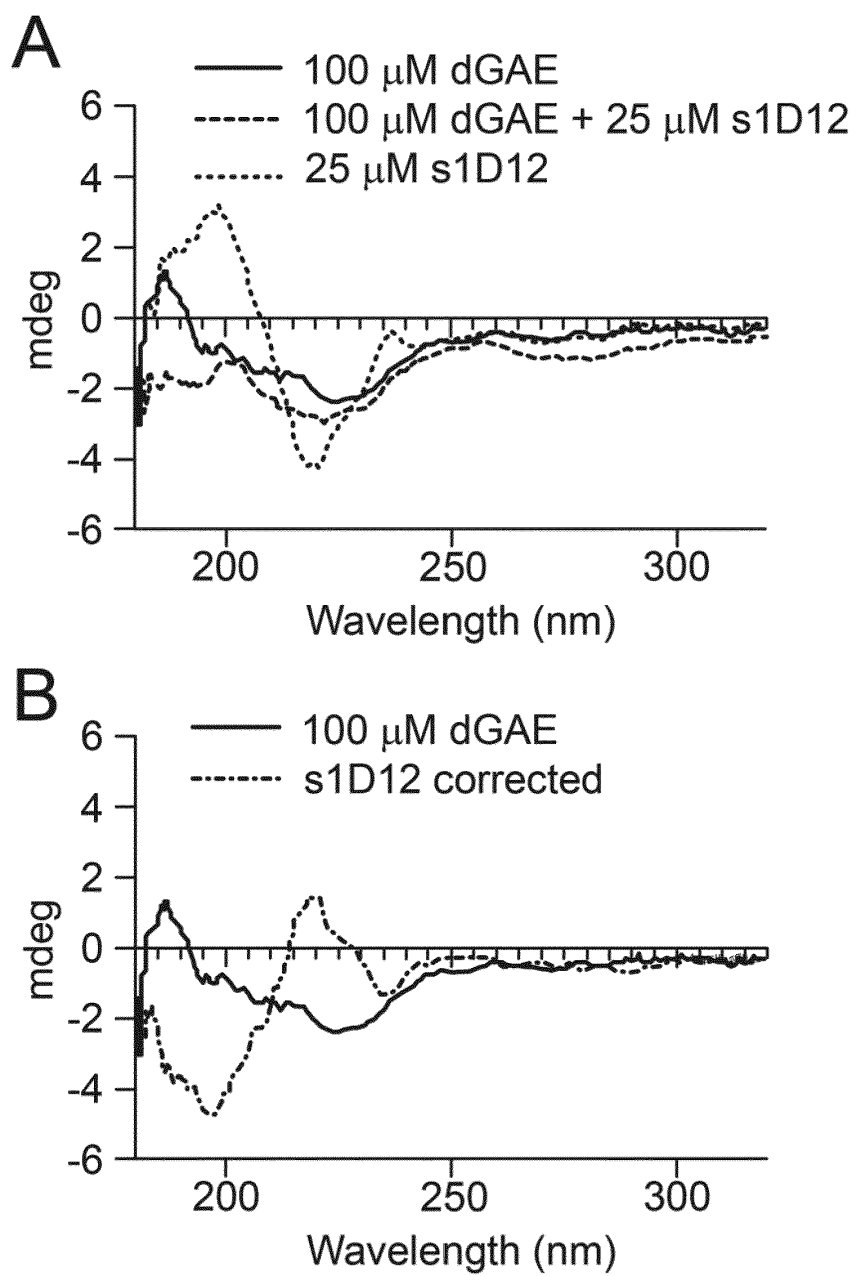


Fig. 60

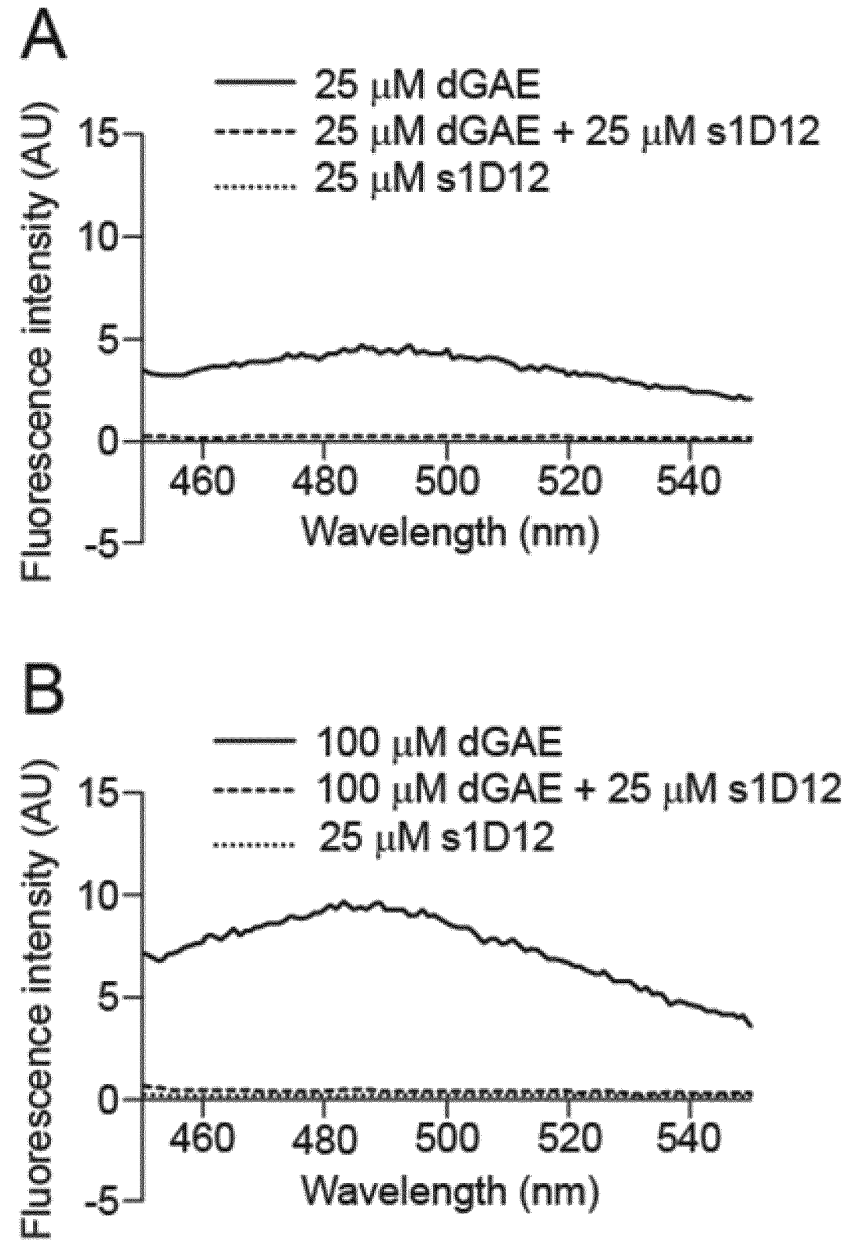


Fig. 61