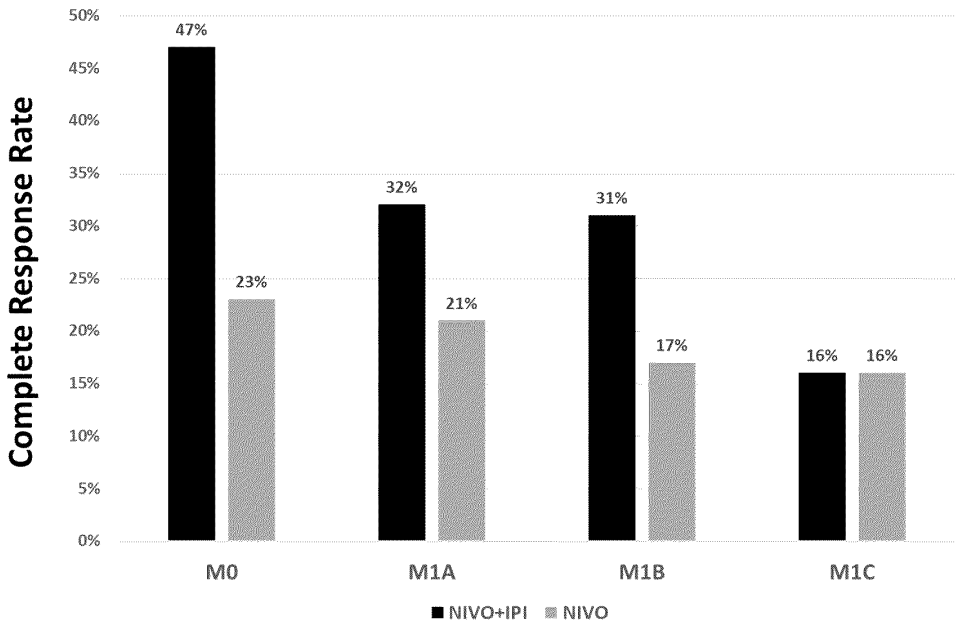
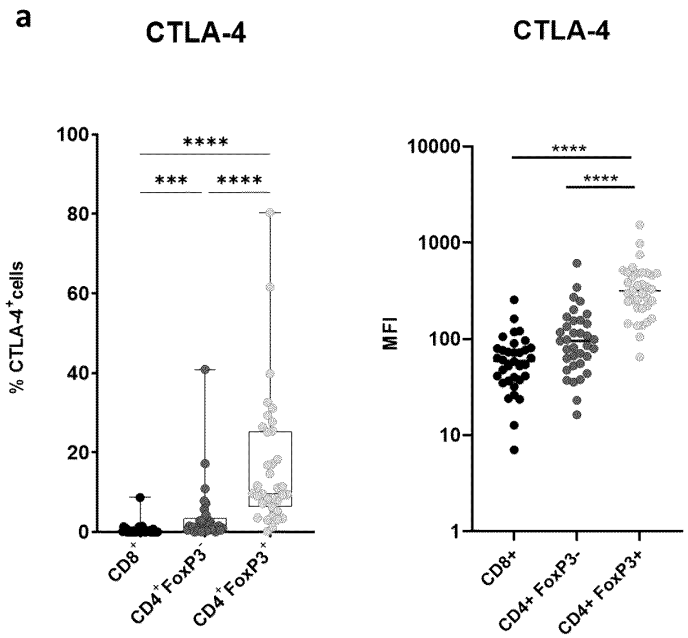

Complete Response Rate
50%
45%
40%
35%
30%
25%
20%
15%
10%
5%
0%
47%
23%
32%
21%
31%
17%
16%
16%
M0
M1A
M1B
M1C
NIVO+IPI
NIVO

FIGURE 1


a
CTLA-4
CTLA-4
% CTLA-4+cells
MFI
****
***
****
****
****
CD8+
CD4+FoxP3-
CD4+FoxP3+
CD8+
CD4+ FoxP3-
CD4+ FoxP3+

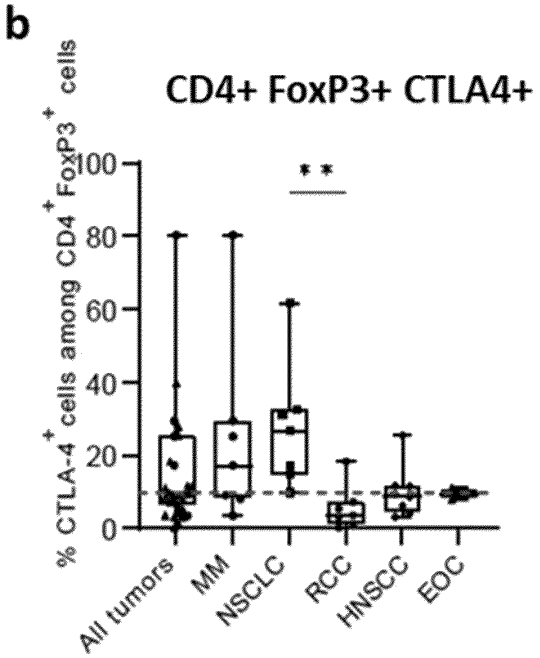

b
CD4+ FoxP3+ CTLA4+
% CTLA-4+ cells among CD4+FoxP3+ cells
**
All tumors
MM
NSCLC
RCC
HNSCC
EOC

FIGURE 2

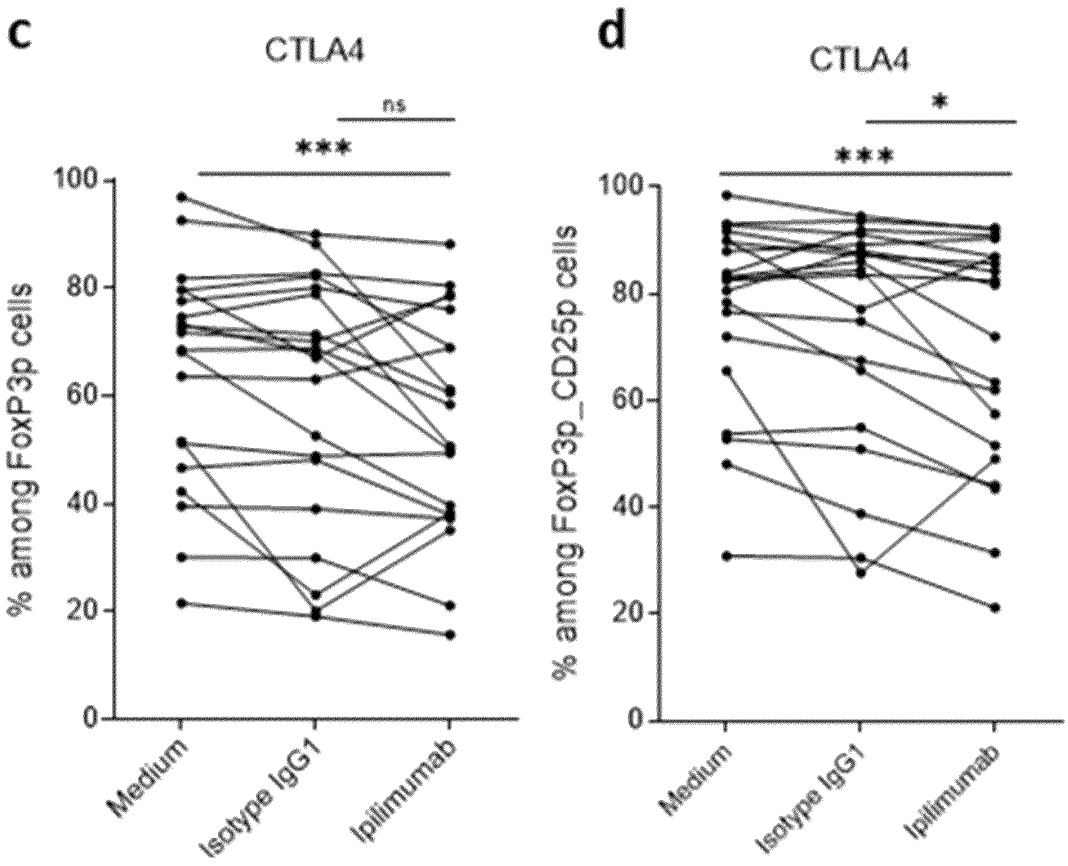


FIGURE 2 (Following)

e

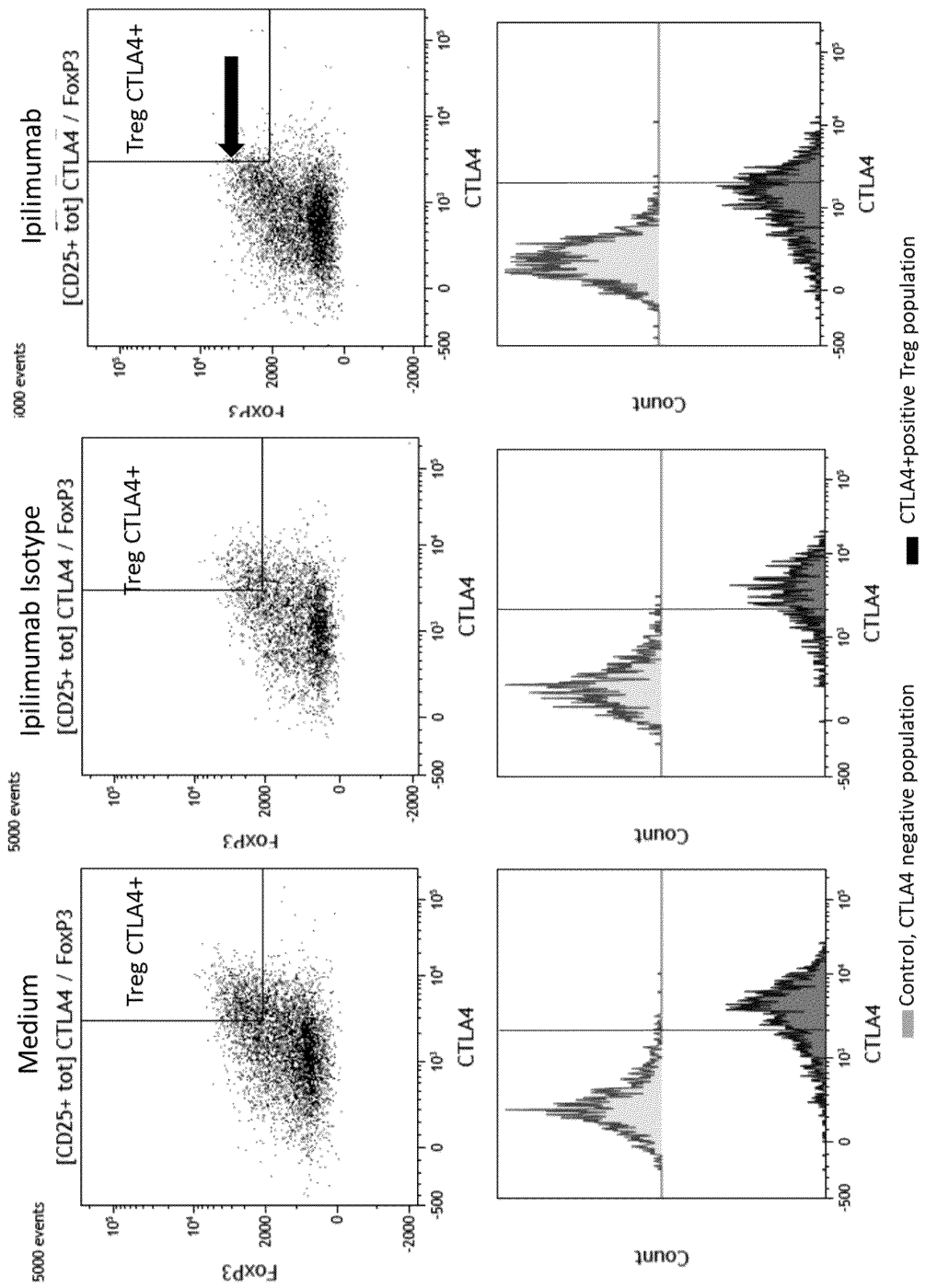


FIGURE 2 (Following)

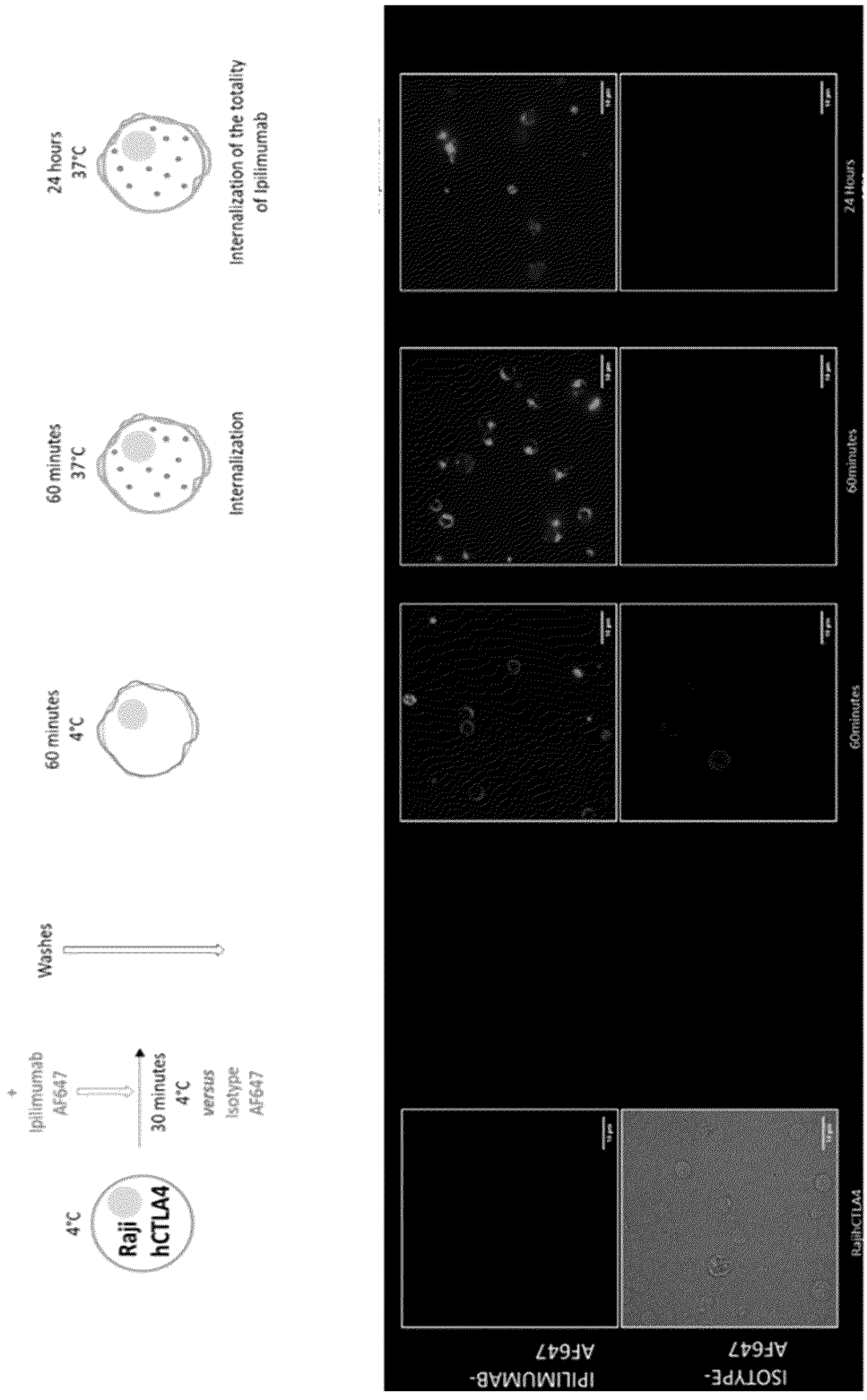

4°C
Raji hCTLA4
+ Ipilimumab AF647
30 minutes 4°C
versus Isotype AF647
Washes
60 minutes 4°C
60 minutes 37°C
Internalization
24 hours 37°C
Internalization of the totality of Ipilimumab
ISOTYPE-AF647
IPILIMUMAB-AF647
RajiHCTLA4
60minutes
60minutes
24 Hours

FIGURE 3

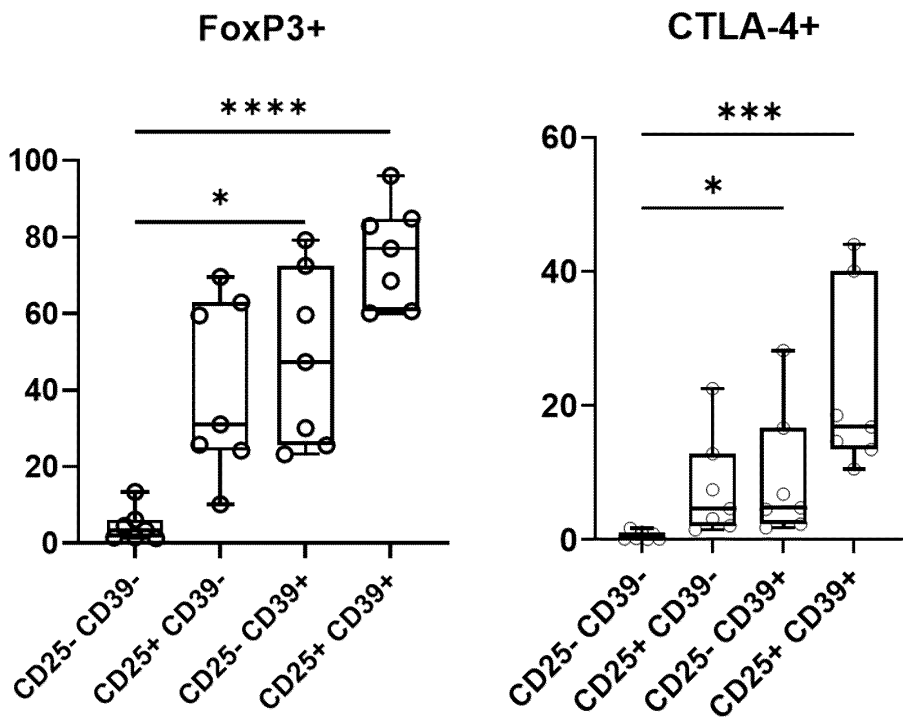

FoxP3+
****
*
100
80
60
40
20
0
CD25- CD39-
CD25+ CD39-
CD25- CD39+
CD25+ CD39+
CTLA-4+
***
*
60
40
20
0
CD25- CD39-
CD25+ CD39-
CD25- CD39+
CD25+ CD39+

FIGURE 4

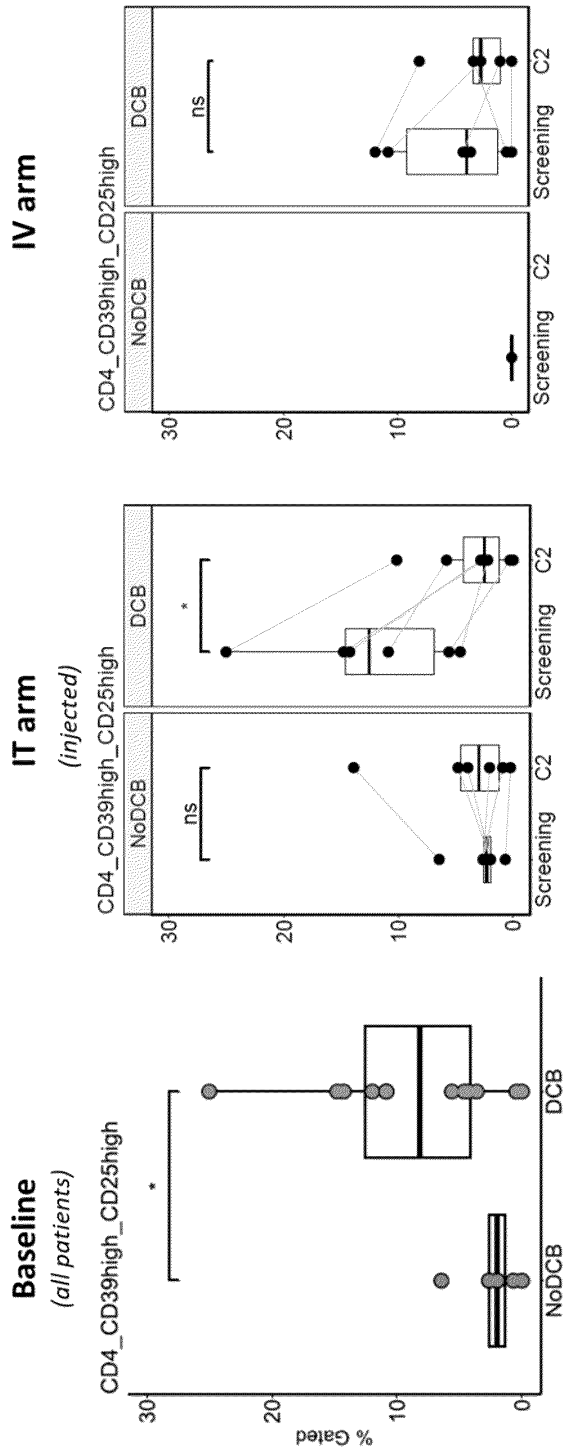

Baseline
(all patients)
IT arm
(injected)
IV arm
CD4_CD39high_CD25high
% Gated
NoDCB
DCB
Screening
C2
ns
*
0
10
20
30

FIGURE 5

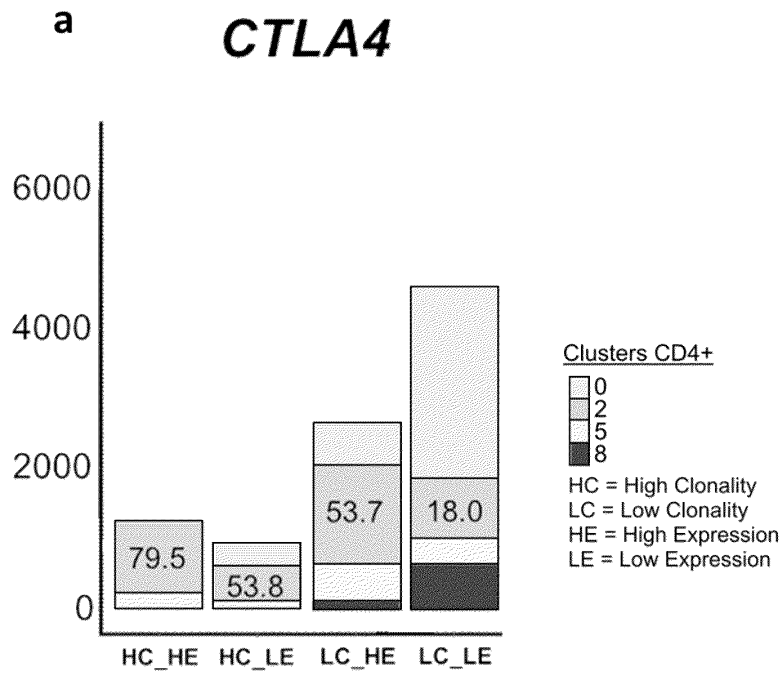
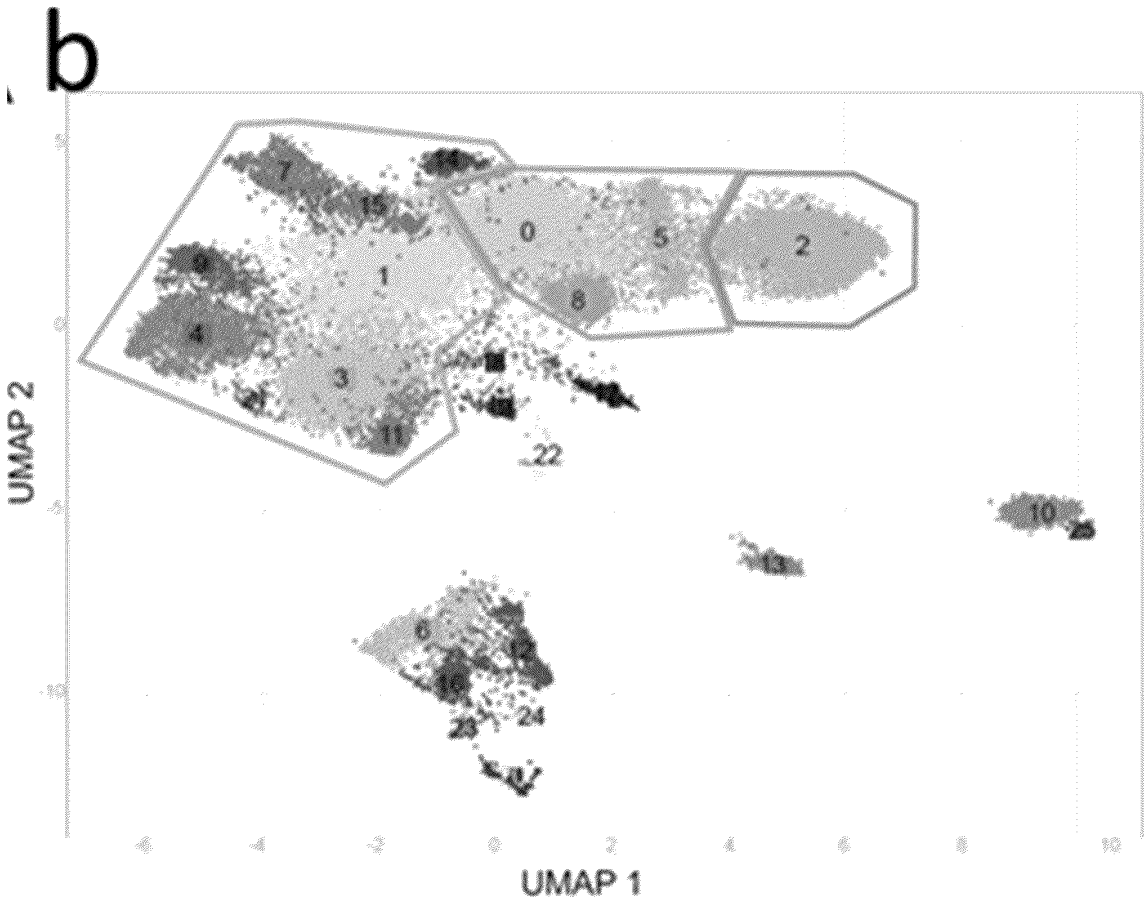

a
CTLA4
6000
4000
2000
0
79.5
53.8
53.7
18.0
HC_HE
HC_LE
LC_HE
LC_LE
Clusters CD4+
0
2
5
8
HC = High Clonality
LC = Low Clonality
HE = High Expression
LE = Low Expression

FIGURE 6



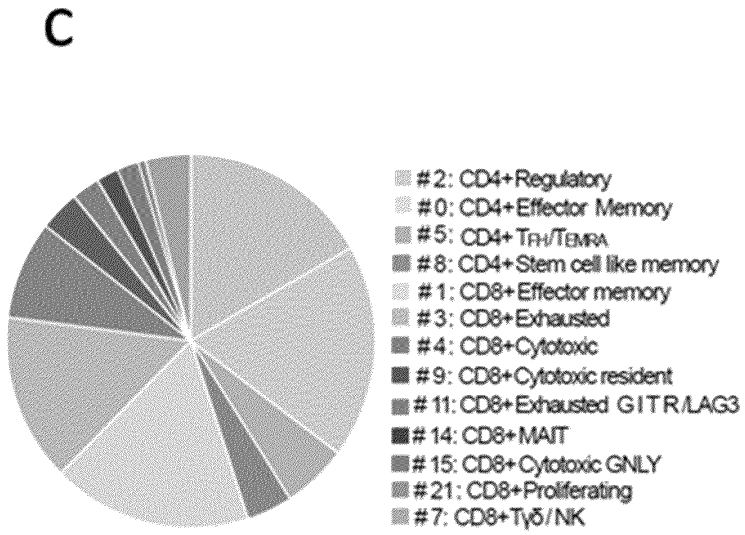


FIGURE 6 (Following)

a

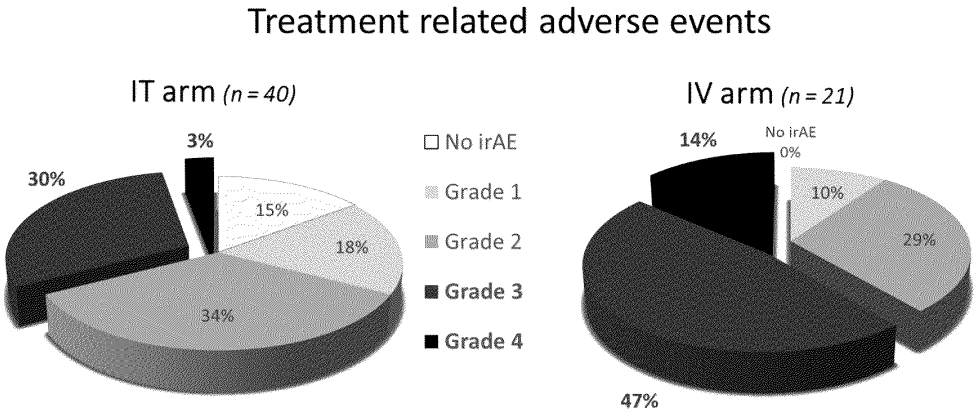

Treatment related adverse events
IT arm (n = 40)
IV arm (n = 21)
3%
30%
15%
18%
34%
No irAE
Grade 1
Grade 2
Grade 3
Grade 4
14%
No irAE
0%
10%
29%
47%

FIGURE 7

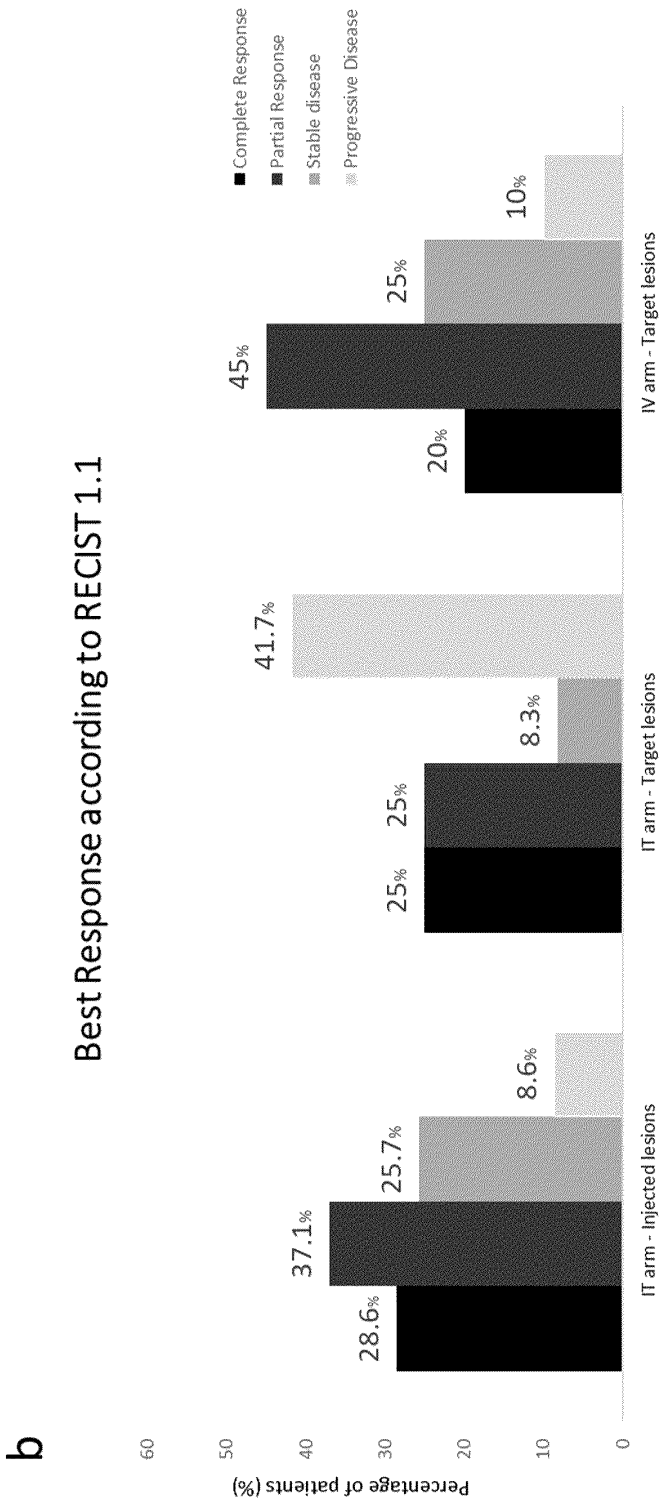


FIGURE 7 (Following)

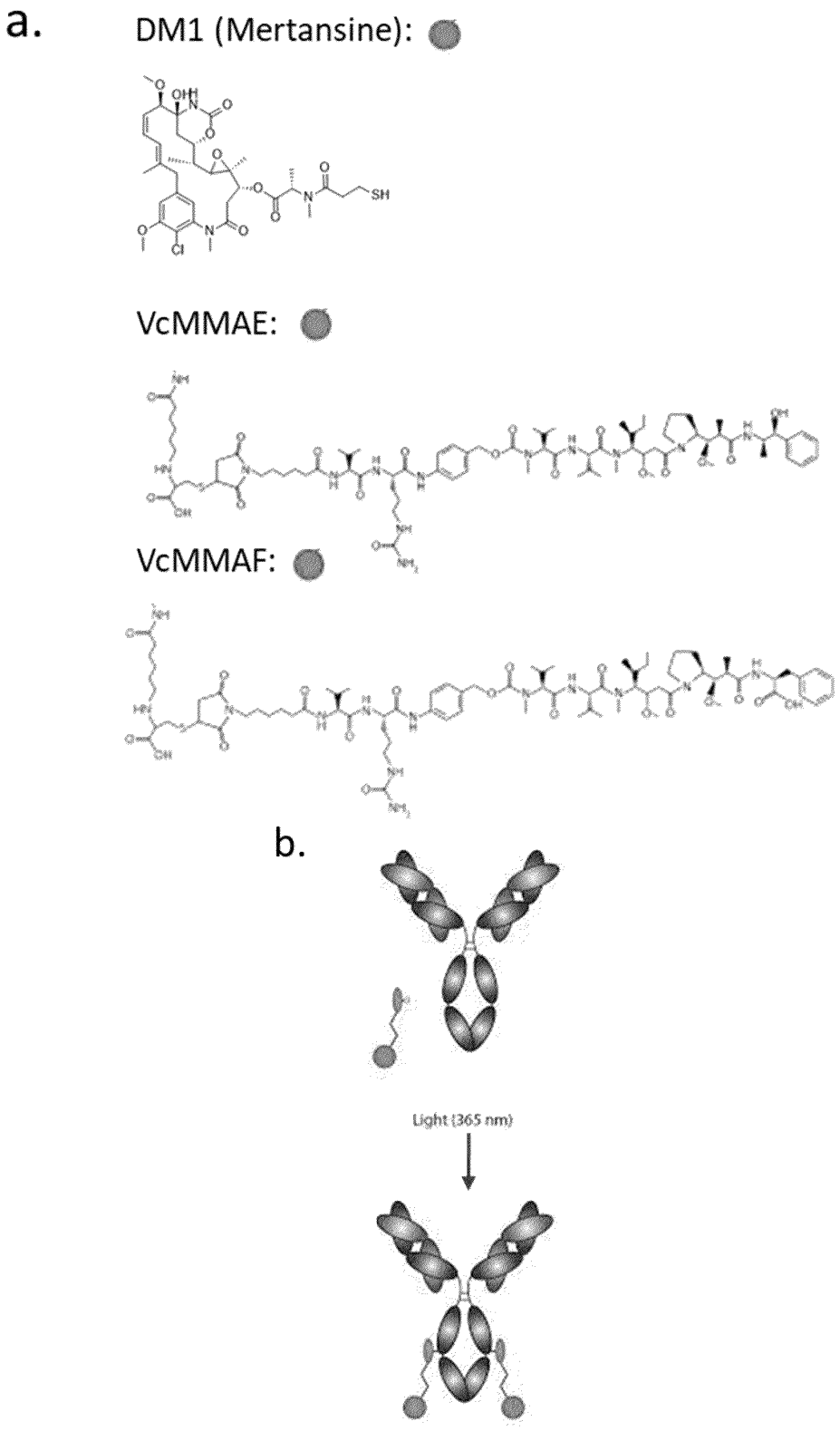


FIGURE 8

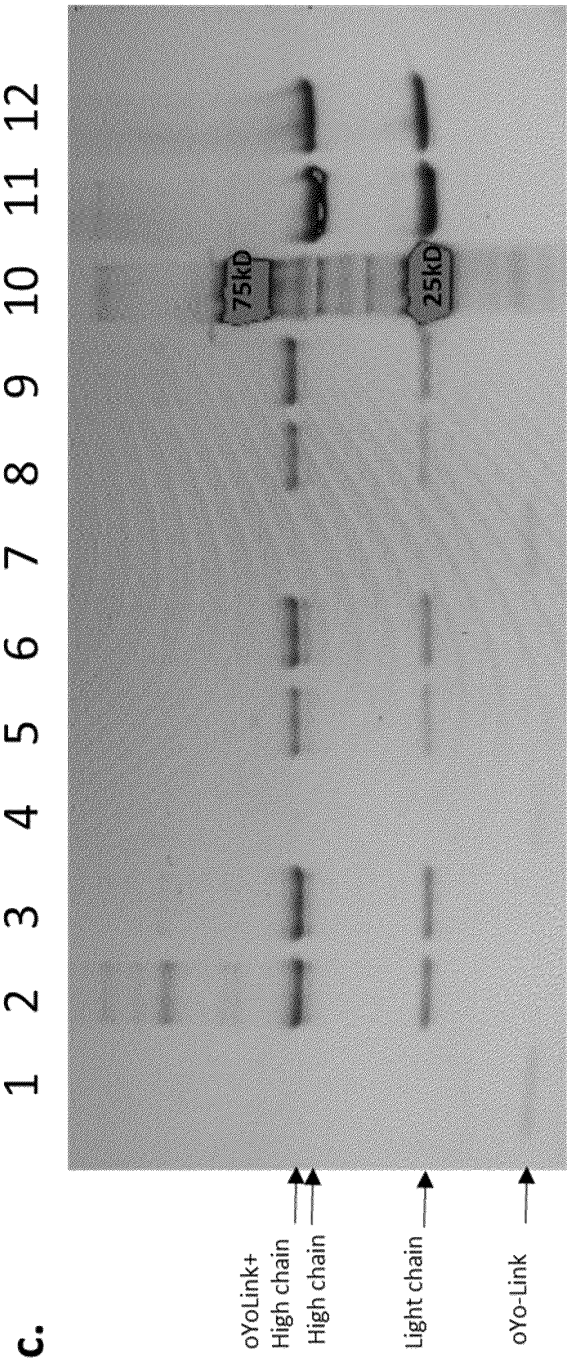
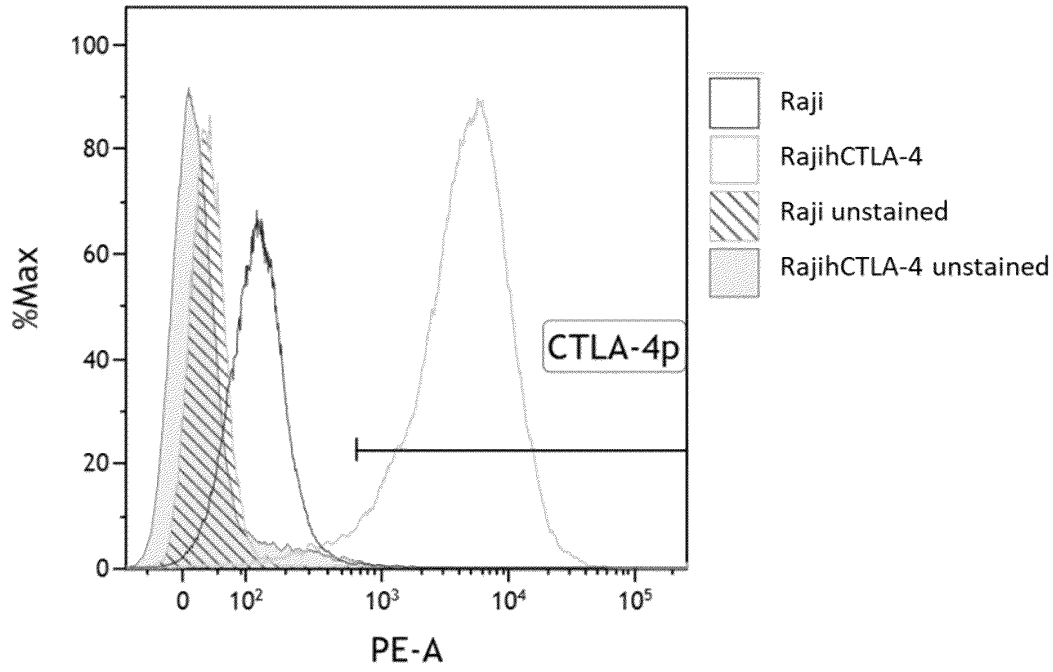


FIGURE 8 (Following)



Marker	%Gated
CTLA-4p	0,55
CTLA-4p	95,55
CTLA-4p	0,12
CTLA-4p	0,58

FIGURE 9

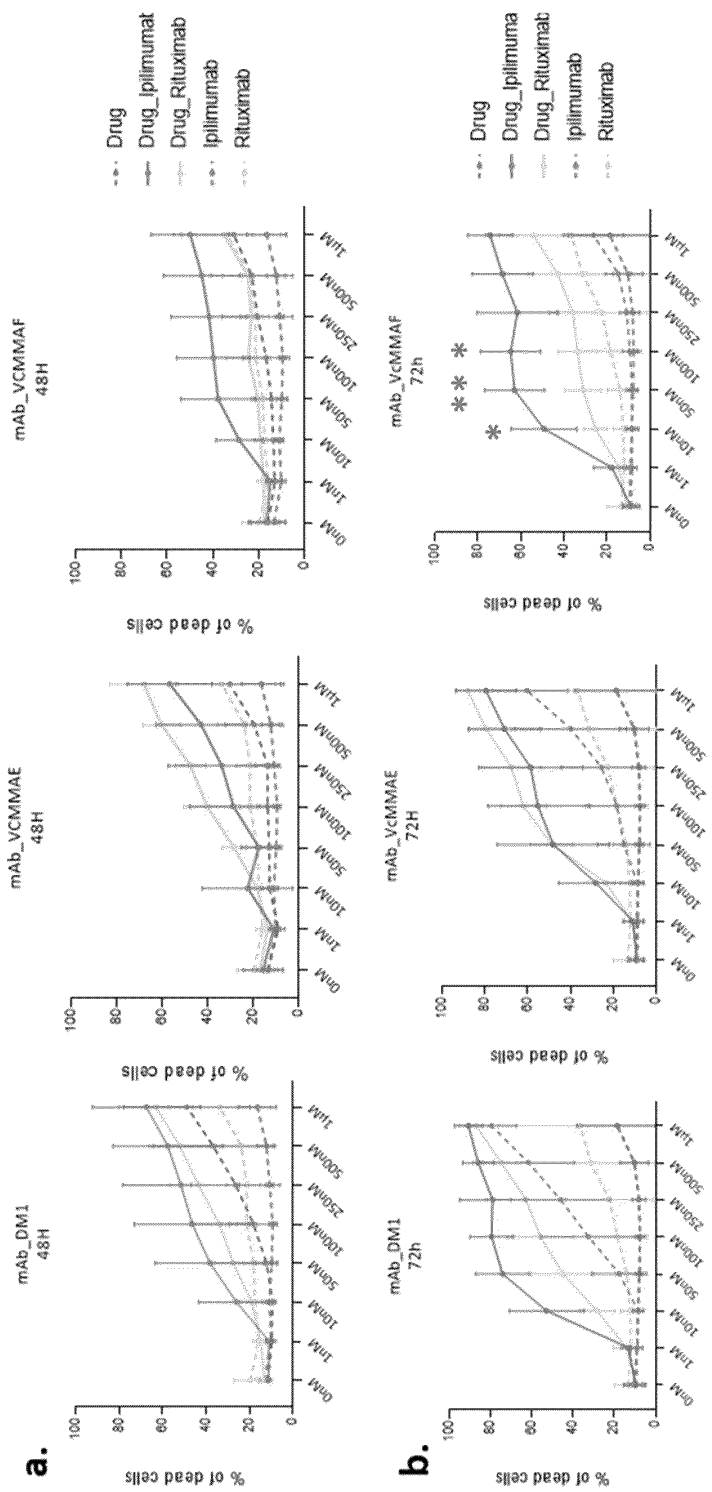

a.
b.
mAb_DM1
48H
mAb_VCMMAE
48H
mAb_VCMMAF
48H
mAb_DM1
72h
mAb_VcMMAE
72H
mAb_VcMMAF
72h
% of dead cells
Drug
Drug_Ipilimumat
Drug_Rituximab
Ipilimumab
Rituximab
Drug_Ipilimuma
Drug_Rituximat
0nM
1nM
10nM
50nM
100nM
250nM
500nM
1µM

FIGURE 10

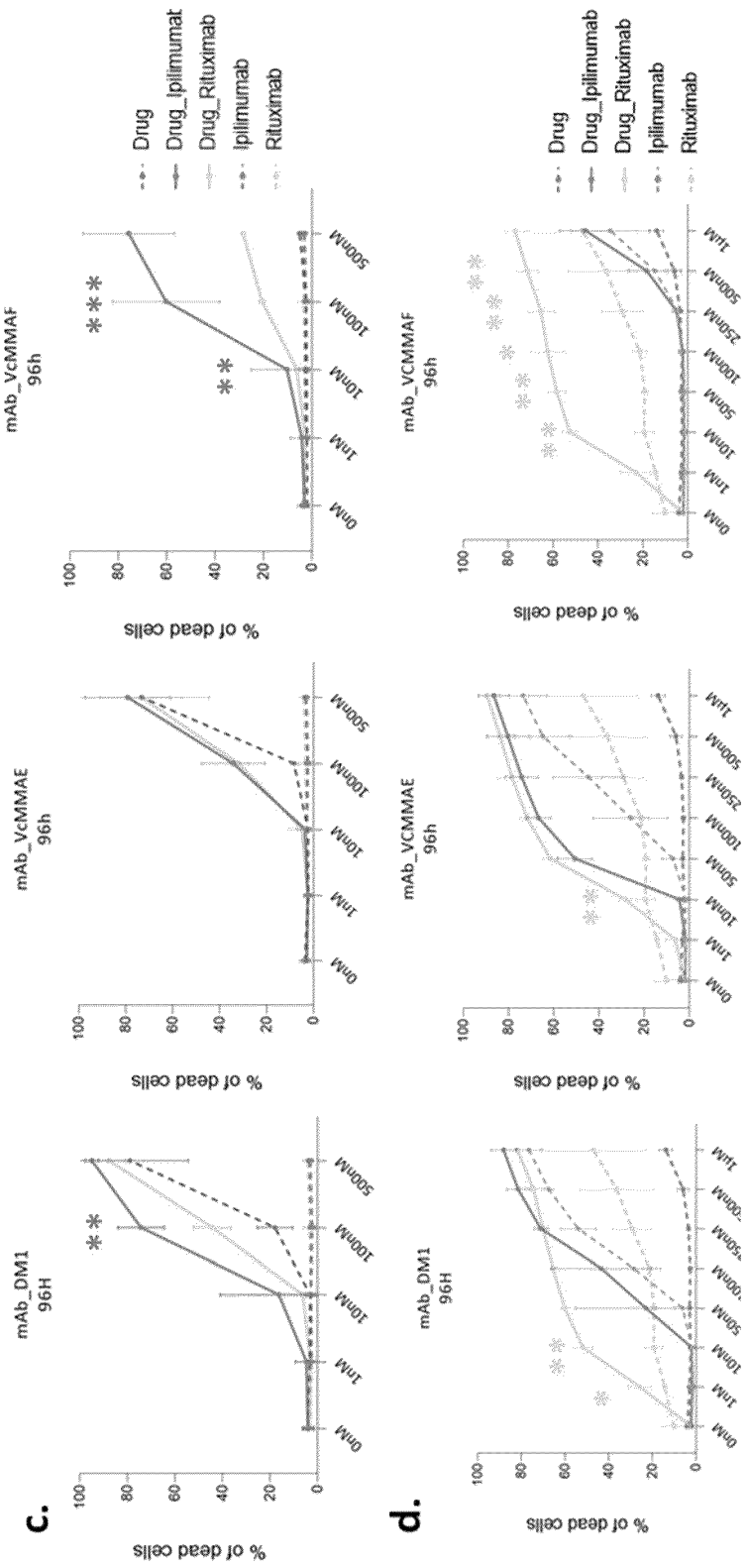
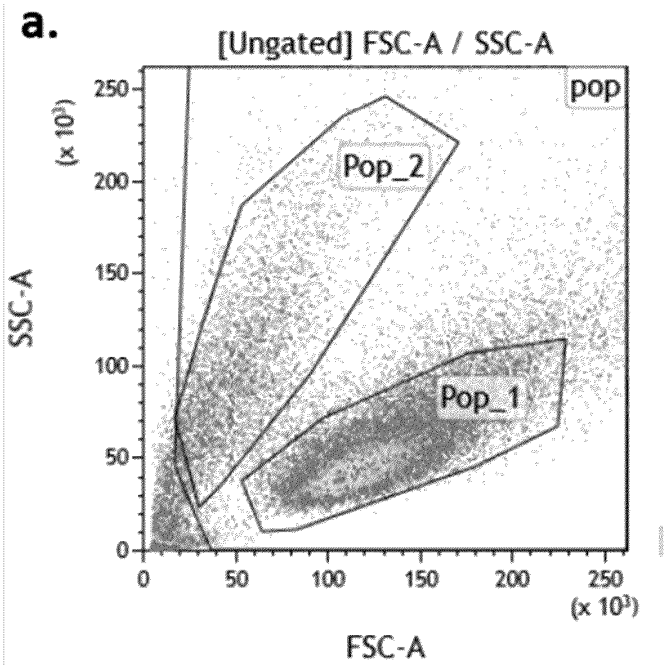


FIGURE 10 (Following)



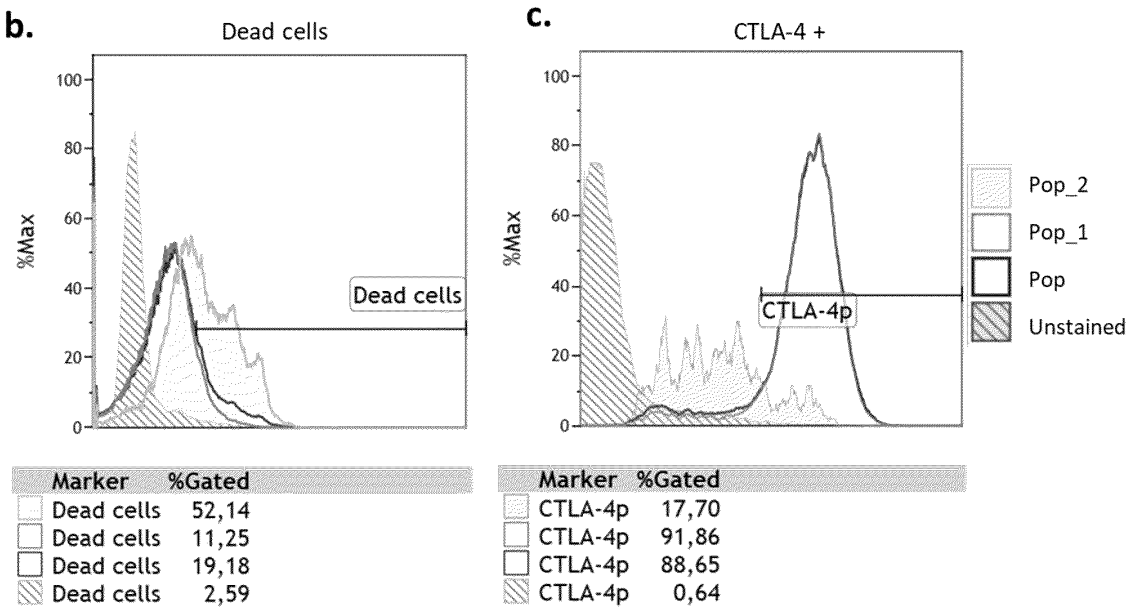


FIGURE 11 (Following)

d.

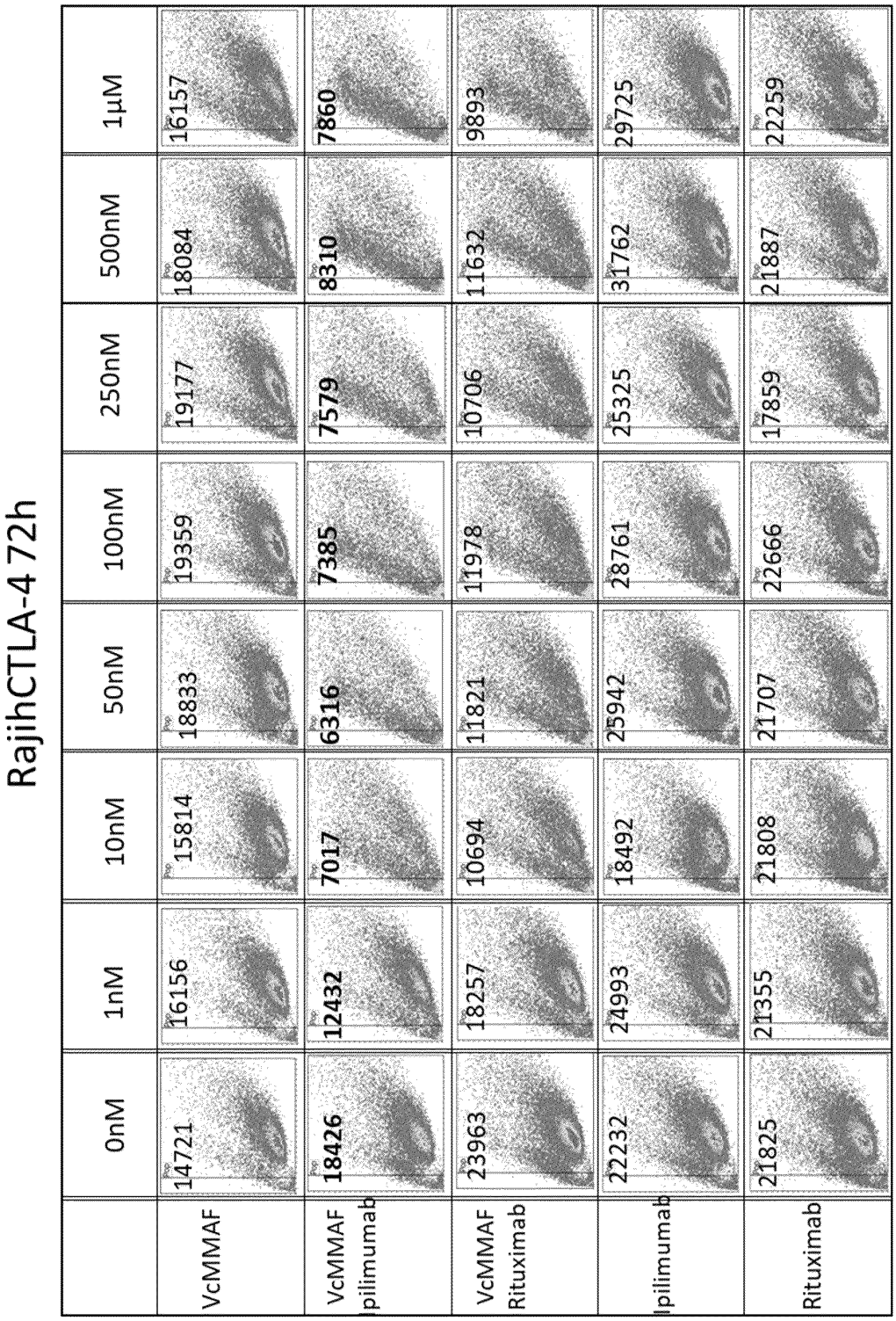


FIGURE 11 (Following)

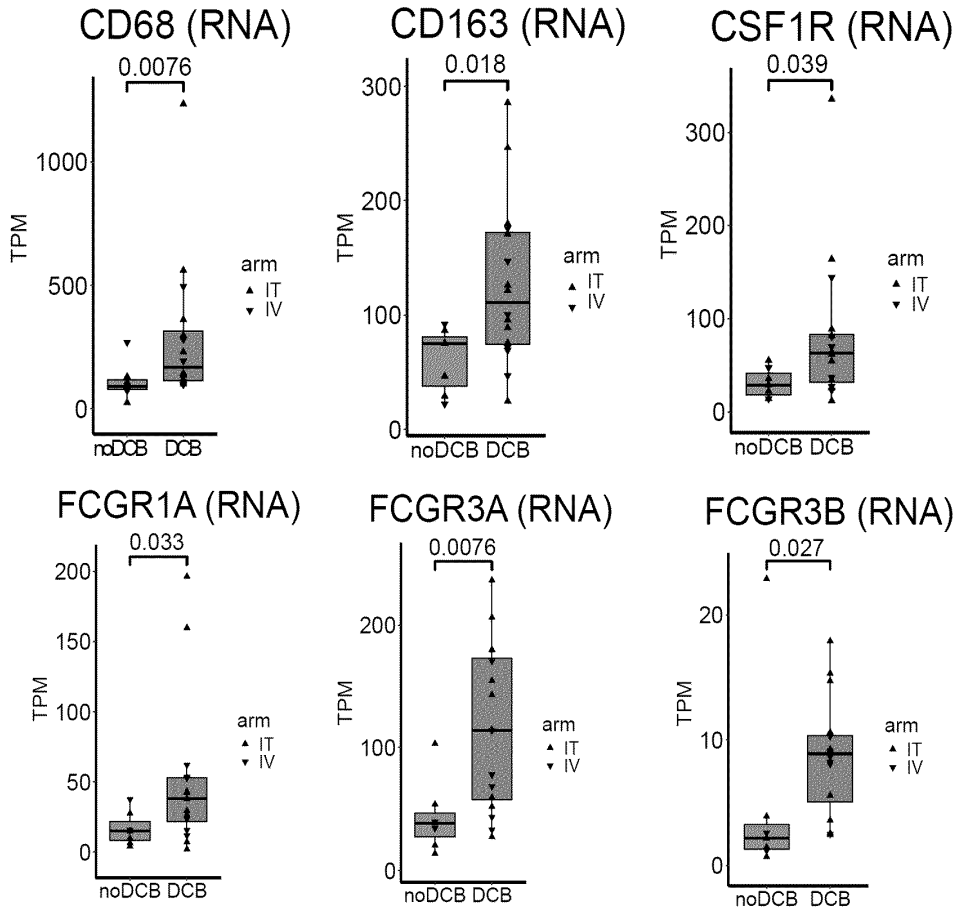

CD68 (RNA)
0.0076
TPM
1000
500
0
noDCB DCB
arm
IT
IV
CD163 (RNA)
0.018
300
200
100
0
TPM
noDCB DCB
arm
IT
IV
CSF1R (RNA)
0.039
300
200
100
0
TPM
noDCB DCB
arm
IT
IV
FCGR1A (RNA)
0.033
200
150
100
50
0
TPM
noDCB DCB
arm
IT
IV
FCGR3A (RNA)
0.0076
200
100
0
TPM
noDCB DCB
arm
IT
IV
FCGR3B (RNA)
0.027
20
10
0
TPM
noDCB DCB
arm
IT
IV

FIGURE 12