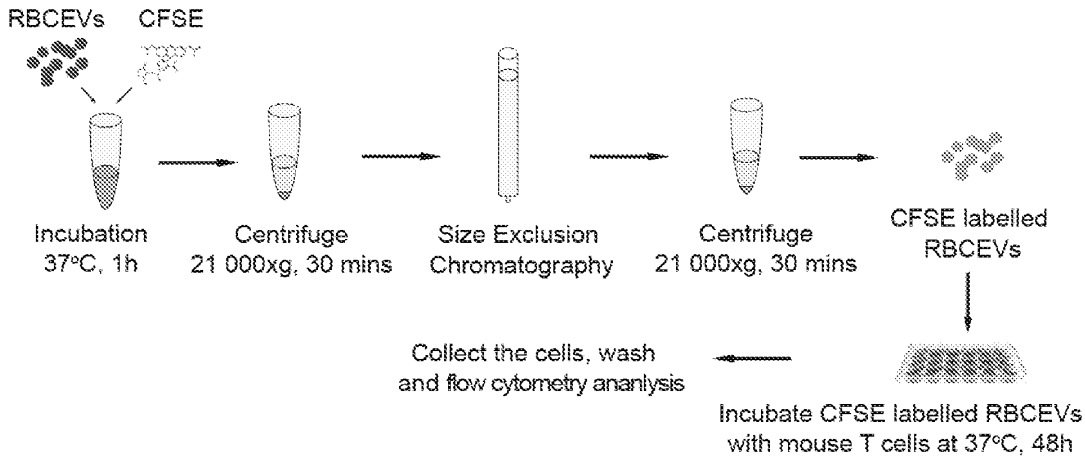

Figure 1A

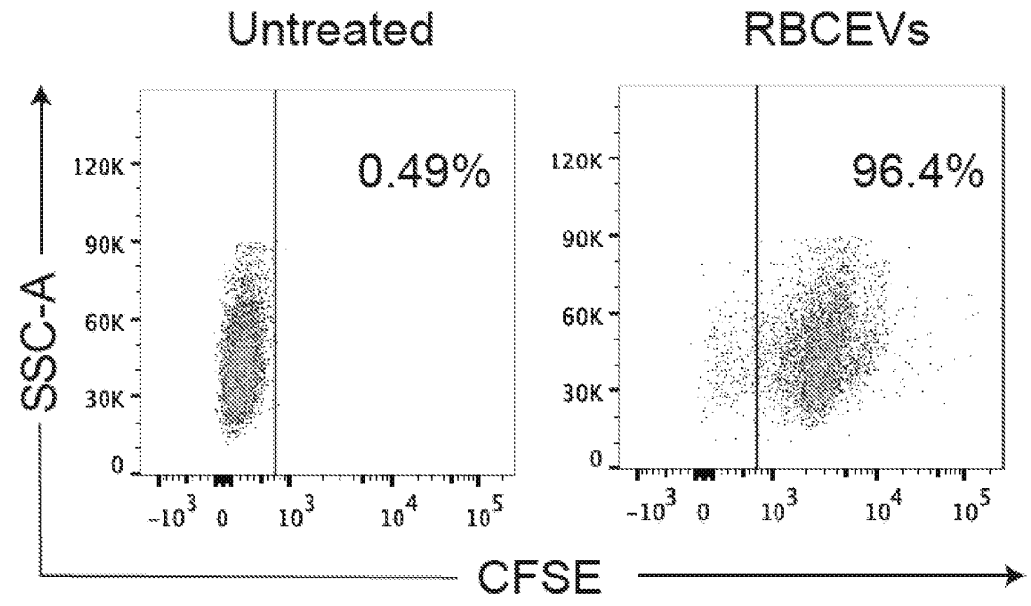

Figure 1B

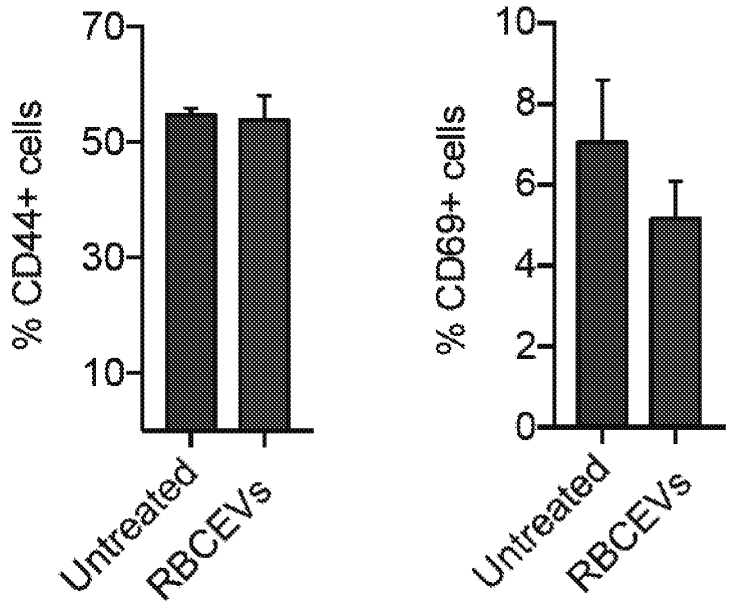

Figure 1C

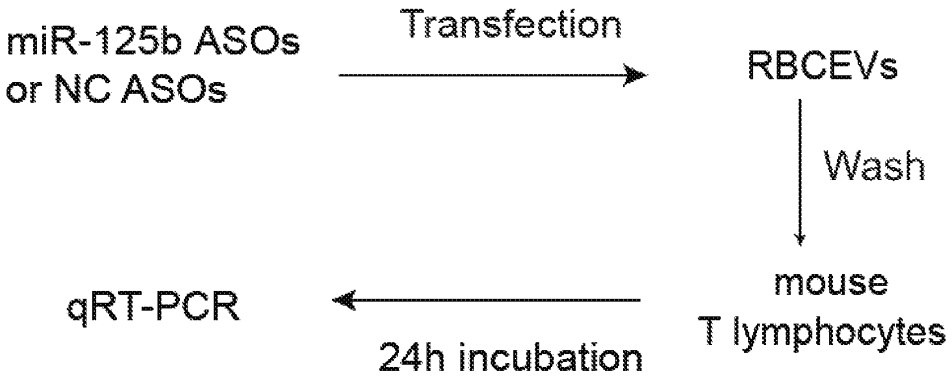

Figure 1D

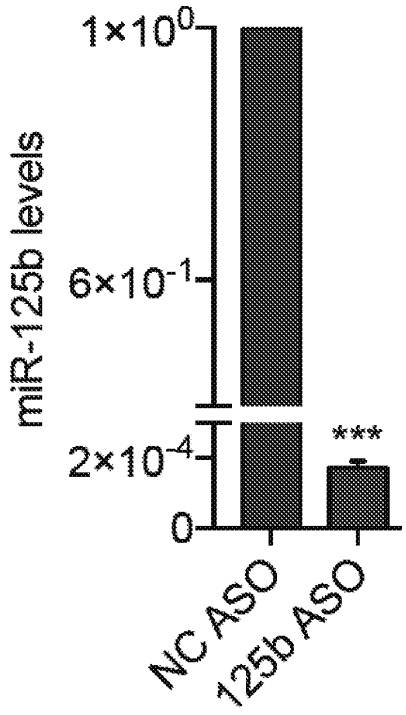

Figure 1E

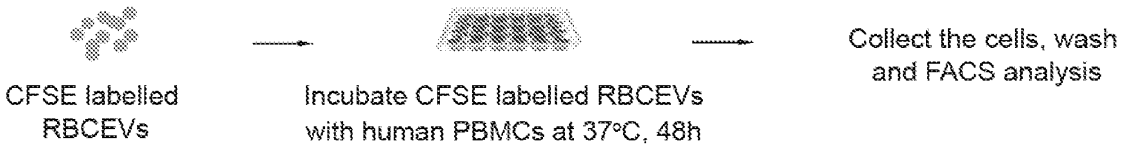

Figure 2A

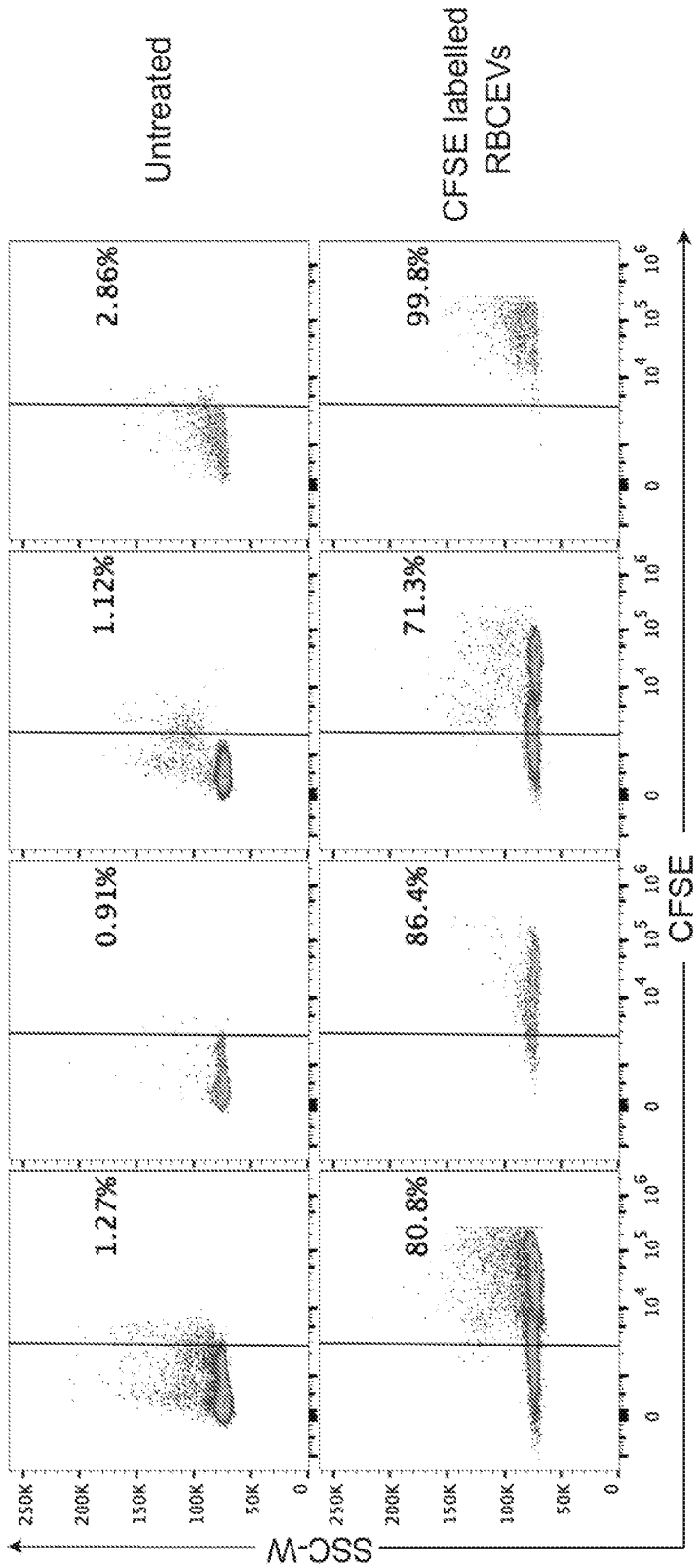
Untreated
CFSE labelled RBCEVs
All cells
CD19+ B cells
CD 3+ T cells
CD14+ monocytes
1.27%
0.91%
1.12%
2.86%
80.8%
86.4%
71.3%
99.8%
SSC-W
CFSE
250K
200K
150K
100K
50K
0
10^4
10^5
10^6

Figure 2B

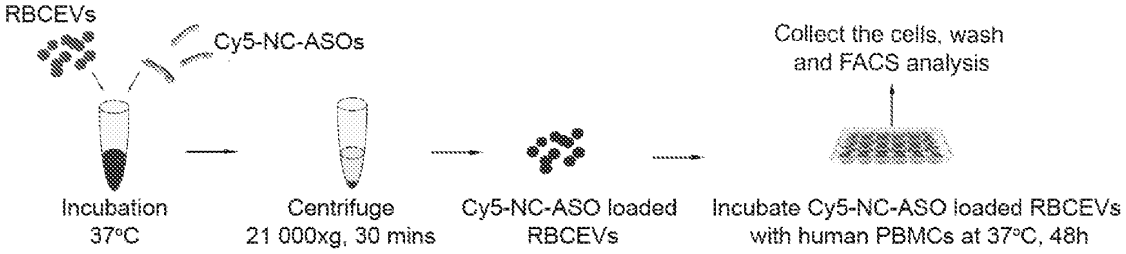

Figure 2C

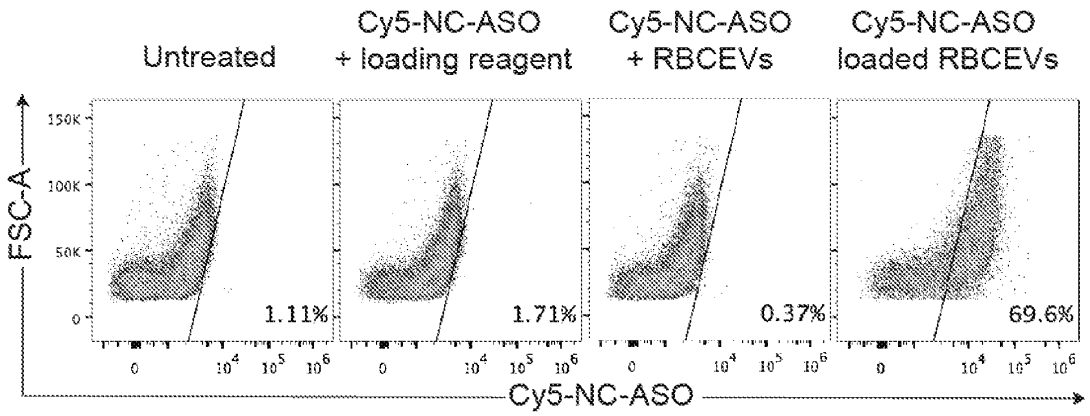

Figure 2D

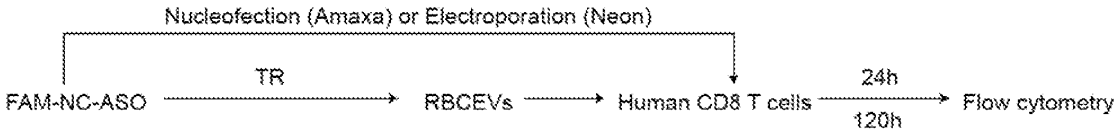

Figure 3A

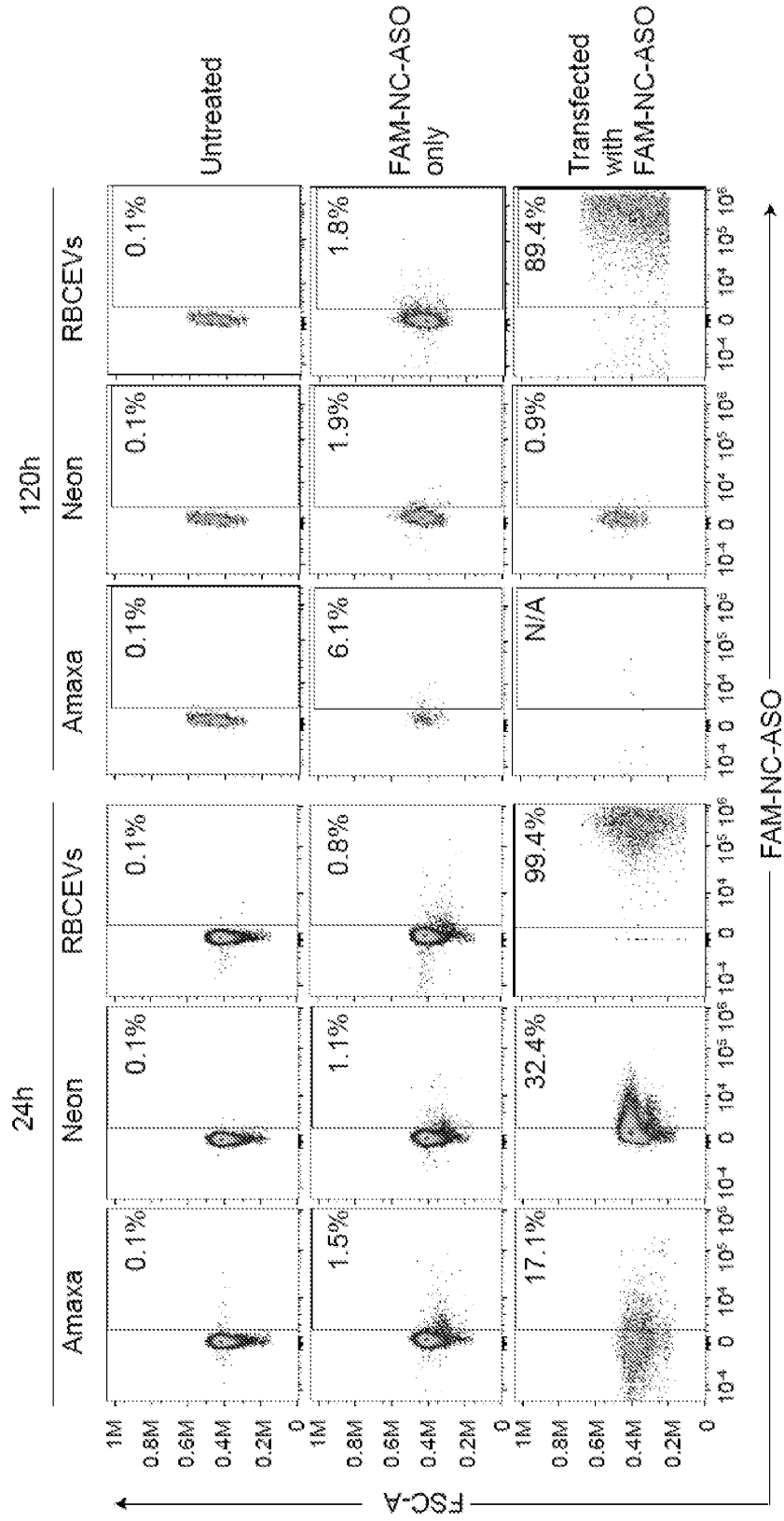
24h
120h
Amaxa
Neon
RBCEVs
Amaxa
Neon
RBCEVs
Untreated
FAM-NC-ASO only
Transfected with FAM-NC-ASO
0.1%
0.1%
0.1%
0.1%
0.1%
0.1%
1.5%
1.1%
0.8%
6.1%
1.9%
1.8%
17.1%
32.4%
99.4%
N/A
0.9%
89.4%
FSC-A
FAM-NC-ASO

Figure 3B

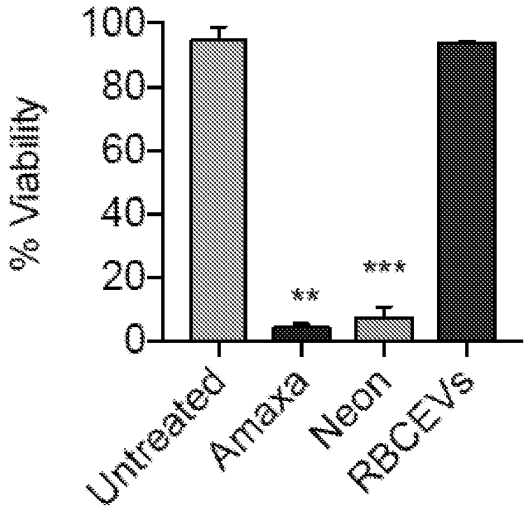

Figure 3C

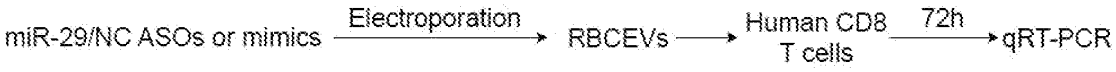

Figure 4A

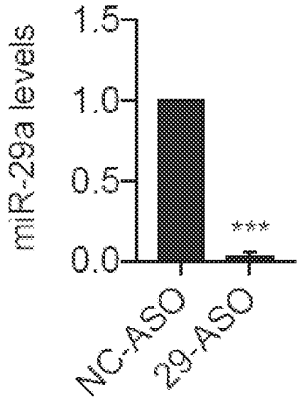

Figure 4B

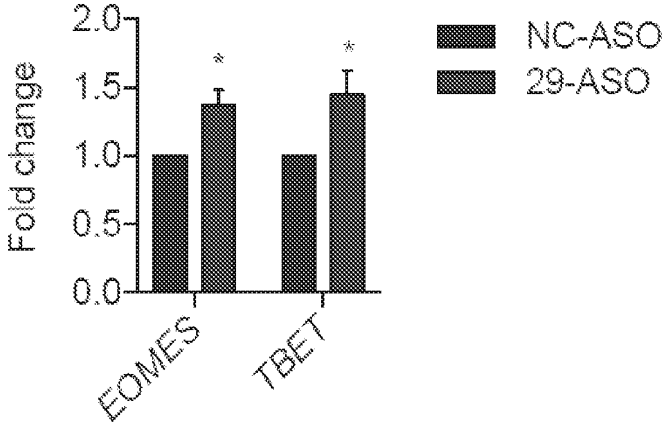

Figure 4C

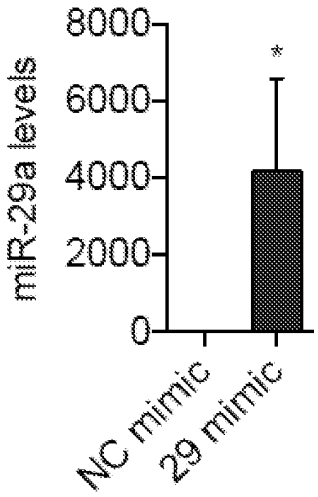

Figure 4D

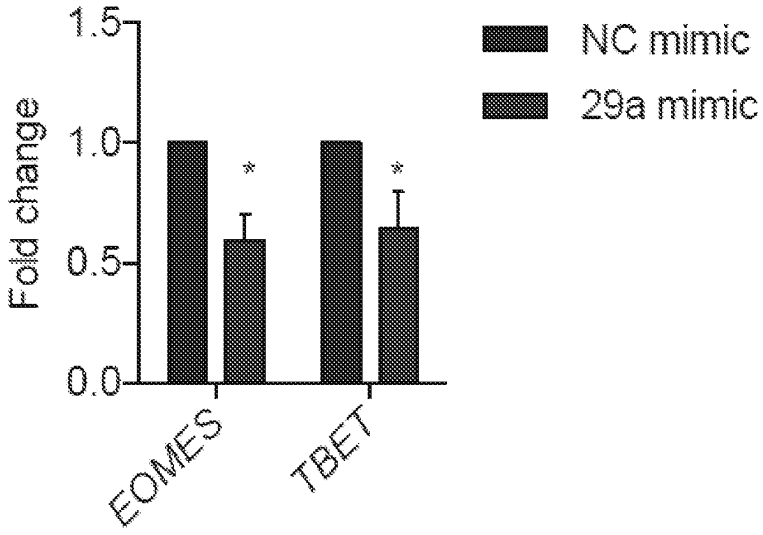

Figure 4E

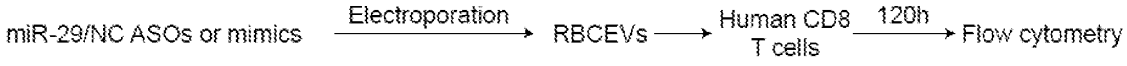

Figure 5A

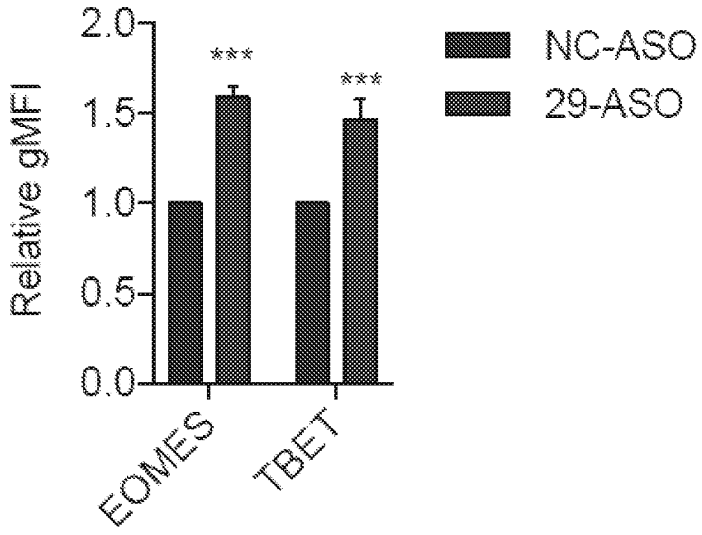

Figure 5B

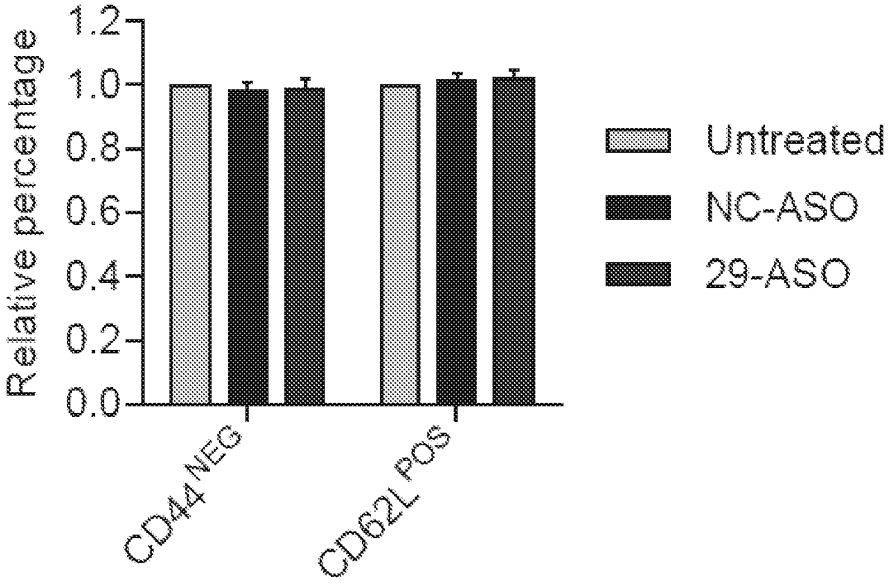

Figure 5C

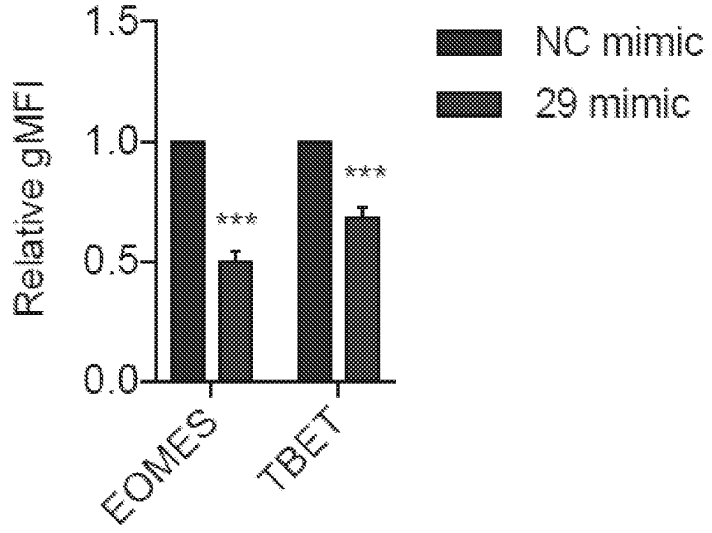

Figure 5D

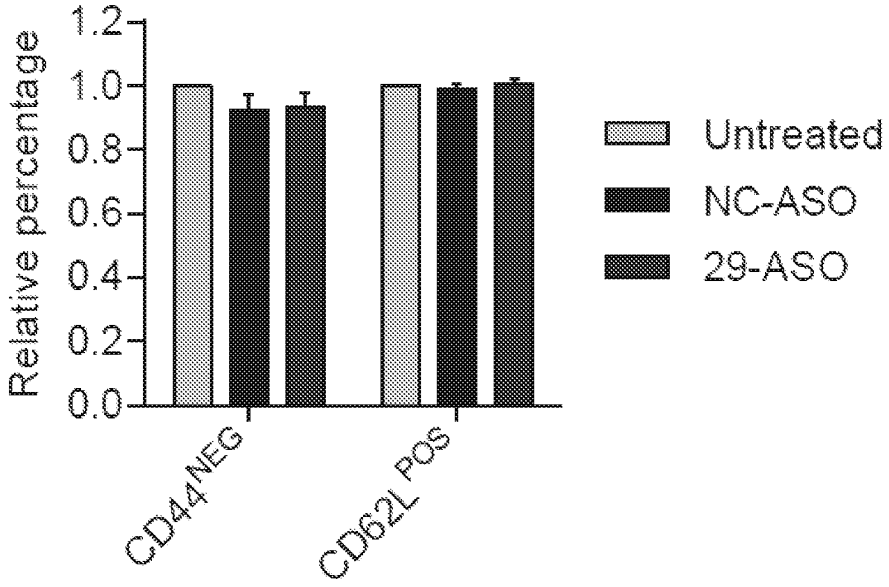

Figure 5E

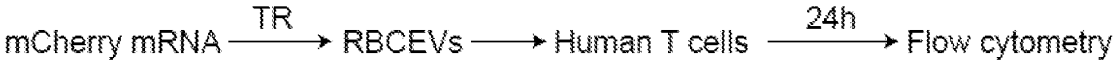

Figure 6A

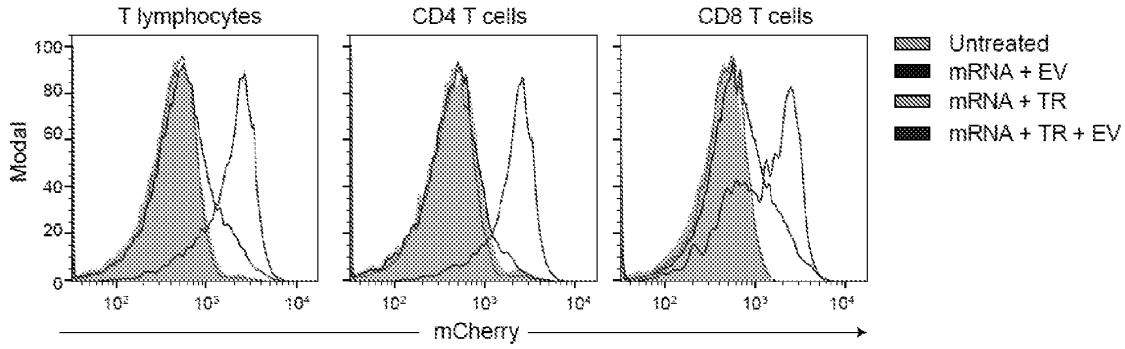

Figure 6B

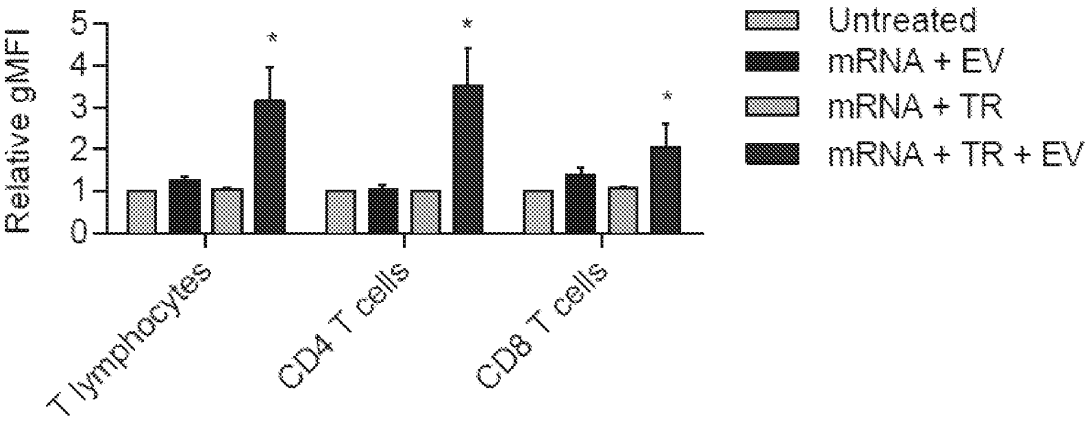

Figure 6C

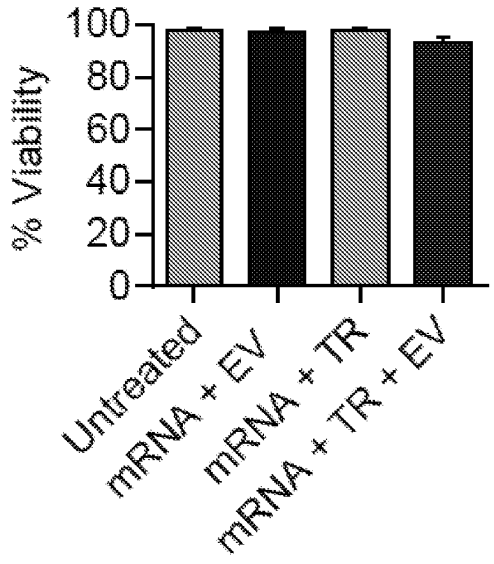

Figure 6D

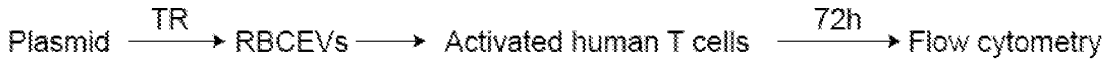

Figure 7A

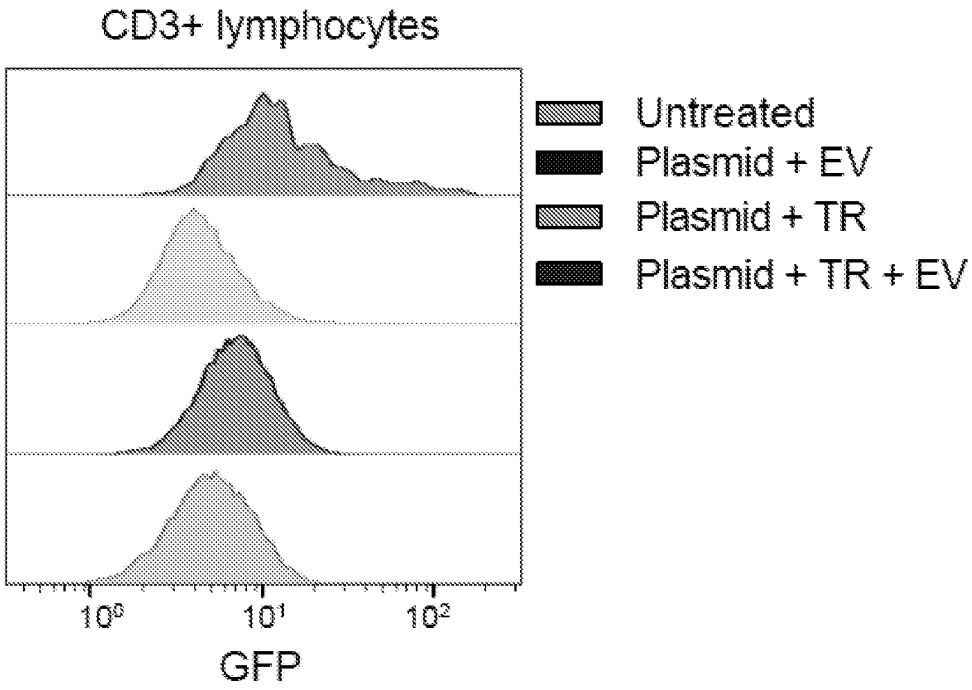

Figure 7B

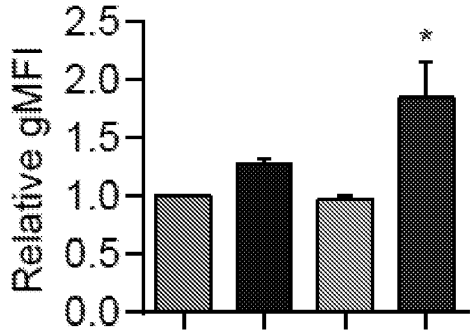

Figure 7C

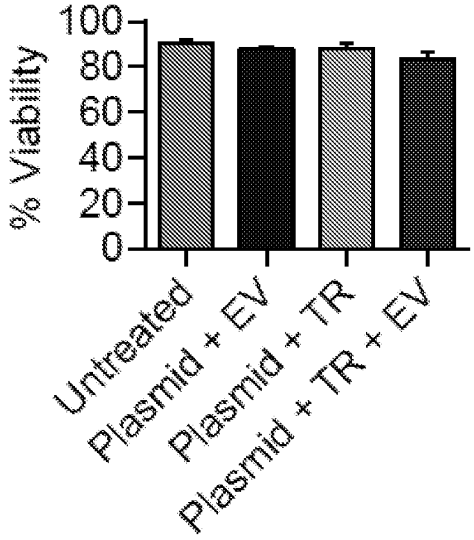

Figure 7D

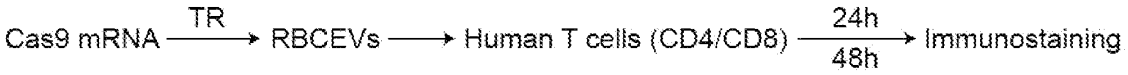

Figure 8A

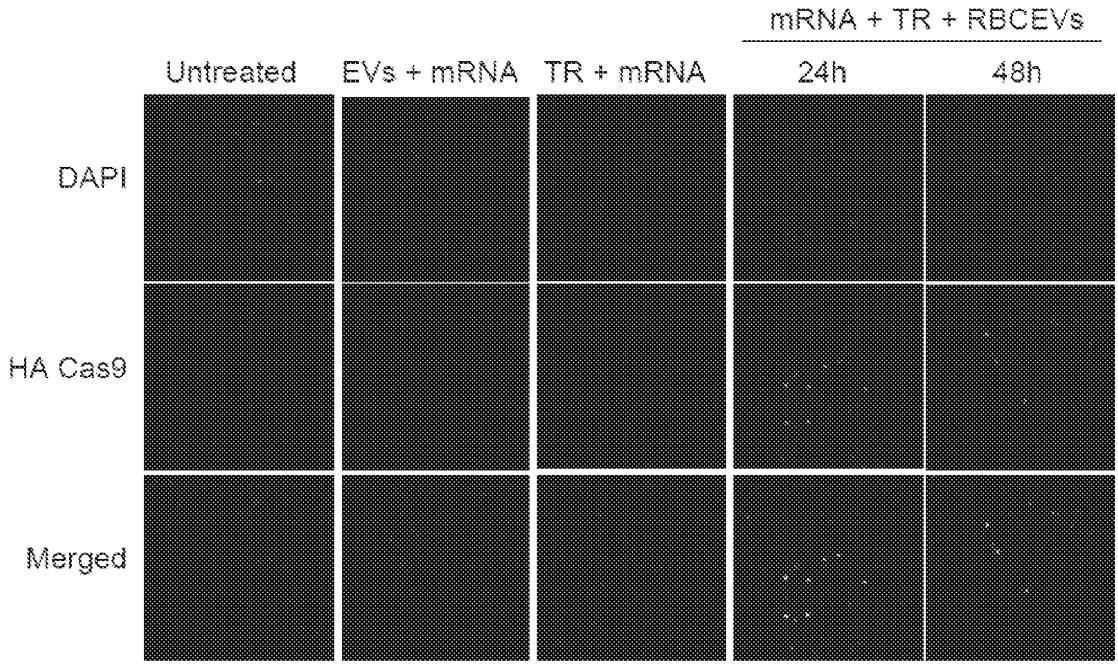

Figure 8B

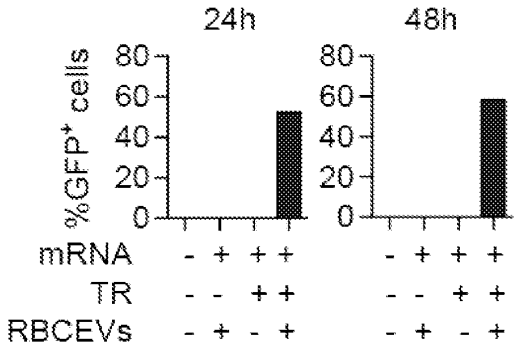

Figure 8C

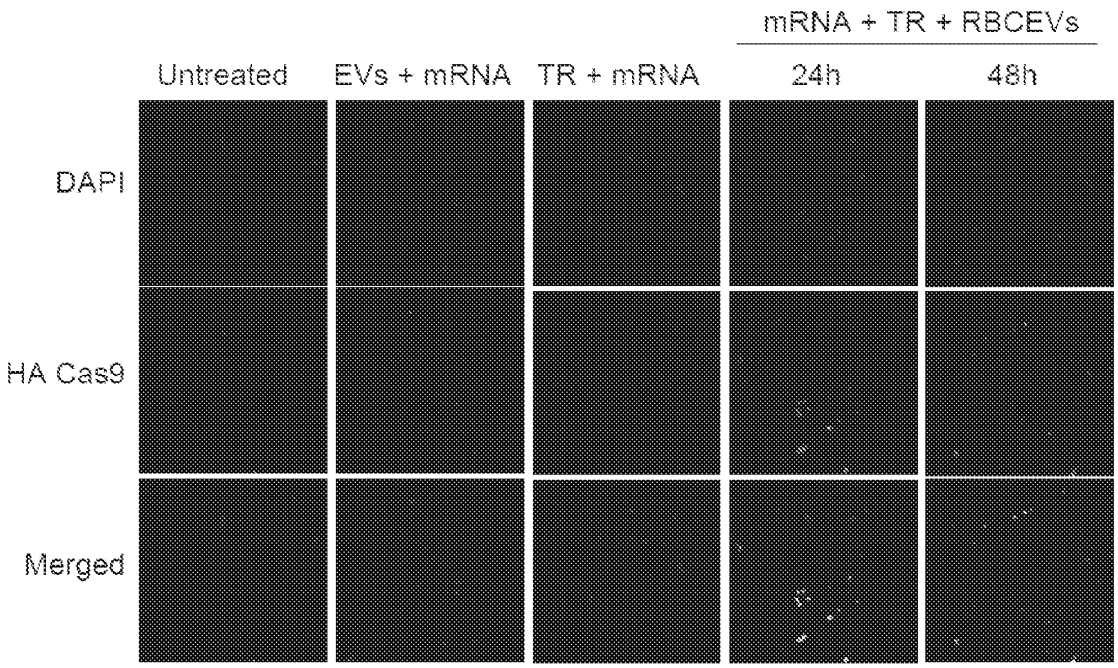

Figure 8D

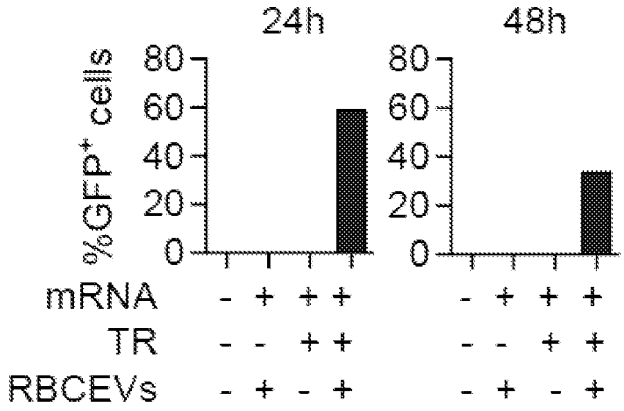

Figure 8E

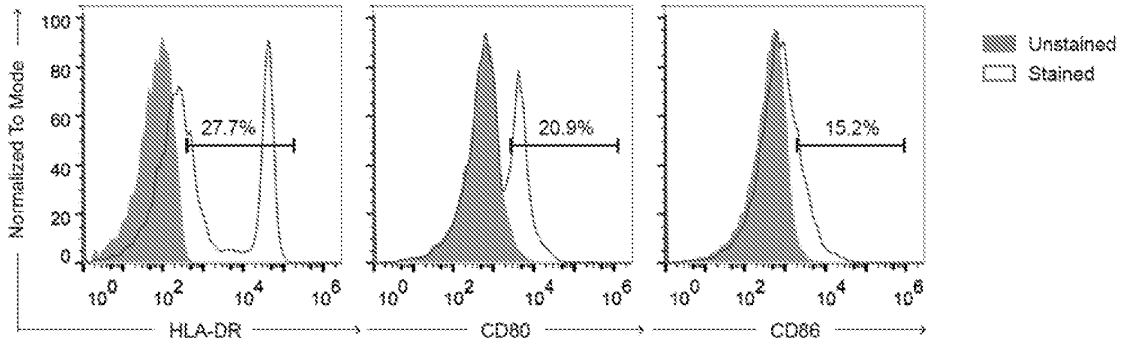

Figure 9A

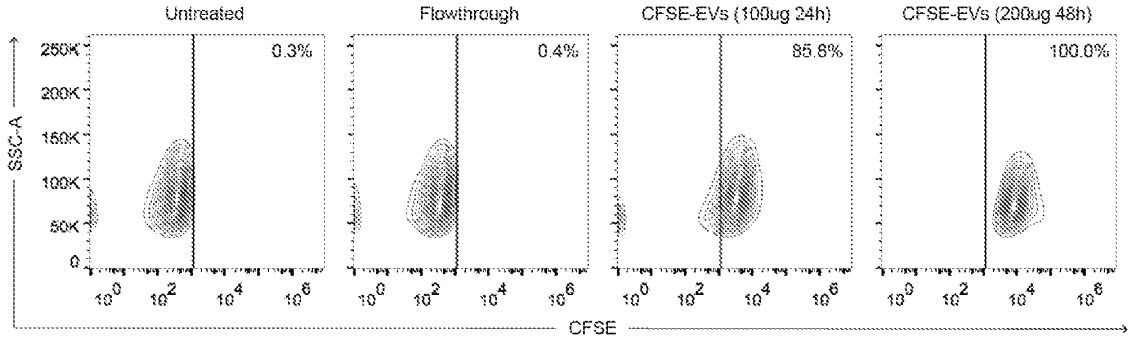

Figure 9B

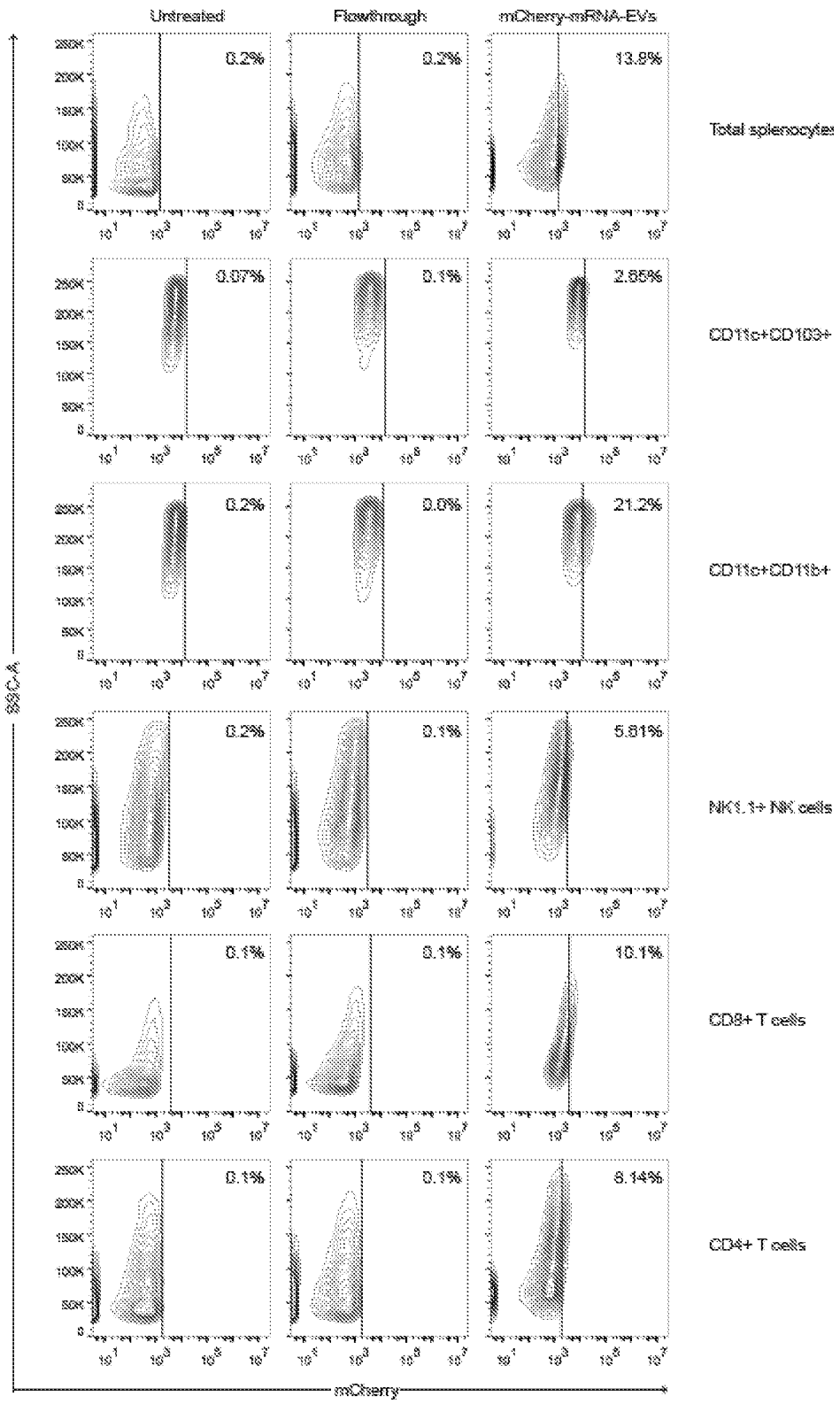
Untreated
Flowthrough
mCherry-mRNA-EVs
0.2%
0.2%
13.8%
Total splenocytes
0.07%
0.1%
2.65%
CD11c+CD103+
0.2%
0.0%
21.2%
CD11c+CD11b+
0.2%
0.1%
5.81%
NK1.1+ NK cells
0.1%
0.1%
10.1%
CD8+ T cells
0.1%
0.1%
8.14%
CD4+ T cells
SSC-A
mCherry

Figure 10

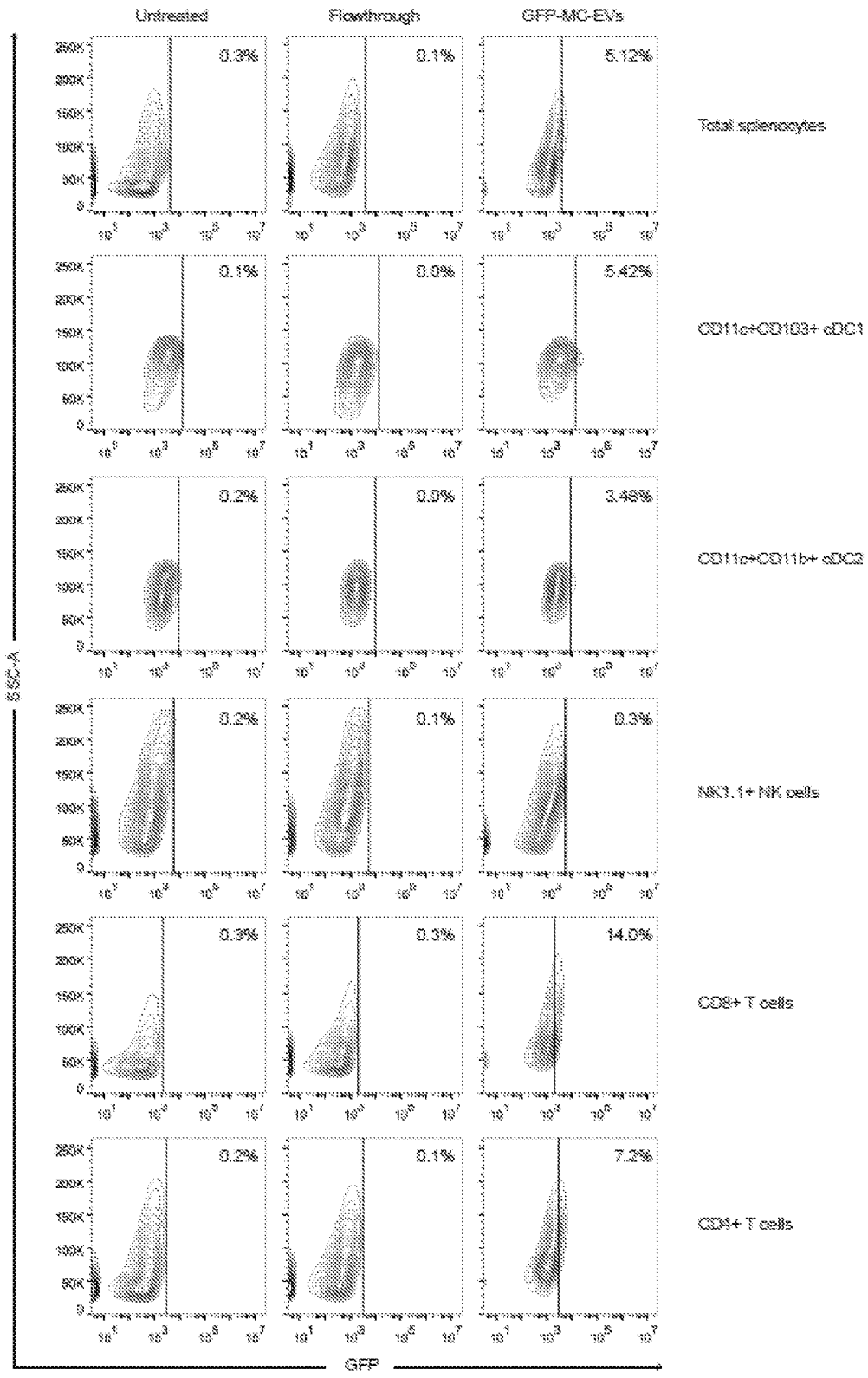
Untreated
Flowthrough
GFP-MC-EVs
0.3%
0.1%
5.12%
Total splenocytes
0.1%
0.0%
5.42%
CD11c+CD103+ cDC1
0.2%
0.0%
3.46%
CD11c+CD11b+ cDC2
0.2%
0.1%
0.3%
NK1.1+ NK cells
0.3%
0.3%
14.0%
CD8+ T cells
0.2%
0.1%
7.2%
CD4+ T cells
SSC-A
GFP

Figure 11

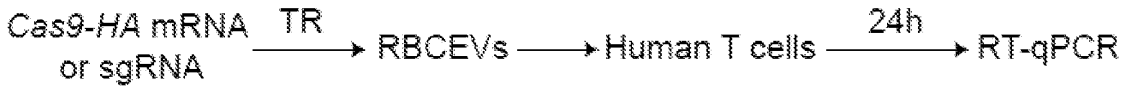

Figure 12A

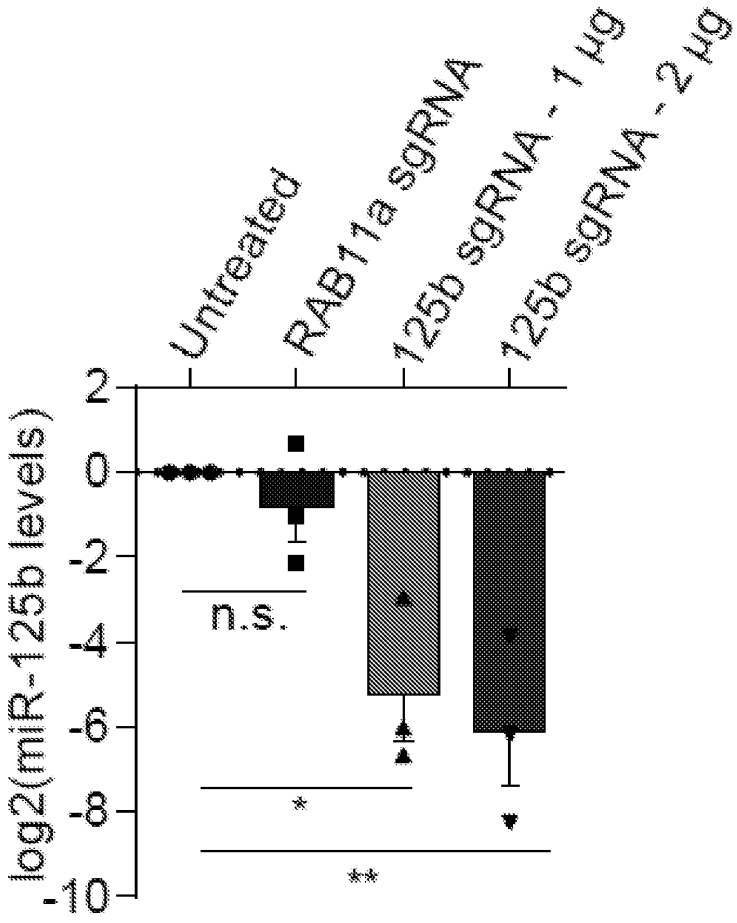

Figure 12B

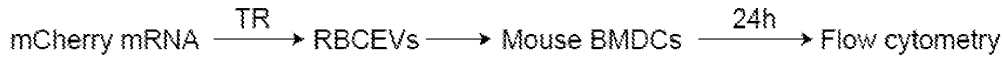

Figure 13A

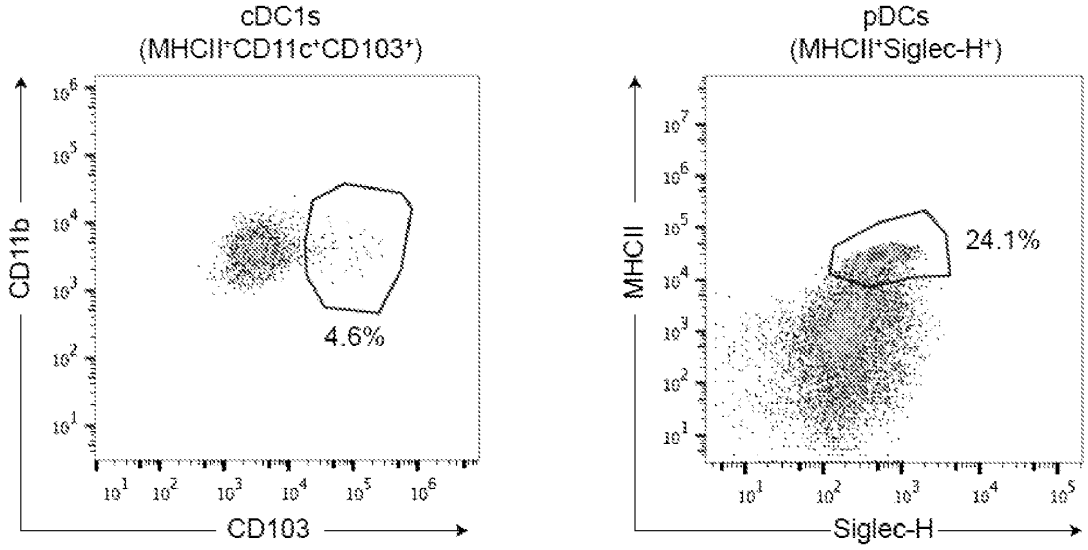

Figure 13B

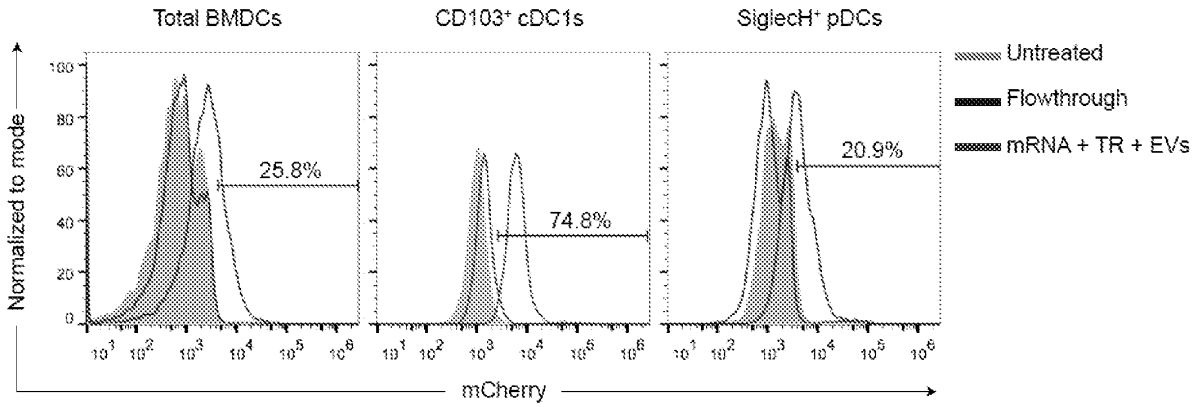

Figure 13C

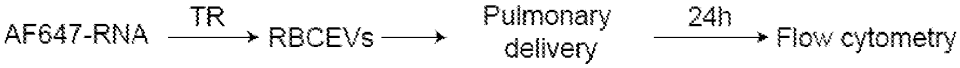

Figure 14A

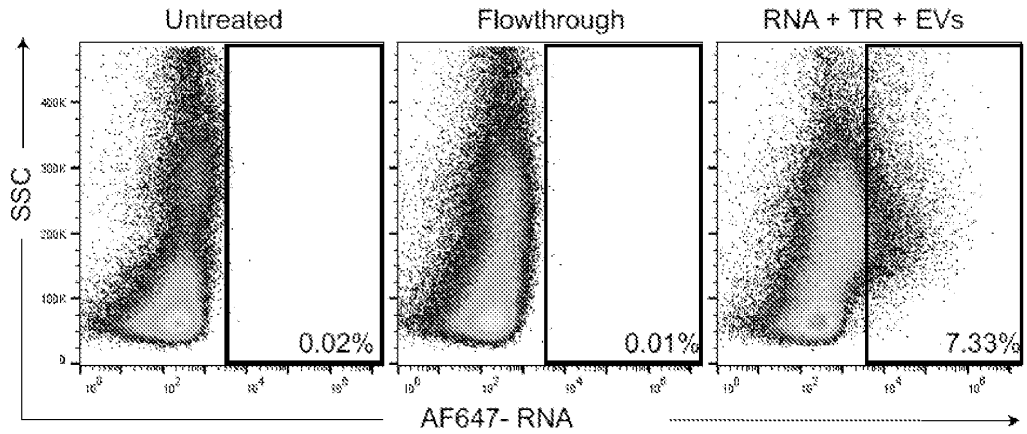

Figure 14B

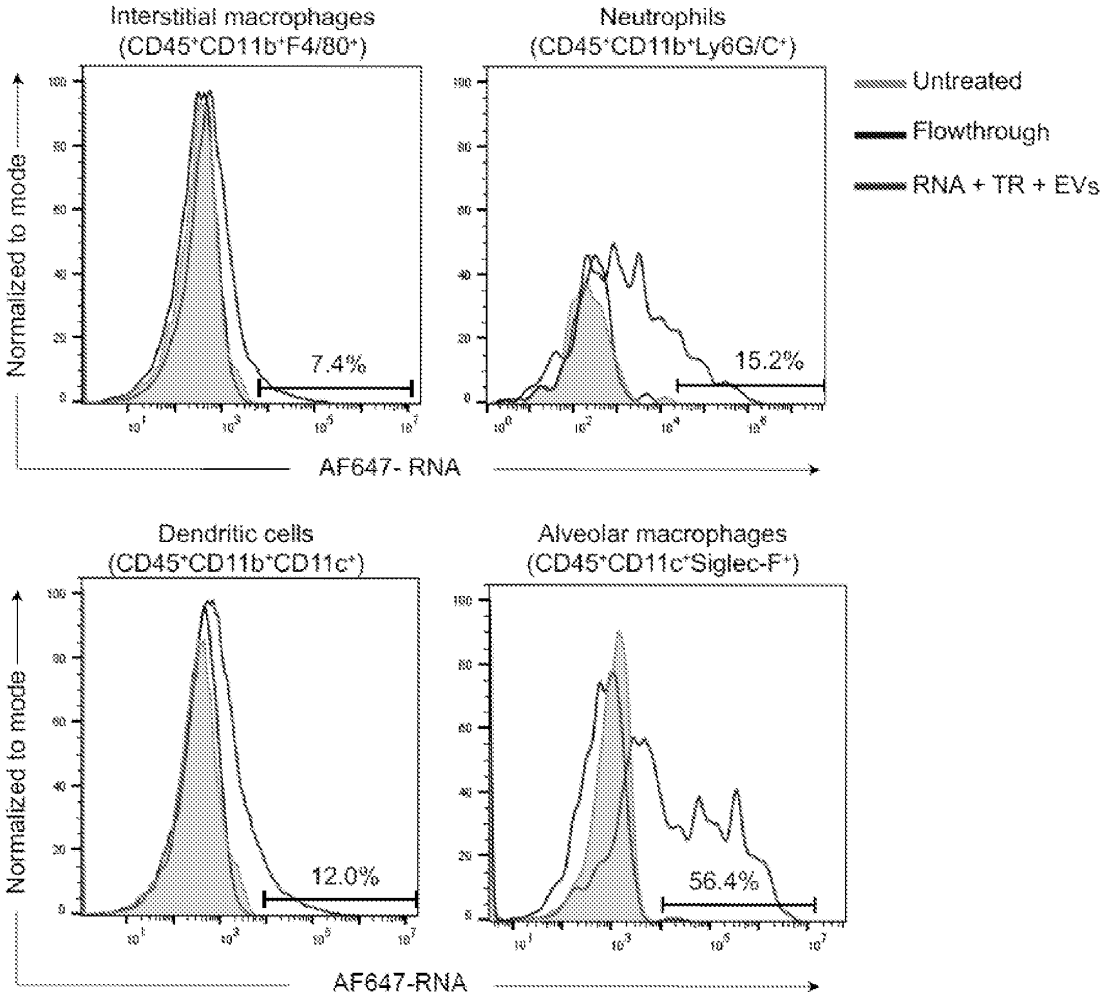
Interstitial macrophages
(CD45+CD11b+F4/80+)
Neutrophils
(CD45+CD11b+Ly6G/C+)
Untreated
Flowthrough
RNA + TR + EVs
Normalized to mode
7.4%
15.2%
AF647- RNA
Dendritic cells
(CD45+CD11b+CD11c+)
Alveolar macrophages
(CD45+CD11c+Siglec-F+)
Normalized to mode
12.0%
56.4%
AF647-RNA

Figure 14C