Figure 1

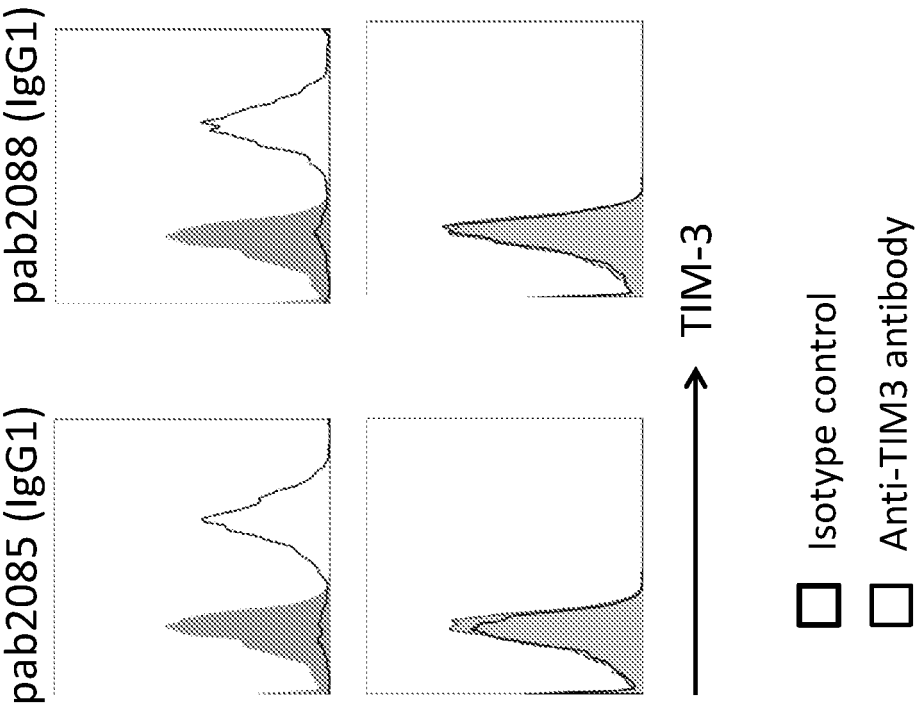

pab2085 (IgG1)
pab2088 (IgG1)
Murine 1624-5 cells expressing human TIM-3
Wild type murine 1624-5 cells
TIM-3
Isotype control
Anti-TIM3 antibody

Figure 2A

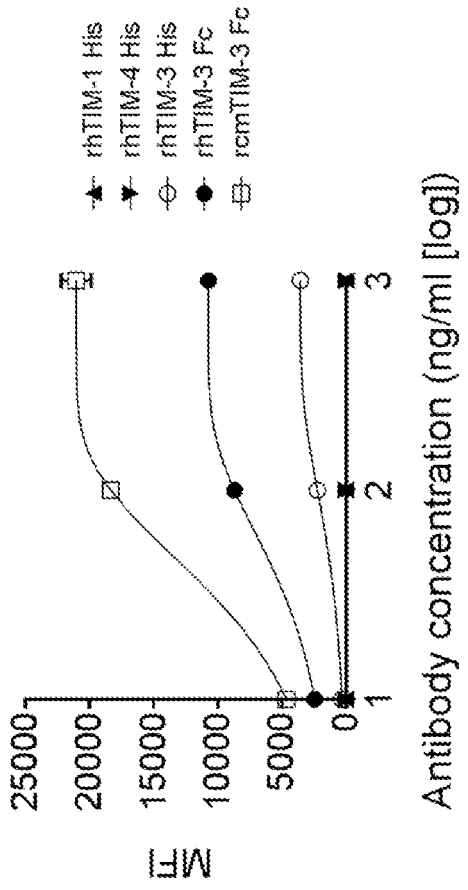


Figure 2B

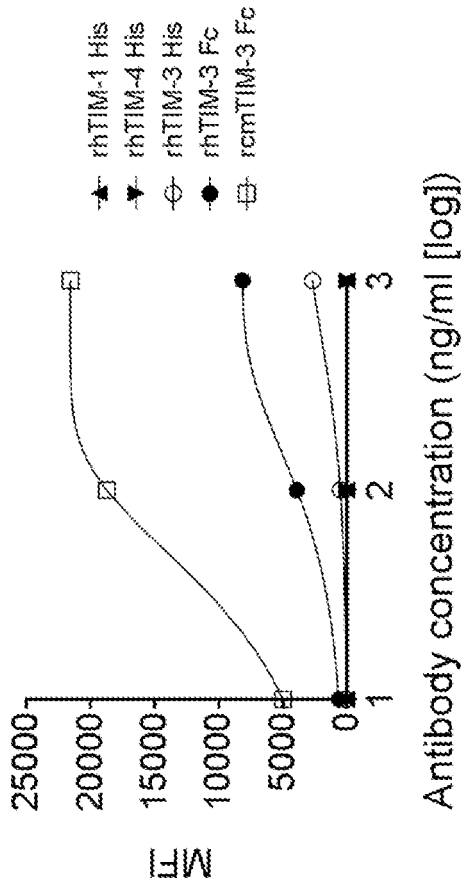


Figure 3A

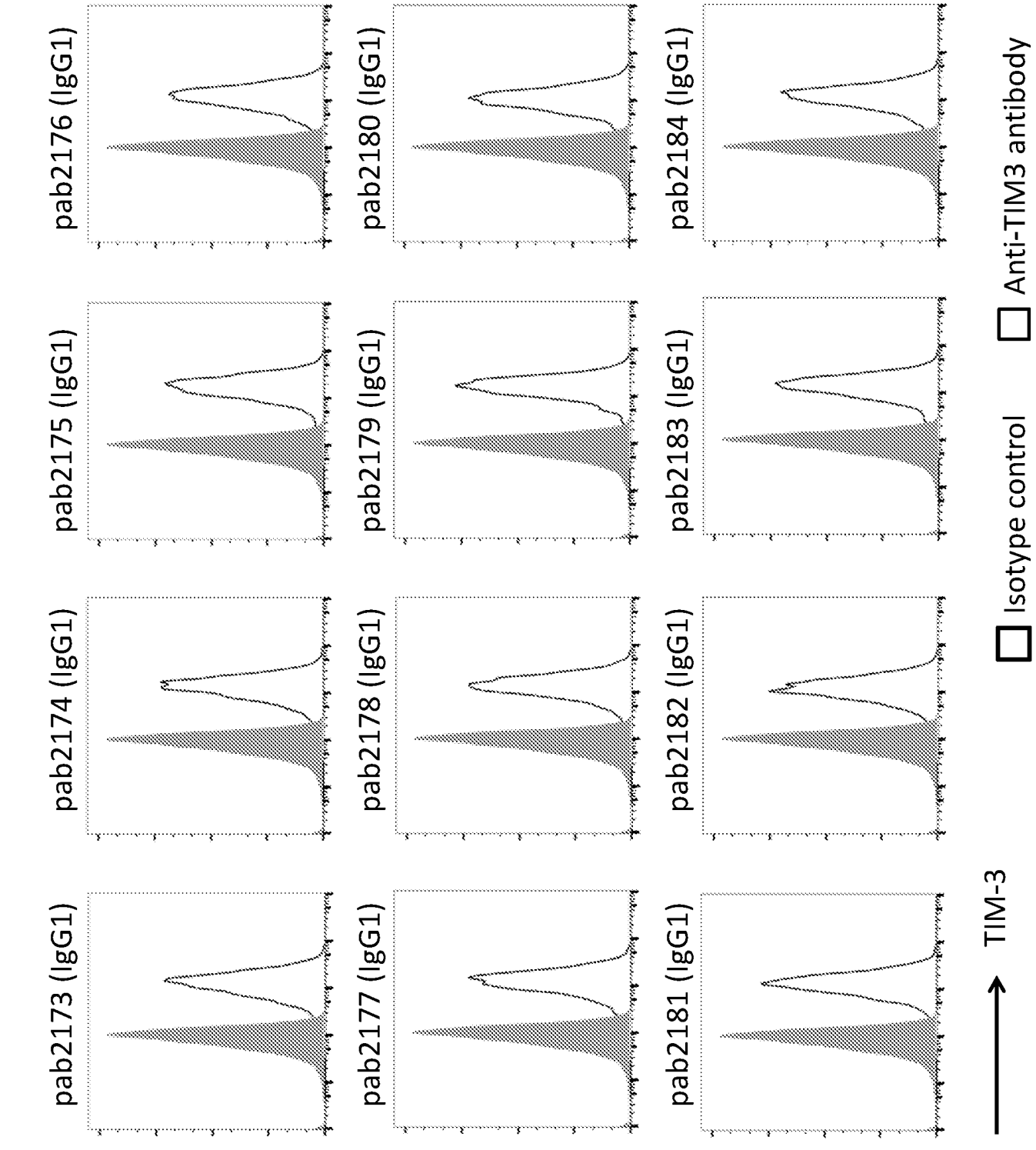

pab2173 (IgG1)
pab2174 (IgG1)
pab2175 (IgG1)
pab2176 (IgG1)
pab2177 (IgG1)
pab2178 (IgG1)
pab2179 (IgG1)
pab2180 (IgG1)
pab2181 (IgG1)
pab2182 (IgG1)
pab2183 (IgG1)
pab2184 (IgG1)
1624-5 cells expressing human TIM-3
TIM-3
Isotype control
Anti-TIM3 antibody

Figure 3B

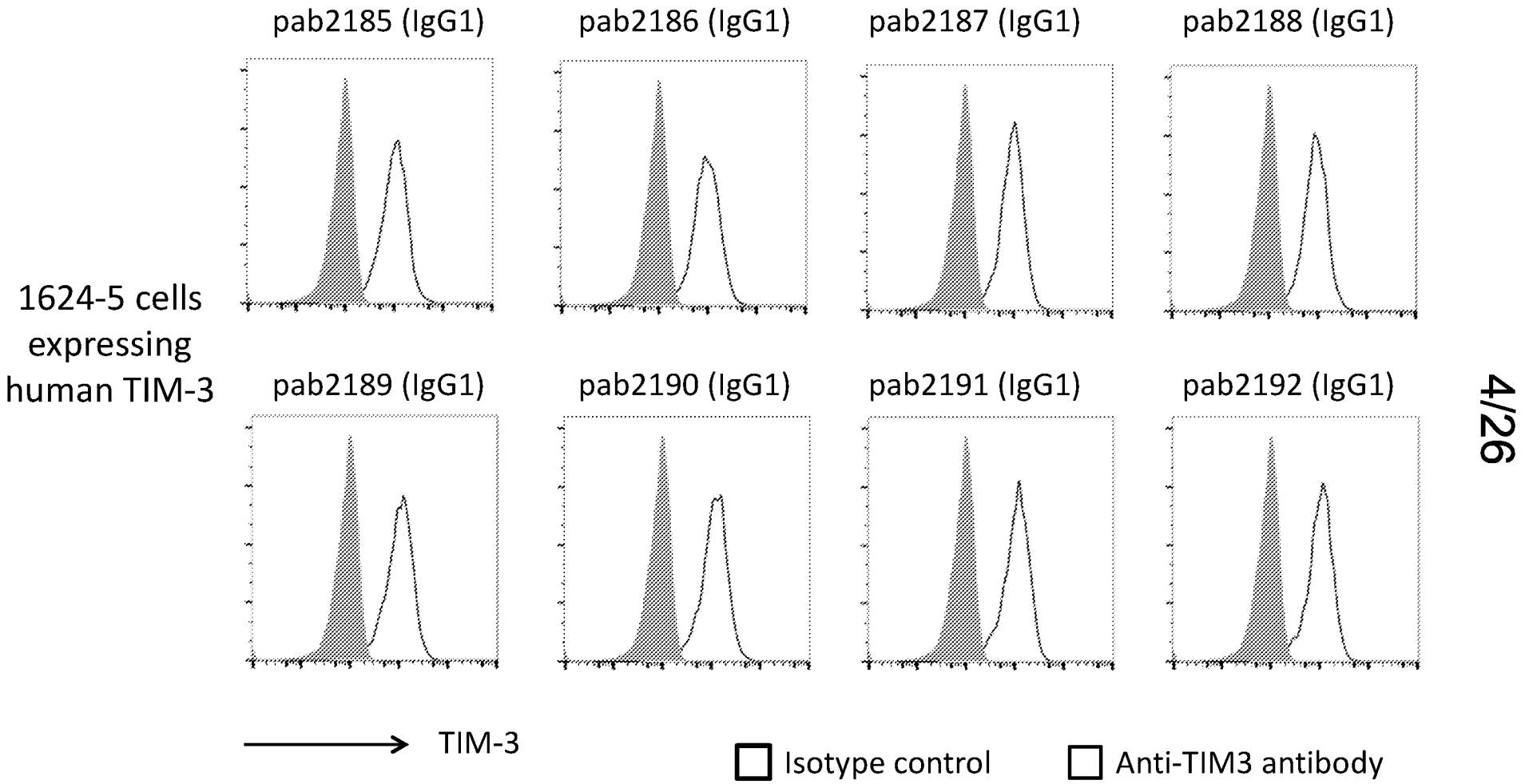

pab2185 (IgG1)
pab2186 (IgG1)
pab2187 (IgG1)
pab2188 (IgG1)
1624-5 cells expressing human TIM-3
pab2189 (IgG1)
pab2190 (IgG1)
pab2191 (IgG1)
pab2192 (IgG1)
TIM-3
Isotype control
Anti-TIM3 antibody

Figure 3C

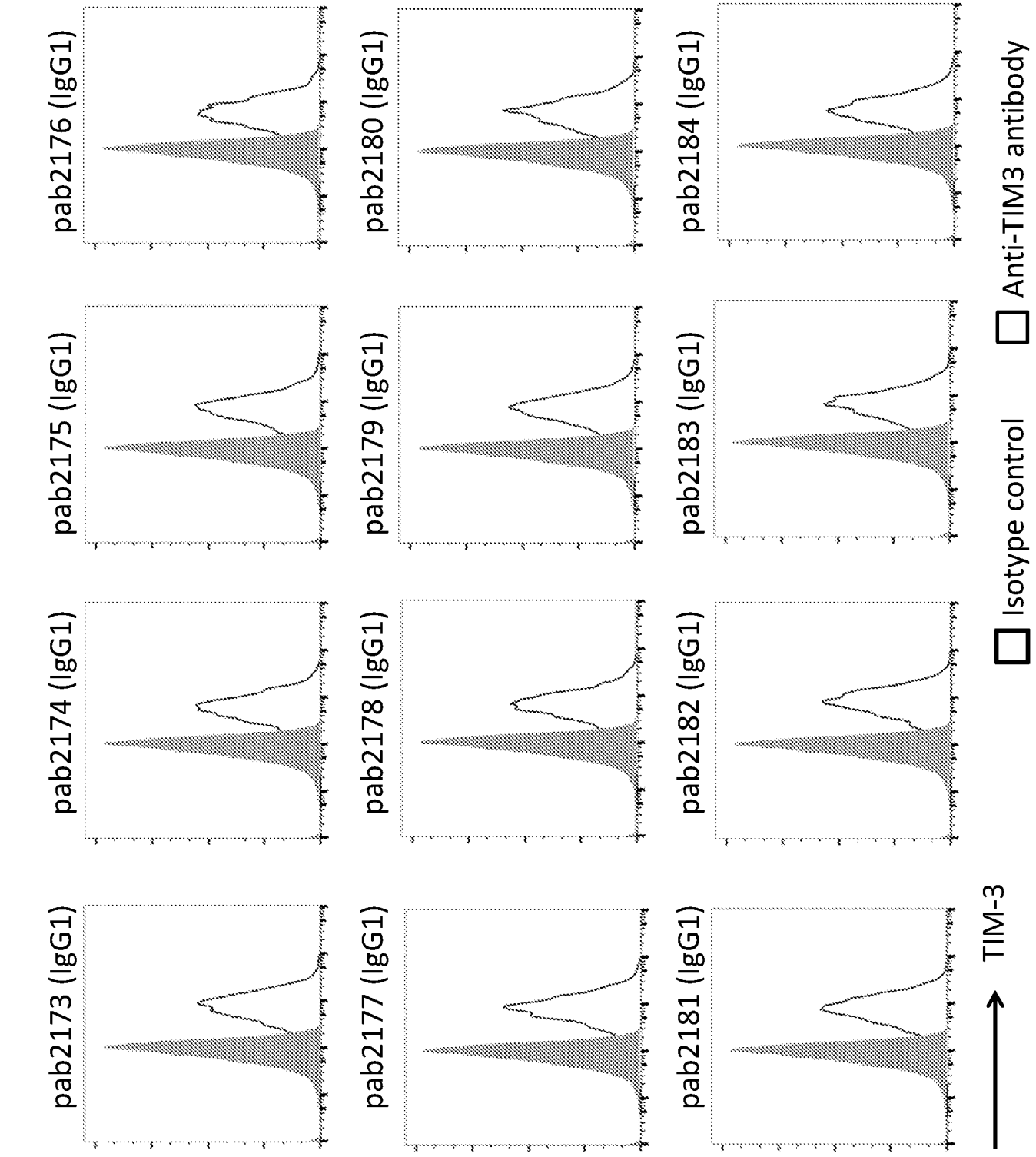

pab2173 (IgG1)
pab2174 (IgG1)
pab2175 (IgG1)
pab2176 (IgG1)
pab2177 (IgG1)
pab2178 (IgG1)
pab2179 (IgG1)
pab2180 (IgG1)
pab2181 (IgG1)
pab2182 (IgG1)
pab2183 (IgG1)
pab2184 (IgG1)
1624-5 cells expressing cynomolgus TIM-3
TIM-3
Isotype control
Anti-TIM3 antibody

Figure 3D

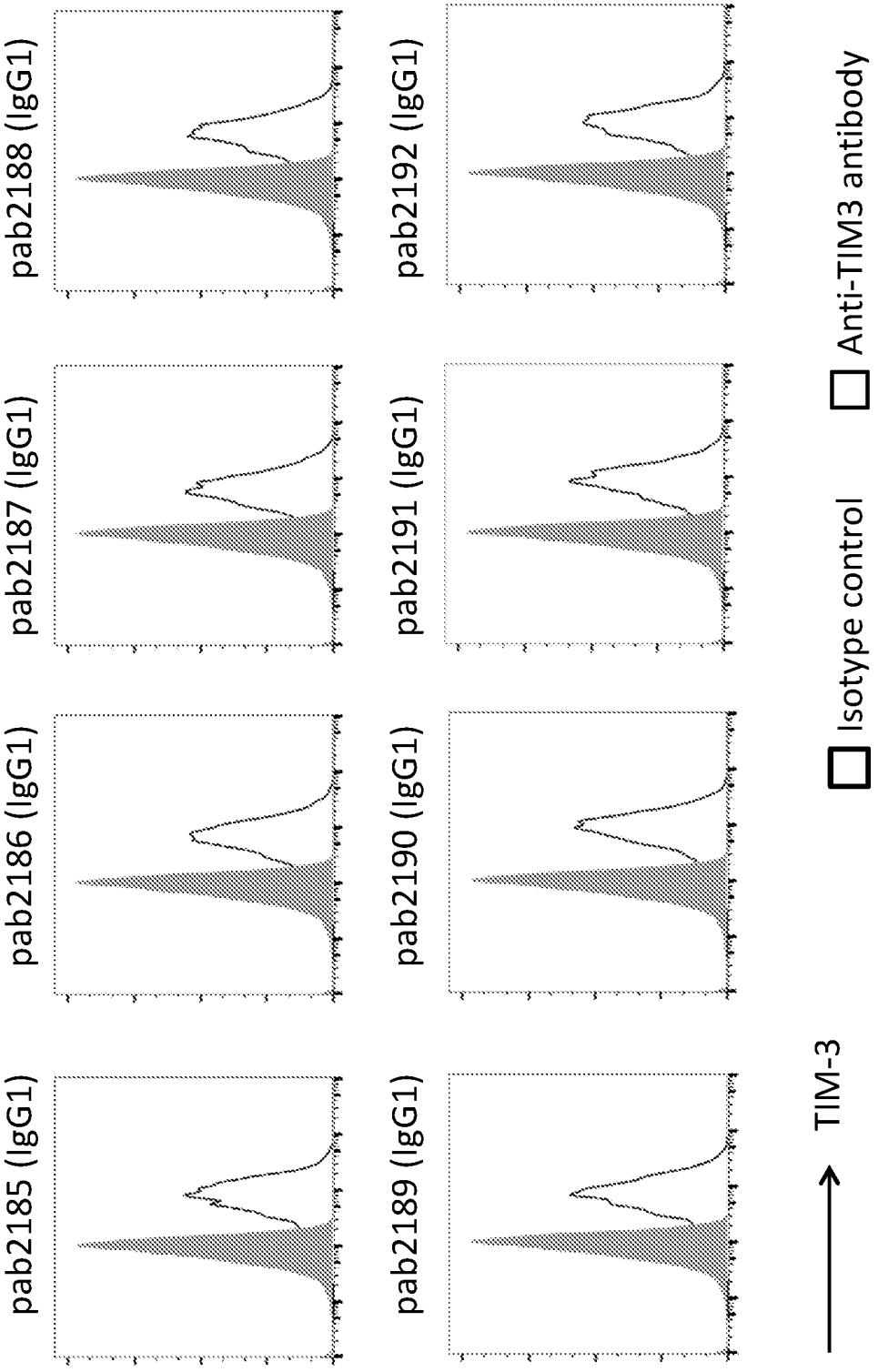

pab2185 (IgG1)
pab2186 (IgG1)
pab2187 (IgG1)
pab2188 (IgG1)
pab2189 (IgG1)
pab2190 (IgG1)
pab2191 (IgG1)
pab2192 (IgG1)
1624-5 cells expressing cynomolgus TIM-3
TIM-3
Isotype control
Anti-TIM3 antibody

Figure 4

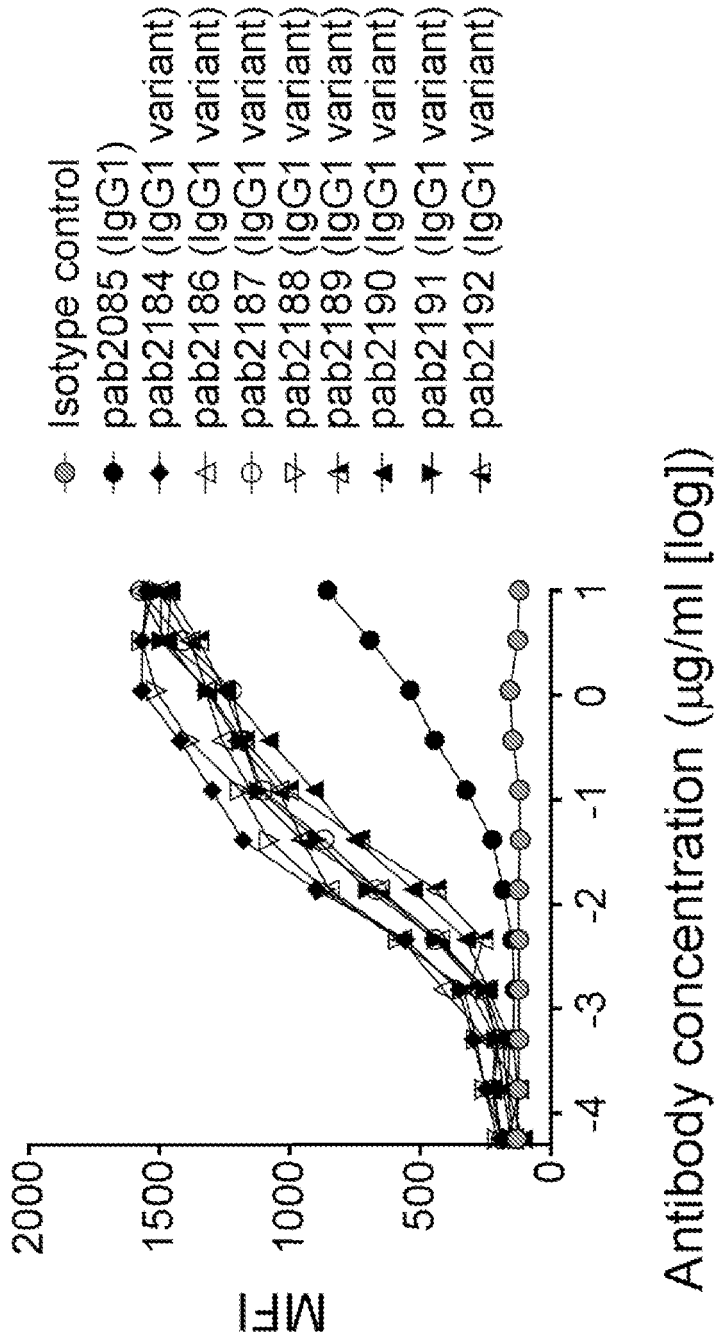

Binding to human CD8+ T cells
Isotype control
pab2085 (IgG1)
pab2184 (IgG1 variant)
pab2186 (IgG1 variant)
pab2187 (IgG1 variant)
pab2188 (IgG1 variant)
pab2189 (IgG1 variant)
pab2190 (IgG1 variant)
pab2191 (IgG1 variant)
pab2192 (IgG1 variant)
MFI
2000
1500
1000
500
0
-4
-3
-2
-1
0
1
Antibody concentration (µg/ml [log])

Figure 5

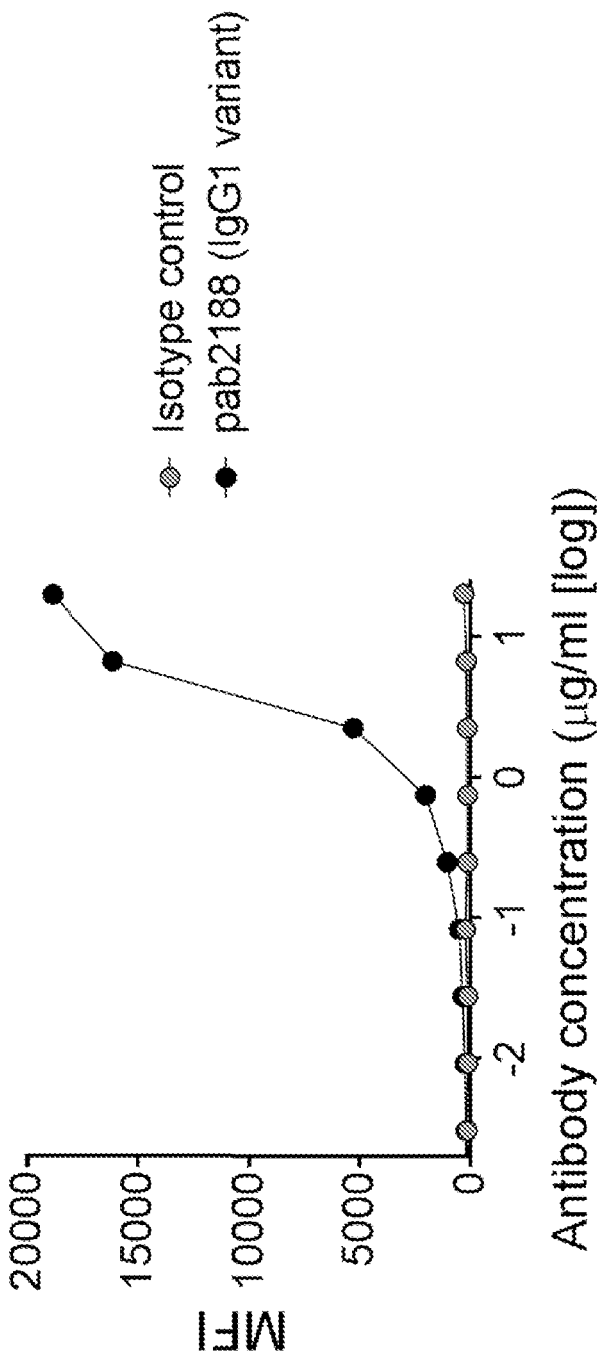

Binding to cynomolgus myeloid cells
Isotype control
pab2188 (IgG1 variant)
MFI
20000
15000
10000
5000
0
-2
-1
0
1
Antibody concentration (µg/ml [log])

Figure 6A

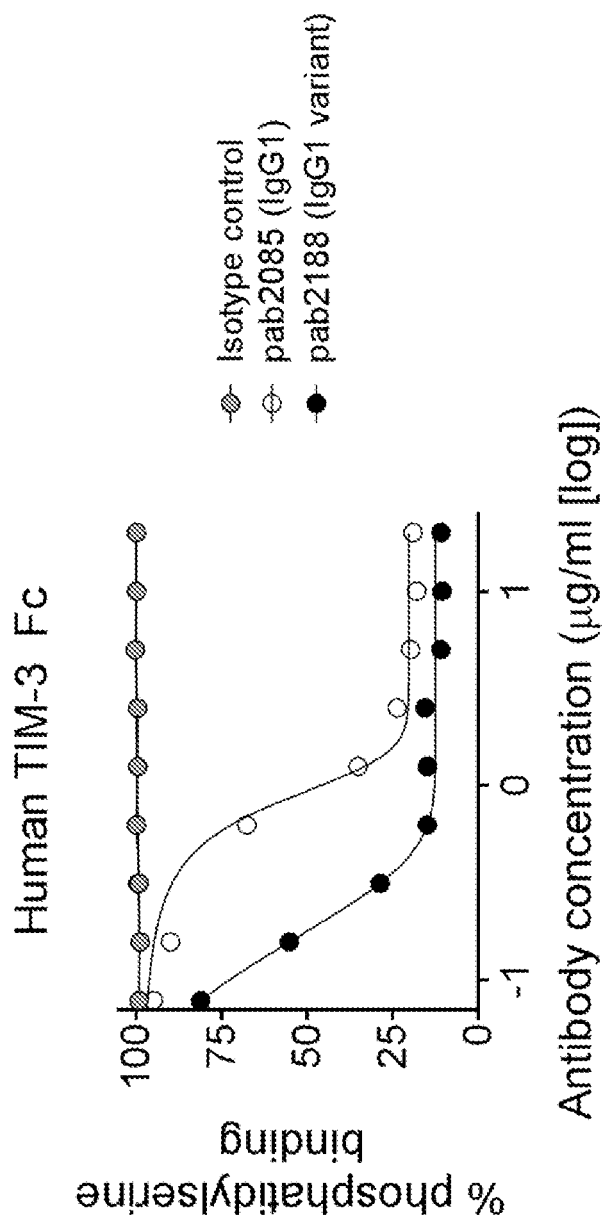
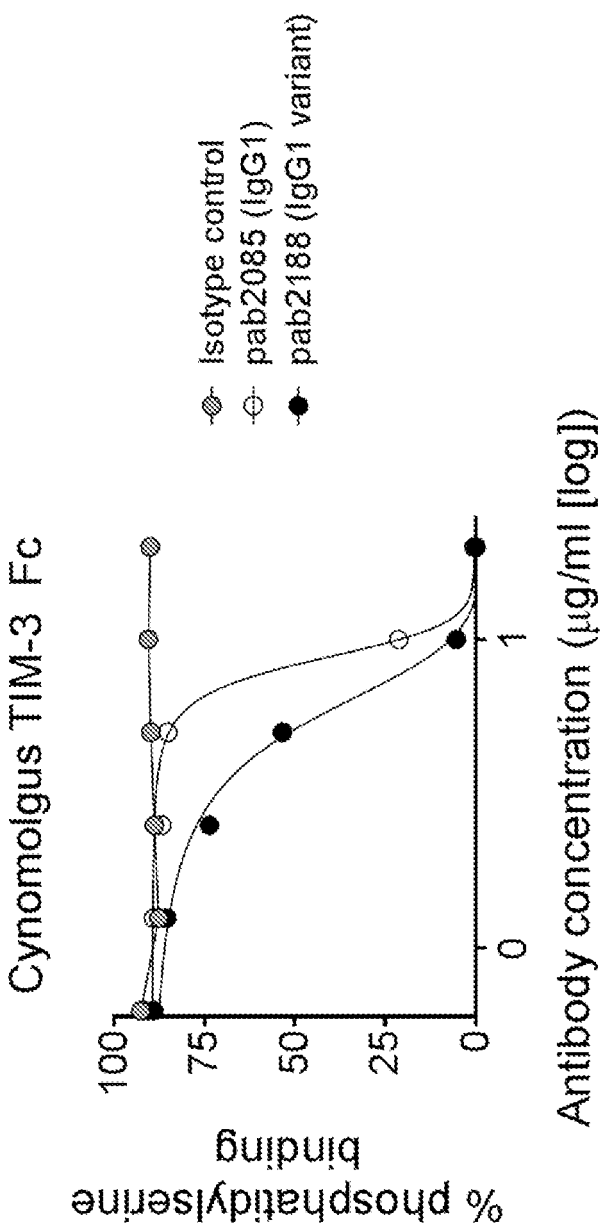


Figure 6B



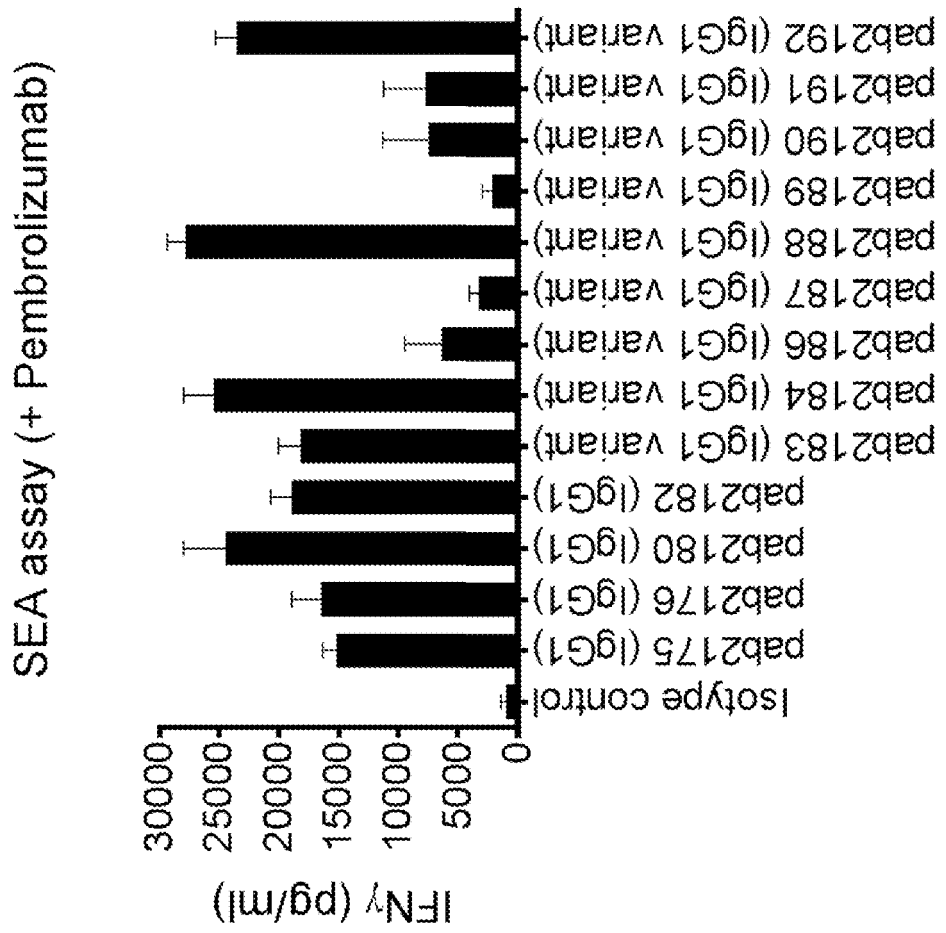

SEA assay (+ Pembrolizumab)
IFNγ (pg/ml)
30000
25000
20000
15000
10000
5000
0
Isotype control
pab2175 (IgG1)
pab2176 (IgG1)
pab2180 (IgG1)
pab2182 (IgG1)
pab2183 (IgG1 variant)
pab2184 (IgG1 variant)
pab2186 (IgG1 variant)
pab2187 (IgG1 variant)
pab2188 (IgG1 variant)
pab2189 (IgG1 variant)
pab2190 (IgG1 variant)
pab2191 (IgG1 variant)
pab2192 (IgG1 variant)

Figure 7

Figure 8A

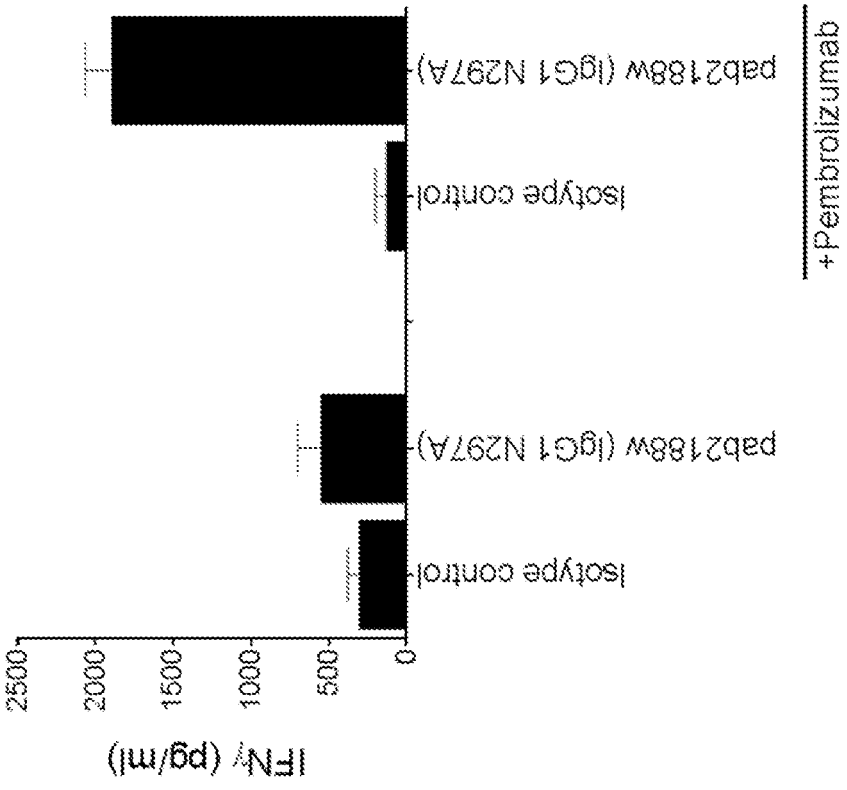


Figure 8B

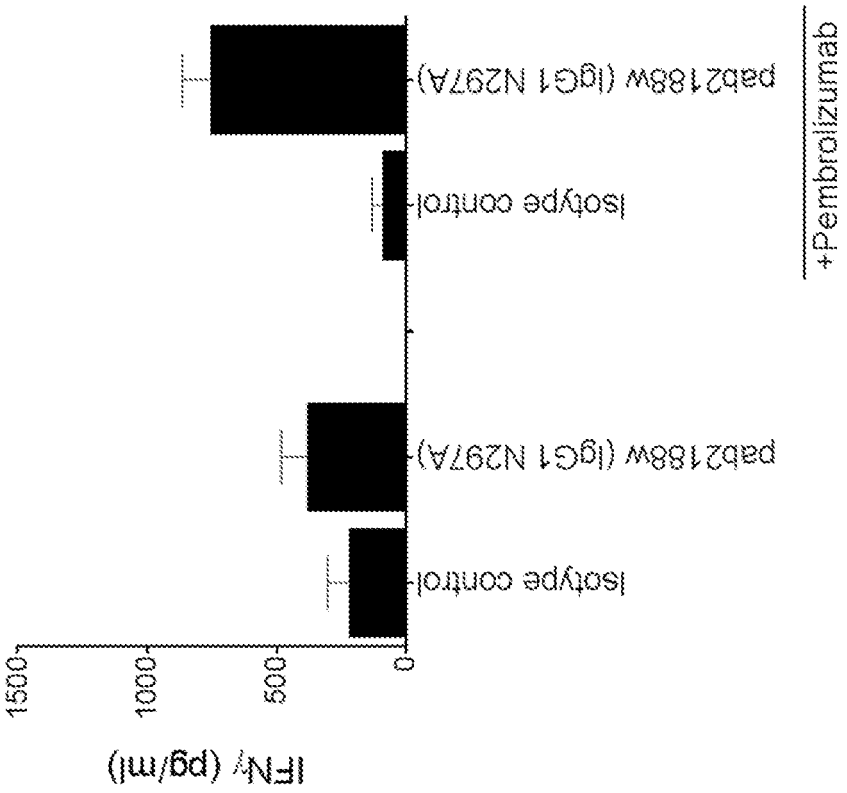


Figure 8C

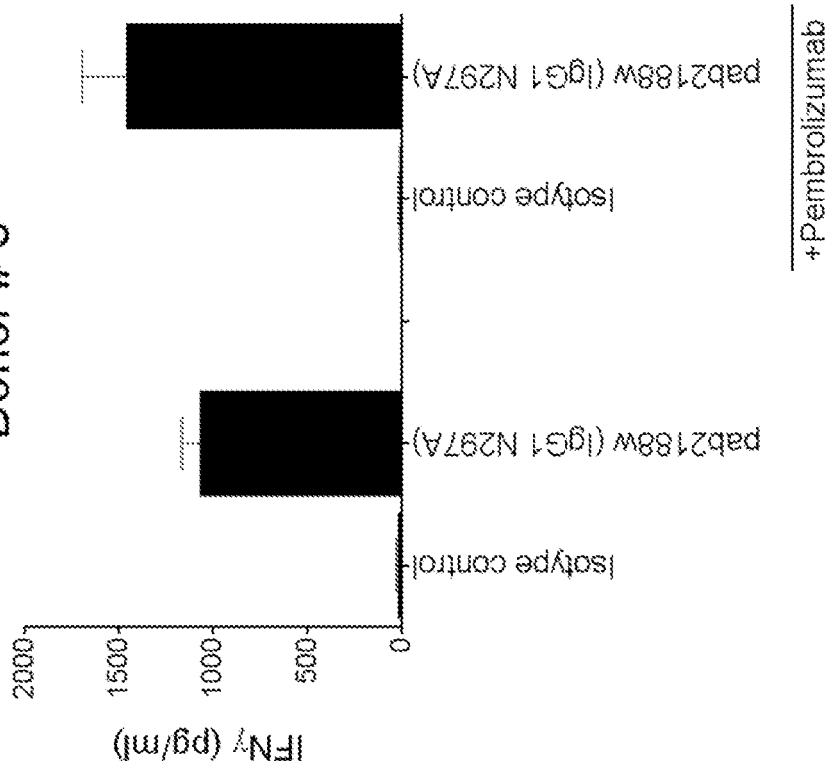


Figure 8D

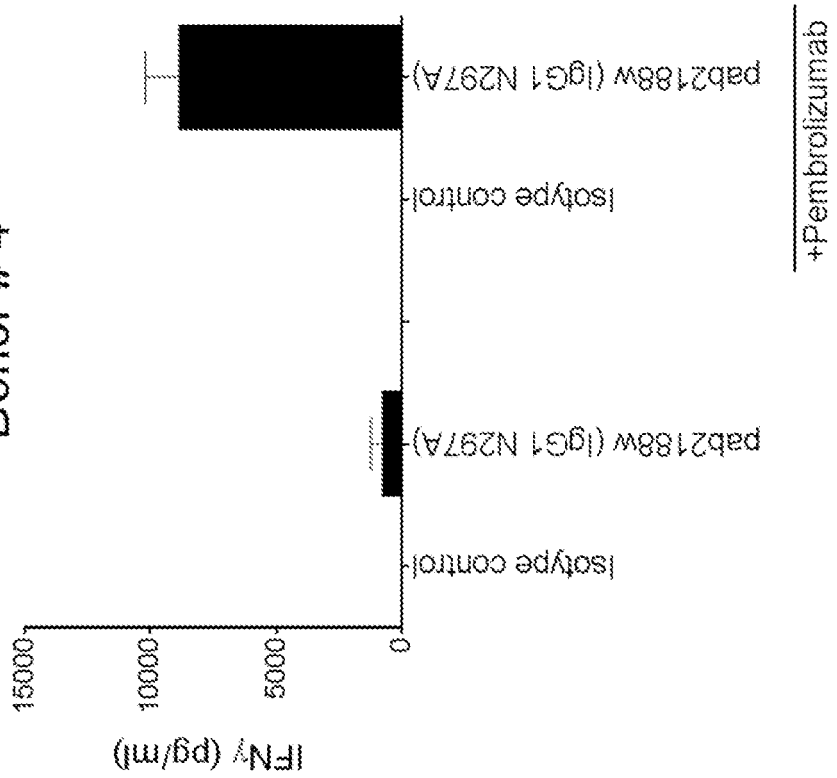


Figure 8F

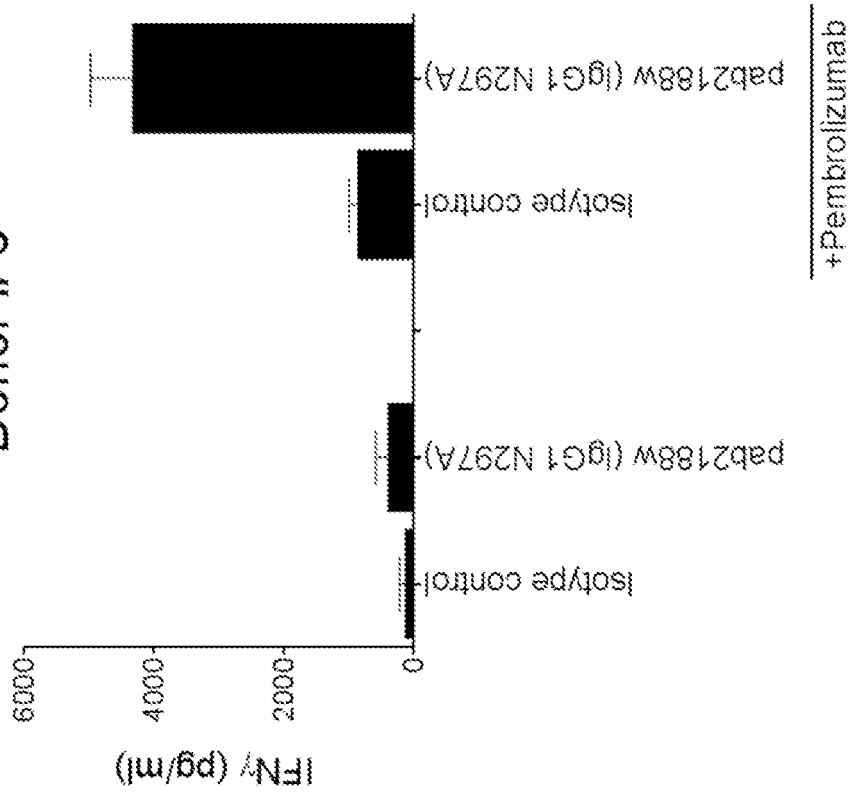


Figure 8E

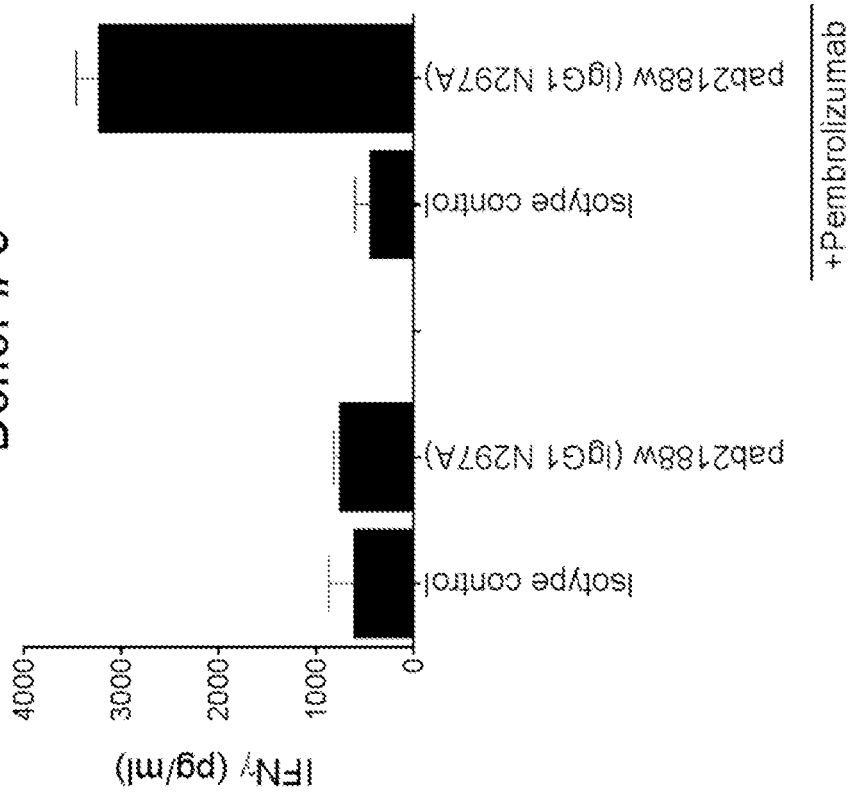


Figure 9A

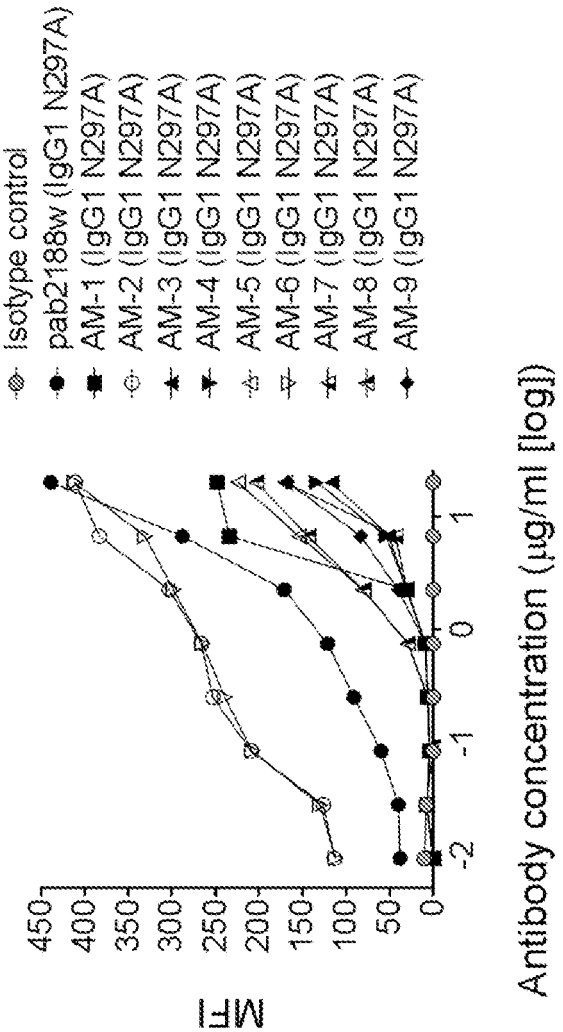


Figure 9B

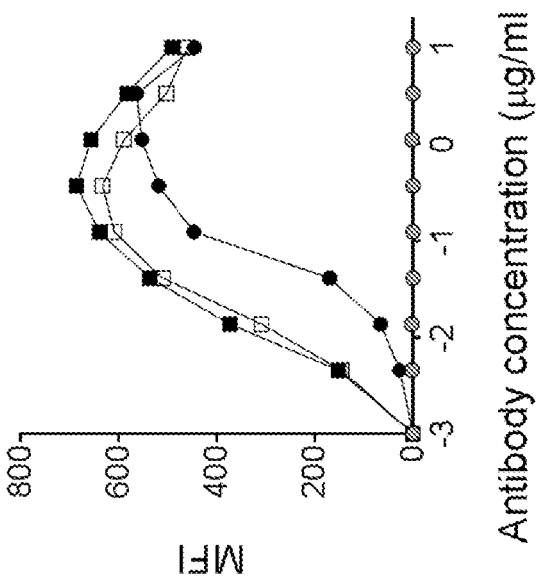


Figure 9C Binding to SEA-stimulated human CD8+ T cells

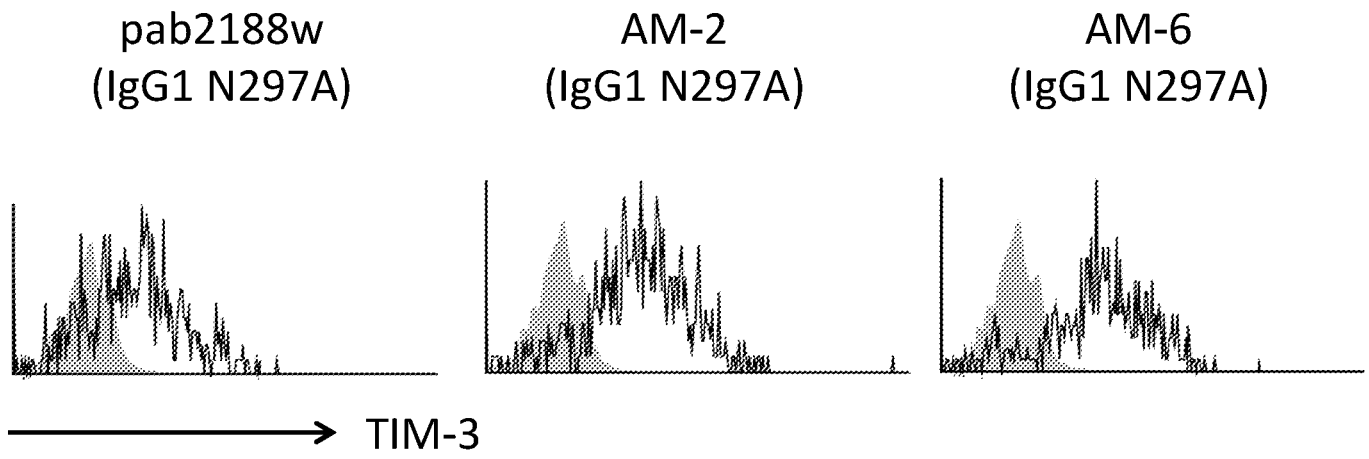


Figure 9D

Binding to SEA-stimulated cynomolgus CD8+ T cells

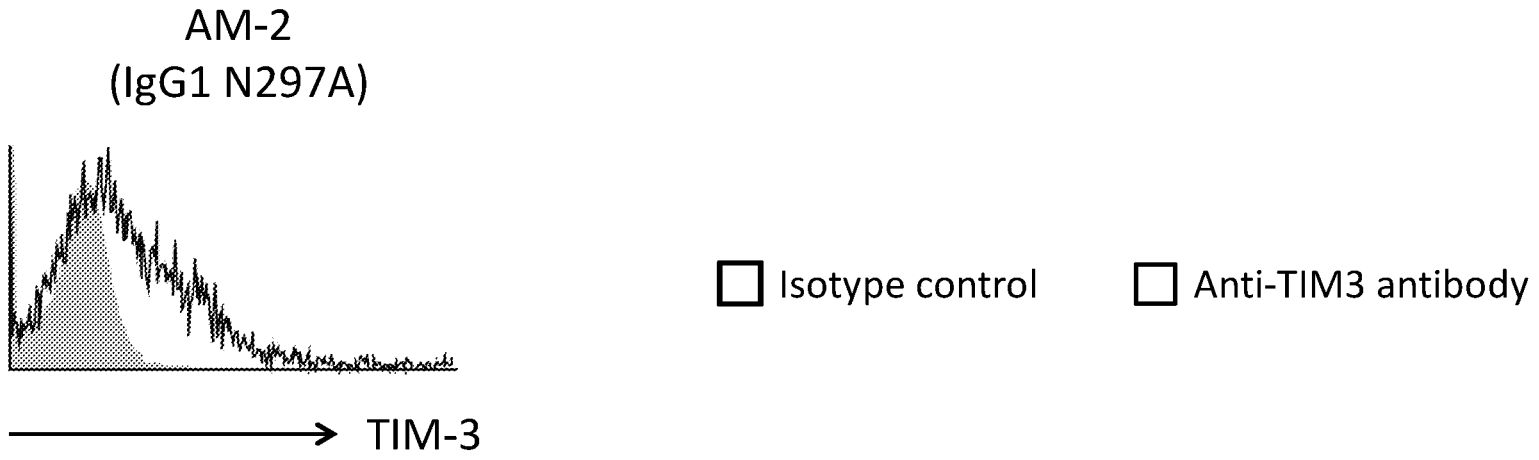


Figure 9E

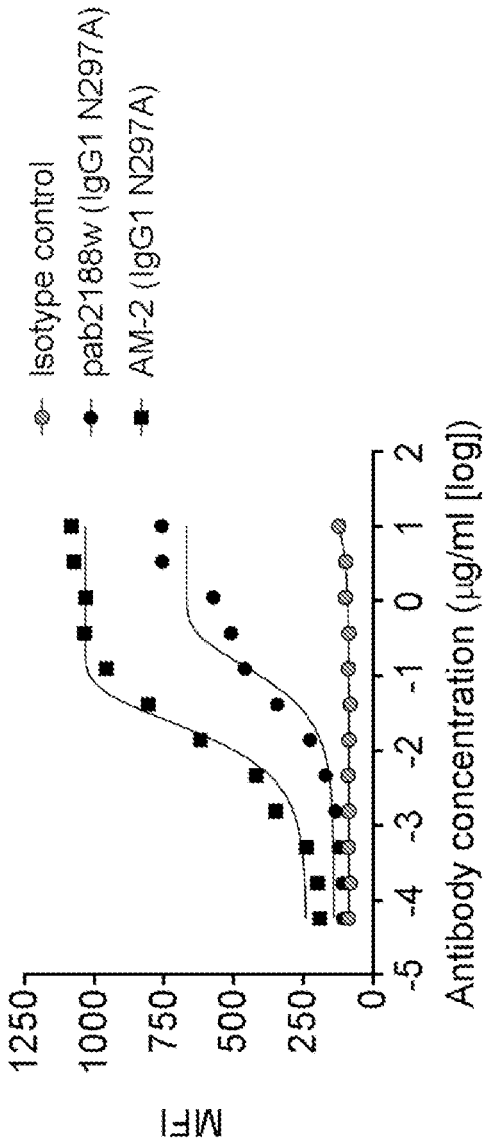


Figure 9F

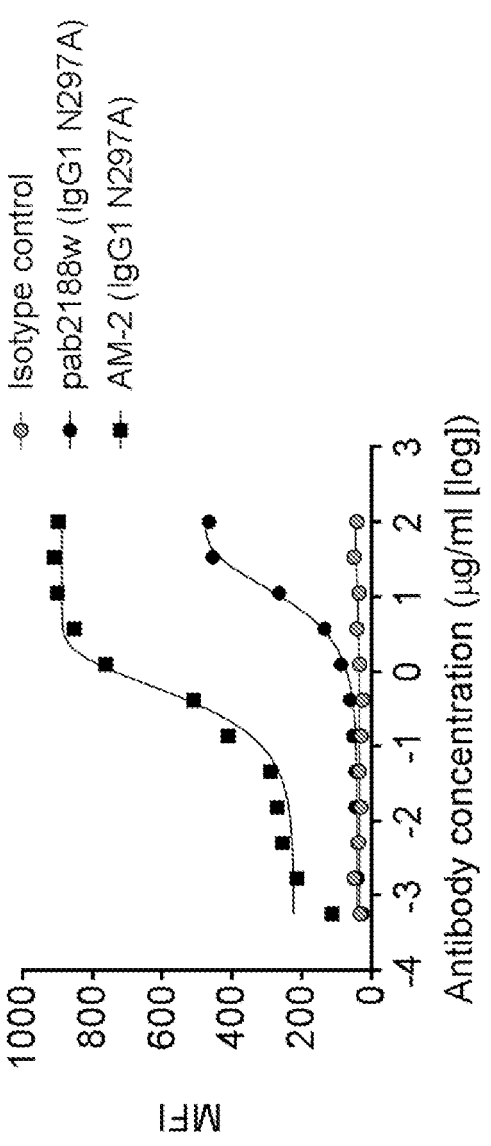


Figure 10A

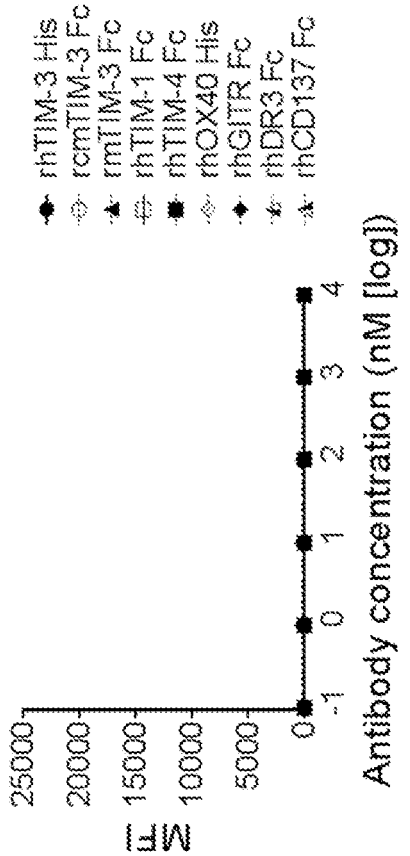


Figure 10B

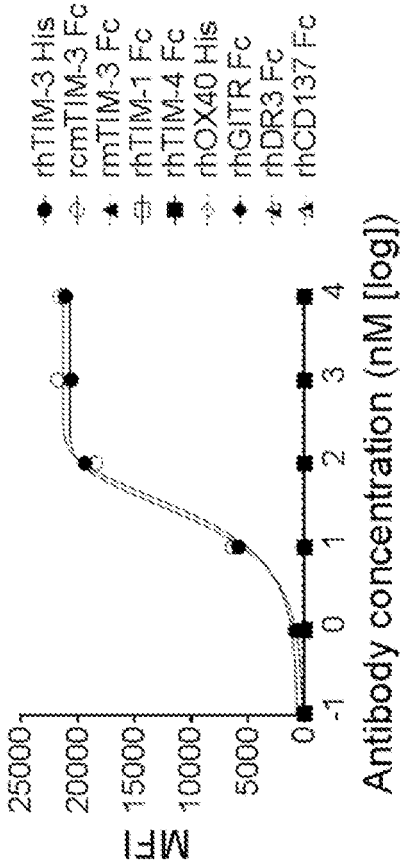


Figure 10C

Figure 10D

Figure 11A

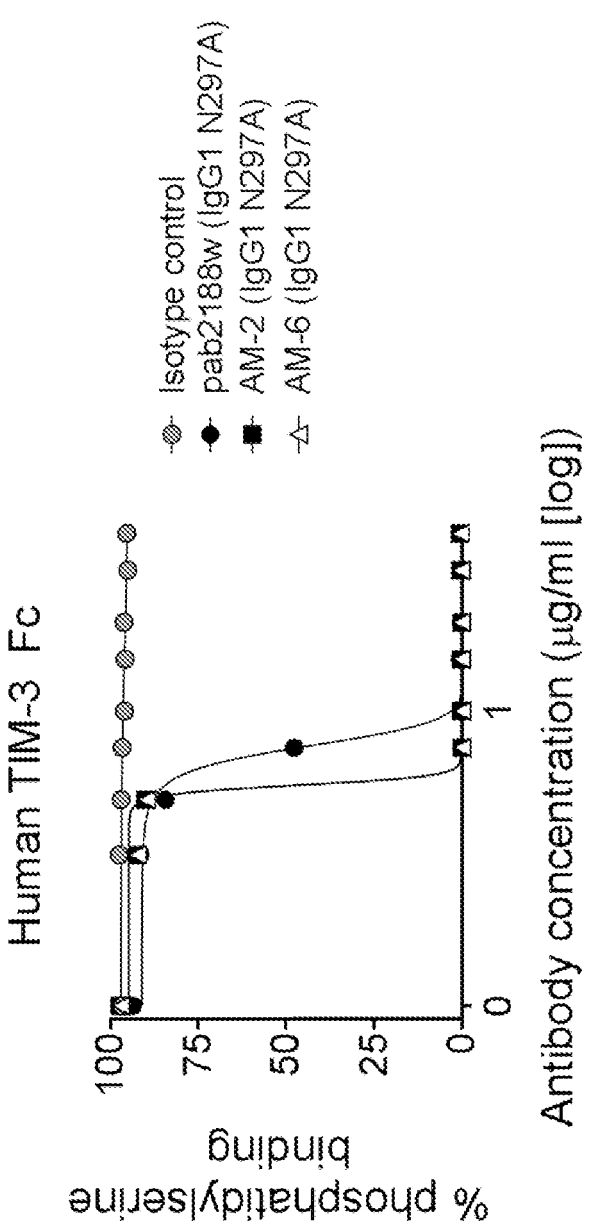
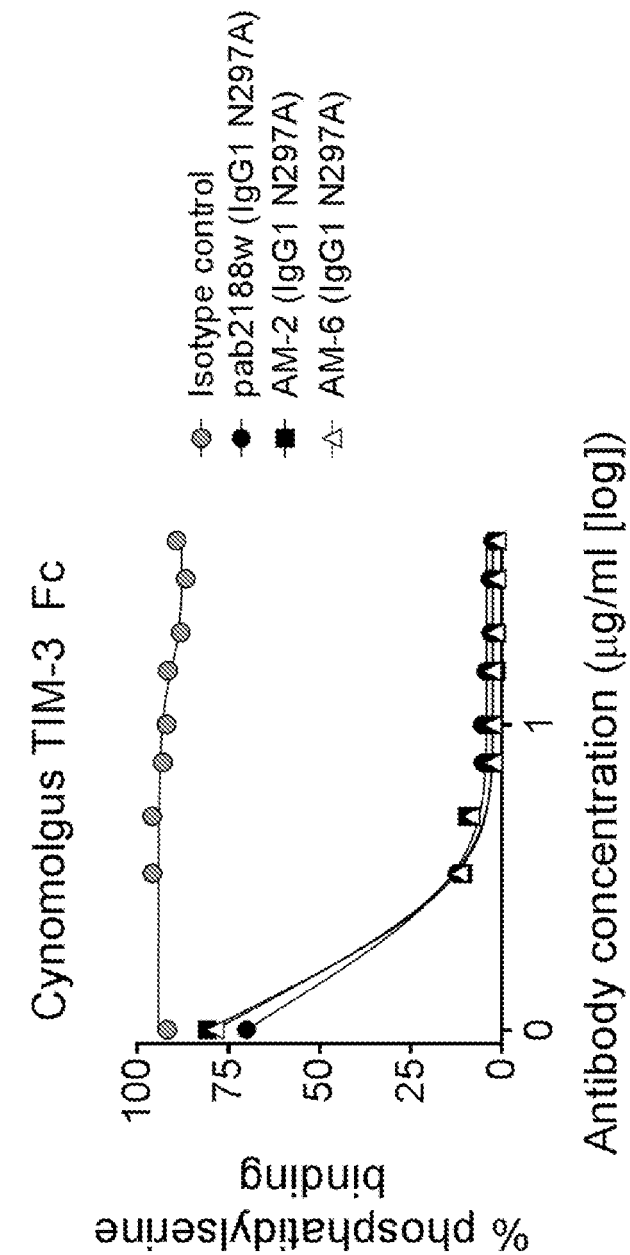


Figure 11B



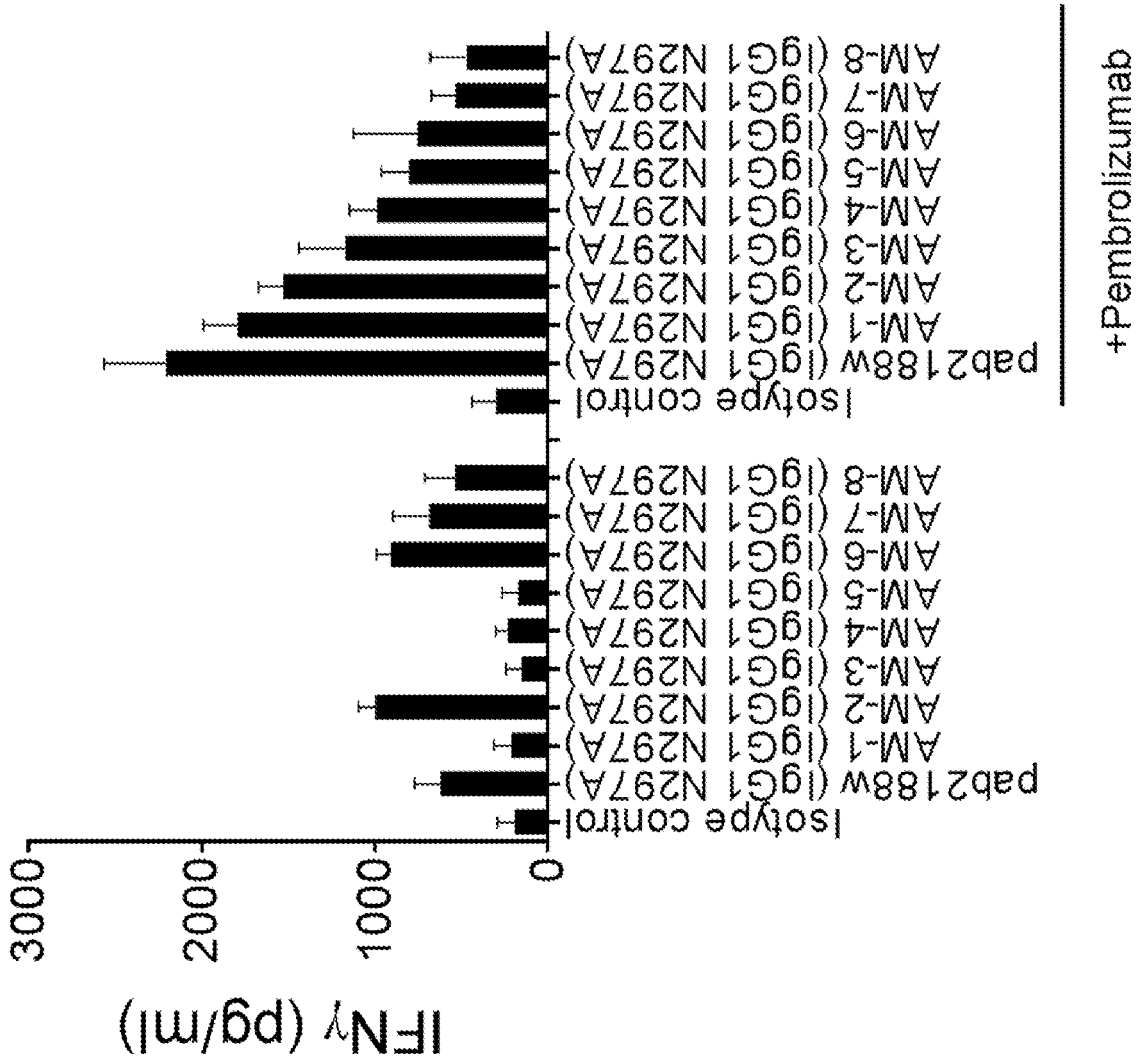

IFNγ (pg/ml)
3000
2000
1000
0
Isotype control
pab2188w (IgG1 N297A)
AM-1 (IgG1 N297A)
AM-2 (IgG1 N297A)
AM-3 (IgG1 N297A)
AM-4 (IgG1 N297A)
AM-5 (IgG1 N297A)
AM-6 (IgG1 N297A)
AM-7 (IgG1 N297A)
AM-8 (IgG1 N297A)
Isotype control
pab2188w (IgG1 N297A)
AM-1 (IgG1 N297A)
AM-2 (IgG1 N297A)
AM-3 (IgG1 N297A)
AM-4 (IgG1 N297A)
AM-5 (IgG1 N297A)
AM-6 (IgG1 N297A)
AM-7 (IgG1 N297A)
AM-8 (IgG1 N297A)
+Pembrolizumab

Figure 12A

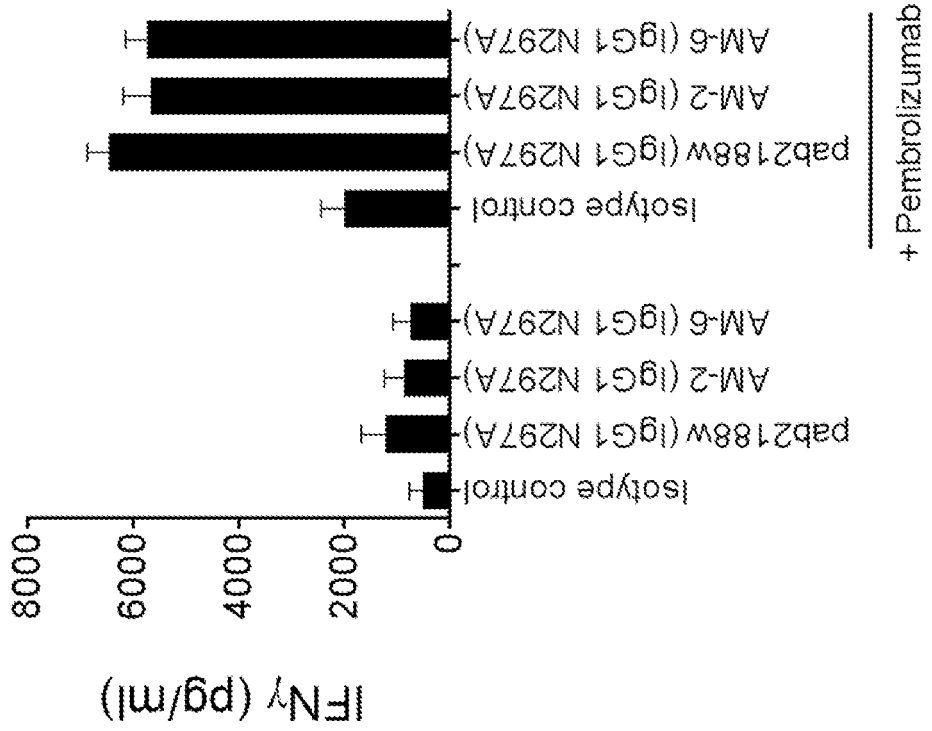

IFNγ (pg/ml)
8000
6000
4000
2000
0
Isotype control
pab2188w (IgG1 N297A)
AM-2 (IgG1 N297A)
AM-6 (IgG1 N297A)
Isotype control
pab2188w (IgG1 N297A)
AM-2 (IgG1 N297A)
AM-6 (IgG1 N297A)
+ Pembrolizumab

Figure 12B

Figure 13A

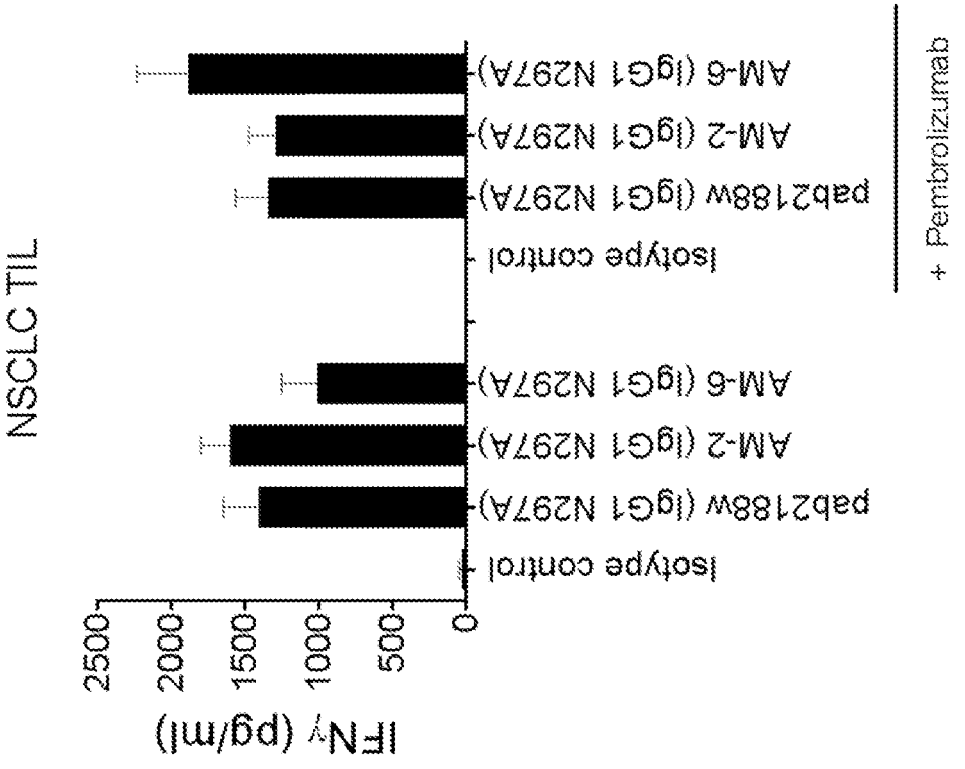


Figure 13B

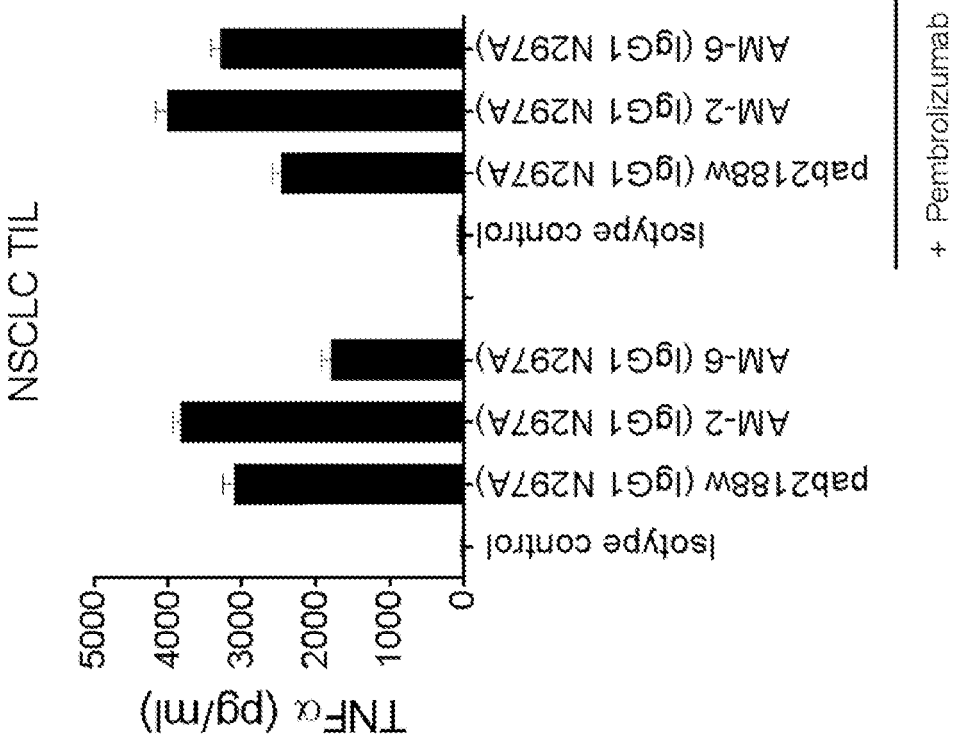


Figure 13C

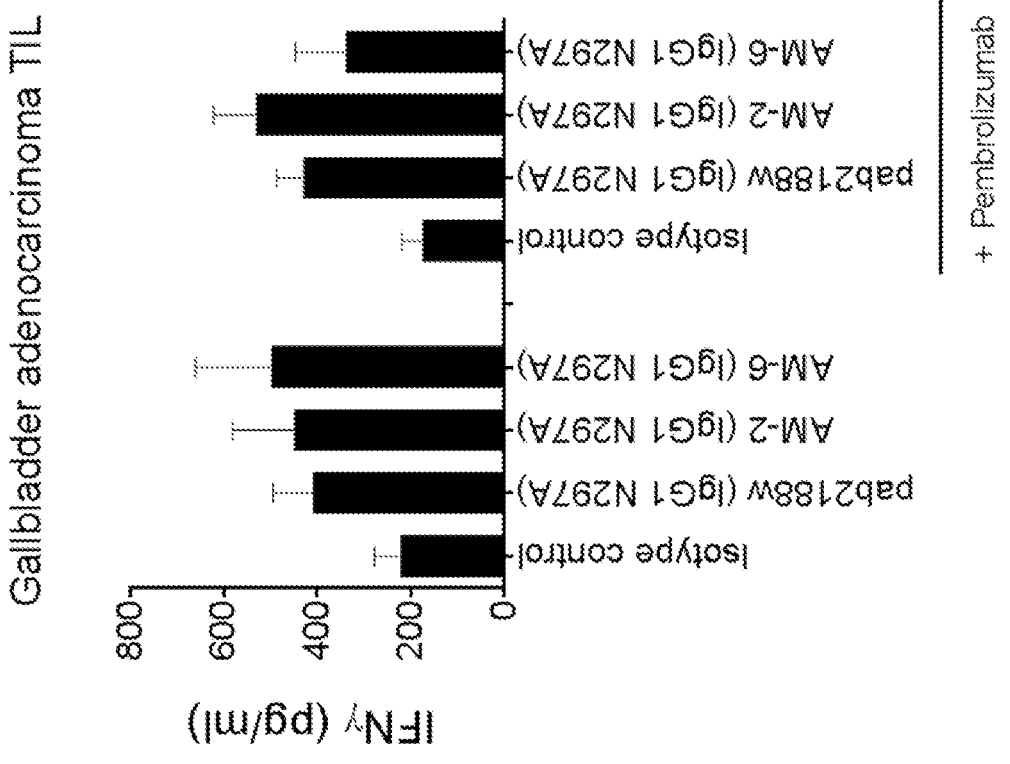


Figure 13D

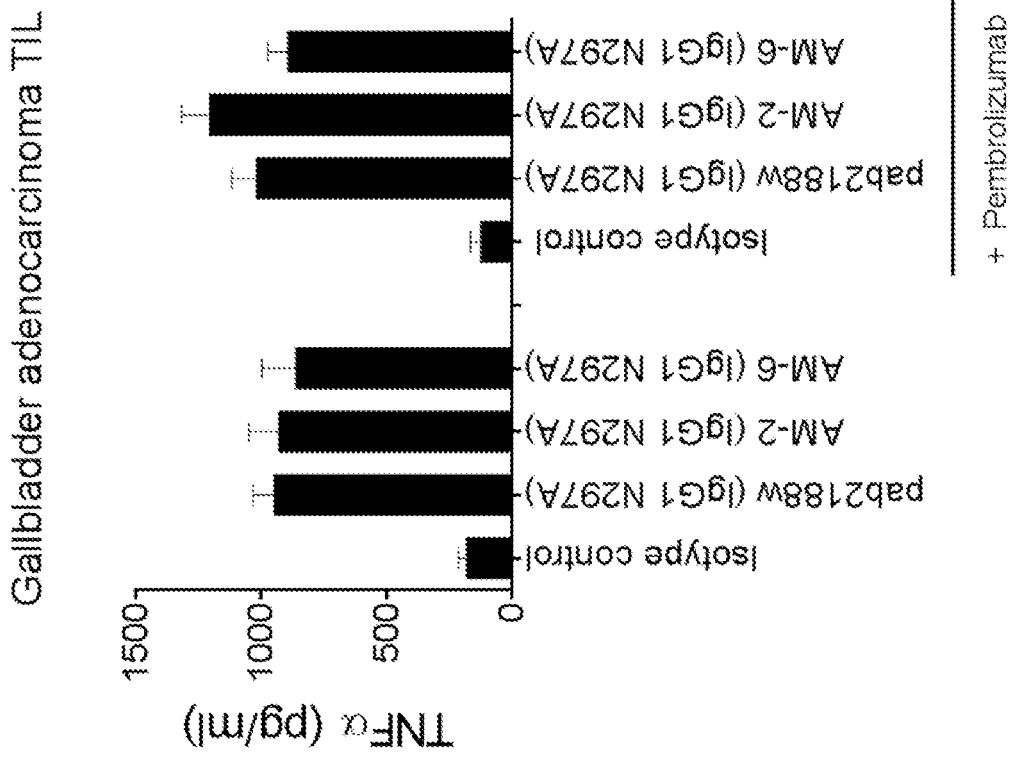


Figure 13E

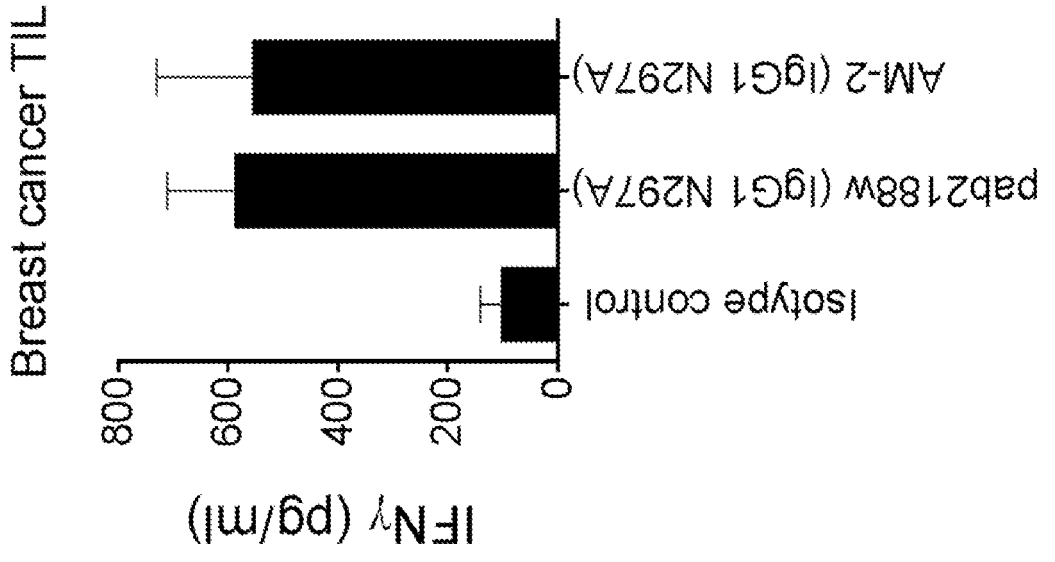


Figure 13F

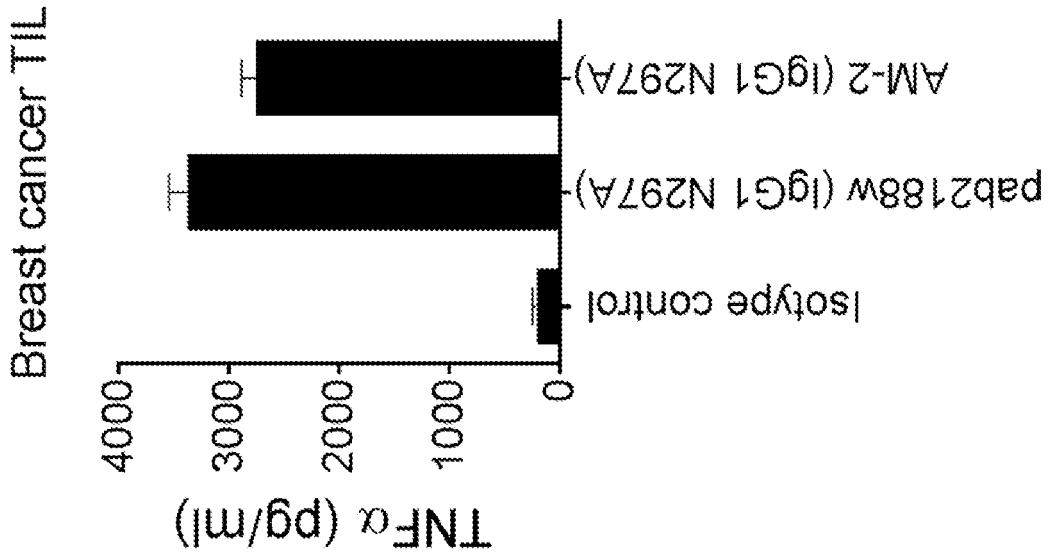


Figure 14A

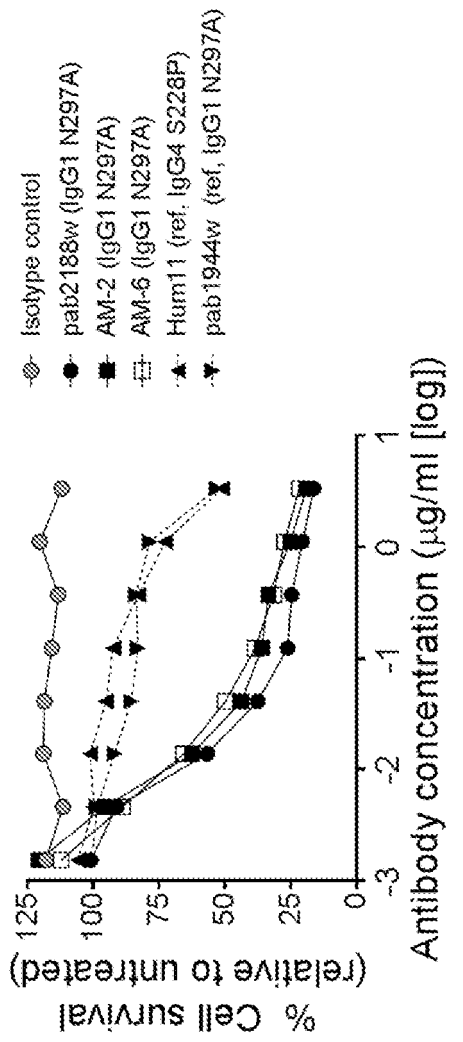


Figure 14B

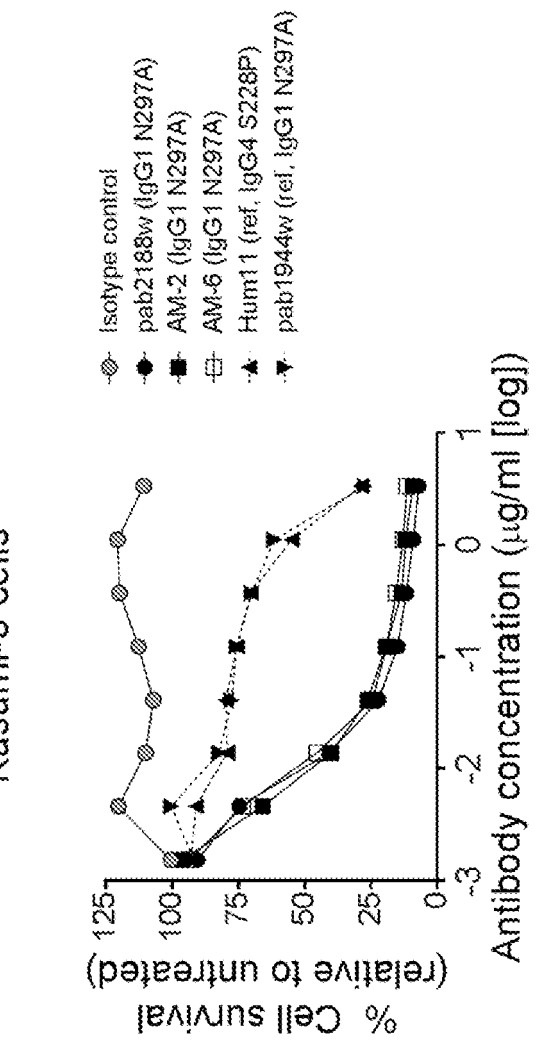


Figure 14C

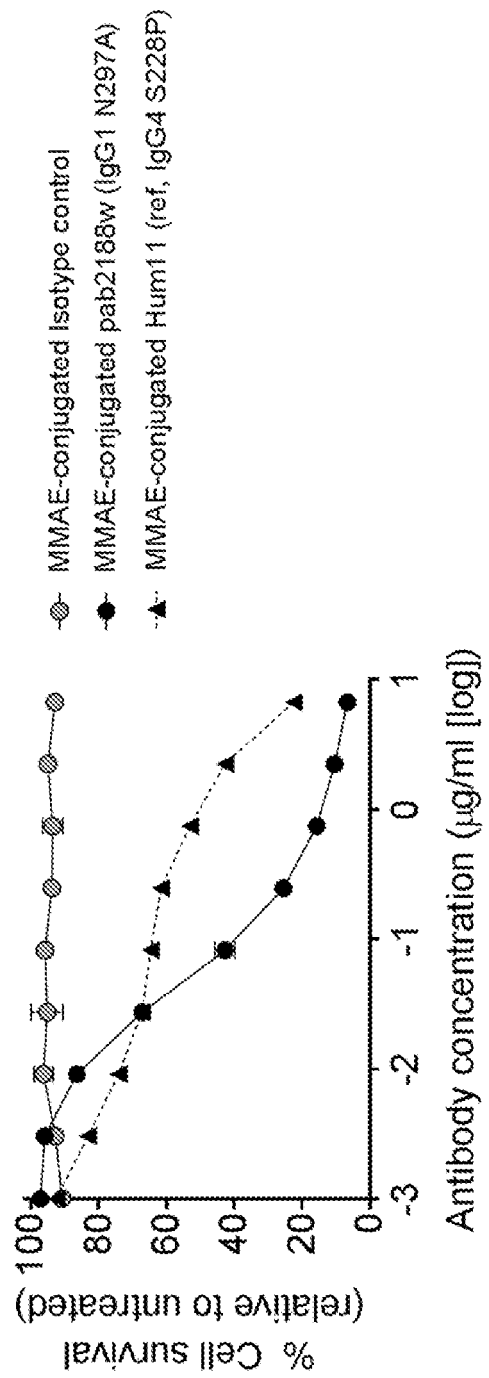
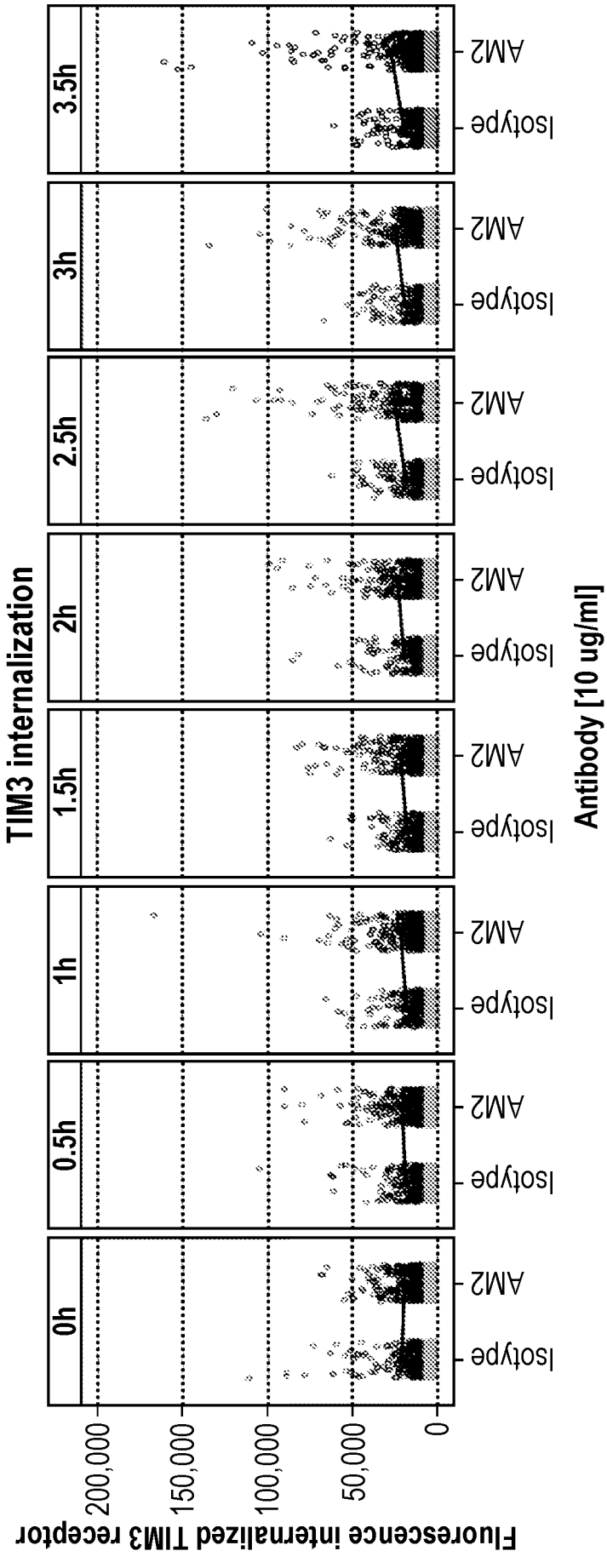

MMAE-conjugated Isotype control
MMAE-conjugated pab2188w (IgG1 N297A)
MMAE-conjugated Hum11 (ref, IgG4 S228P)
% Cell survival (relative to untreated)
100
80
60
40
20
0
-3
-2
-1
0
1
Antibody concentration (µg/ml [log])

Figure 15


TIM3 internalization
0h
0.5h
1h
1.5h
2h
2.5h
3h
3.5h
Isotype
AM2
Fluorescence internalized TIM3 receptor
200,000
150,000
100,000
50,000
0
Antibody [10 ug/ml]