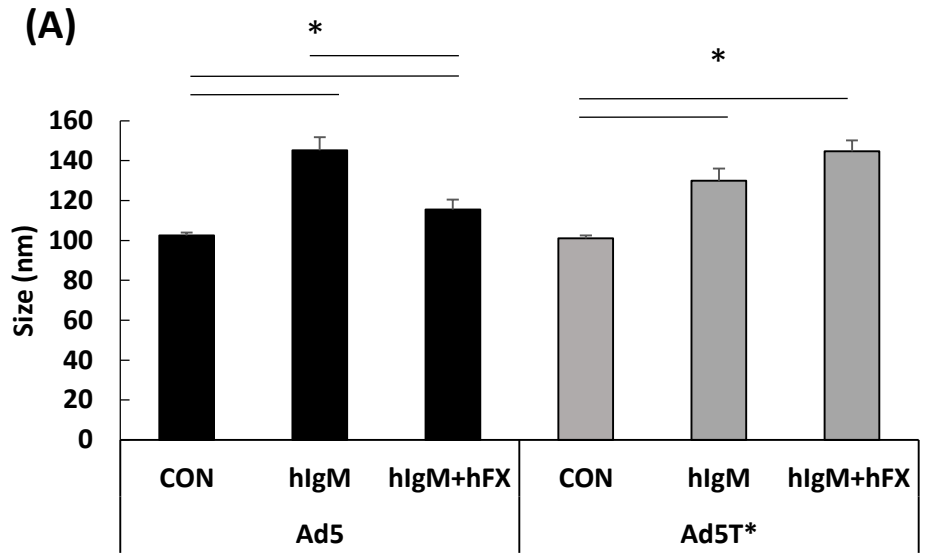
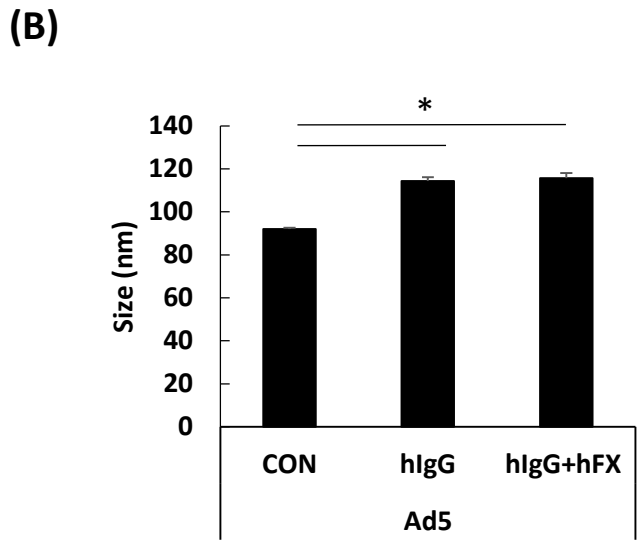
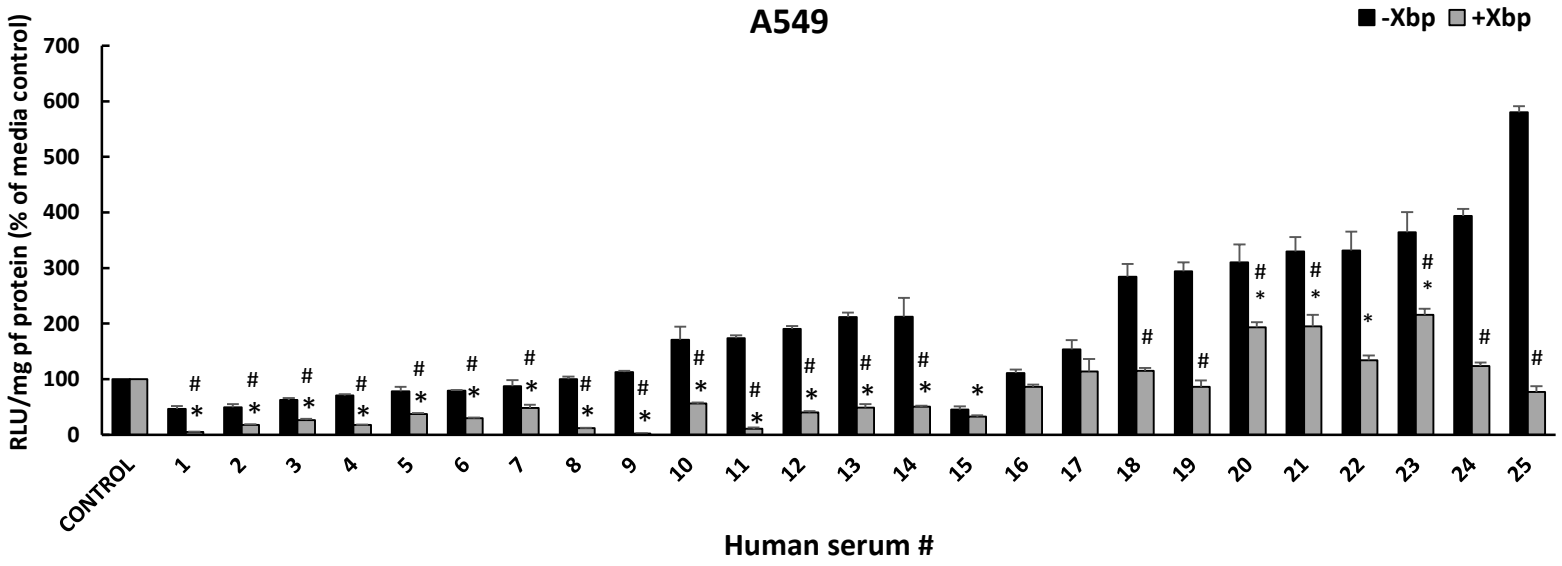
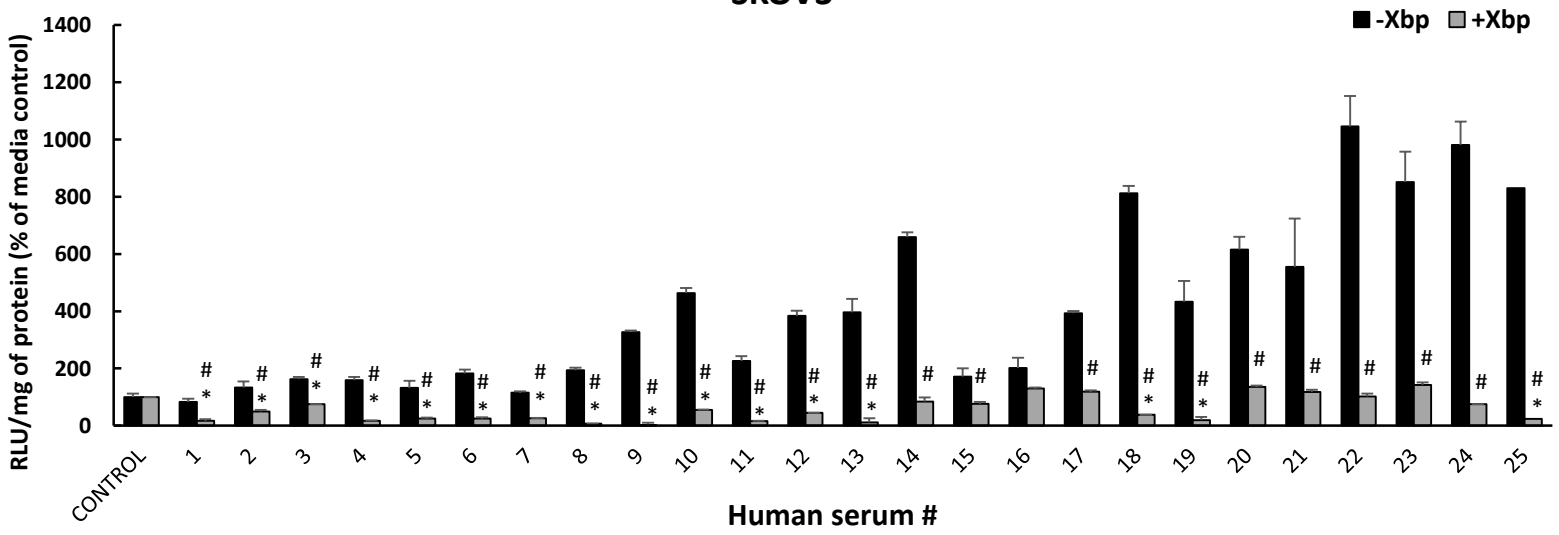

(A)
*
*
160
140
120
100
80
60
40
20
0
Size (nm)
CON
hIgM
hIgM+hFX
CON
hIgM
hIgM+hFX
Ad5
Ad5T*


(B)
*
140
120
100
80
60
40
20
0
Size (nm)
CON
hIgG
hIgG+hFX
Ad5

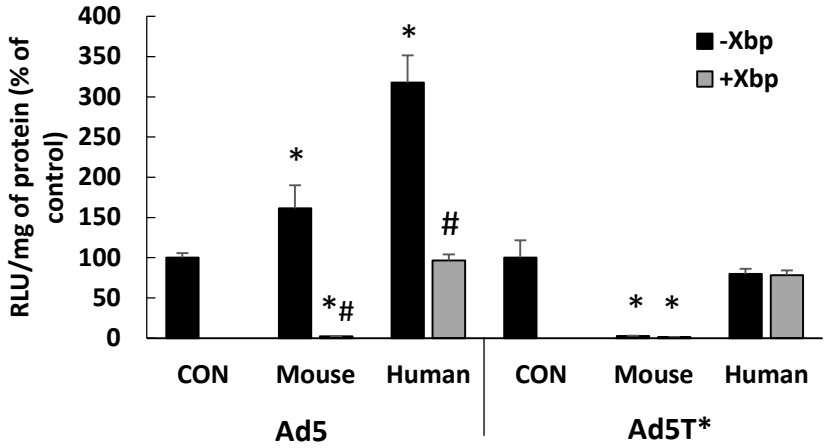
(A)

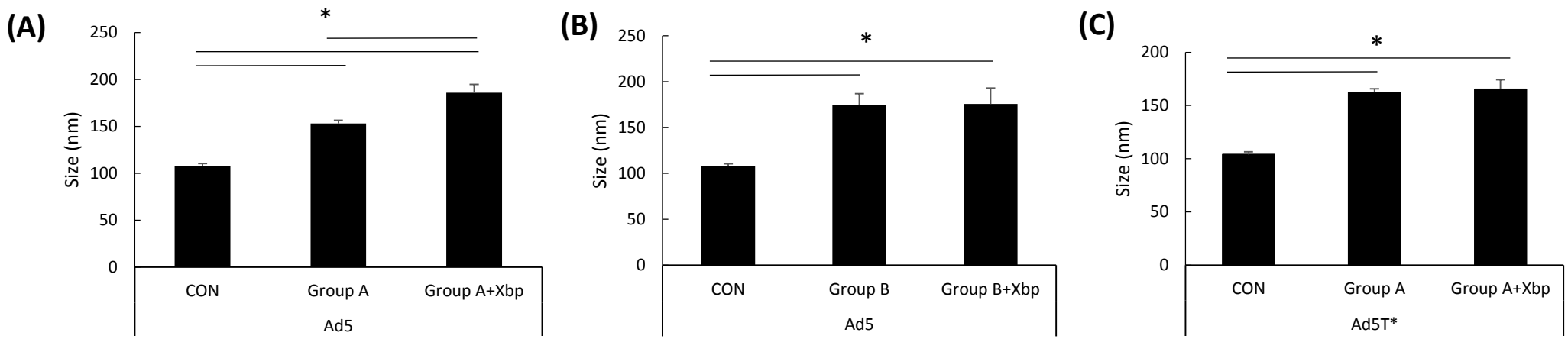

A549
-Xbp
+Xbp
RLU/mg pf protein (% of media control)
700
600
500
400
300
200
100
0
CONTROL
1
2
3
4
5
6
7
8
9
10
11
12
13
14
15
16
17
18
19
20
21
22
23
24
25
Human serum #

(B)

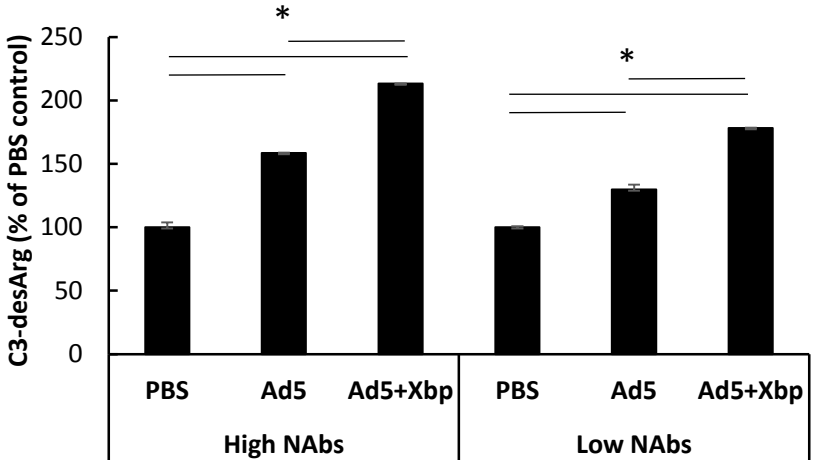

SKOV3
-Xbp
+Xbp
RLU/mg of protein (% of media control)
1400
1200
1000
800
600
400
200
0
CONTROL
1
2
3
4
5
6
7
8
9
10
11
12
13
14
15
16
17
18
19
20
21
22
23
24
25
Human serum #

(C)

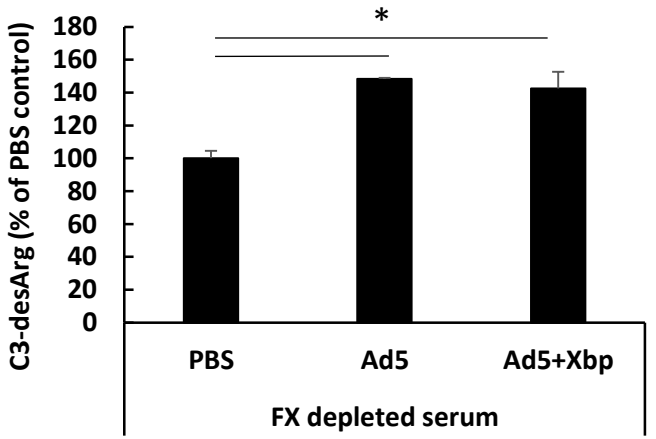

-Xbp
+Xbp
RLU/mg of protein (% of control)
400
350
300
250
200
150
100
50
0
CON
Mouse
Human
CON
Mouse
Human
Ad5
Ad5T*


(A)
*
250
200
150
100
50
0
Size (nm)
CON
Group A
Group A+Xbp
Ad5
(B)
*
250
200
150
100
50
0
Size (nm)
CON
Group B
Group B+Xbp
Ad5
(C)
*
200
150
100
50
0
Size (nm)
CON
Group A
Group A+Xbp
Ad5T*

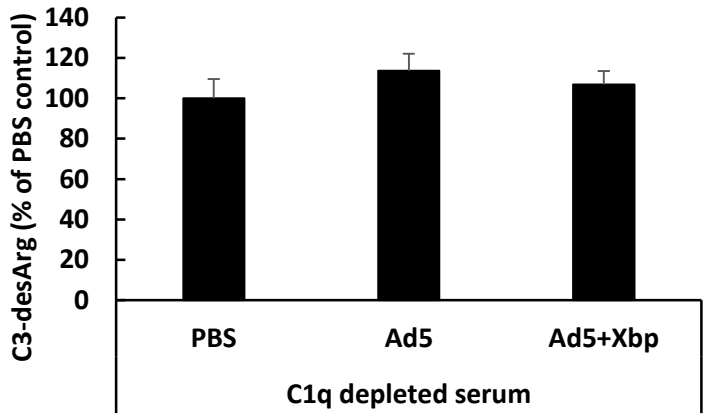
(A)

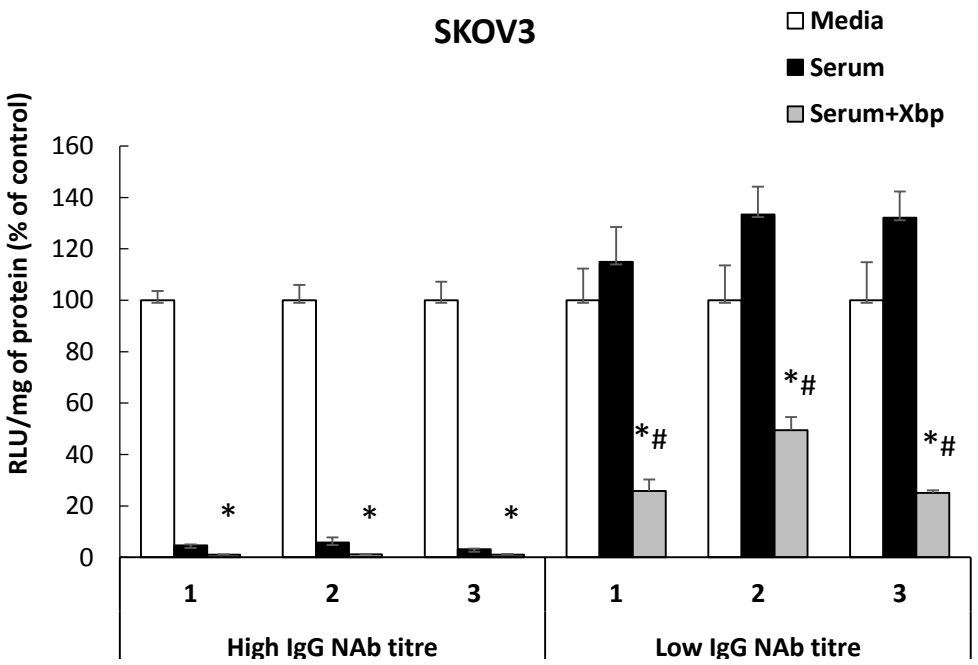

C3-desArg (% of PBS control)
250
200
150
100
50
0
*
*
PBS
Ad5
Ad5+Xbp
PBS
Ad5
Ad5+Xbp
High NAbs
Low NAbs

(B)


C3-desArg (% of PBS control)
180
160
140
120
100
80
60
40
20
0
*
PBS
Ad5
Ad5+Xbp
FX depleted serum

(C)


C3-desArg (% of PBS control)
140
120
100
80
60
40
20
0
PBS
Ad5
Ad5+Xbp
C1q depleted serum



Supplementary Figure 1. Ad5 ($2x10^{10}$ vp/mL) was incubated with media, human serum sample -/+ 40 µg/mL Xbp for 30 min at 37°C. Virus suspensions were diluted 200-fold in serum-free media and 100 µL added to cells for 2 h at 37°C, then replaced with media with 2% FCS. Transgene expression was quantified ~16 h post-transduction and relative light units (RLU) were normalized to mg total protein. The human serum samples were divided into those with high pre-existing neutralising IgG titres (high NAbs) and sera samples with low IgG titres which exhibited a dependence on FX for protection (low NAbs Group A). Samples were pooled accordingly and tested in the C3a ELISA as shown in Figure 4A . Media control (*$p<0.05$) or matched serum –Xbp conditions (#$p<0.05$) versus serum+Xbp .