FIGURE 1

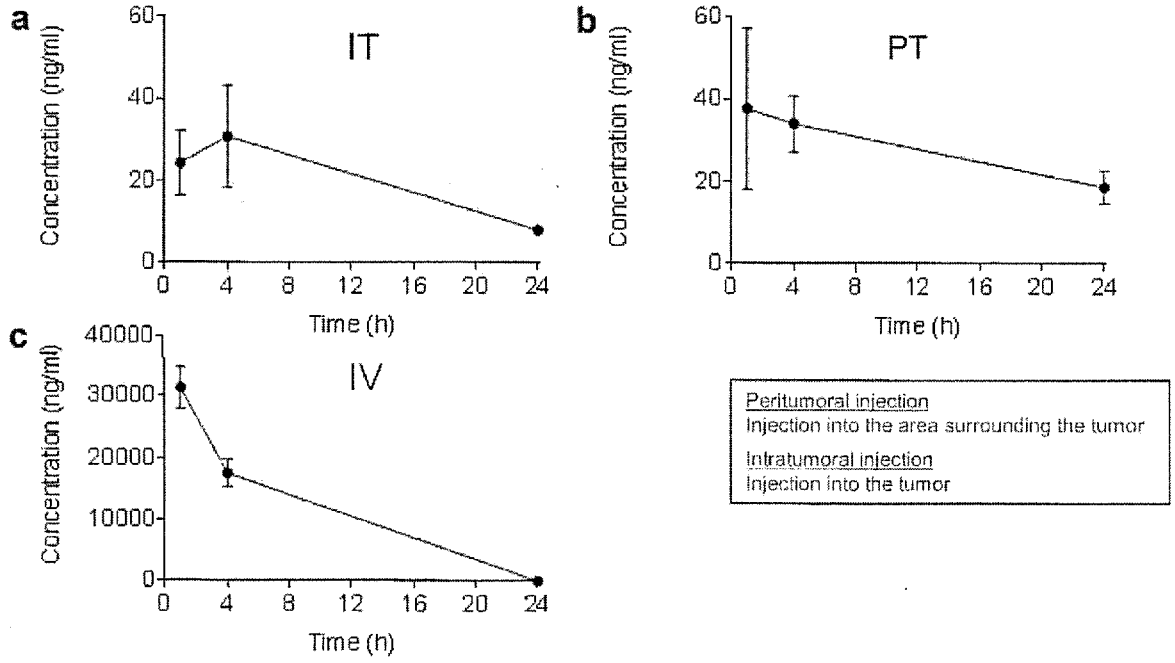

a
IT
Concentration (ng/ml)
60
40
20
0
0 4 8 12 16 20 24
Time (h)
b
PT
Concentration (ng/ml)
60
40
20
0
0 4 8 12 16 20 24
Time (h)
c
IV
Concentration (ng/ml)
40000
30000
20000
10000
0
0 4 8 12 16 20 24
Time (h)
Peritumoral injection
Injection into the area surrounding the tumor
Intratumoral injection
Injection into the tumor

FIGURE 2

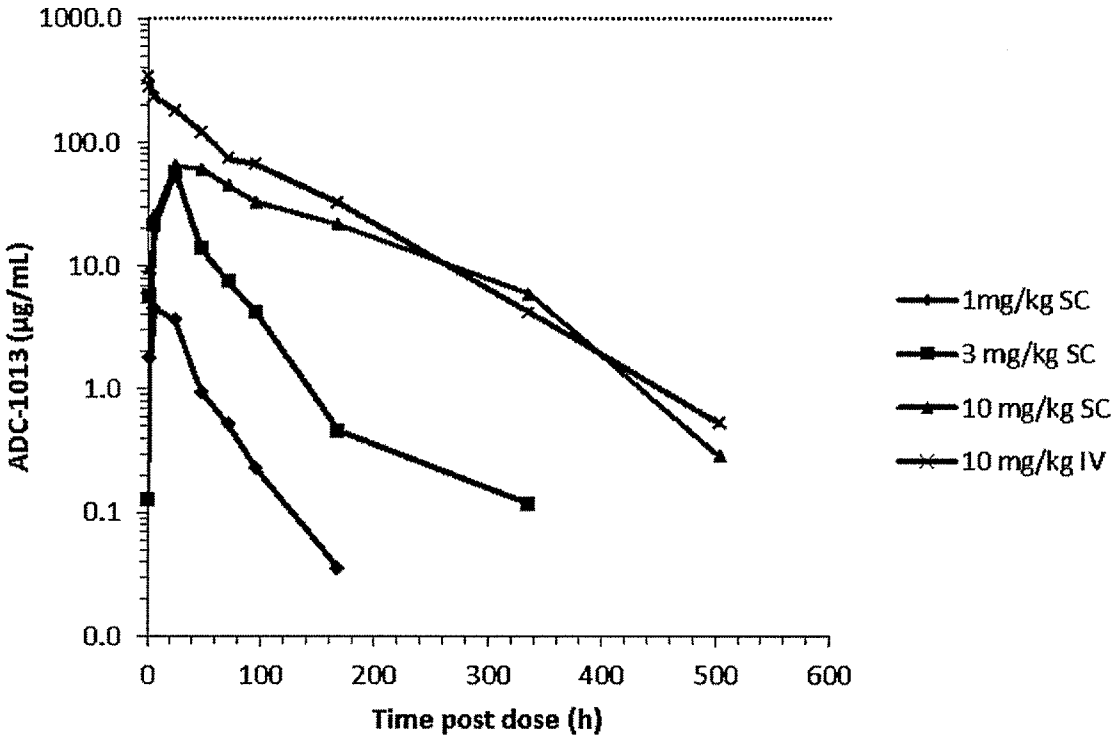
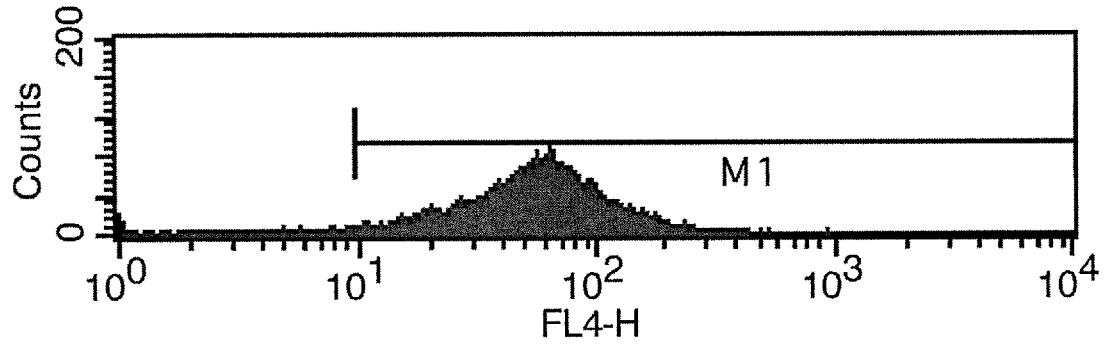

1000.0
100.0
10.0
1.0
0.1
0.0
ADC-1013 (µg/mL)
0
100
200
300
400
500
600
Time post dose (h)
1mg/kg SC
3 mg/kg SC
10 mg/kg SC
10 mg/kg IV

FIGURE 3

A


Counts
200
0
M1
10^0
10^1
10^2
10^3
10^4
FL4-H

B

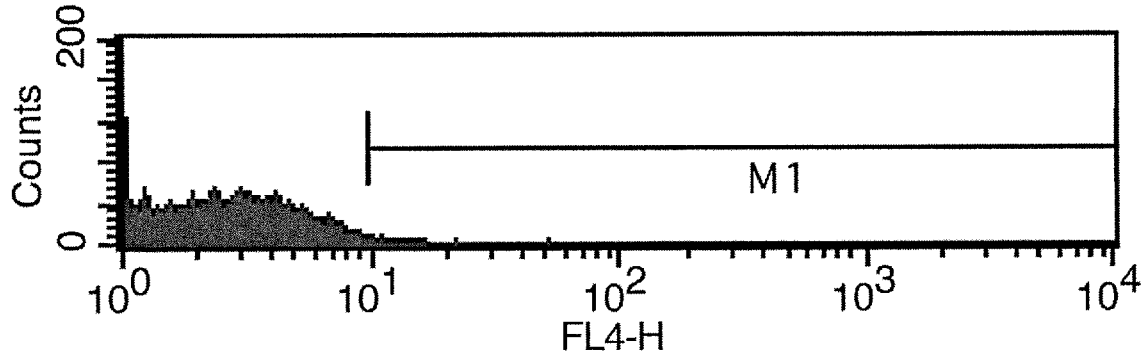

Counts
200
0
M1
10^0
10^1
10^2
10^3
10^4
FL4-H

FIGURE 4

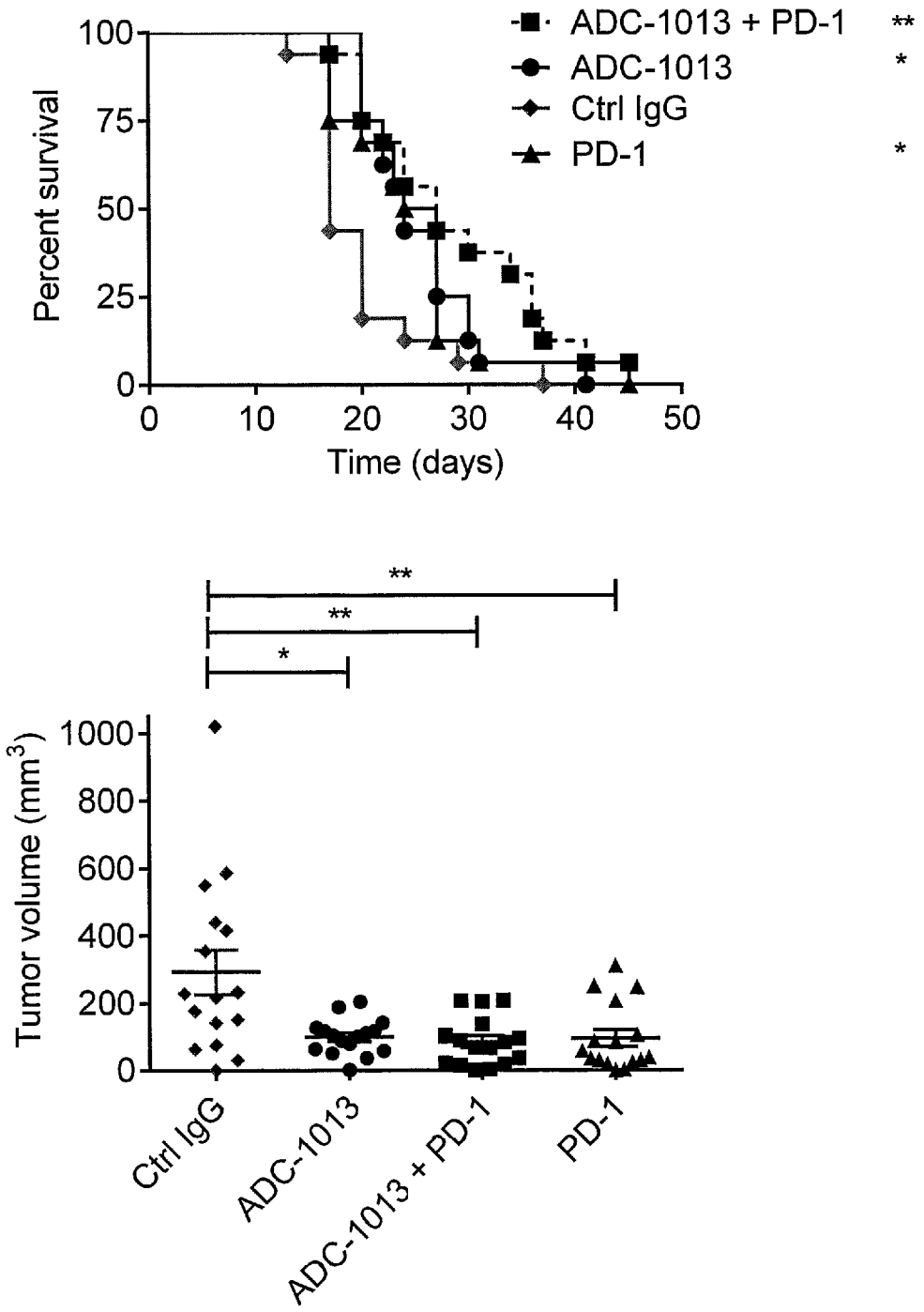

ADC-1013 + PD-1 **
ADC-1013 *
Ctrl IgG
PD-1 *
Percent survival
100
75
50
25
0
0
10
20
30
40
50
Time (days)
**
**
*
Tumor volume (mm3)
1000
800
600
400
200
0
Ctrl IgG
ADC-1013
ADC-1013 + PD-1
PD-1

FIGURE 5A

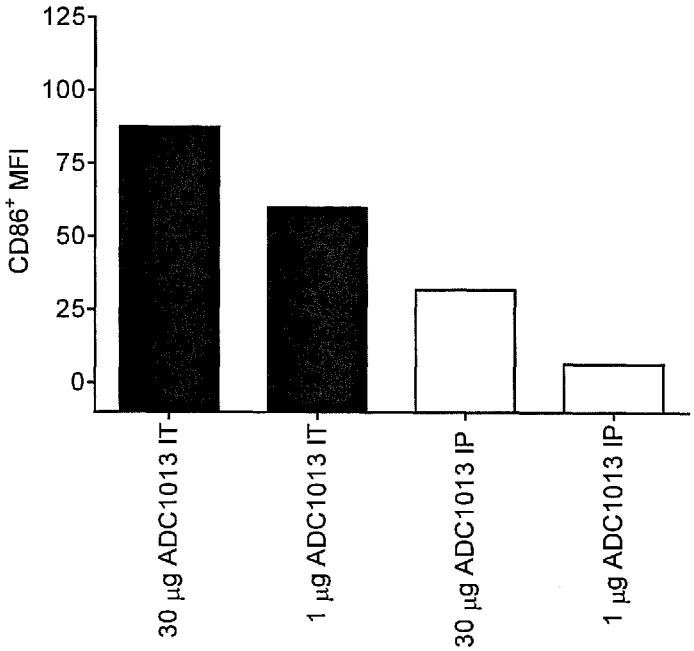


FIGURE 5B

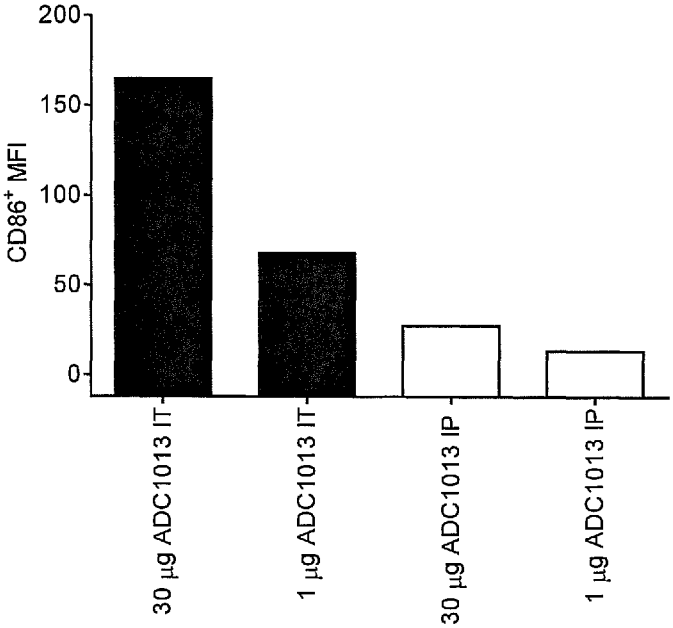


FIGURE 6

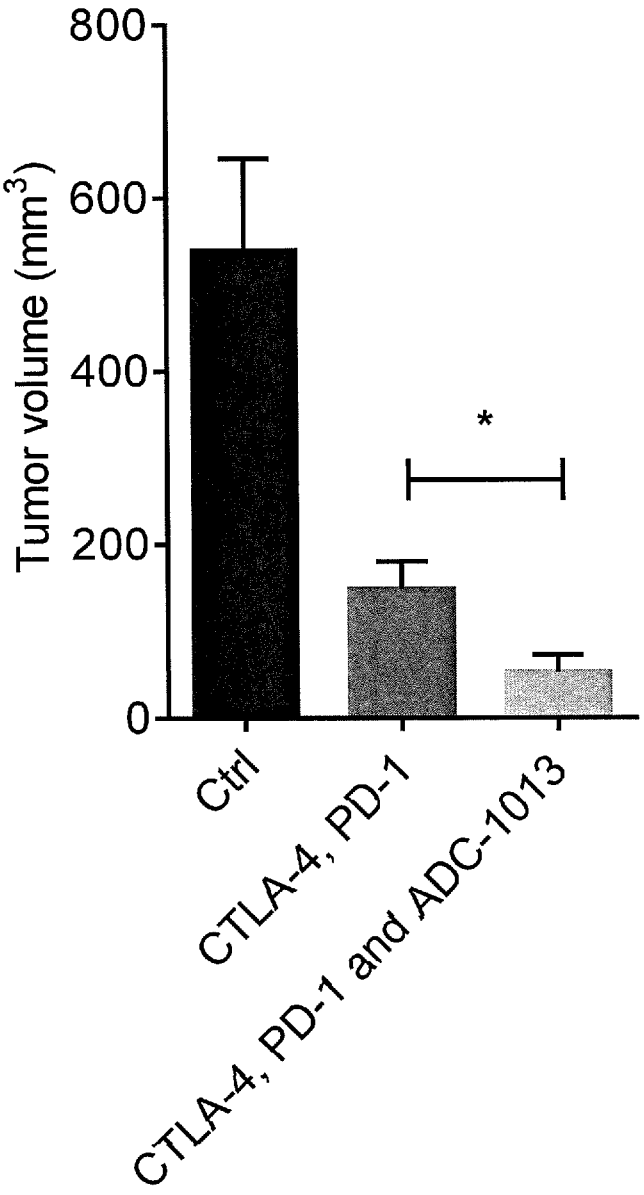

800
600
400
200
0
Tumor volume (mm3)
*
Ctrl
CTLA-4, PD-1
CTLA-4, PD-1 and ADC-1013

FIGURE 7

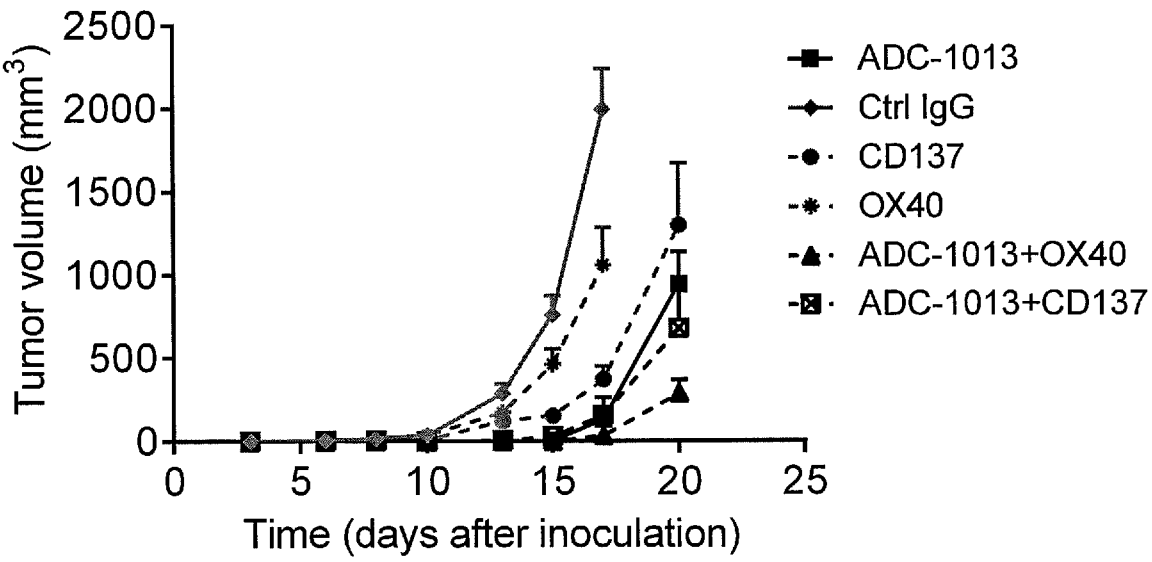
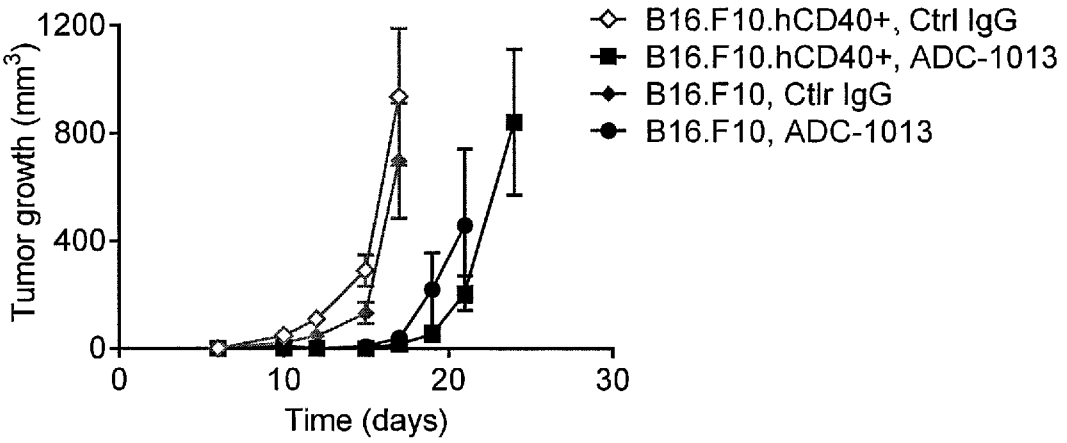

Tumor volume (mm3)
2500
2000
1500
1000
500
0
0
5
10
15
20
25
Time (days after inoculation)
ADC-1013
Ctrl IgG
CD137
OX40
ADC-1013+OX40
ADC-1013+CD137

FIGURE 8


Tumor growth (mm3)
1200
800
400
0
0
10
20
30
Time (days)
B16.F10.hCD40+, Ctrl IgG
B16.F10.hCD40+, ADC-1013
B16.F10, Ctlr IgG
B16.F10, ADC-1013

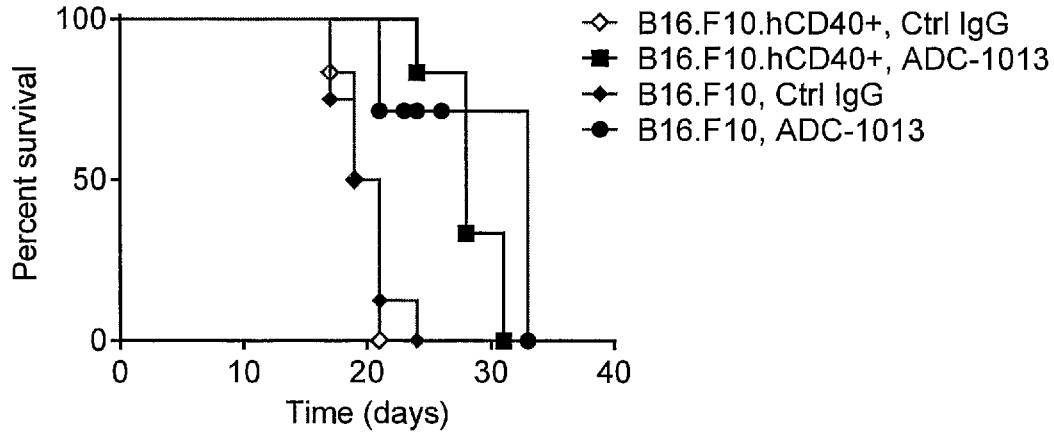
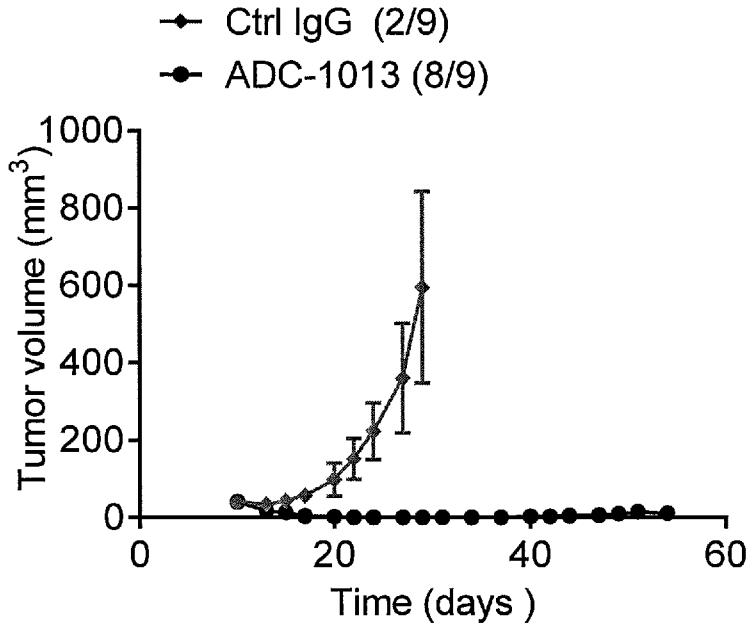

Percent survival
100
50
0
0
10
20
30
40
Time (days)
B16.F10.hCD40+, Ctrl IgG
B16.F10.hCD40+, ADC-1013
B16.F10, Ctrl IgG
B16.F10, ADC-1013

FIGURE 9


Ctrl IgG (2/9)
ADC-1013 (8/9)
Tumor volume (mm3)
1000
800
600
400
200
0
0
20
40
60
Time (days)

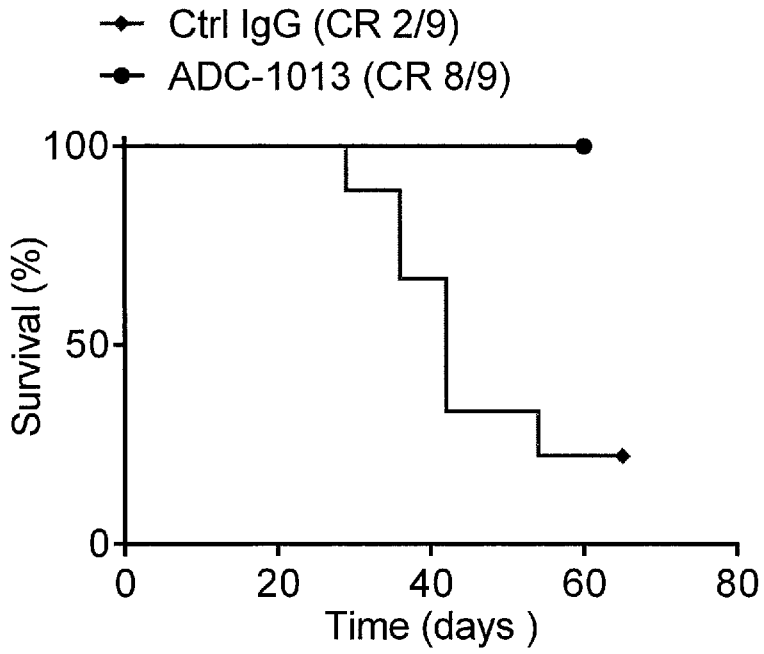

Ctrl IgG (CR 2/9)
ADC-1013 (CR 8/9)
Survival (%)
100
50
0
0
20
40
60
80
Time (days)

FIGURE 10

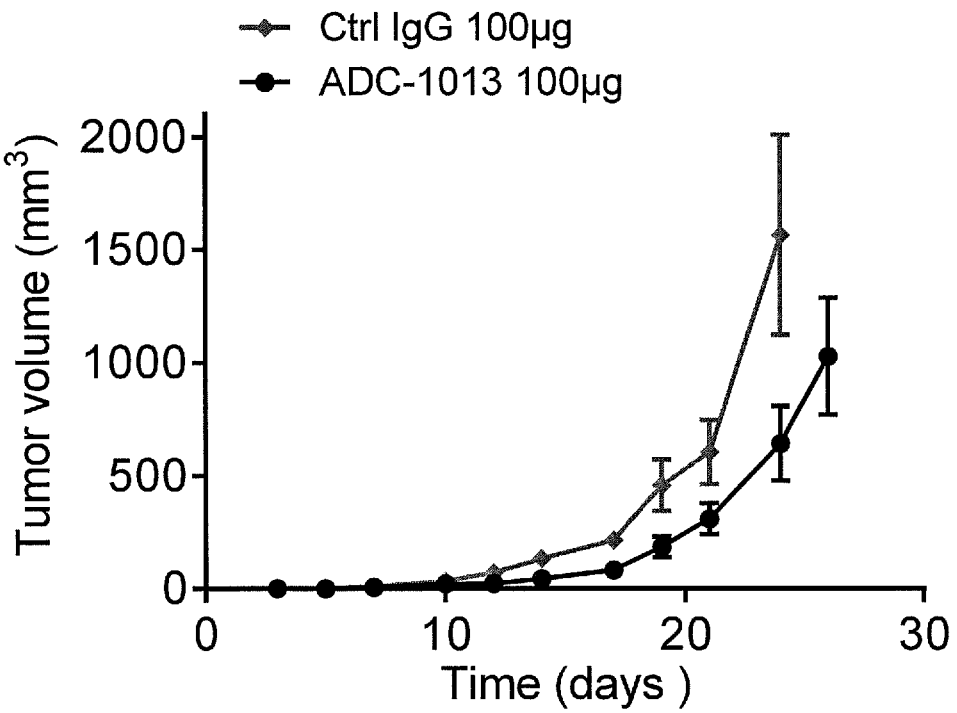

Ctrl IgG 100µg
ADC-1013 100µg
Tumor volume (mm3)
2000
1500
1000
500
0
0
10
20
30
Time (days)

FIGURE 11

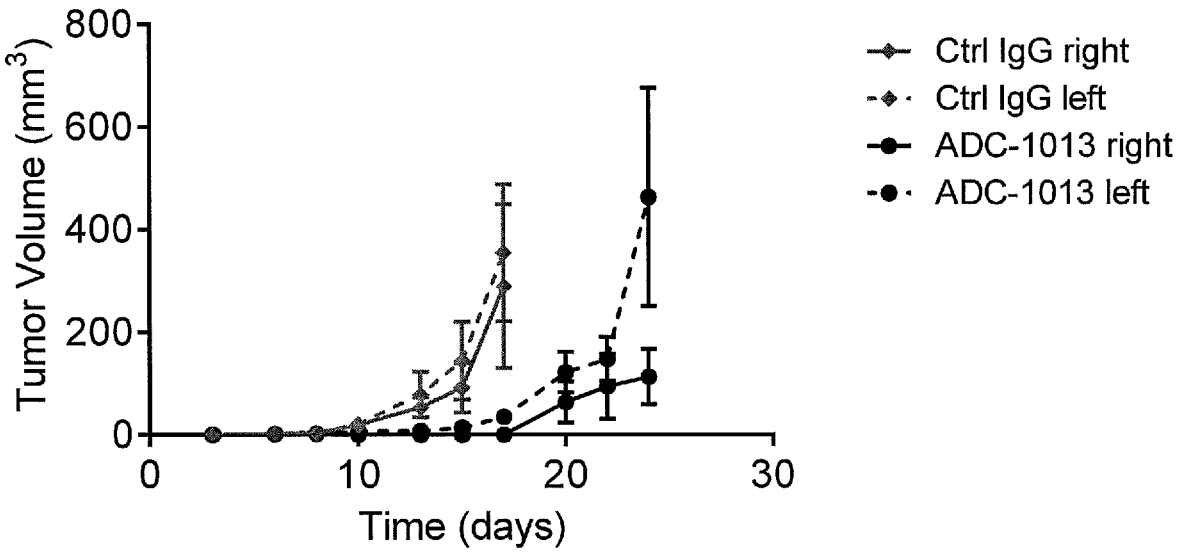

Tumor Volume (mm3)
800
600
400
200
0
0
10
20
30
Time (days)
Ctrl IgG right
Ctrl IgG left
ADC-1013 right
ADC-1013 left

FIGURE 12

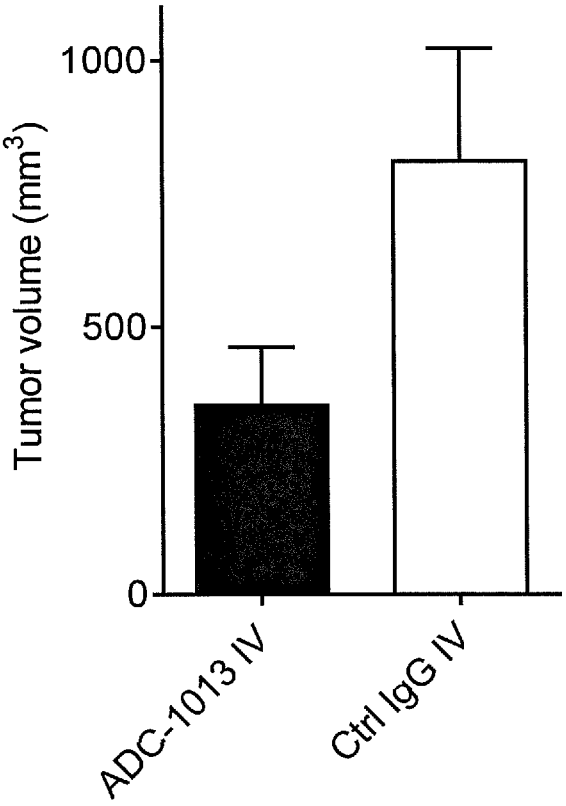
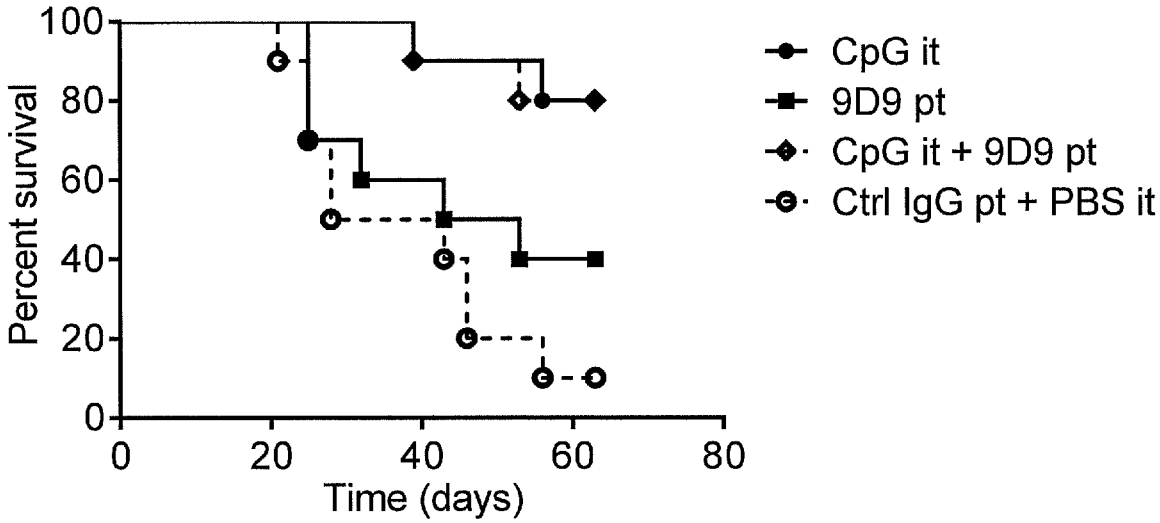


FIGURE 13


100
80
60
40
20
0
Percent survival
0
20
40
60
80
Time (days)
CpG it
9D9 pt
CpG it + 9D9 pt
Ctrl IgG pt + PBS it