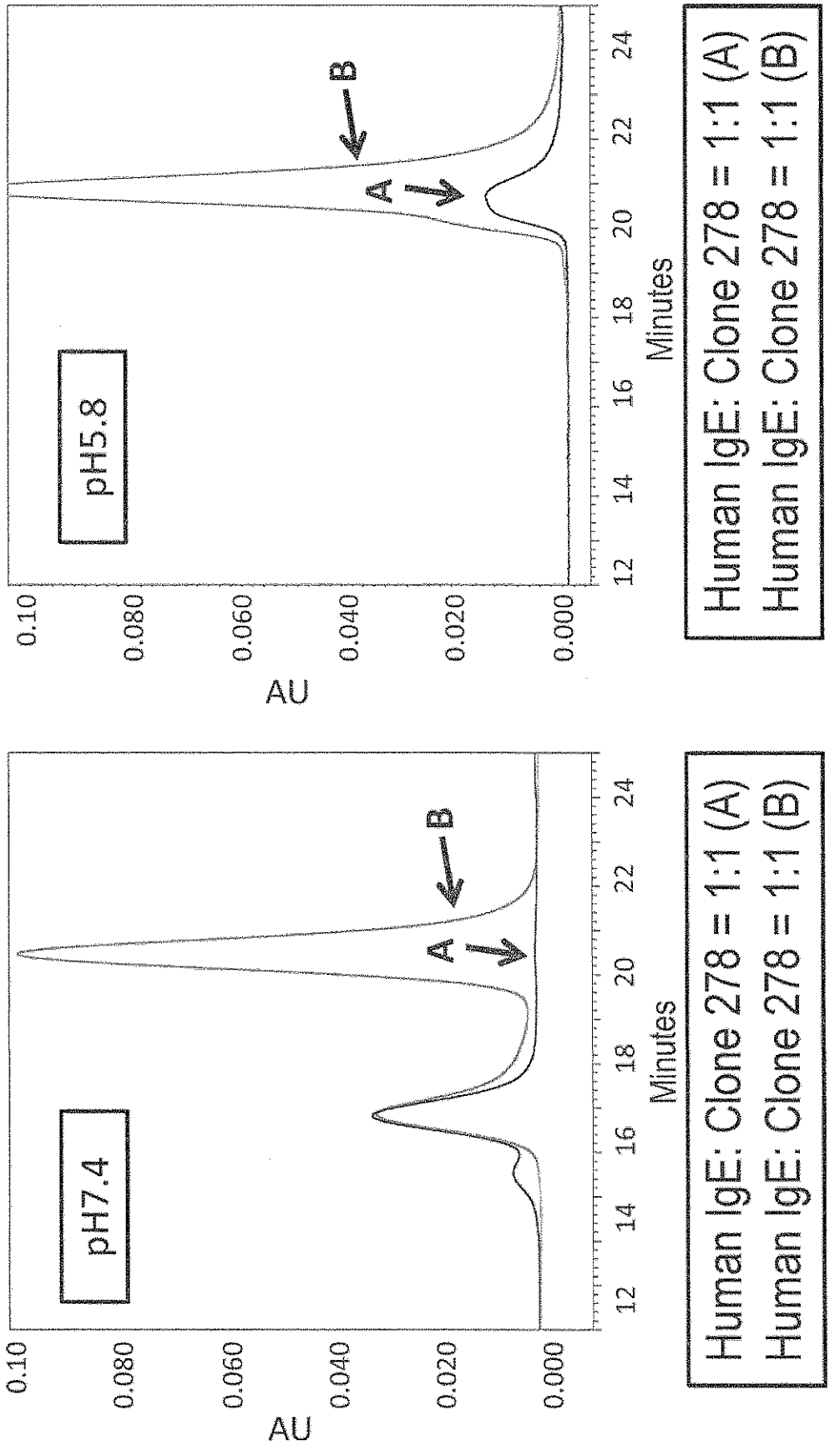

pH5.8
pH7.4
AU
Minutes
A
B
0.10
0.080
0.060
0.040
0.020
0.000
12
14
16
18
20
22
24
Human IgE: Clone 278 = 1:1 (A)
Human IgE: Clone 278 = 1:1 (B)
Human IgE: Clone 278 = 1:1 (A)
Human IgE: Clone 278 = 1:1 (B)

FIG. 1

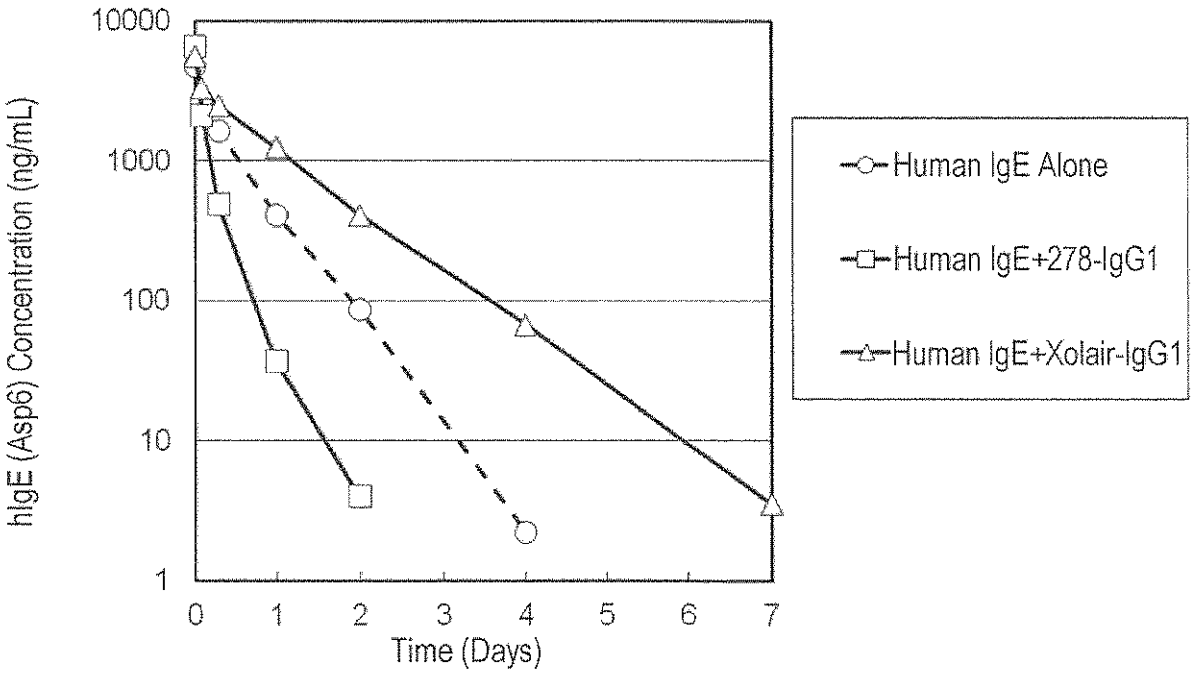

10000
1000
100
10
1
hIgE (Asp6) Concentration (ng/mL)
0
1
2
3
4
5
6
7
Time (Days)
Human IgE Alone
Human IgE+278-IgG1
Human IgE+Xolair-IgG1

FIG. 2

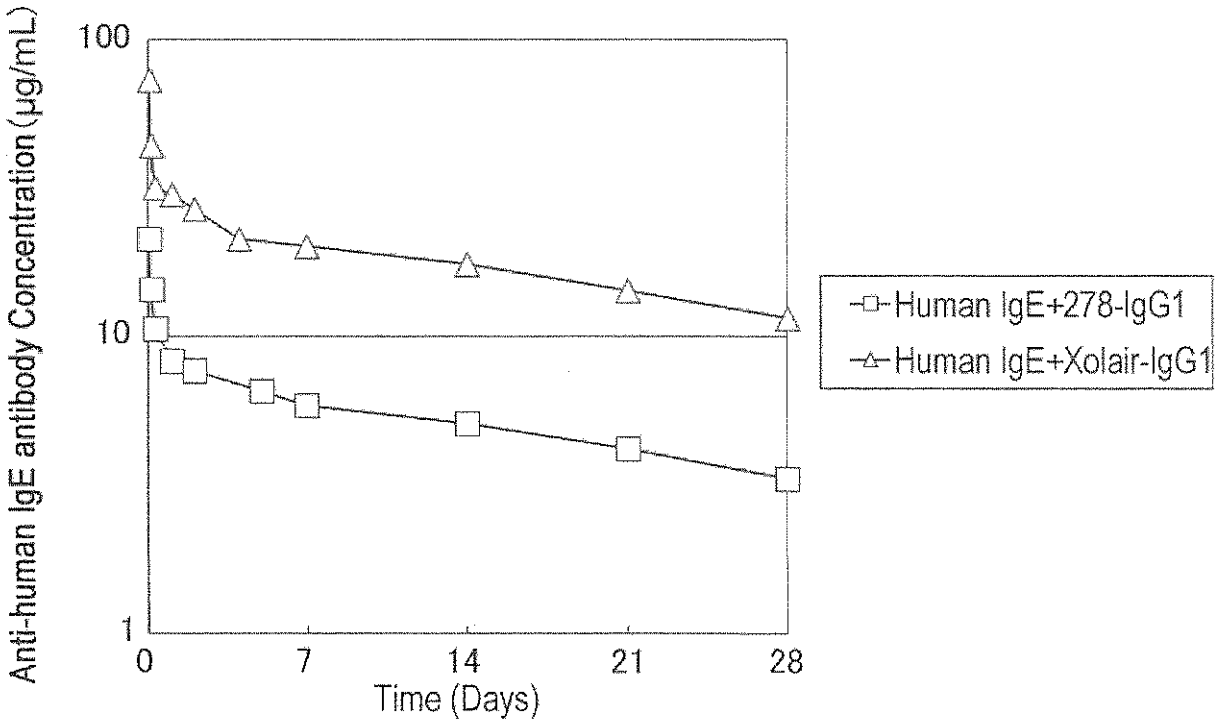

Anti-human IgE antibody Concentration (μg/mL)
100
10
1
0
7
14
21
28
Time (Days)
Human IgE+278-IgG1
Human IgE+Xolair-IgG1

FIG. 3

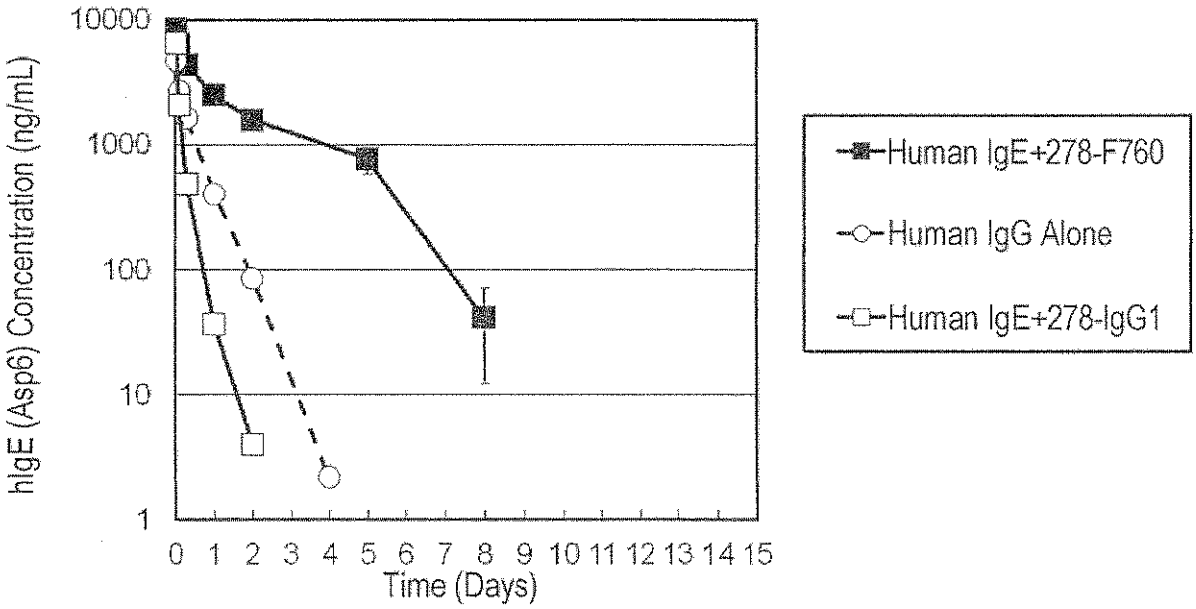

hIgE (Asp6) Concentration (ng/mL)
10000
1000
100
10
1
0 1 2 3 4 5 6 7 8 9 10 11 12 13 14 15
Time (Days)
Human IgE+278-F760
Human IgG Alone
Human IgE+278-IgG1

FIG. 4

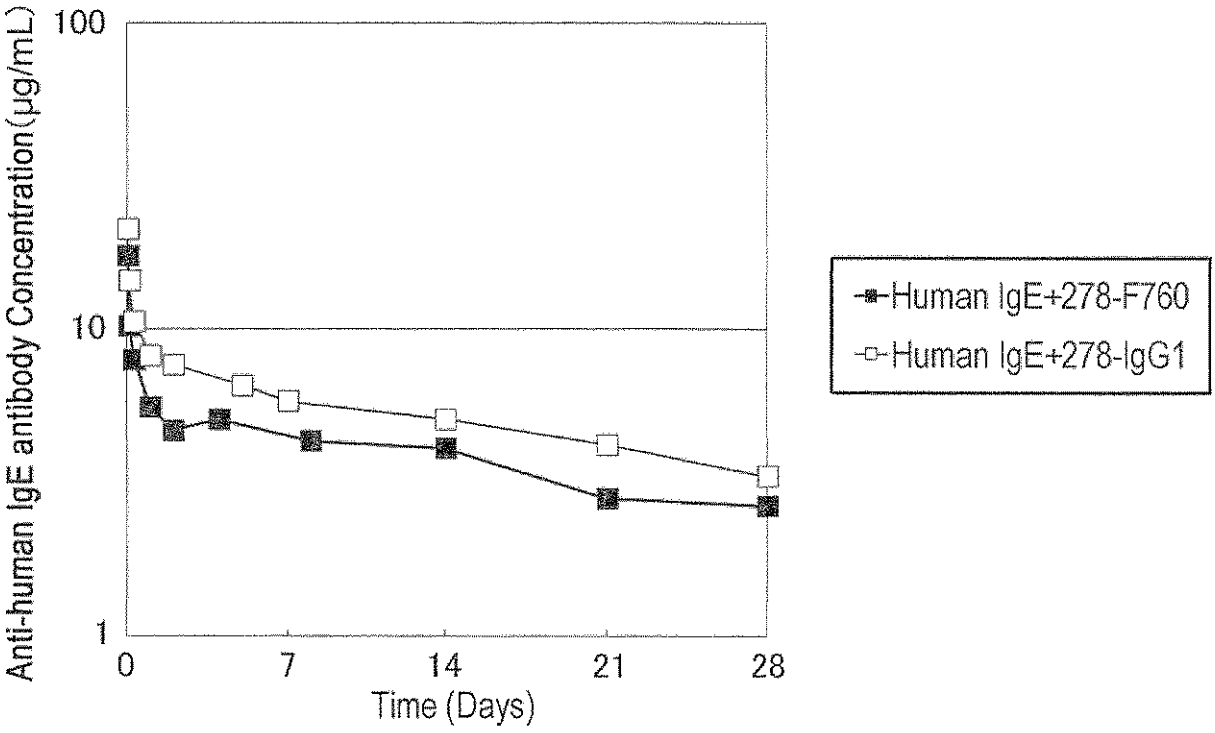

Anti-human IgE antibody Concentration(μg/mL)
100
10
1
0
7
14
21
28
Time (Days)
Human IgE+278-F760
Human IgE+278-IgG1

FIG. 5

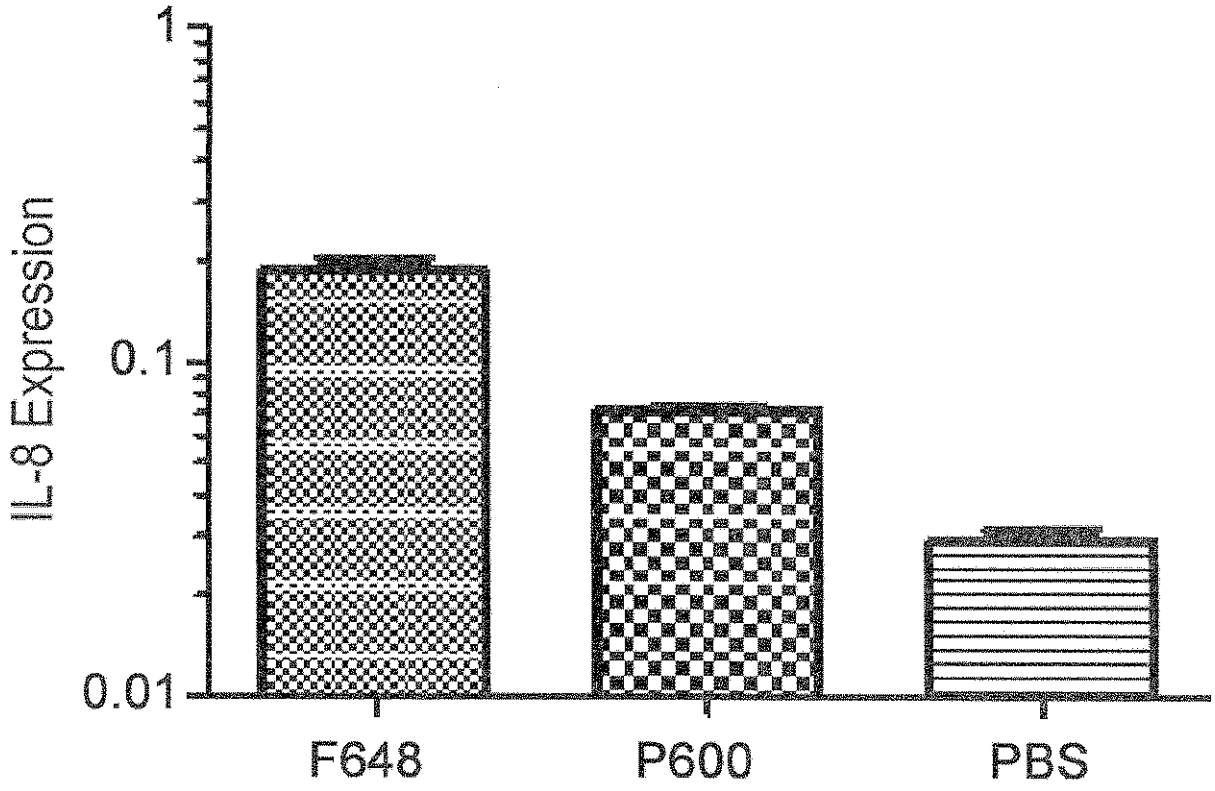


FIG. 6

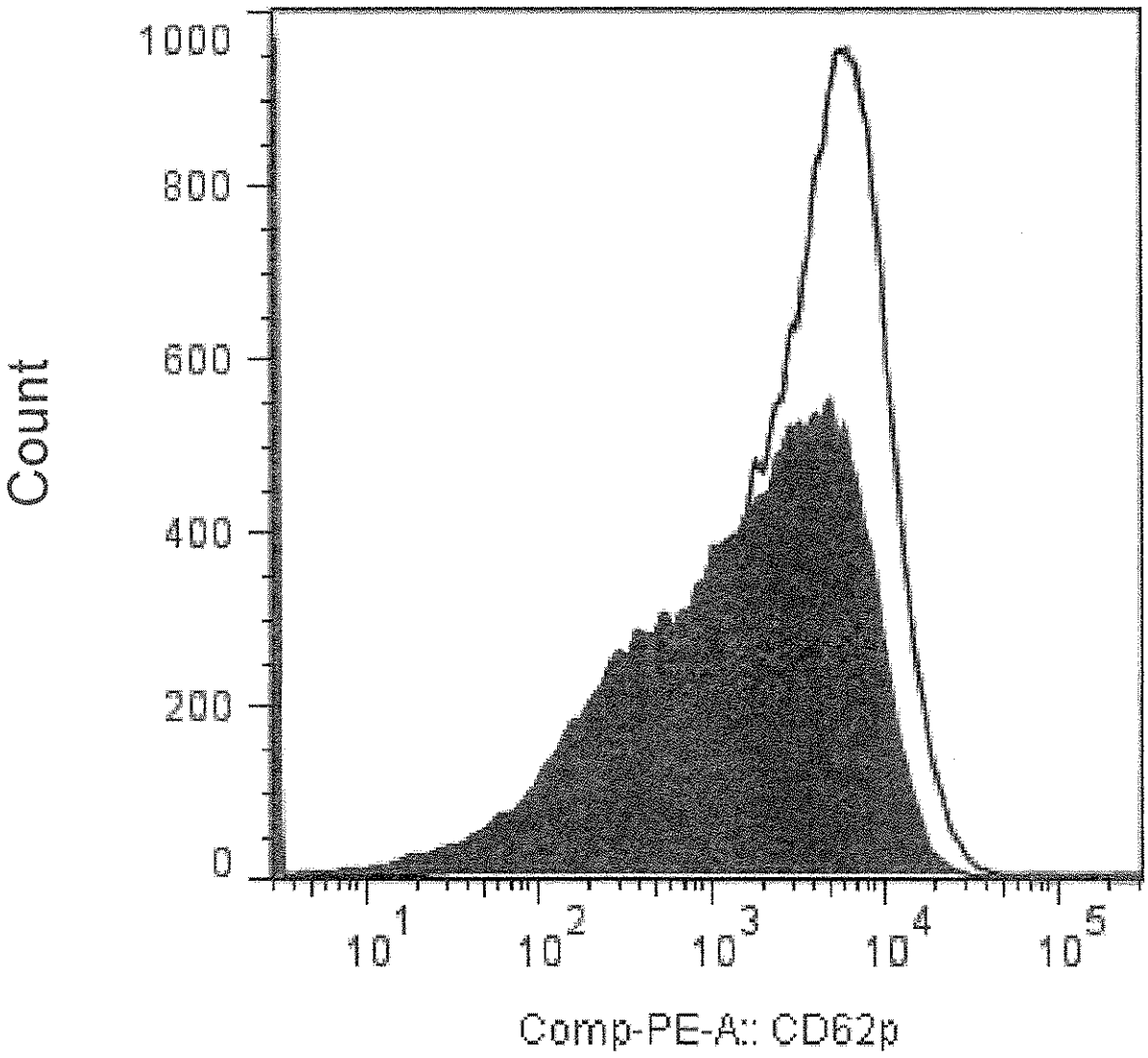


FIG. 7

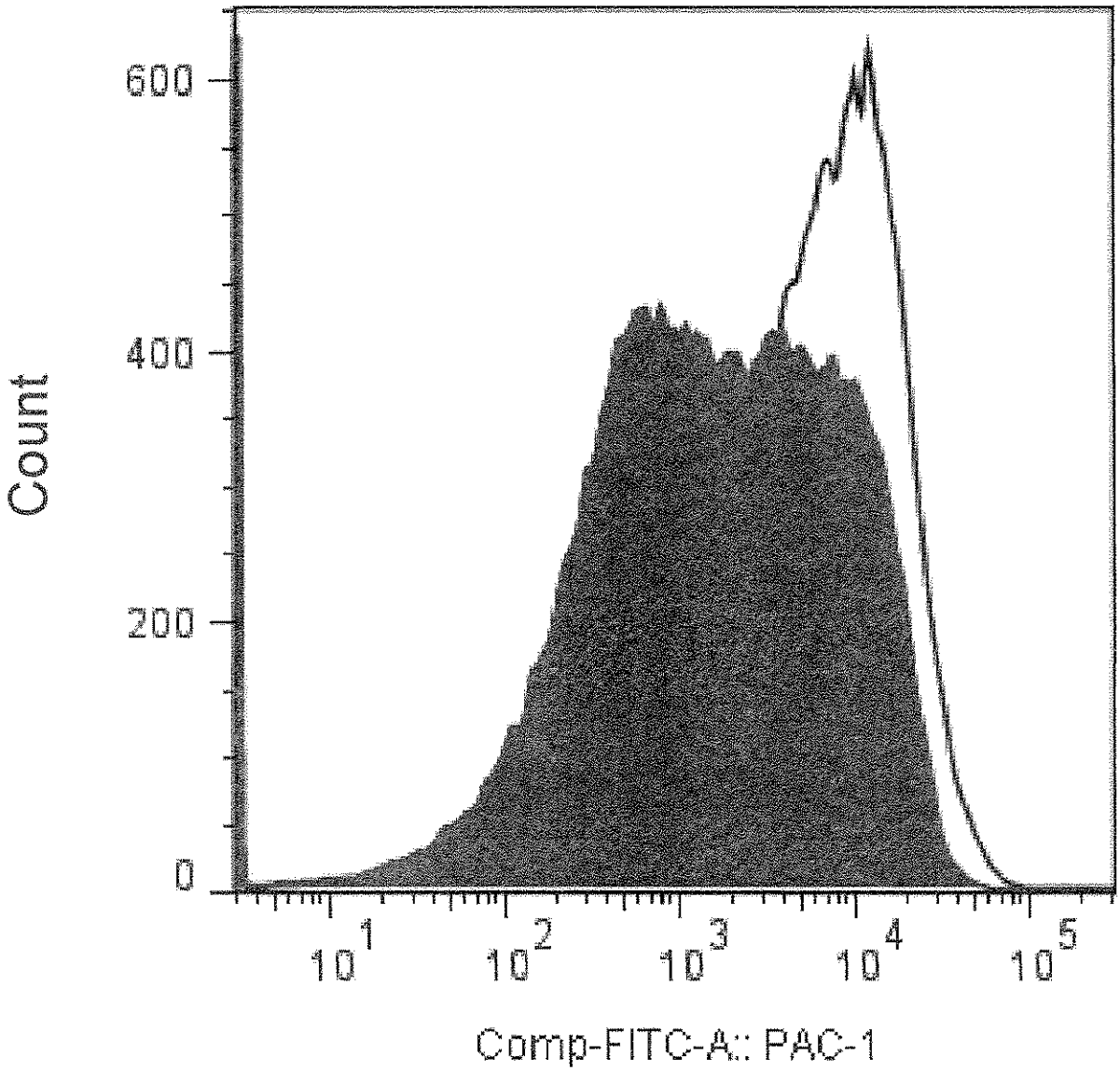


FIG. 8

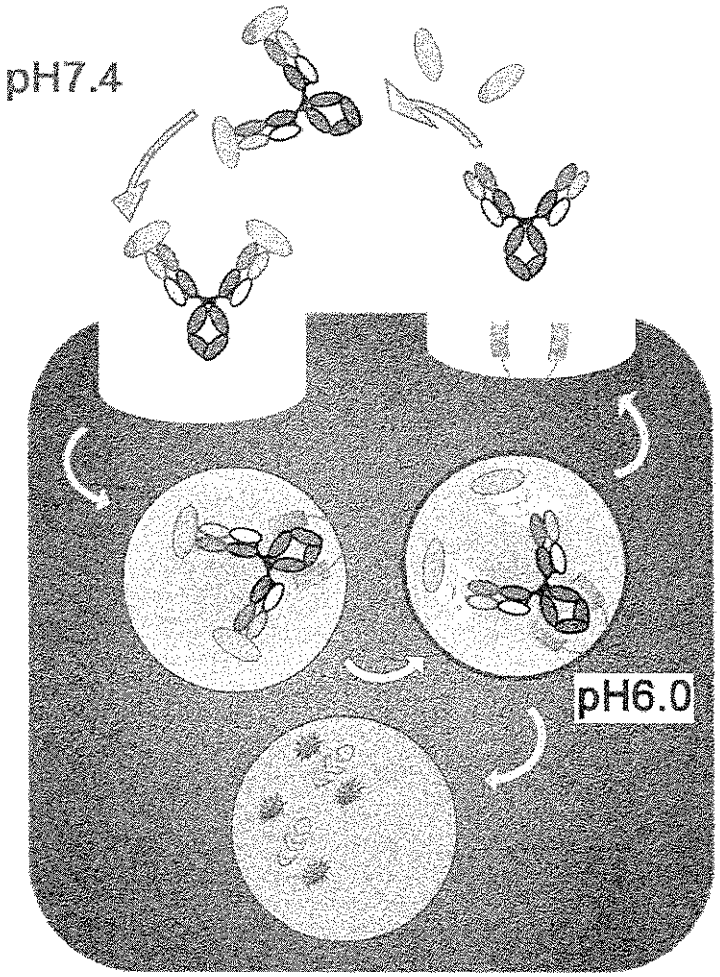


FIG. 9

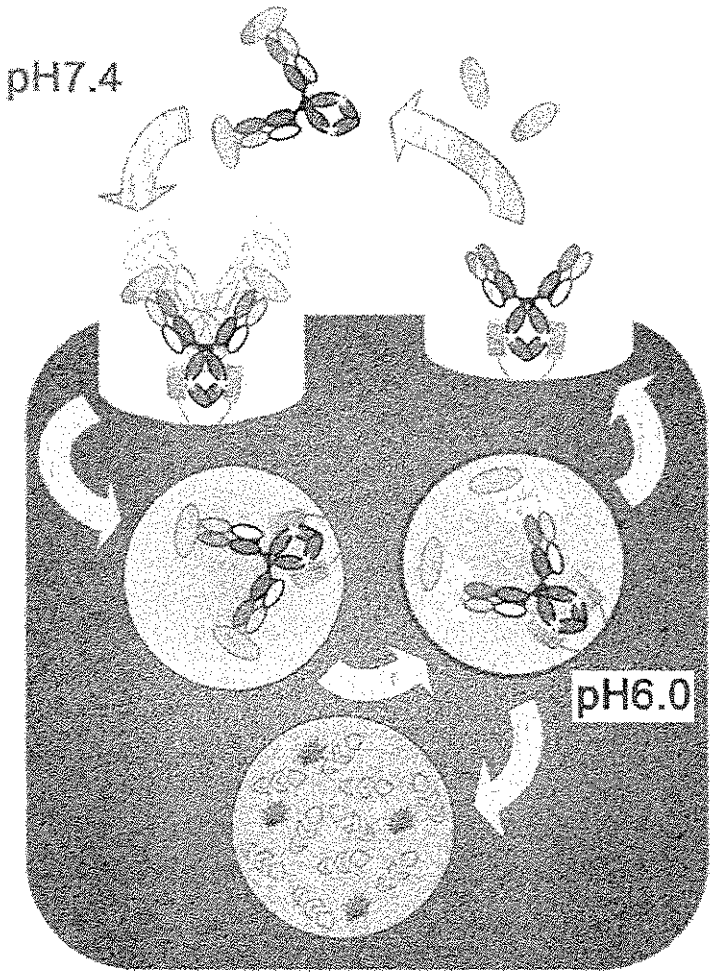


FIG. 10

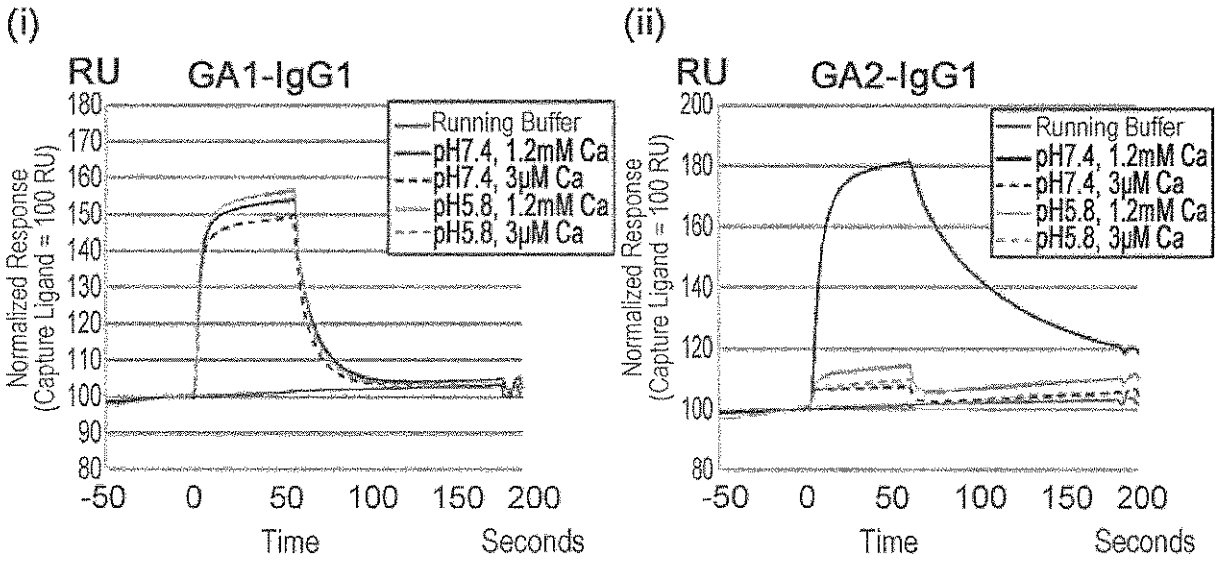

(i)
RU
GA1-IgG1
Normalized Response
(Capture Ligand = 100 RU)
180
170
160
150
140
130
120
110
100
90
80
-50
0
50
100
150
200
Time
Seconds
Running Buffer
pH7.4, 1.2mM Ca
pH7.4, 3μM Ca
pH5.8, 1.2mM Ca
pH5.8, 3μM Ca
(ii)
RU
GA2-IgG1
Normalized Response
(Capture Ligand = 100 RU)
200
180
160
140
120
100
80
-50
0
50
100
150
200
Time
Seconds
Running Buffer
pH7.4, 1.2mM Ca
pH7.4, 3μM Ca
pH5.8, 1.2mM Ca
pH5.8, 3μM Ca

FIG. 11

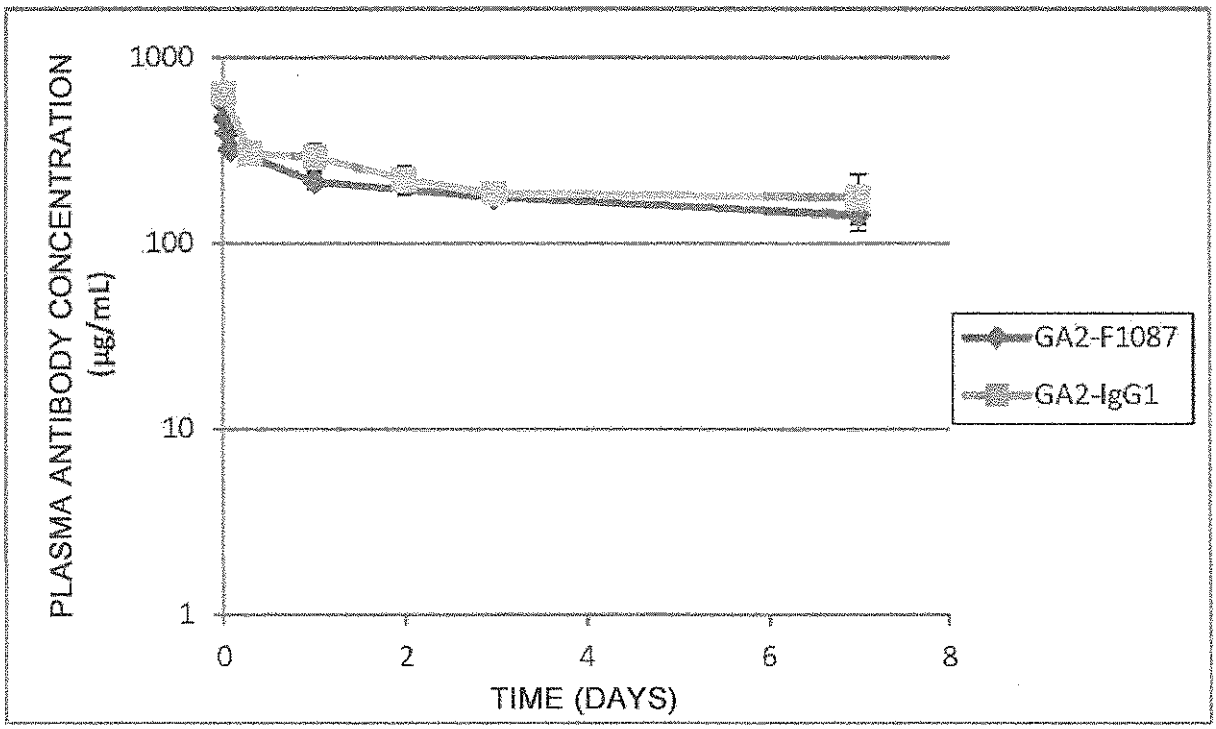

PLASMA ANTIBODY CONCENTRATION (µg/mL)
1000
100
10
1
0
2
4
6
8
TIME (DAYS)
GA2-F1087
GA2-IgG1

FIG. 12

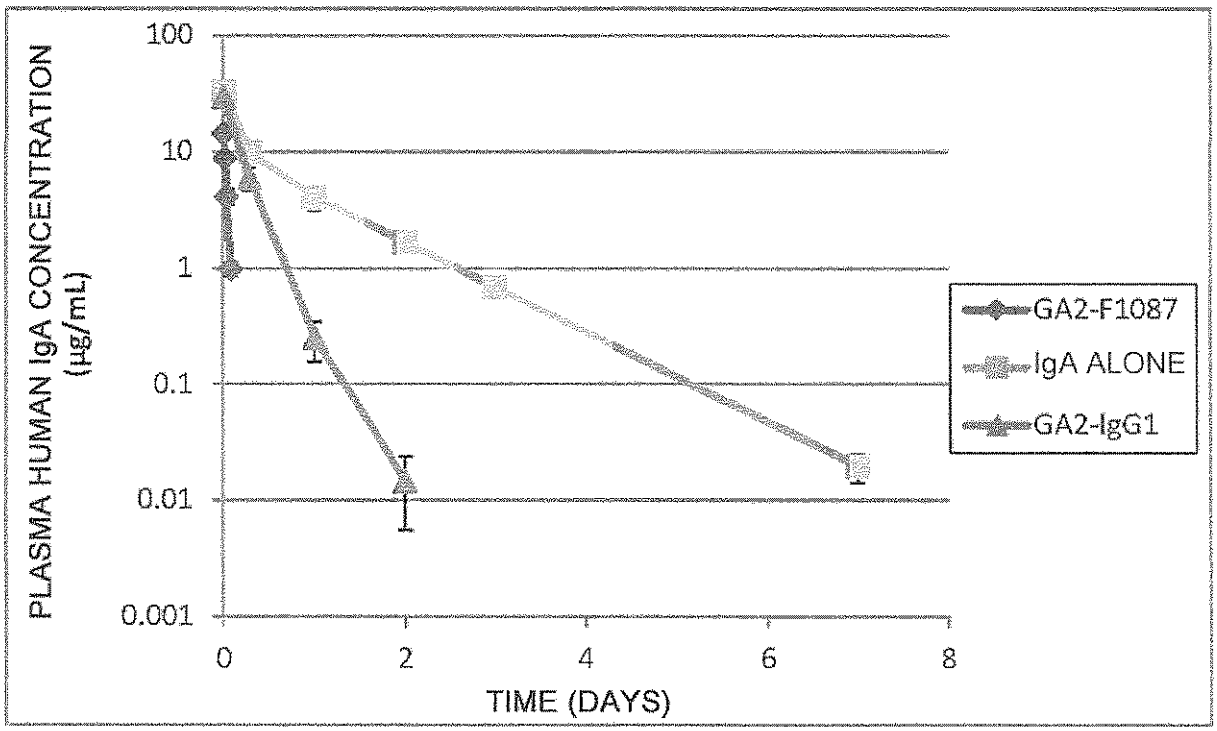

PLASMA HUMAN IgA CONCENTRATION (µg/mL)
100
10
1
0.1
0.01
0.001
0
2
4
6
8
TIME (DAYS)
GA2-F1087
IgA ALONE
GA2-IgG1

FIG. 13

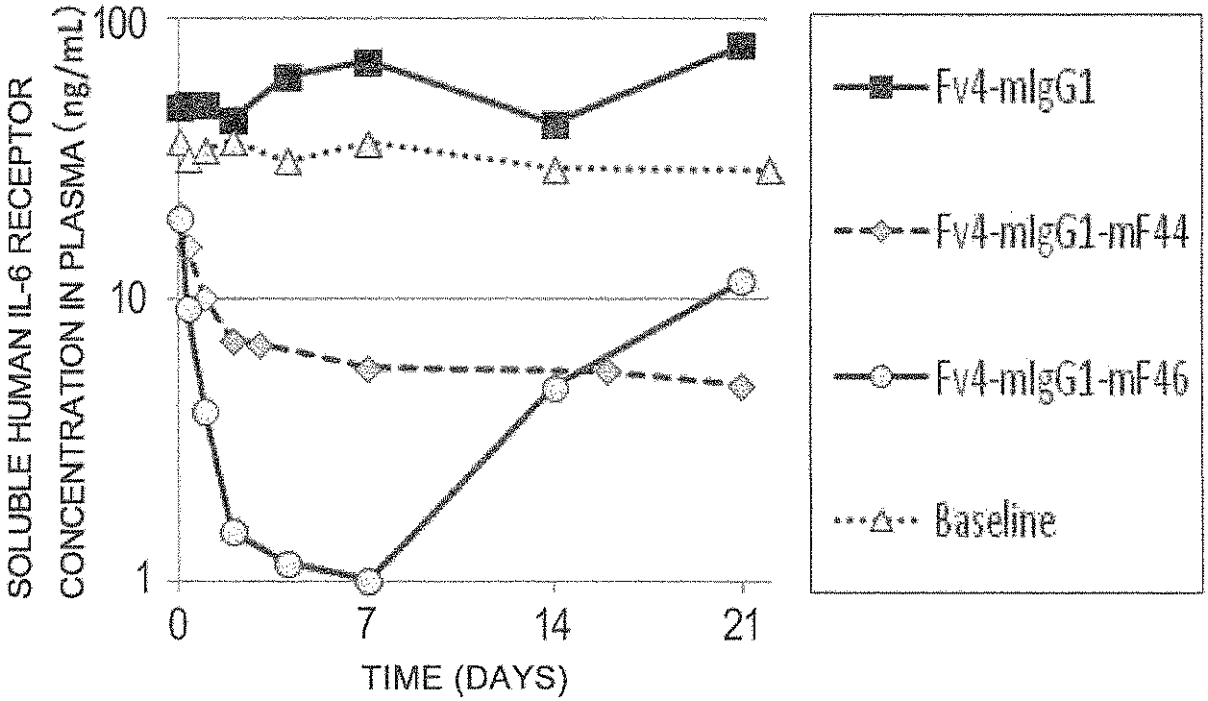

SOLUBLE HUMAN IL-6 RECEPTOR CONCENTRATION IN PLASMA (ng/mL)
100
10
1
0
7
14
21
TIME (DAYS)
Fv4-mIgG1
Fv4-mIgG1-mF44
Fv4-mIgG1-mF46
Baseline

FIG. 14

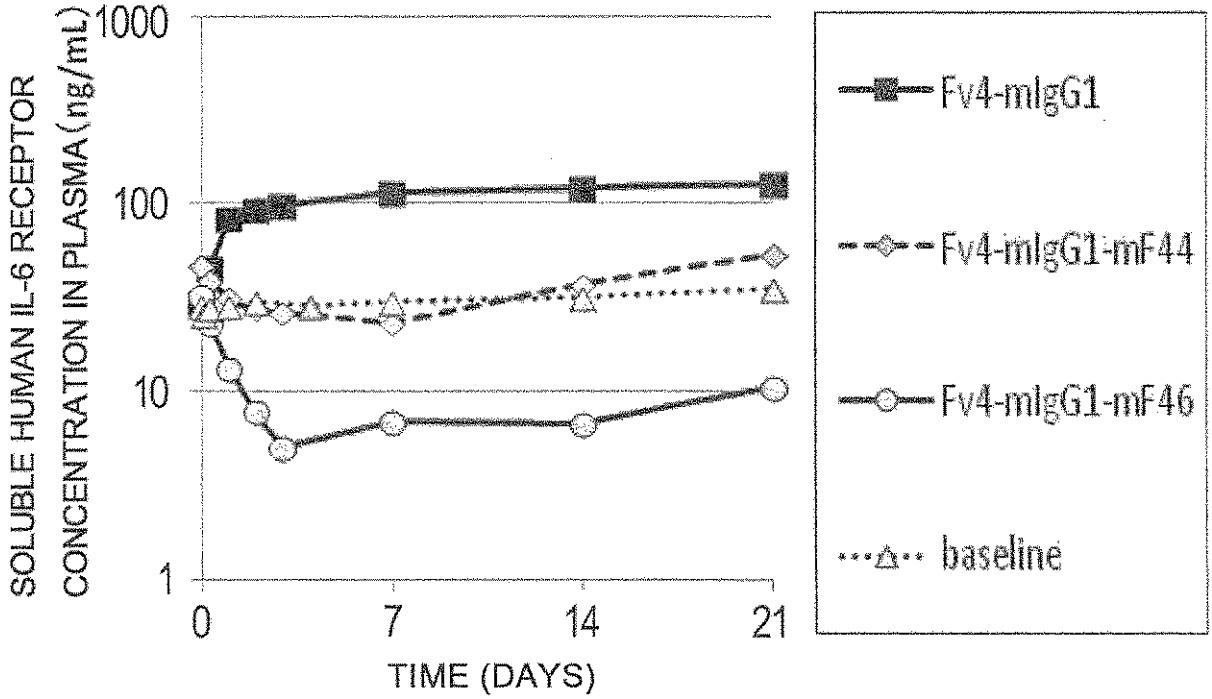

SOLUBLE HUMAN IL-6 RECEPTOR CONCENTRATION IN PLASMA (ng/mL)
1000
100
10
1
0
7
14
21
TIME (DAYS)
Fv4-mIgG1
Fv4-mIgG1-mF44
Fv4-mIgG1-mF46
baseline

FIG. 15

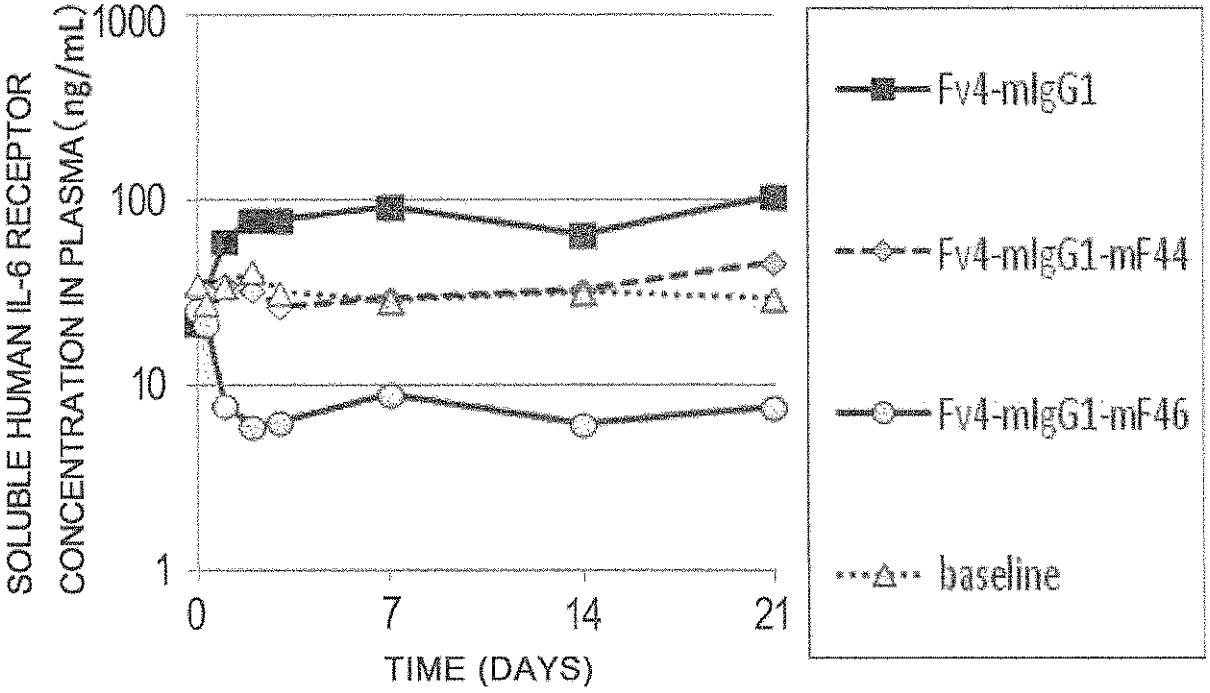

SOLUBLE HUMAN IL-6 RECEPTOR CONCENTRATION IN PLASMA (ng/mL)
1000
100
10
1
0
7
14
21
TIME (DAYS)
Fv4-mIgG1
Fv4-mIgG1-mF44
Fv4-mIgG1-mF46
baseline

FIG. 16

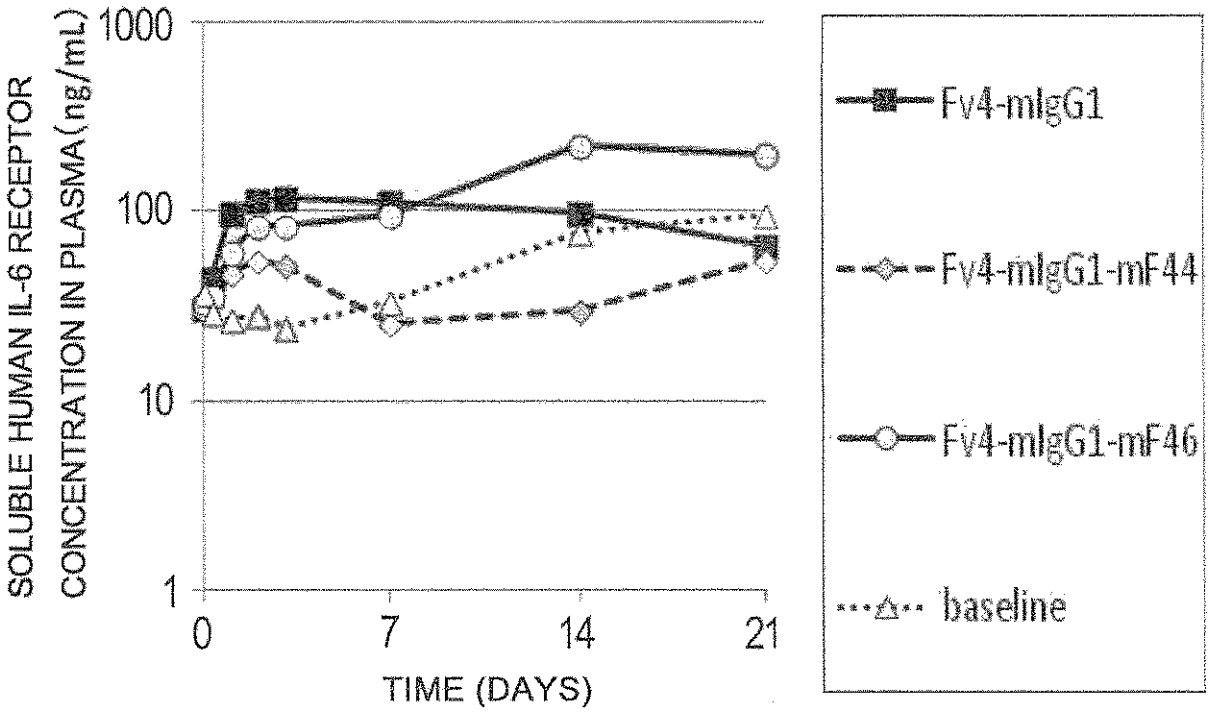

SOLUBLE HUMAN IL-6 RECEPTOR CONCENTRATION IN PLASMA(ng/mL)
1000
100
10
1
0
7
14
21
TIME (DAYS)
Fv4-mIgG1
Fv4-mIgG1-mF44
Fv4-mIgG1-mF46
baseline

FIG. 17

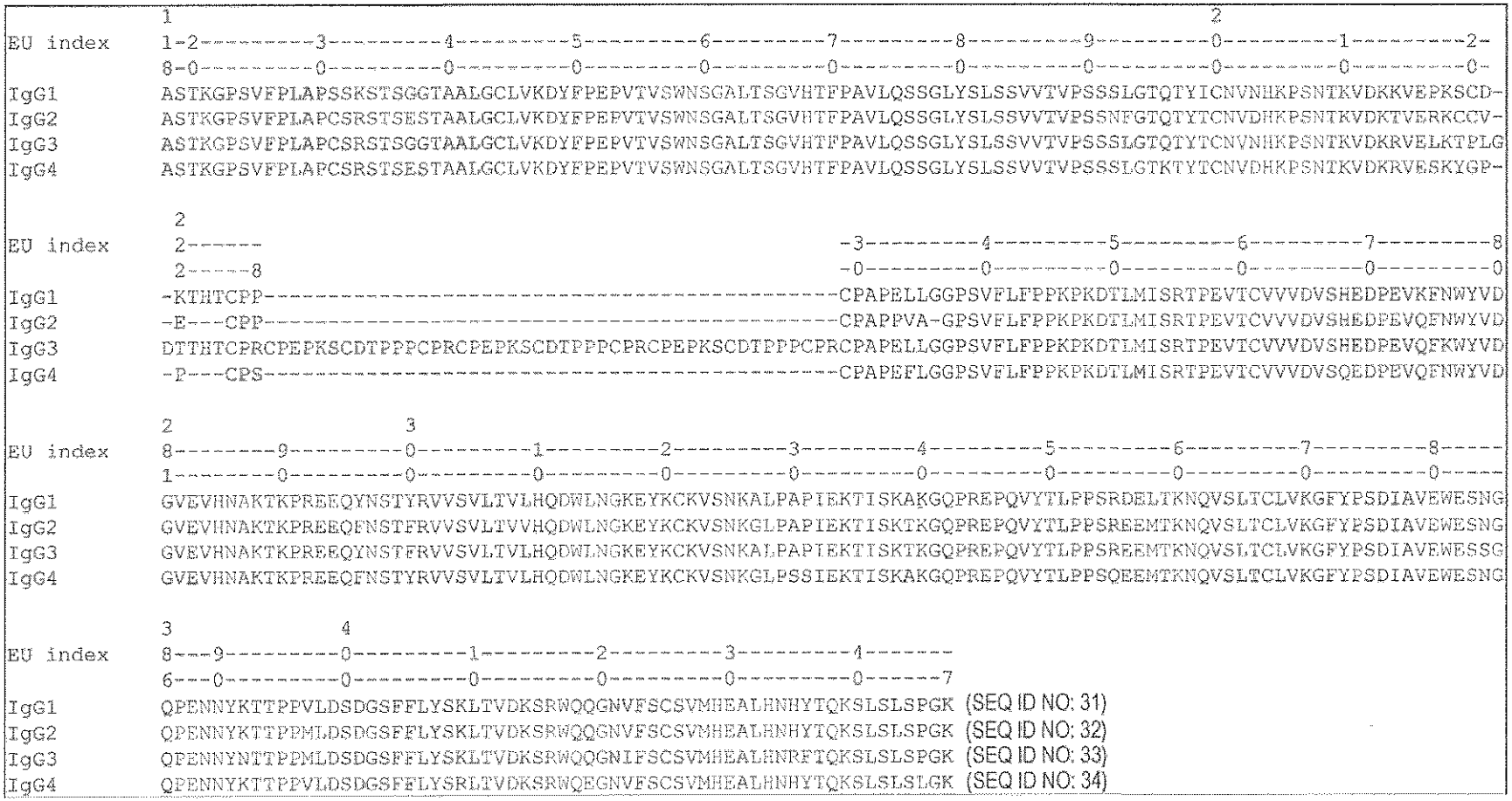

EU index
IgG1 ASTKGPSVFPLAPSSKSTSGGTAALGCLVKDYFPEPVTVSWNSGALTSGVHTFPAVLQSSGLYSLSSVVTVPSSSLGTQTYICNVNHKPSNTKVDKKVEPKSCD-
IgG2 ASTKGPSVFPLAPCSRSTSESTAALGCLVKDYFPEPVTVSWNSGALTSGVHTFPAVLQSSGLYSLSSVVTVPSSNFGTQTYTCNVDHKPSNTKVDKTVERKCCV-
IgG3 ASTKGPSVFPLAPCSRSTSGGTAALGCLVKDYFPEPVTVSWNSGALTSGVHTFPAVLQSSGLYSLSSVVTVPSSSLGTQTYTCNVNHKPSNTKVDKRVELKTPLG
IgG4 ASTKGPSVFPLAPCSRSTSESTAALGCLVKDYFPEPVTVSWNSGALTSGVHTFPAVLQSSGLYSLSSVVTVPSSSLGTKTYTCNVDHKPSNTKVDKRVESKYGP-
EU index
IgG1 -KTHTCPP----------CPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVD
IgG2 -E---CPP----------CPAPPVA-GPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVQFNWYVD
IgG3 DTTHTCPRCPEPKSCDTPPPCPRCPEPKSCDTPPPCPRCPEPKSCDTPPPCPRCPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVQFKWYVD
IgG4 -P---CPS----------CPAPEFLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSQEDPEVQFNWYVD
EU index
IgG1 GVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIEKTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNG
IgG2 GVEVHNAKTKPREEQFNSTFRVVSVLTVVHQDWLNGKEYKCKVSNKGLPAPIEKTISKTKGQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNG
IgG3 GVEVHNAKTKPREEQYNSTFRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIEKTISKTKGQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESSG
IgG4 GVEVHNAKTKPREEQFNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKGLPSSIEKTISKAKGQPREPQVYTLPPSQEEMTKNQVSLTCLVKGFYPSDIAVEWESNG
EU index
IgG1 QPENNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLSPGK (SEQ ID NO: 31)
IgG2 QPENNYKTTPPMLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLSPGK (SEQ ID NO: 32)
IgG3 QPENNYNTTPPMLDSDGSFFLYSKLTVDKSRWQQGNIFSCSVMHEALHNRFTQKSLSLSPGK (SEQ ID NO: 33)
IgG4 QPENNYKTTPPVLDSDGSFFLYSRLTVDKSRWQEGNVFSCSVMHEALHNHYTQKSLSLSLGK (SEQ ID NO: 34)

FIG. 18

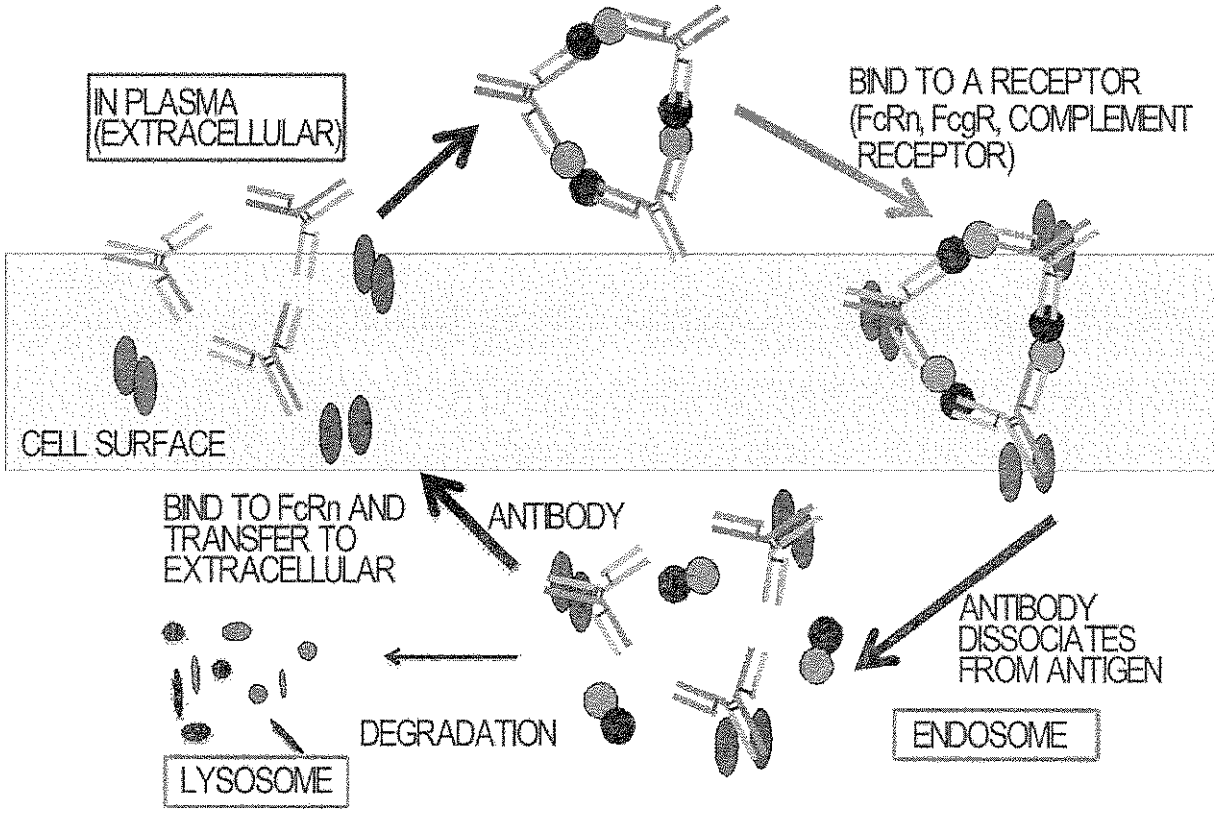

IN PLASMA
(EXTRACELLULAR)
BIND TO A RECEPTOR
(FcRn, FcgR, COMPLEMENT
RECEPTOR)
CELL SURFACE
BIND TO FcRn AND
TRANSFER TO
EXTRACELLULAR
ANTIBODY
ANTIBODY
DISSOCIATES
FROM ANTIGEN
DEGRADATION
LYSOSOME
ENDOSOME

FIG. 19