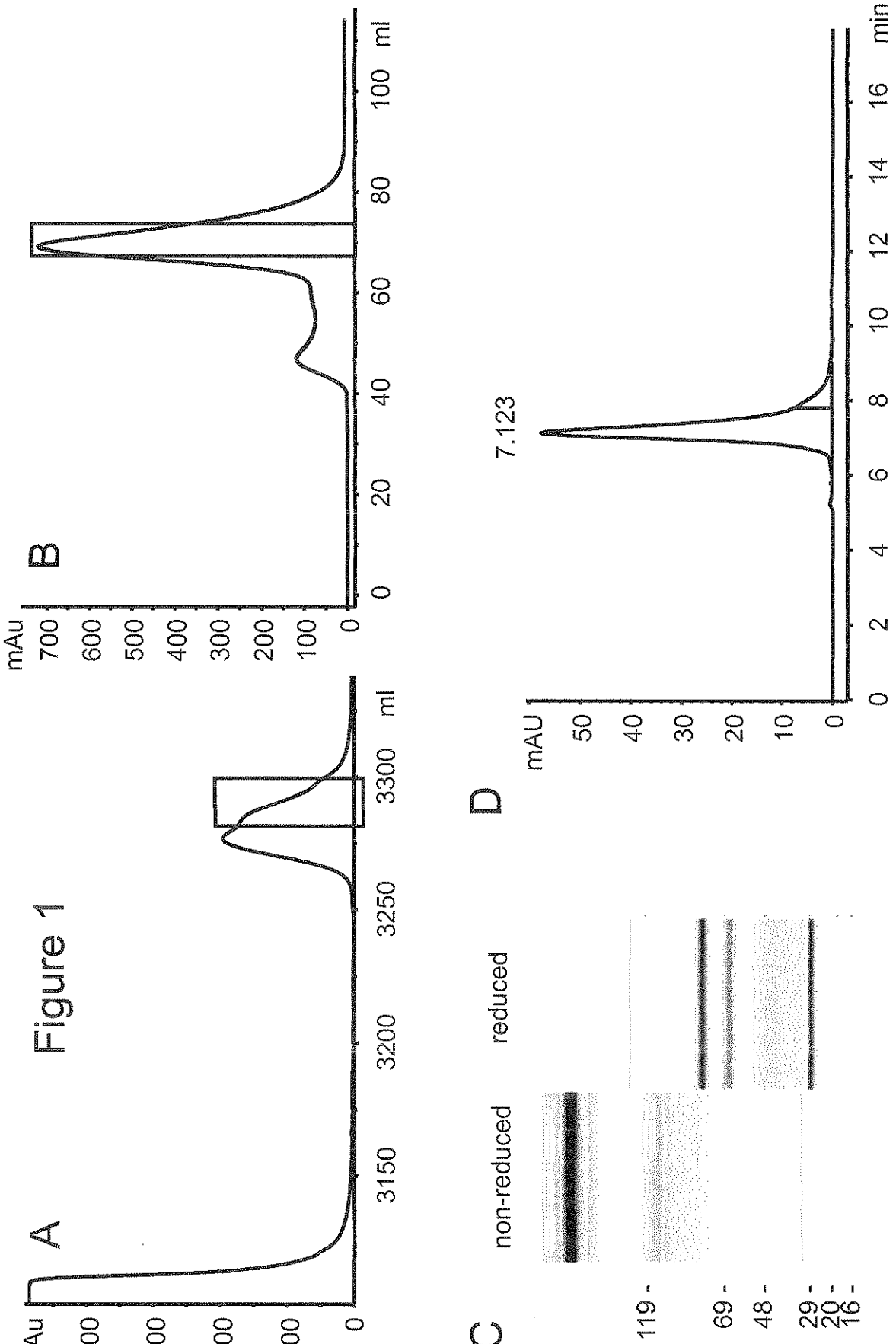
Figure 1


A
mAu
2000
1500
1000
500
0
3150
3200
3250
3300
ml
B
mAu
700
600
500
400
300
200
100
0
0
20
40
60
80
100
ml
C
non-reduced
reduced
119 -
69 -
48 -
29 -
20 -
16 -
D
7.123
mAU
50
40
30
20
10
0
0
2
4
6
8
10
12
14
16
min

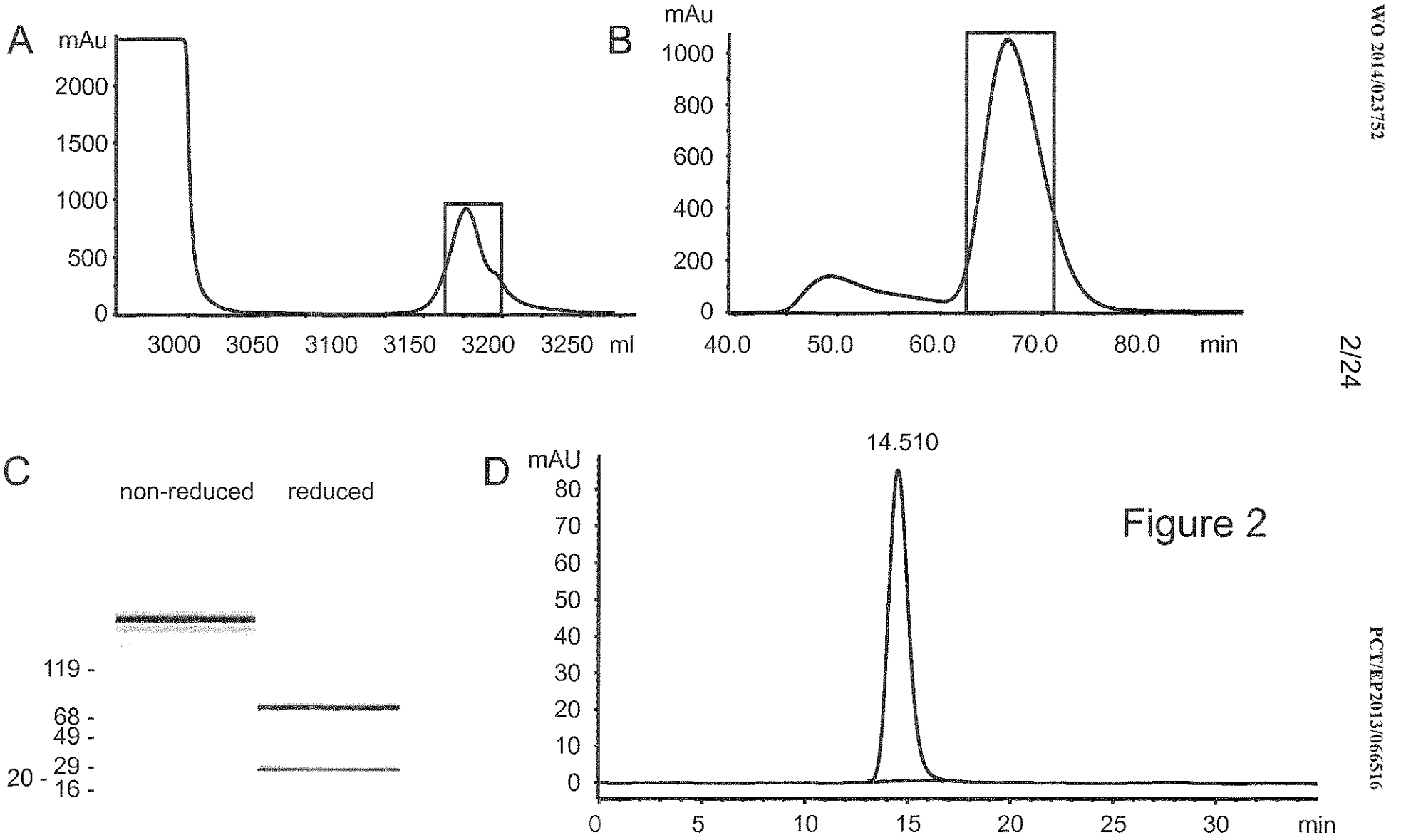

A
mAu
2000
1500
1000
500
0
3000
3050
3100
3150
3200
3250
ml
B
mAu
1000
800
600
400
200
0
40.0
50.0
60.0
70.0
80.0
min
C
non-reduced
reduced
119 -
68 -
49 -
29 -
20 -
16 -
D
14.510
mAU
80
70
60
50
40
30
20
10
0
0
5
10
15
20
25
30
min

Figure 2

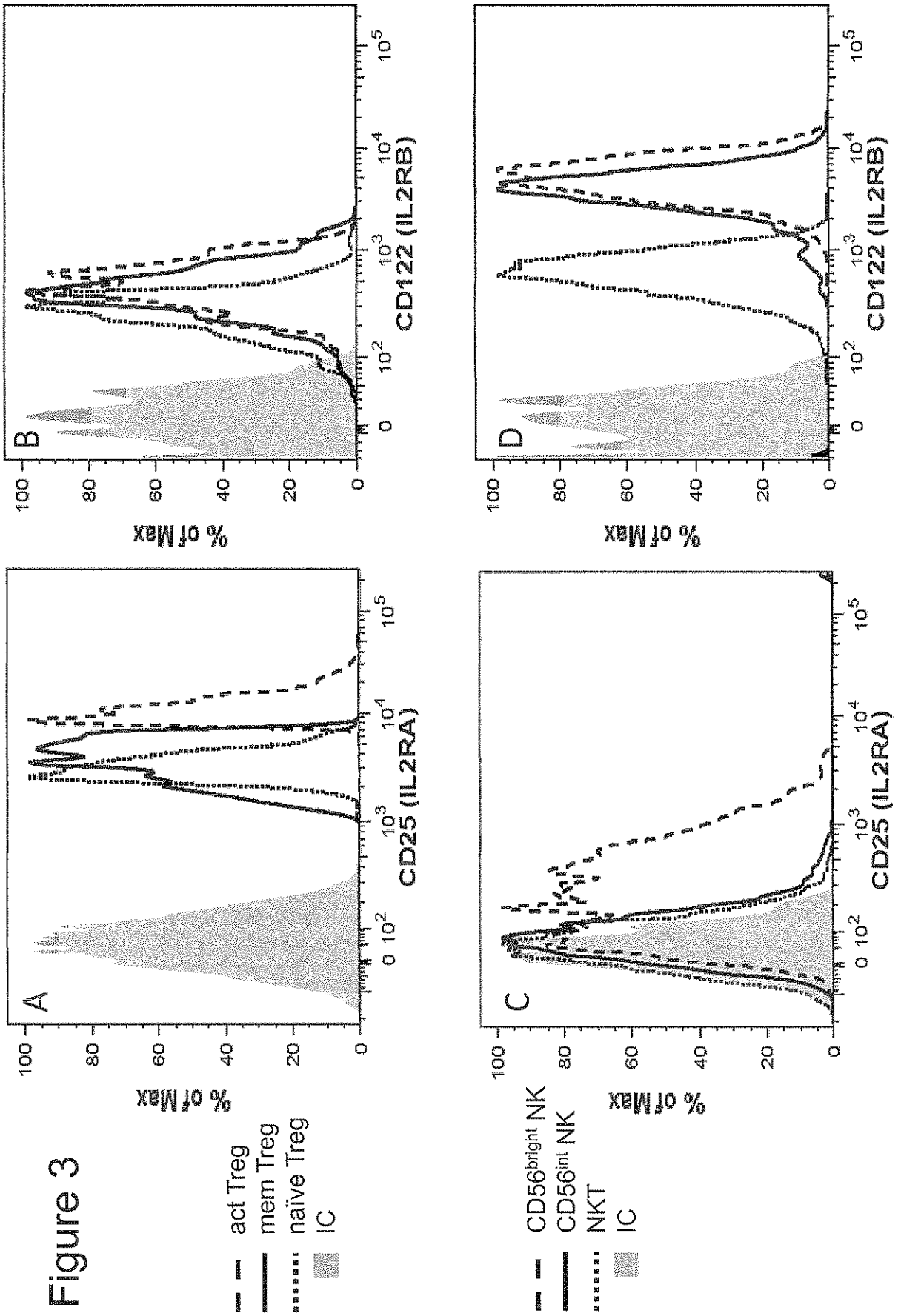

A
B
C
D
% of Max
CD25 (IL2RA)
CD122 (IL2RB)
act Treg
mem Treg
naïve Treg
IC
CD56bright NK
CD56int NK
NKT
IC

Figure 3

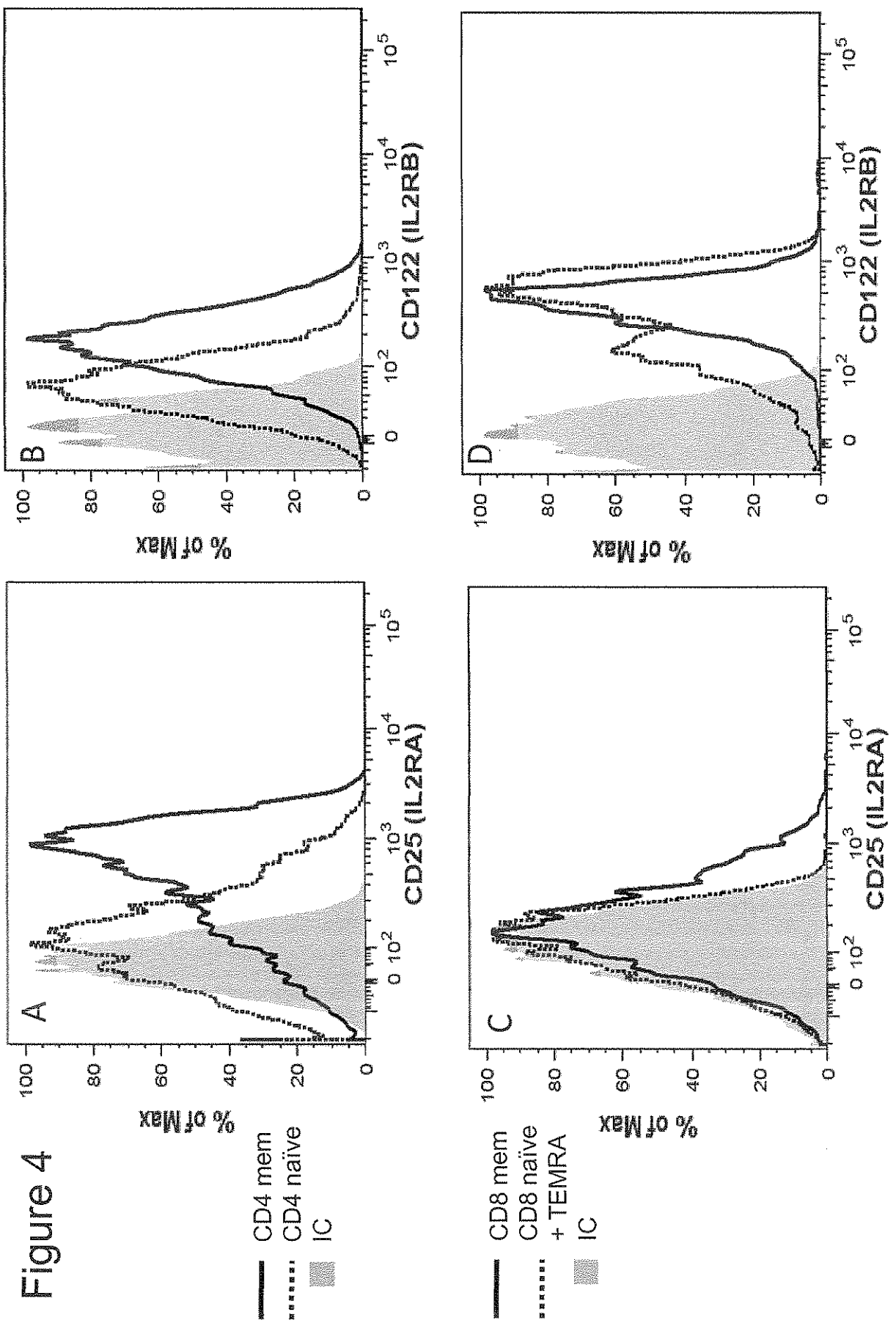
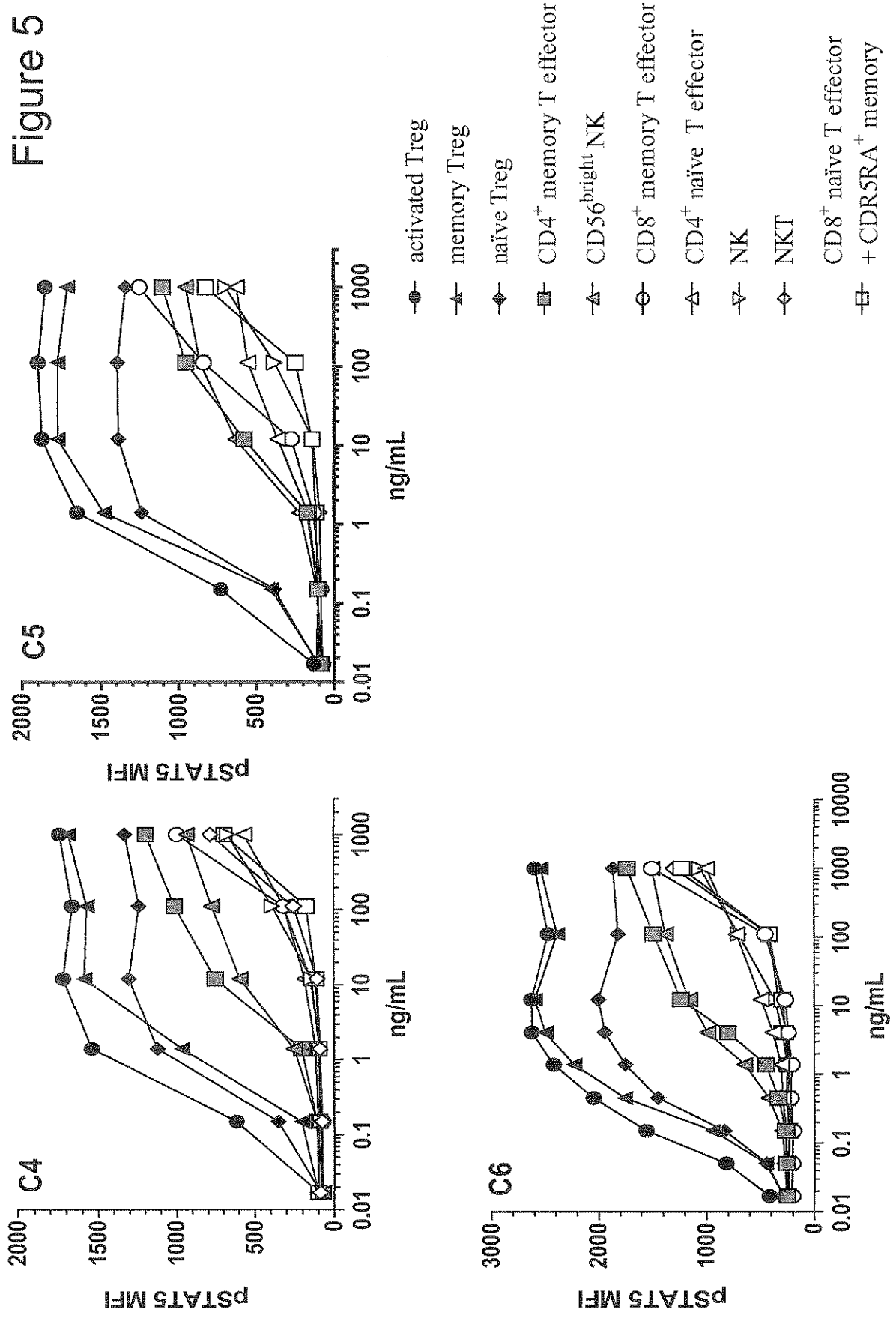

A
B
C
D
% of Max
CD25 (IL2RA)
CD122 (IL2RB)
CD4 mem
CD4 naïve
IC
CD8 mem
CD8 naïve
+ TEMRA
IC

Figure 4

Figure 5


C4
C5
C6
pSTAT5 MFI
ng/mL
0
500
1000
1500
2000
3000
0.01
0.1
1
10
100
1000
10000
activated Treg
memory Treg
naïve Treg
CD4+ memory T effector
CD56bright NK
CD8+ memory T effector
CD4+ naïve T effector
NK
NKT
CD8+ naïve T effector
+ CDR5RA+ memory

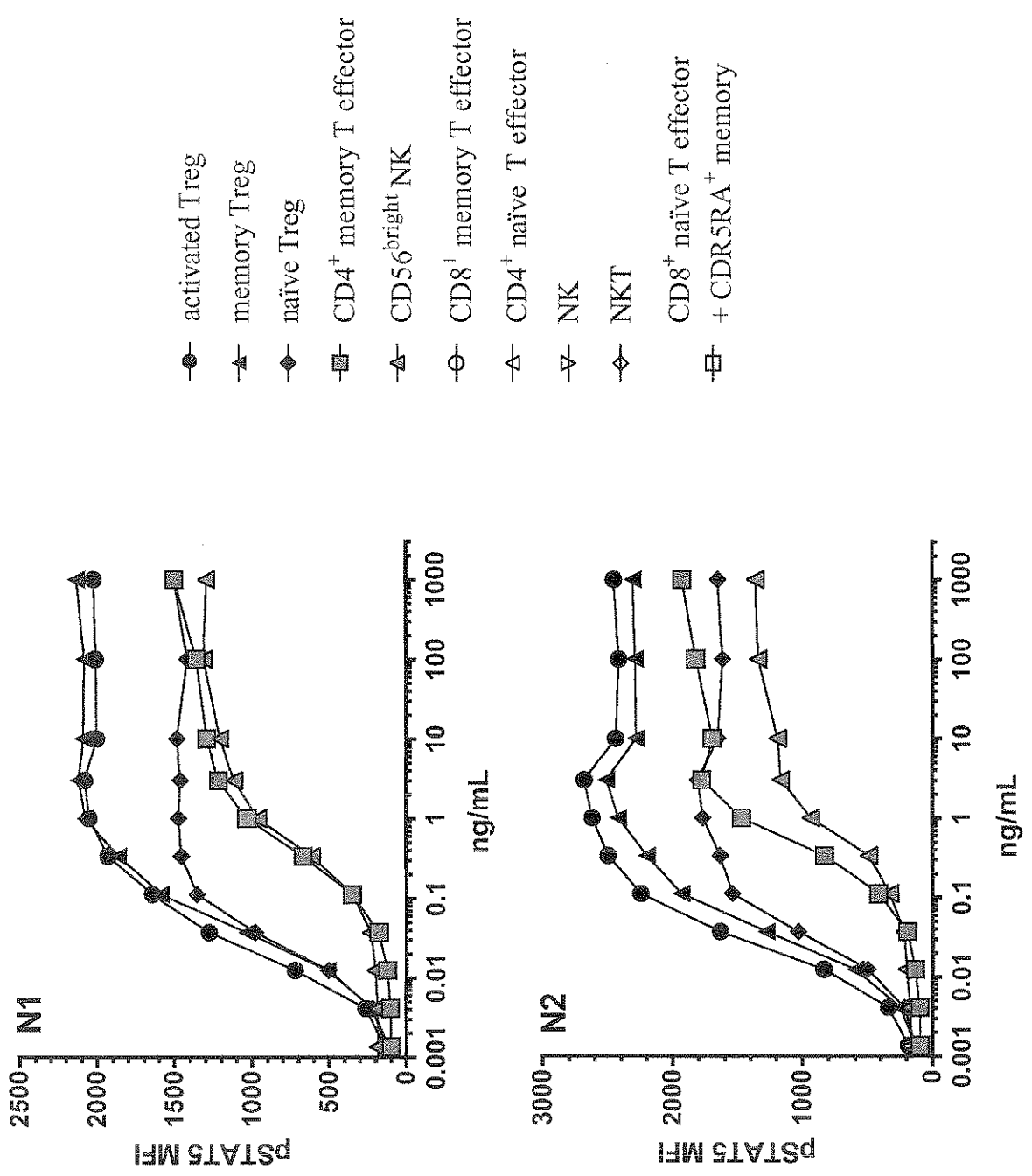

activated Treg
memory Treg
naïve Treg
CD4+ memory T effector
CD56bright NK
CD8+ memory T effector
CD4+ naïve T effector
NK
NKT
CD8+ naïve T effector
+ CDR5RA+ memory
N1
N2
pSTAT5 MFI
ng/mL
0
500
1000
1500
2000
2500
3000
0.001
0.01
0.1
1
10
100
1000

Figure 6

Figure 6

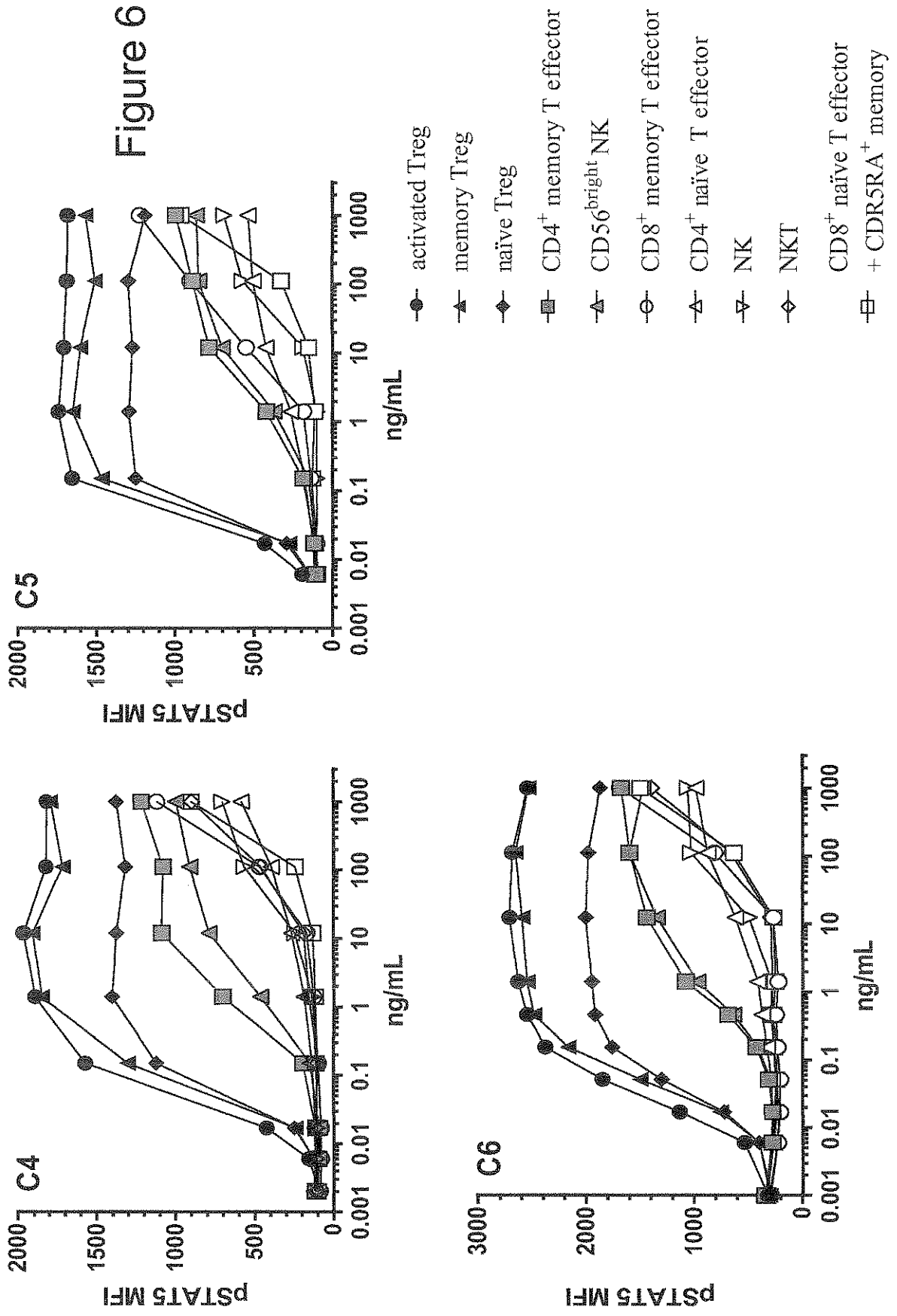

C4
C5
C6
pSTAT5 MFI
ng/mL
0
500
1000
1500
2000
3000
0.001
0.01
0.1
1
10
100
1000
activated Treg
memory Treg
naïve Treg
CD4+ memory T effector
CD56bright NK
CD8+ memory T effector
CD4+ naïve T effector
NK
NKT
CD8+ naïve T effector
+ CDR5RA+ memory

Figure 7

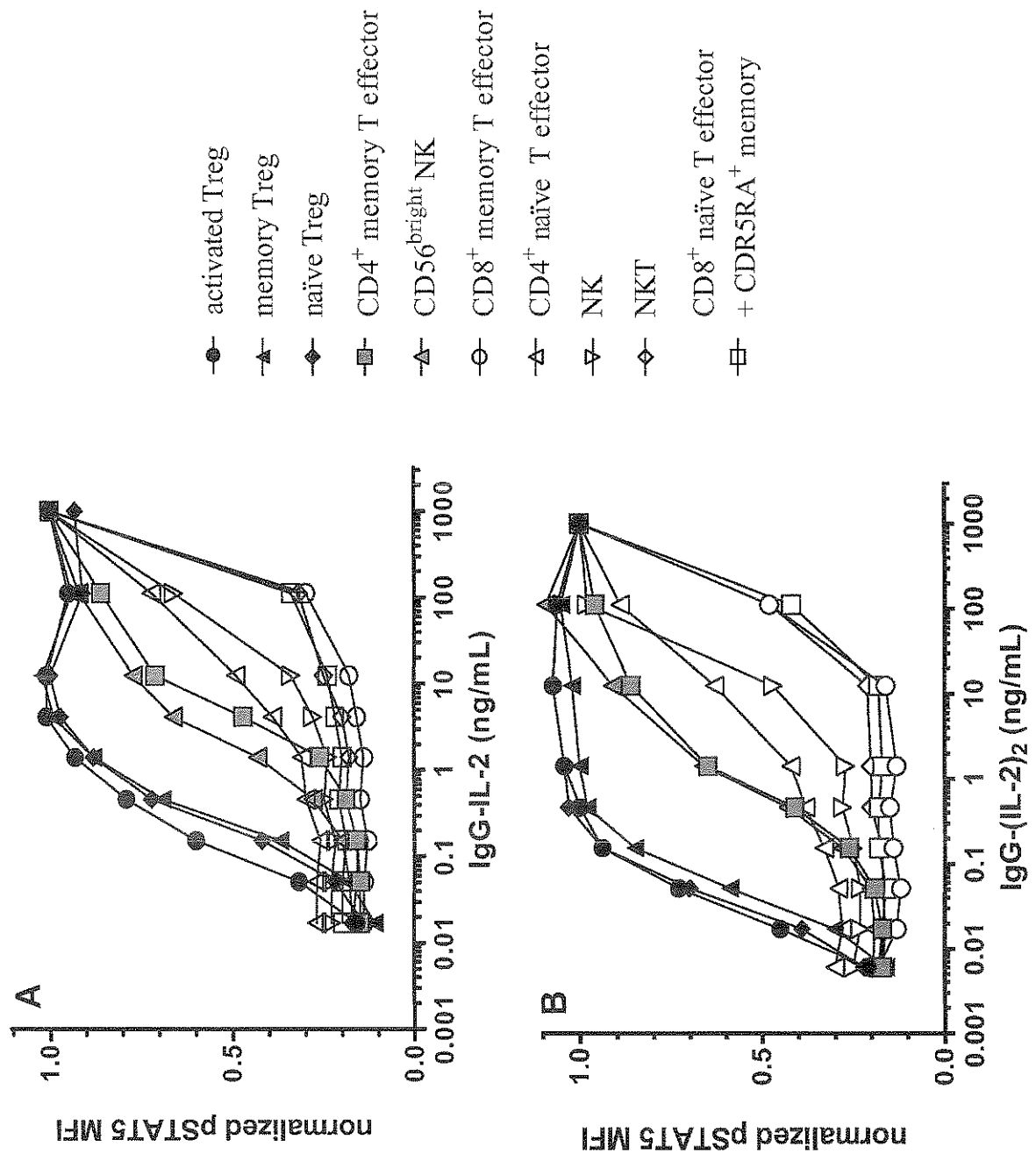

activated Treg
memory Treg
naïve Treg
CD4+ memory T effector
CD56bright NK
CD8+ memory T effector
CD4+ naïve T effector
NK
NKT
CD8+ naïve T effector
+ CDR5RA+ memory
A
B
normalized pSTAT5 MFI
1.0
0.5
0.0
0.001
0.01
0.1
1
10
100
1000
IgG-IL-2 (ng/mL)
IgG-(IL-2)2 (ng/mL)

Figure 8

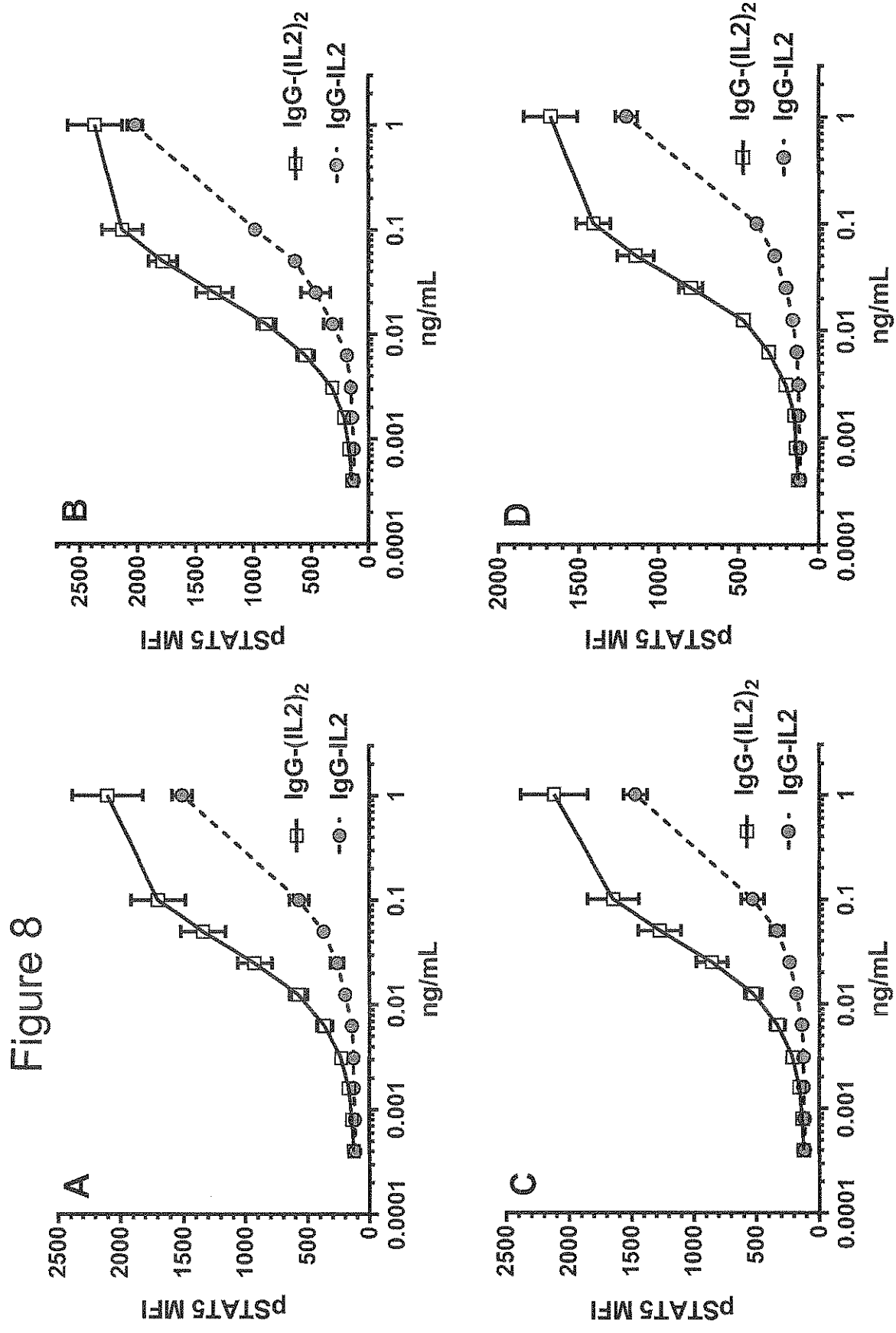

A
B
C
D
pSTAT5 MFI
ng/mL
IgG-(IL2)2
IgG-IL2
0
500
1000
1500
2000
2500
0.0001
0.001
0.01
0.1
1

Figure 9

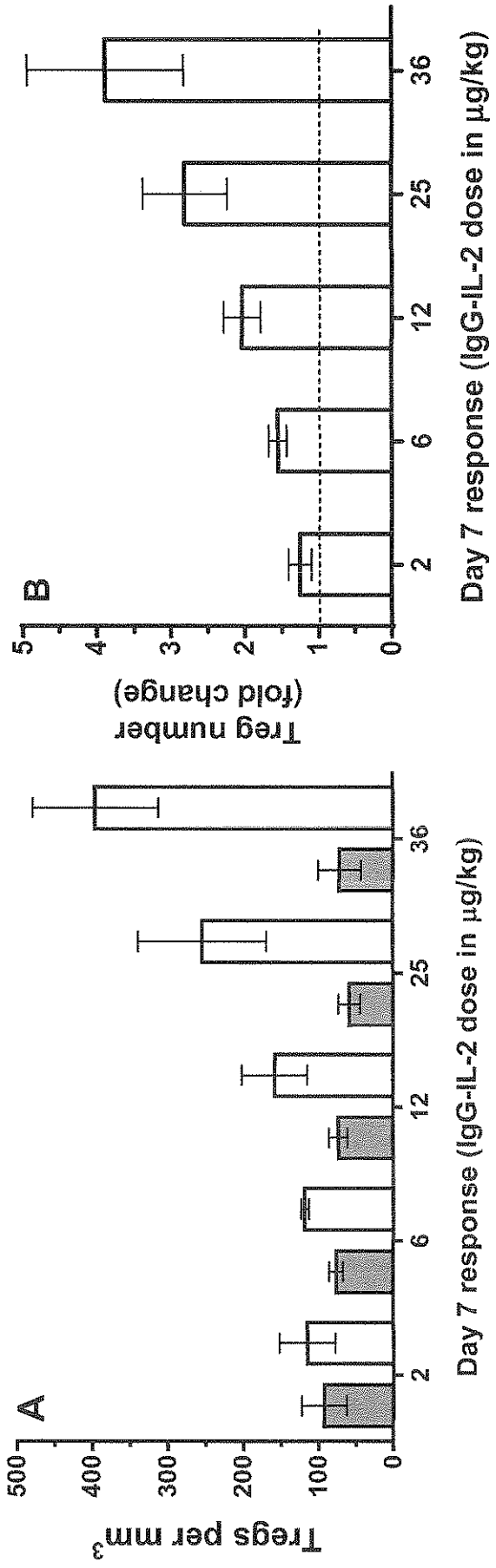

A
Tregs per mm3
500
400
300
200
100
0
2
6
12
25
36
Day 7 response (IgG-IL-2 dose in µg/kg)
B
Treg number
(fold change)
5
4
3
2
1
0
2
6
12
25
36
Day 7 response (IgG-IL-2 dose in µg/kg)

Figure 10

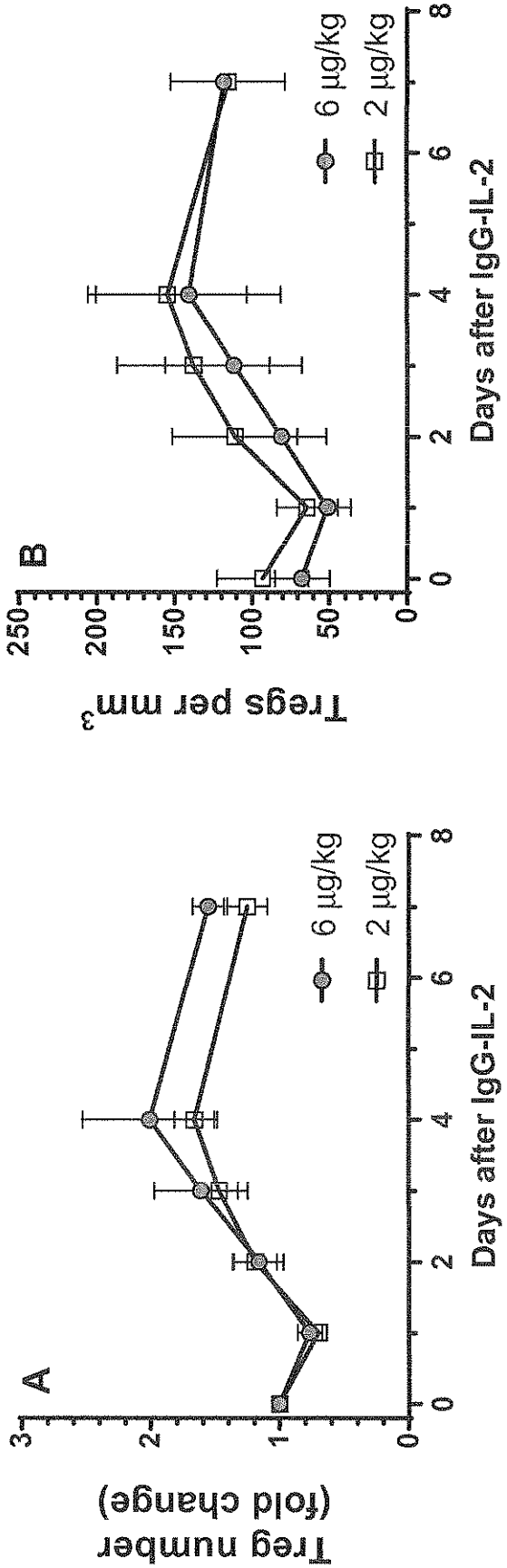
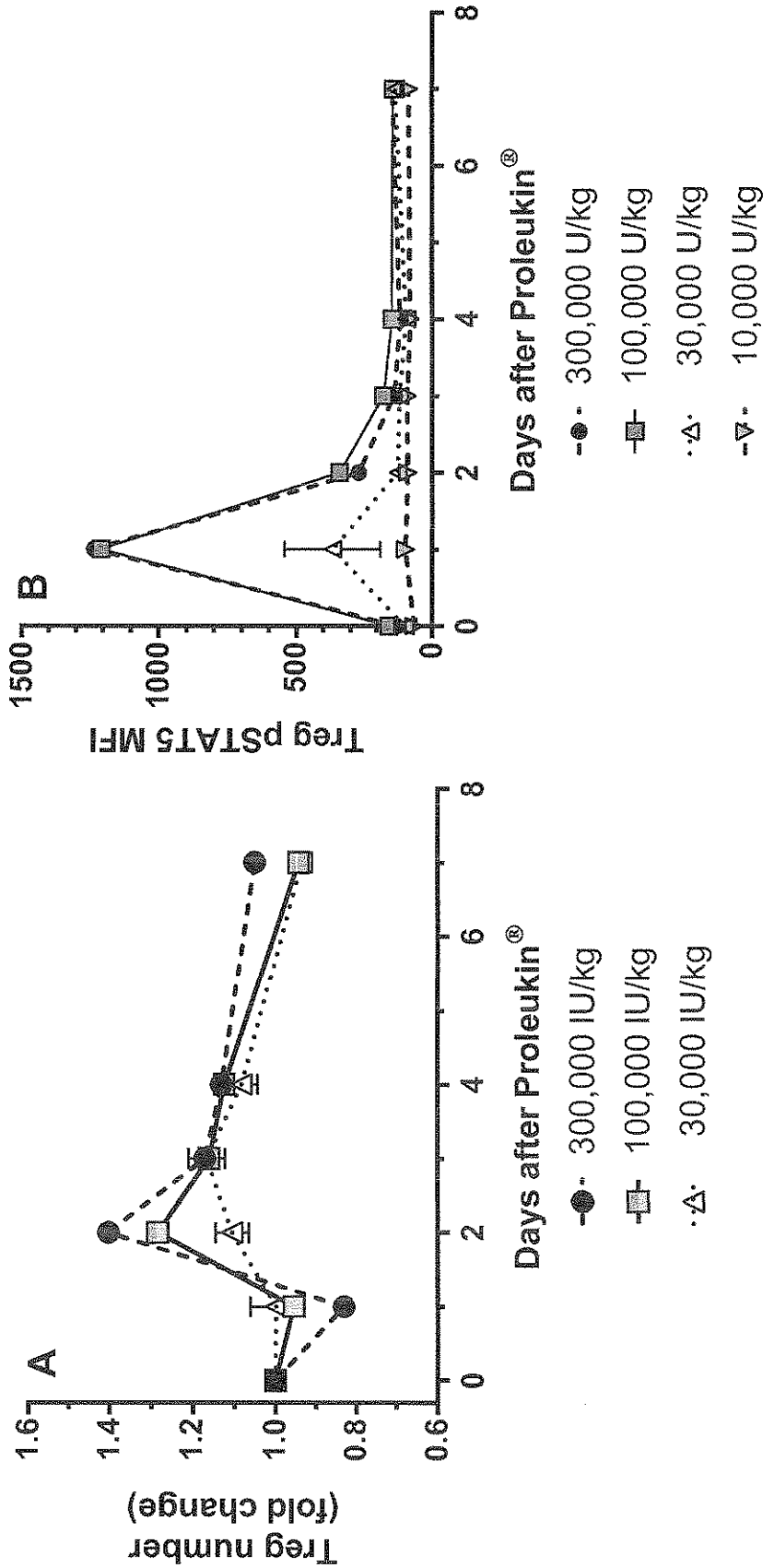
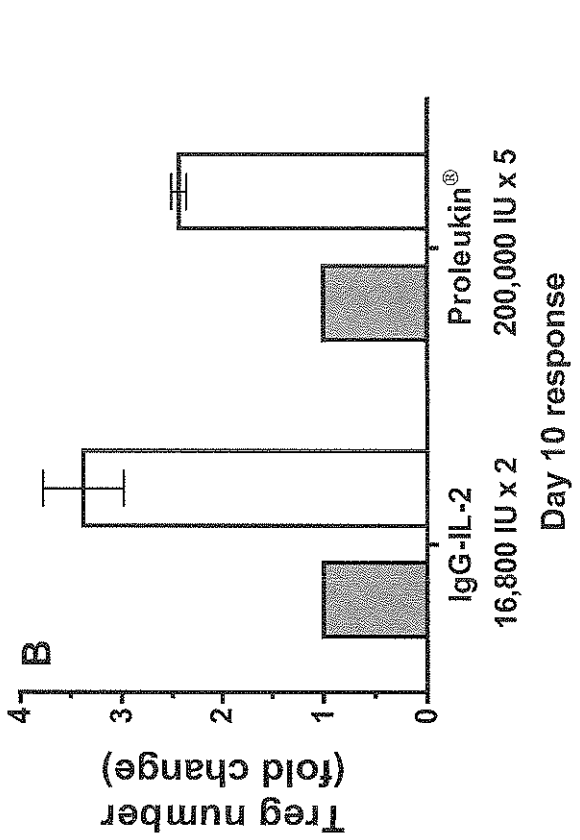
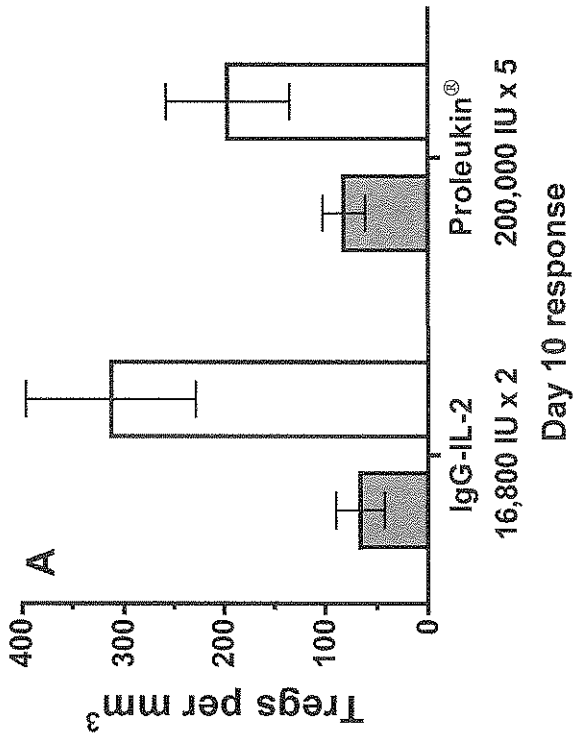

A
Treg number (fold change)
Days after IgG-IL-2
6 µg/kg
2 µg/kg
B
Tregs per mm3
Days after IgG-IL-2
6 µg/kg
2 µg/kg

Figure 11


A
Treg number (fold change)
1.6
1.4
1.2
1.0
0.8
0.6
0
2
4
6
8
Days after Proleukin®
300,000 IU/kg
100,000 IU/kg
30,000 IU/kg
B
Treg pSTAT5 MFI
1500
1000
500
0
0
2
4
6
8
Days after Proleukin®
300,000 U/kg
100,000 U/kg
30,000 U/kg
10,000 U/kg


B
Treg number (fold change)
0
1
2
3
4
IgG-IL-2 16,800 IU x 2
Proleukin® 200,000 IU x 5
Day 10 response


A
Tregs per mm3
0
100
200
300
400
IgG-IL-2 16,800 IU x 2
Proleukin® 200,000 IU x 5
Day 10 response

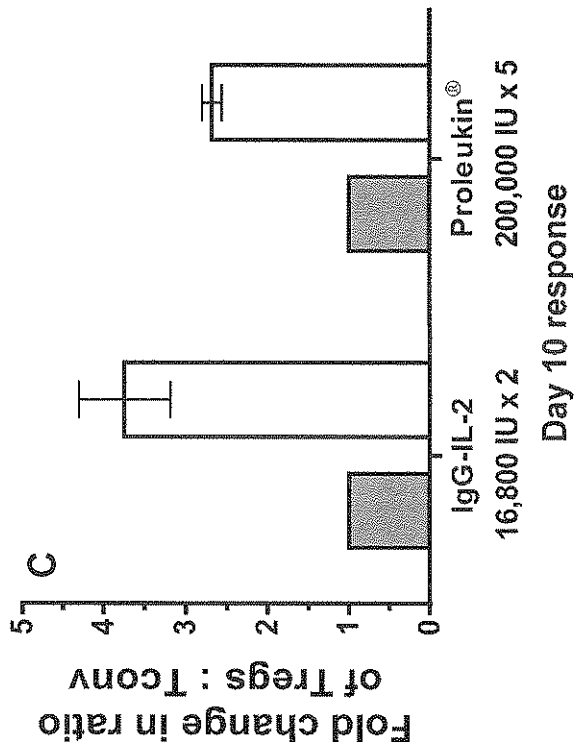

C
Fold change in ratio of Tregs : Tconv
0
1
2
3
4
5
IgG-IL-2 16,800 IU x 2
Proleukin® 200,000 IU x 5
Day 10 response

Figure 12

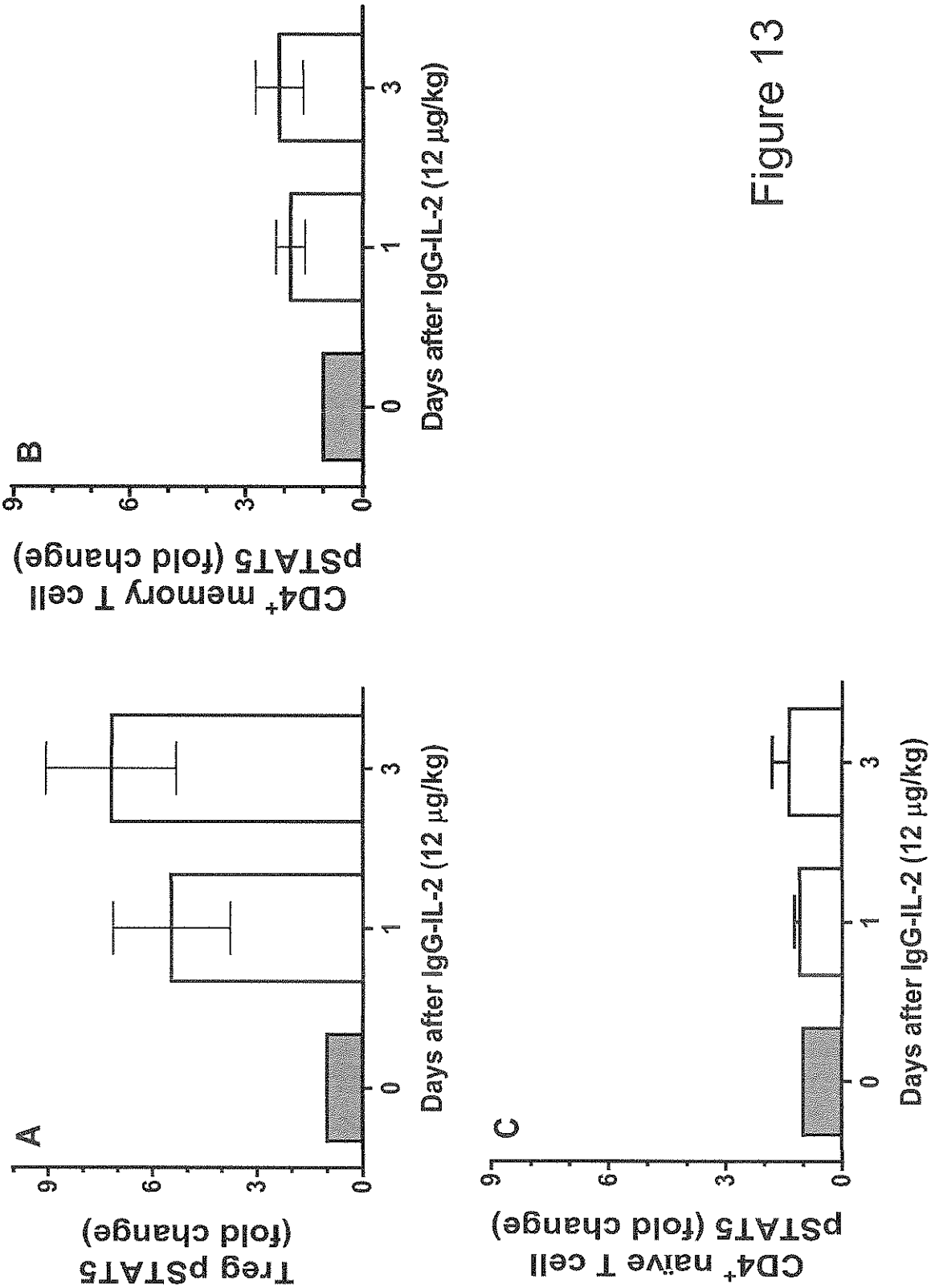

A
Treg pSTAT5 (fold change)
Days after IgG-IL-2 (12 µg/kg)
B
CD4+ memory T cell pSTAT5 (fold change)
Days after IgG-IL-2 (12 µg/kg)
C
CD4+ naïve T cell pSTAT5 (fold change)
Days after IgG-IL-2 (12 µg/kg)

Figure 13

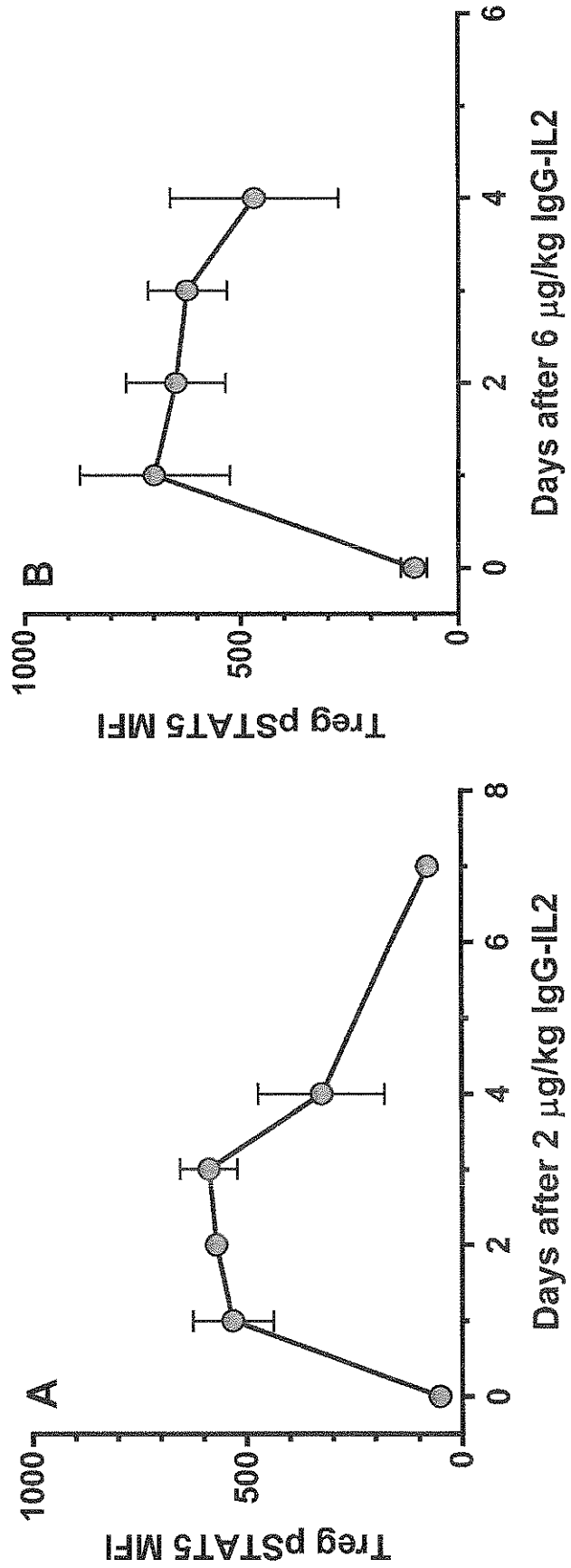

A
B
Treg pSTAT5 MFI
1000
500
0
0
2
4
6
8
Days after 2 µg/kg IgG-IL2
Days after 6 µg/kg IgG-IL2

Figure 14

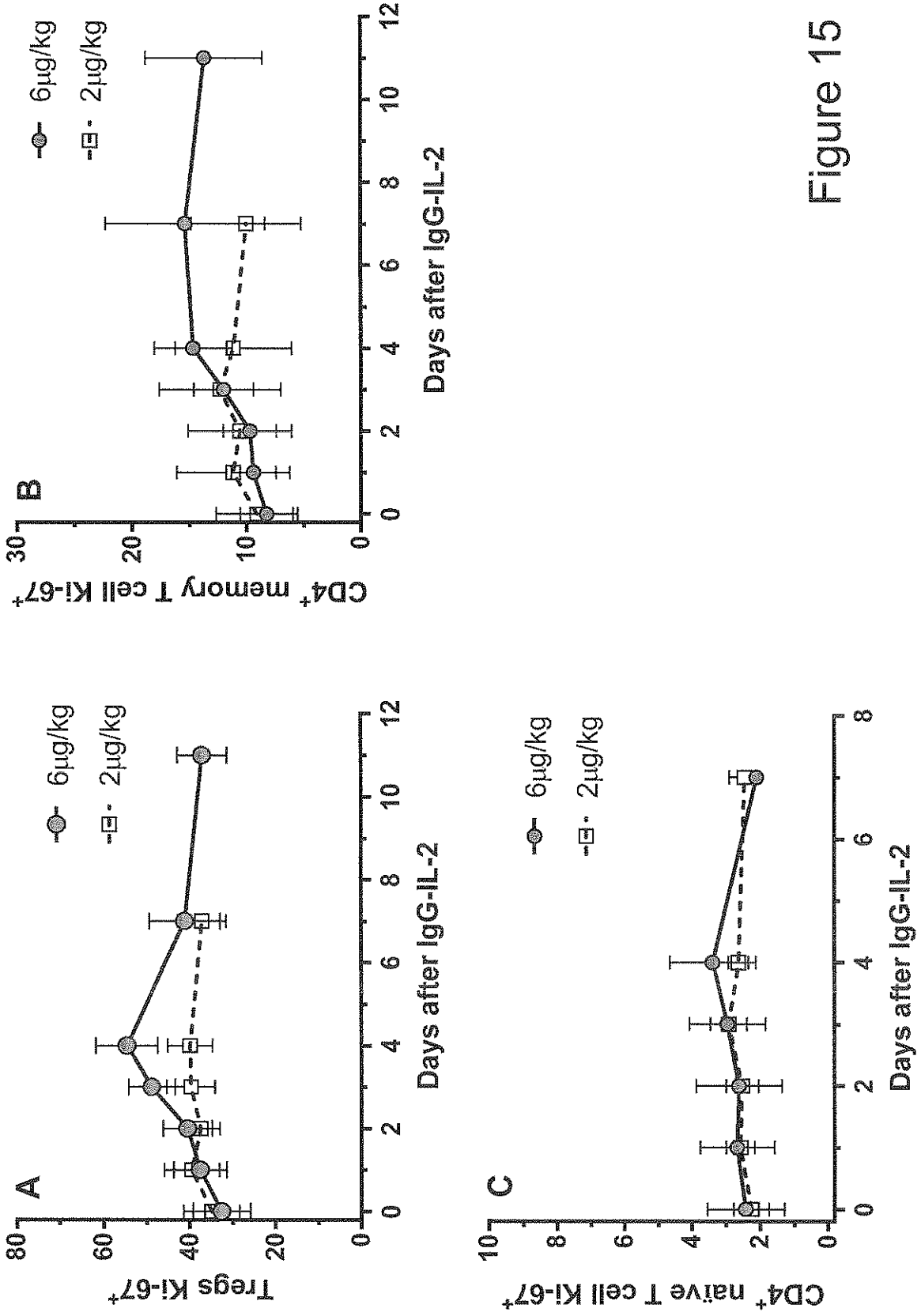

A
Tregs Ki-67+
80
60
40
20
0
0
2
4
6
8
10
12
Days after IgG-IL-2
6µg/kg
2µg/kg
B
CD4+ memory T cell Ki-67+
30
20
10
0
C
CD4+ naive T cell Ki-67+
10
8
6
4
2
0
8

Figure 15

Figure 16

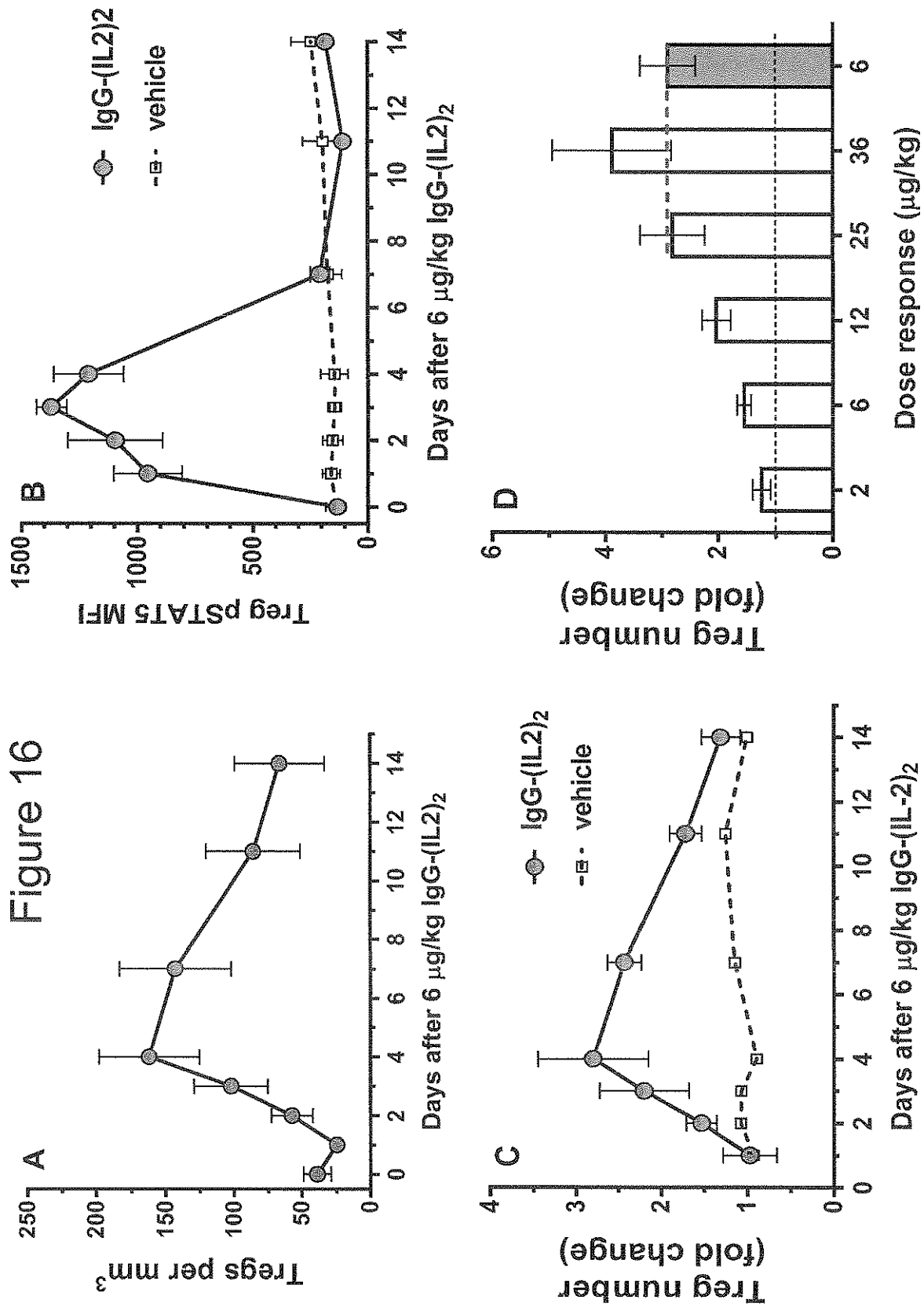
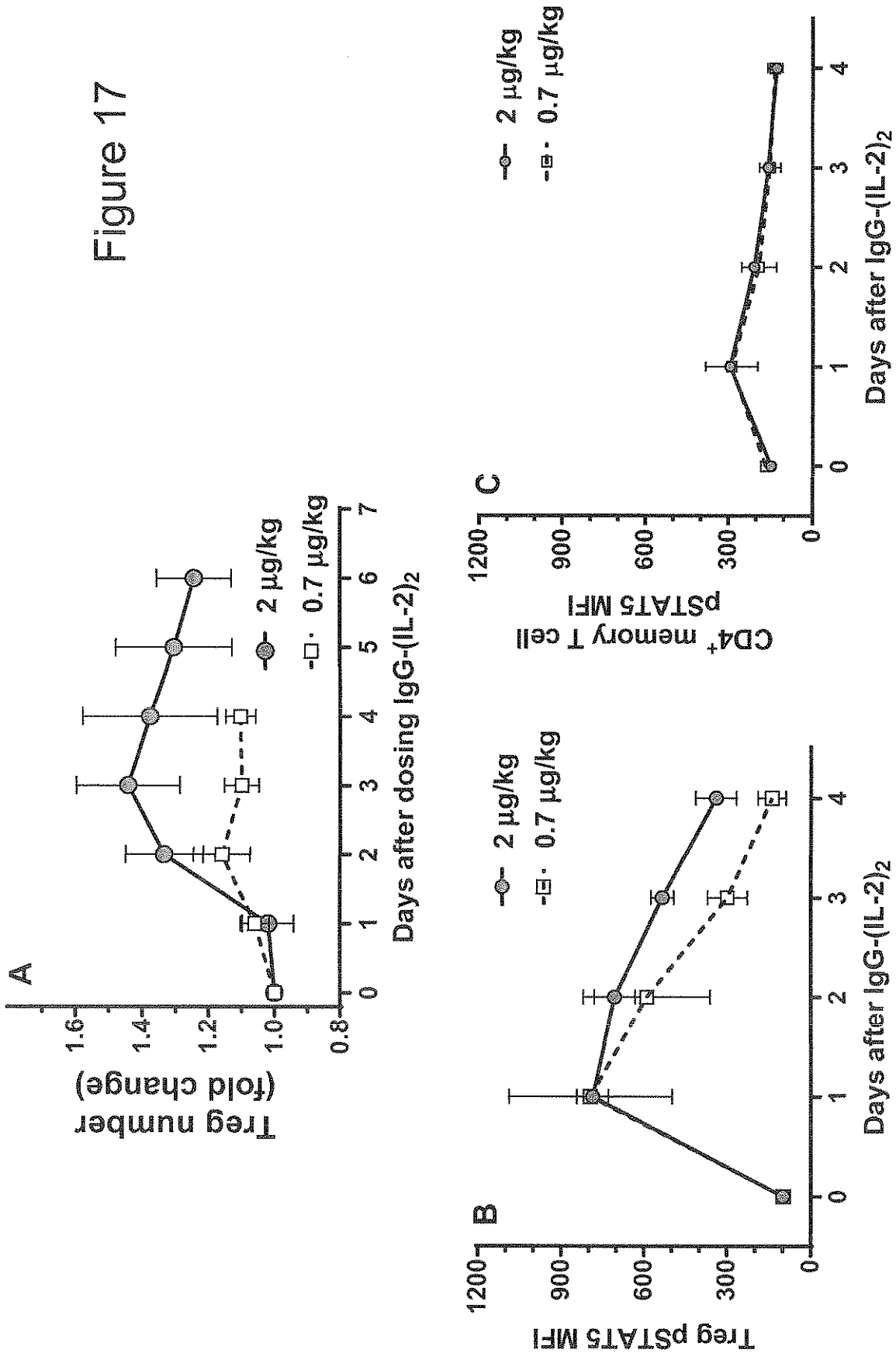

A
Tregs per mm3
Days after 6 µg/kg IgG-(IL2)2
B
Treg pSTAT5 MFI
IgG-(IL2)2
vehicle
Days after 6 µg/kg IgG-(IL2)2
C
Treg number (fold change)
IgG-(IL2)2
vehicle
Days after 6 µg/kg IgG-(IL-2)2
D
Treg number (fold change)
Dose response (µg/kg)

Figure 17


A
Treg number (fold change)
1.6
1.4
1.2
1.0
0.8
0
1
2
3
4
5
6
7
Days after dosing IgG-(IL-2)2
2 µg/kg
0.7 µg/kg
B
Treg pSTAT5 MFI
1200
900
600
300
0
Days after IgG-(IL-2)2
C
CD4+ memory T cell pSTAT5 MFI

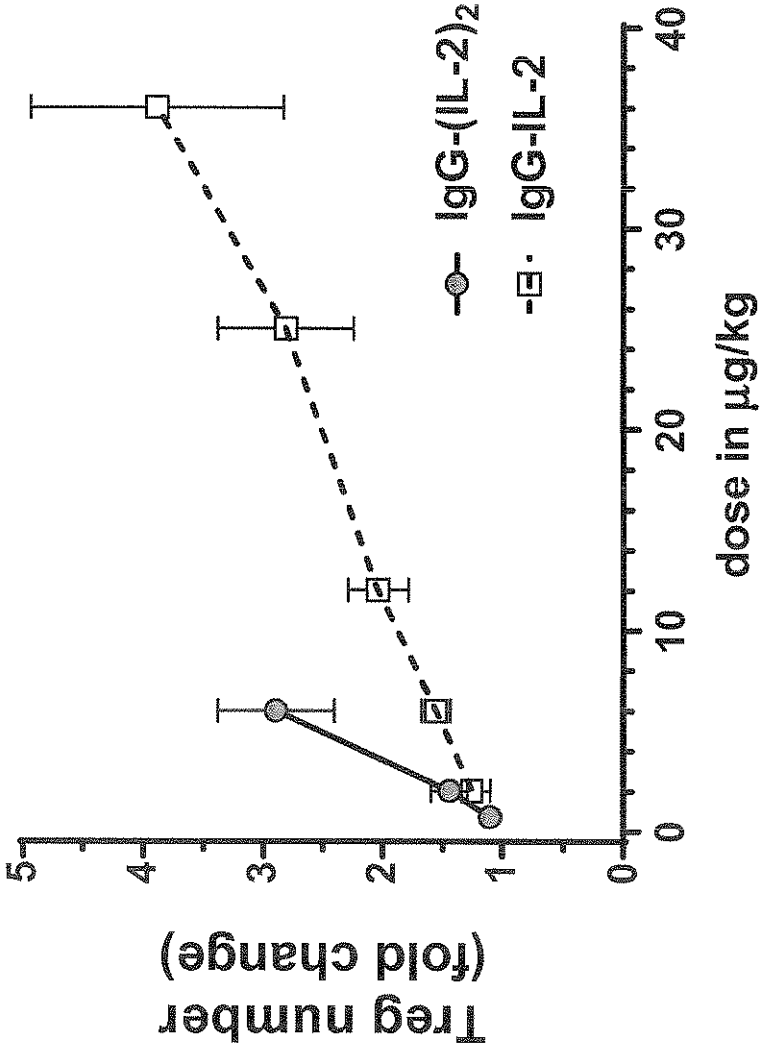
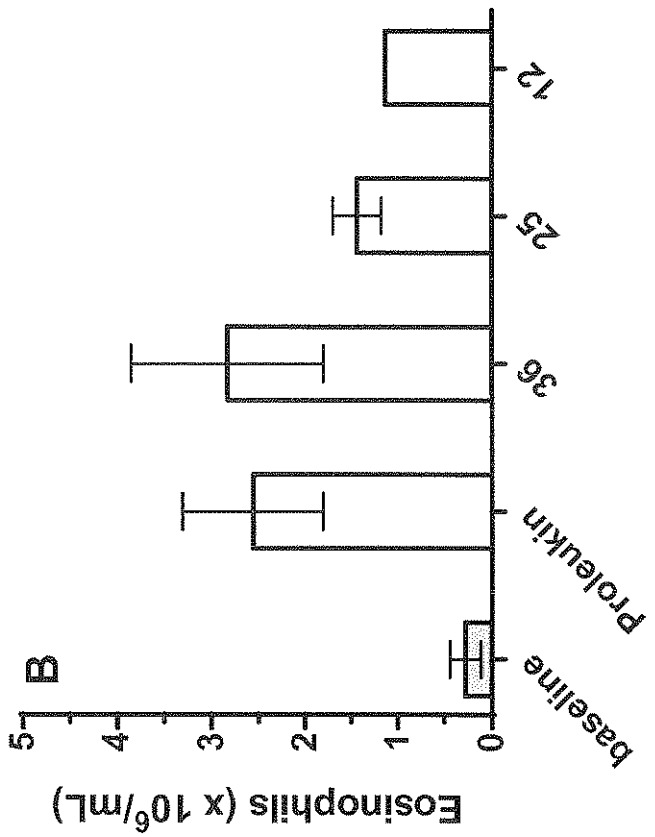

Treg number
(fold change)
0
1
2
3
4
5
0
10
20
30
40
dose in µg/kg
IgG-(IL-2)2
IgG-IL-2

Figure 18


B
Eosinophils (x 10^6/mL)
0
1
2
3
4
5
baseline
Proleukin
36
25
12

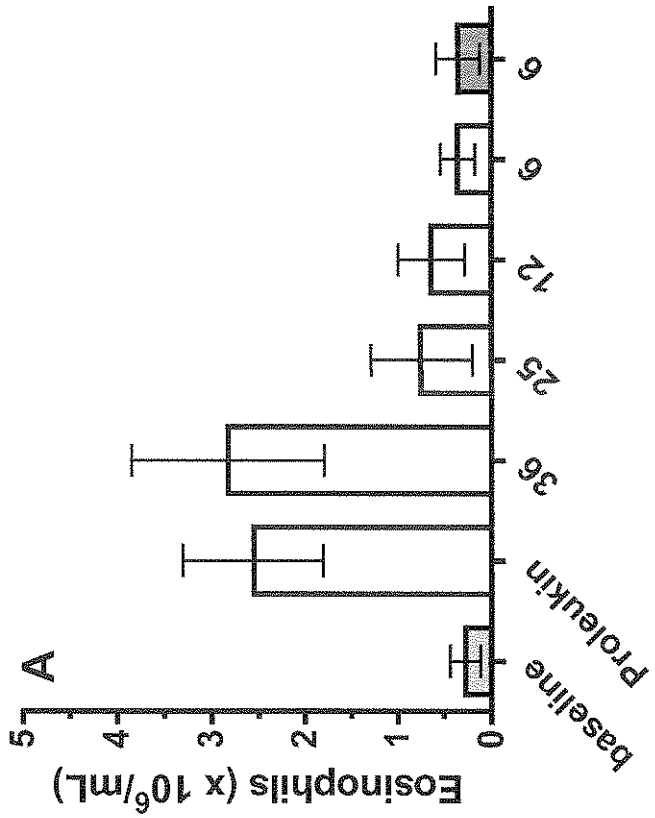

A
Eosinophils (x 10^6/mL)
0
1
2
3
4
5
baseline
Proleukin
36
25
12
6
6

Figure 19

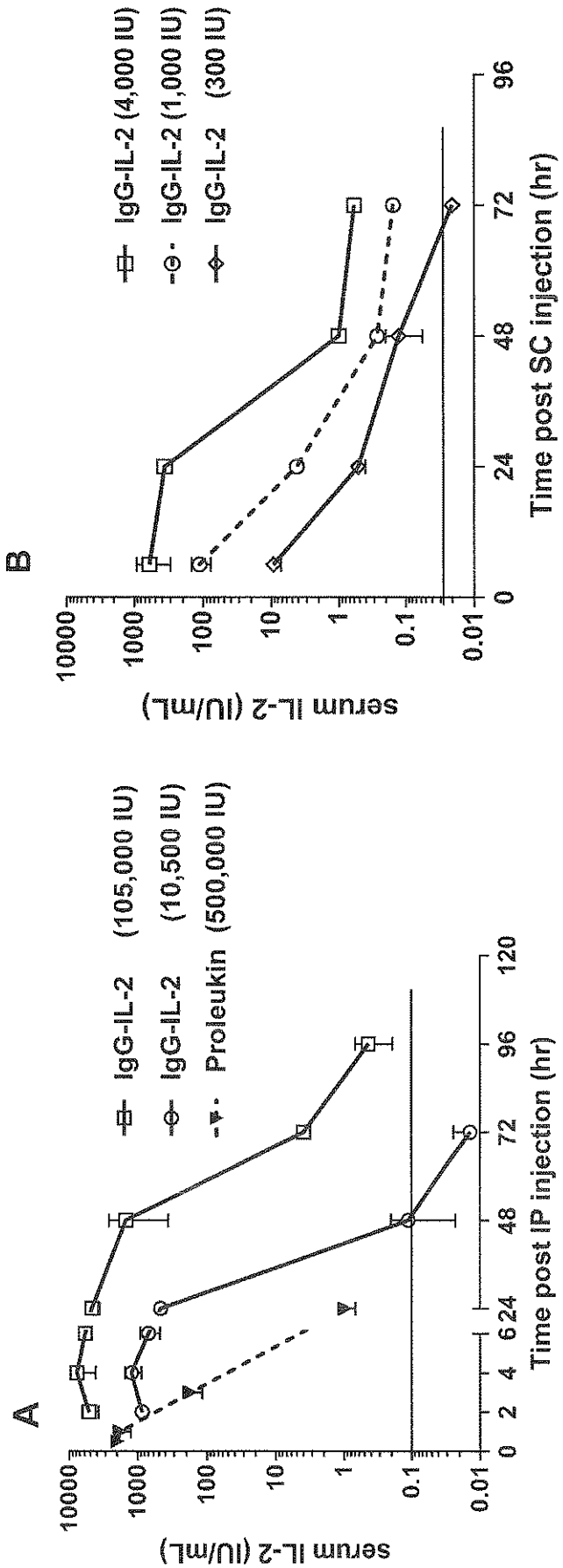

A
B
serum IL-2 (IU/mL)
Time post IP injection (hr)
Time post SC injection (hr)
IgG-IL-2 (105,000 IU)
IgG-IL-2 (10,500 IU)
Proleukin (500,000 IU)
IgG-IL-2 (4,000 IU)
IgG-IL-2 (1,000 IU)
IgG-IL-2 (300 IU)
10000
1000
100
10
1
0.1
0.01
0
2
4
6
24
48
72
96
120

Figure 20

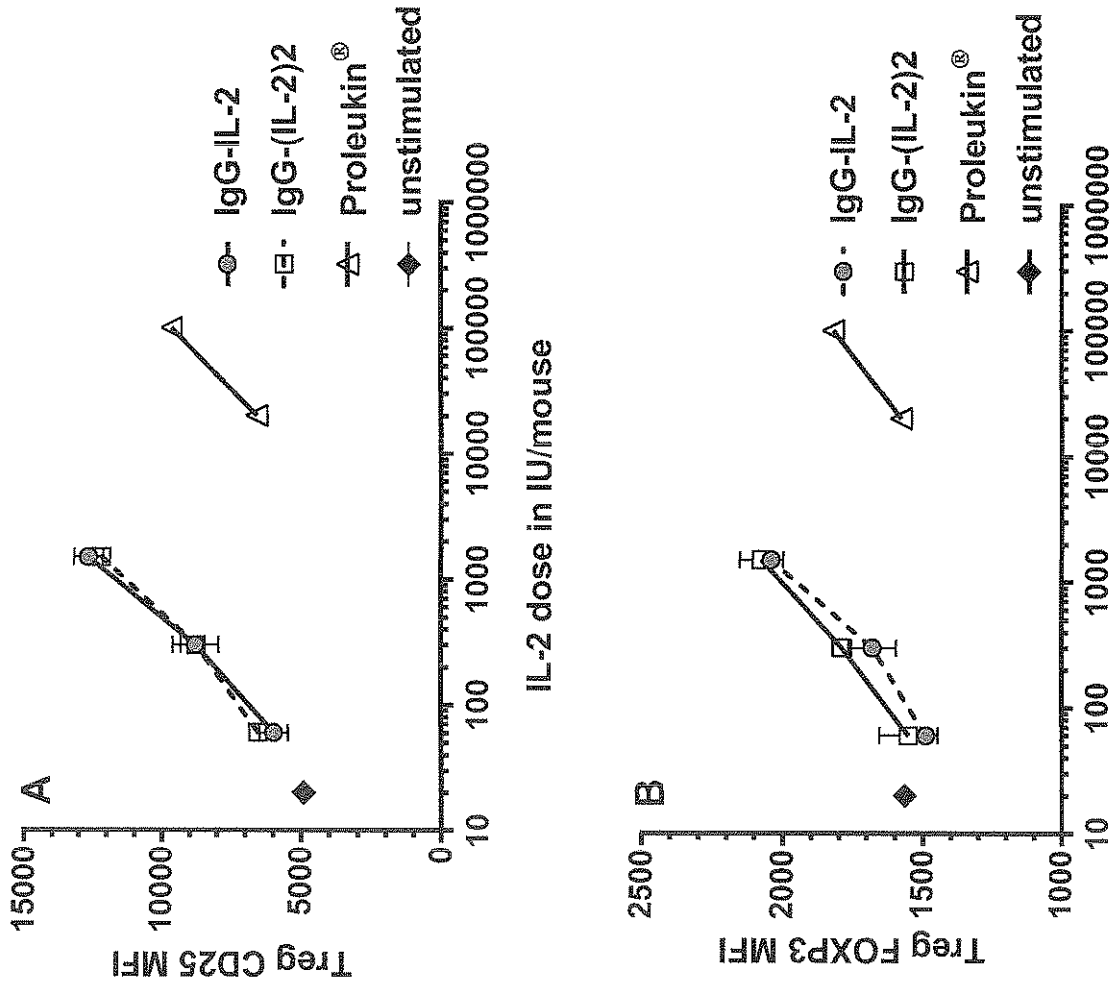

A
Treg CD25 MFI
15000
10000
5000
0
10
100
1000
10000
100000
1000000
IL-2 dose in IU/mouse
IgG-IL-2
IgG-(IL-2)2
Proleukin®
unstimulated
B
Treg FOXP3 MFI
2500
2000
1500
1000
10
100
1000
10000
100000
1000000
IL-2 dose in IU/mouse
IgG-IL-2
IgG-(IL-2)2
Proleukin®
unstimulated

Figure 21

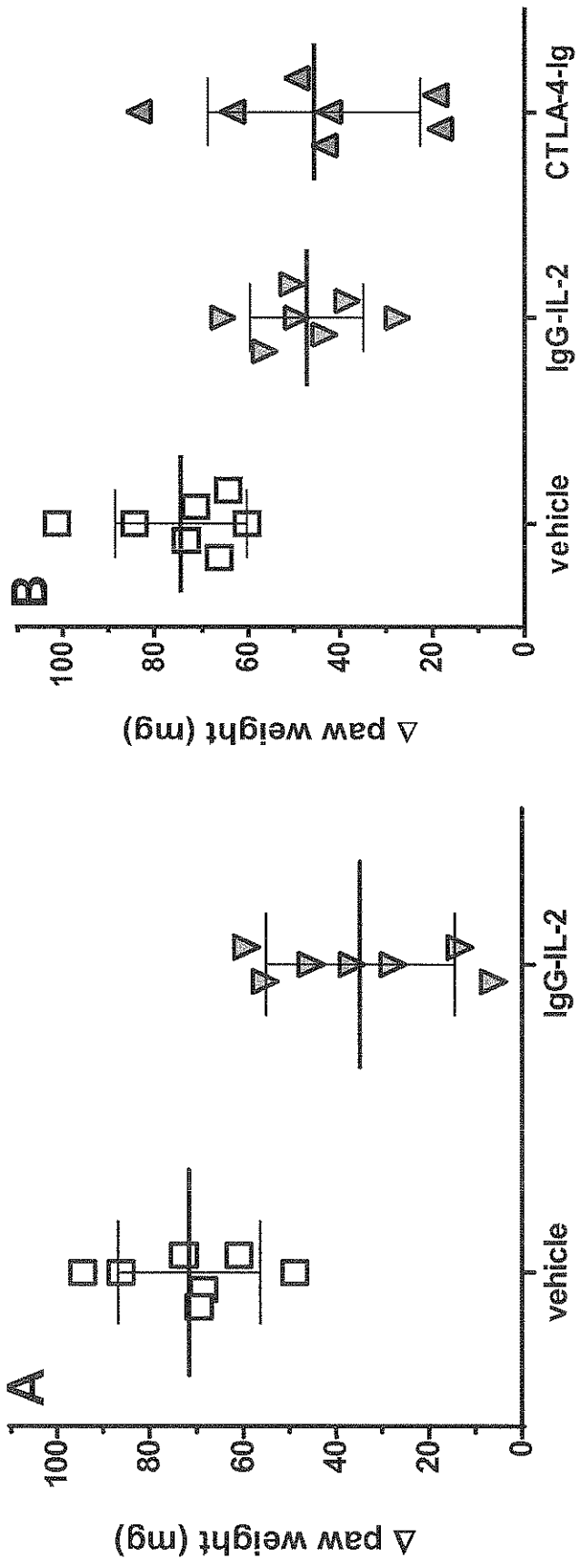


Figure 22

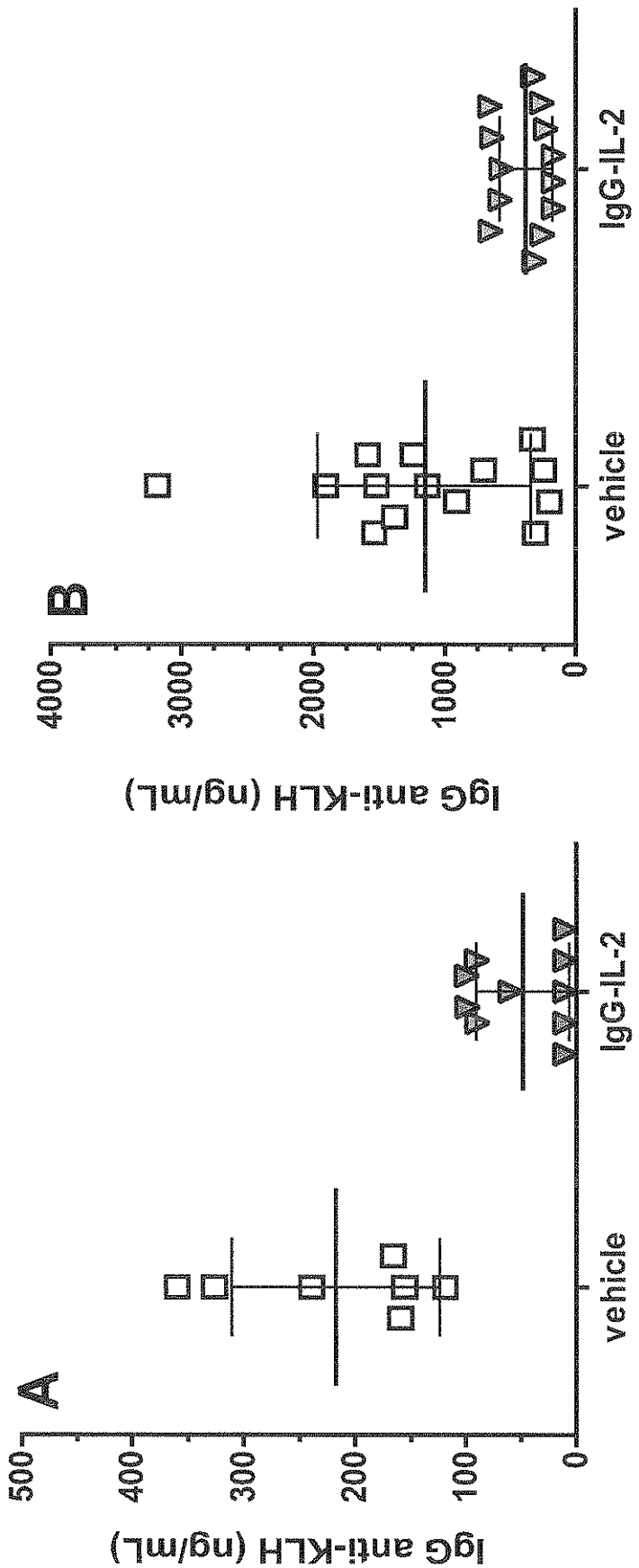

A
B
IgG anti-KLH (ng/mL)
500
400
300
200
100
0
4000
3000
2000
1000
vehicle
IgG-IL-2

Figure 23