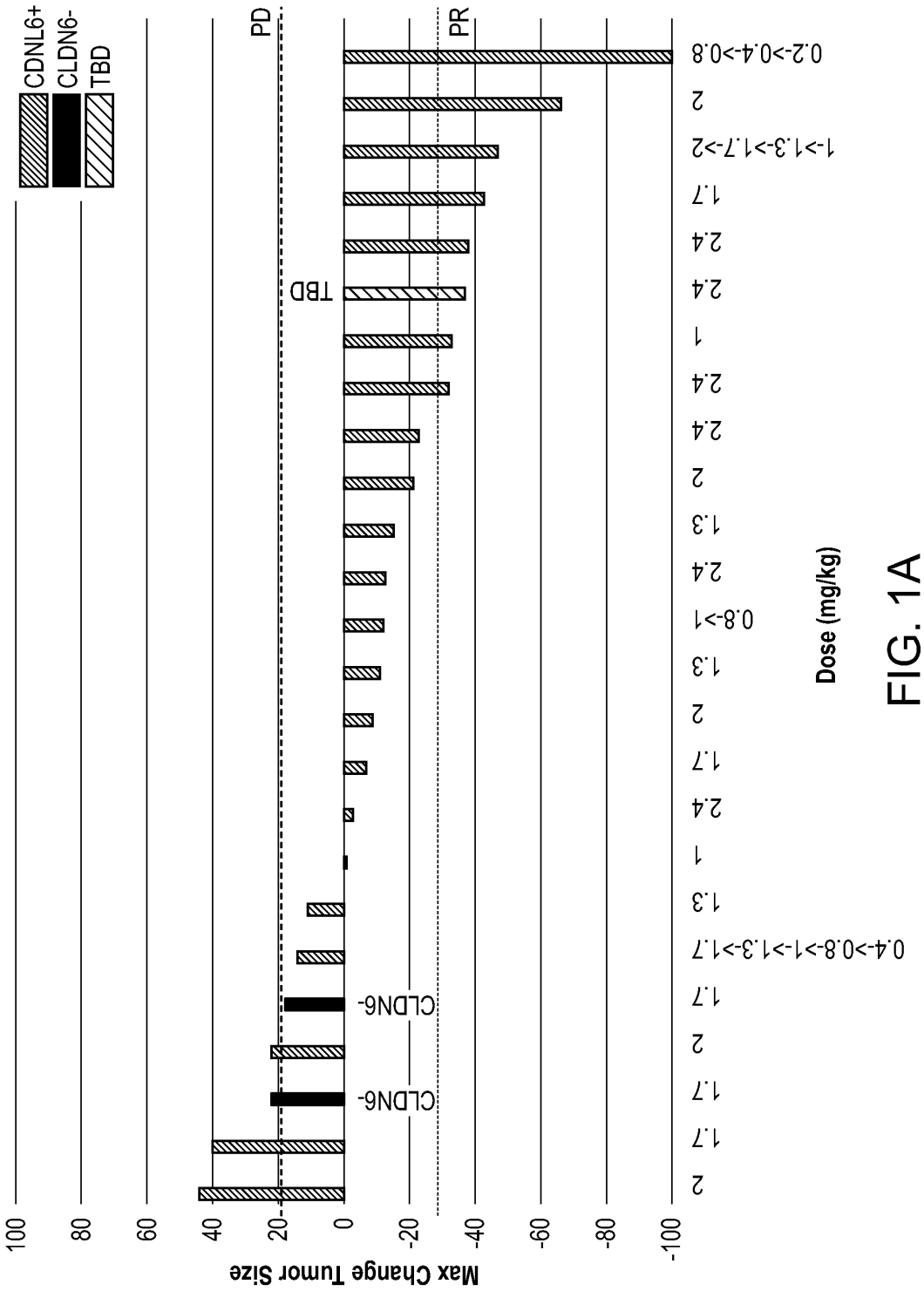

CDNL6+
CLDN6-
TBD
PD
PR
Max Change Tumor Size
100
80
60
40
20
0
-20
-40
-60
-80
-100
CLDN6-
CLDN6-
TBD
2
1.7
1.7
2
1.7
0.4->0.8->1->1.3->1.7
1.3
1
2.4
1.7
2
1.3
0.8->1
2.4
1.3
2
2.4
2.4
1
2.4
2.4
1.7
1->1.3->1.7->2
2
0.2->0.4->0.8
Dose (mg/kg)

FIG. 1A

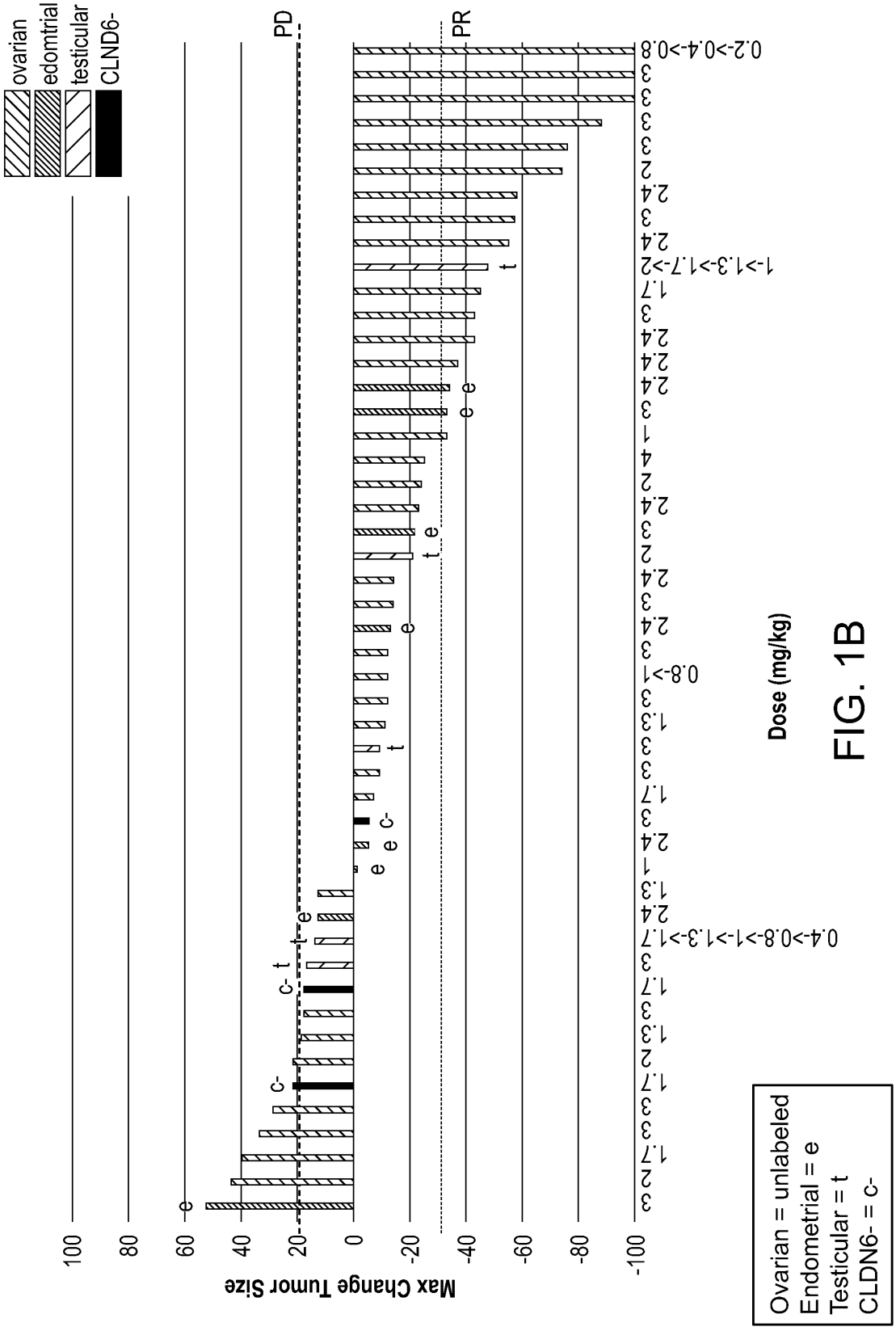

ovarian
edomtrial
testicular
CLND6-
PD
PR
Max Change Tumor Size
100
80
60
40
20
0
-20
-40
-60
-80
-100
0.2->0.4->0.8
1->1.3->1.7->2
0.4->0.8->1->1.3->1.7
0.8->1
Dose (mg/kg)
Ovarian = unlabeled
Endometrial = e
Testicular = t
CLDN6- = c-

FIG. 1B

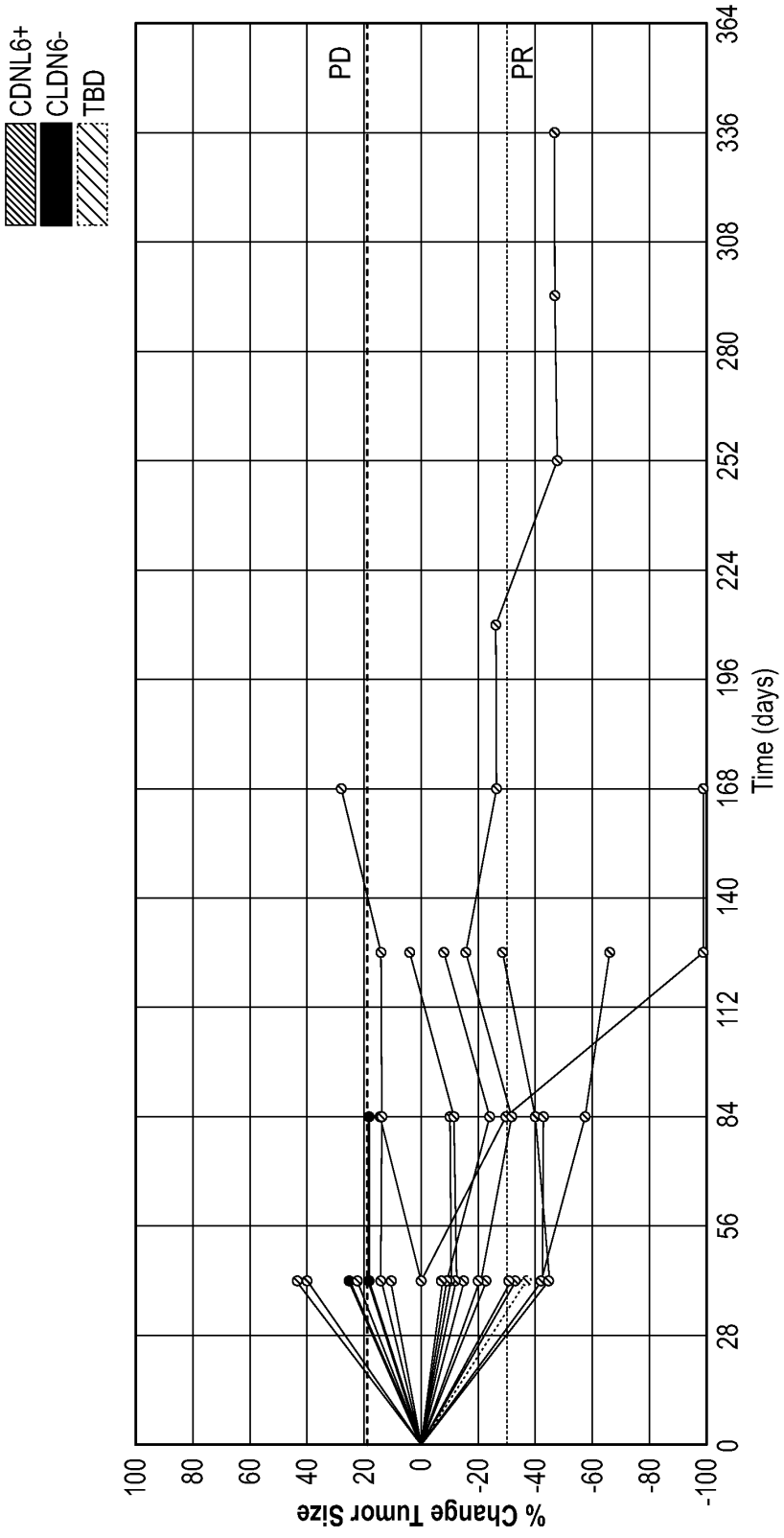

CDNL6+
CLDN6-
TBD
PD
PR
% Change Tumor Size
100
80
60
40
20
0
-20
-40
-60
-80
-100
0
28
56
84
112
140
168
196
224
252
280
308
336
364
Time (days)

FIG. 2A

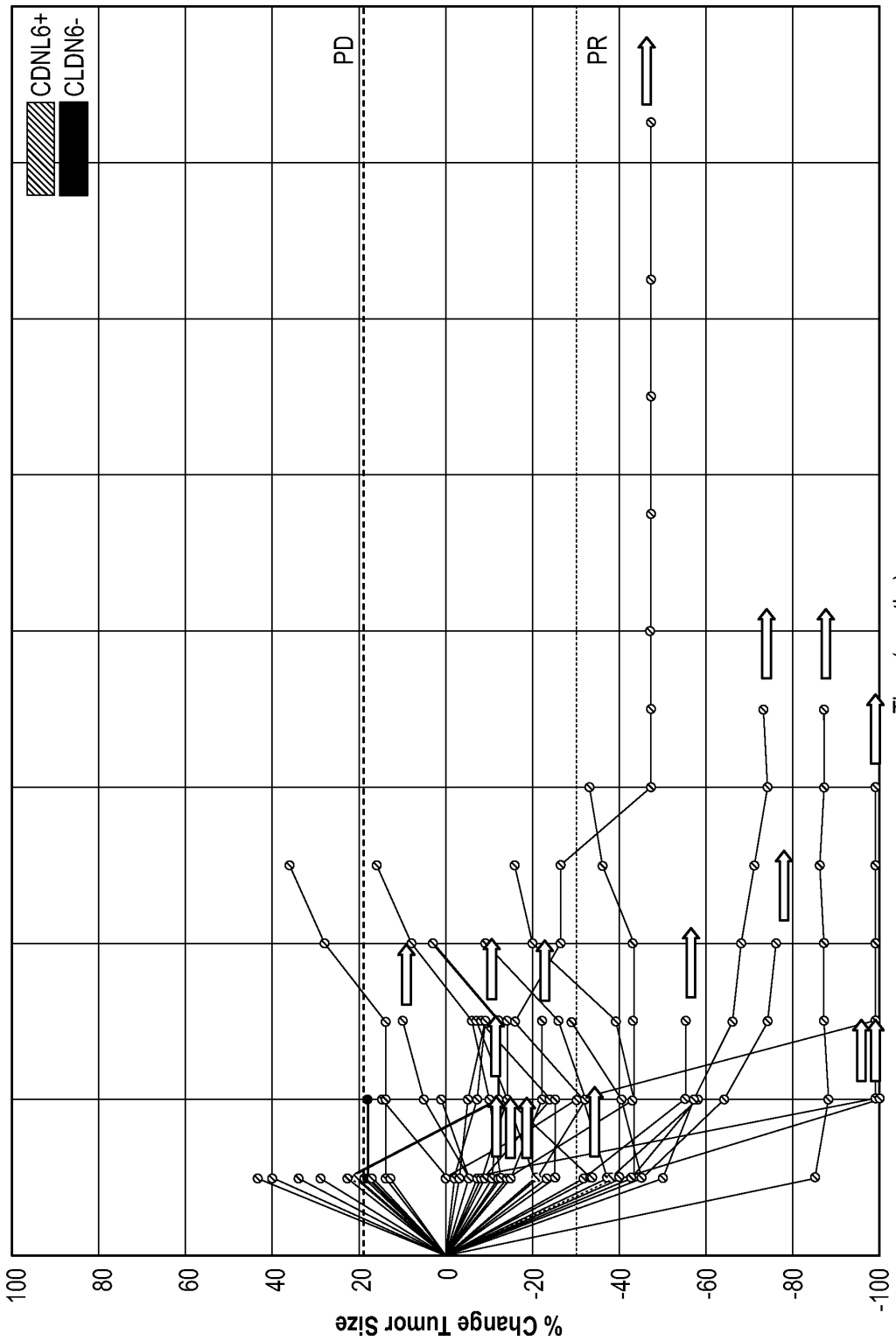
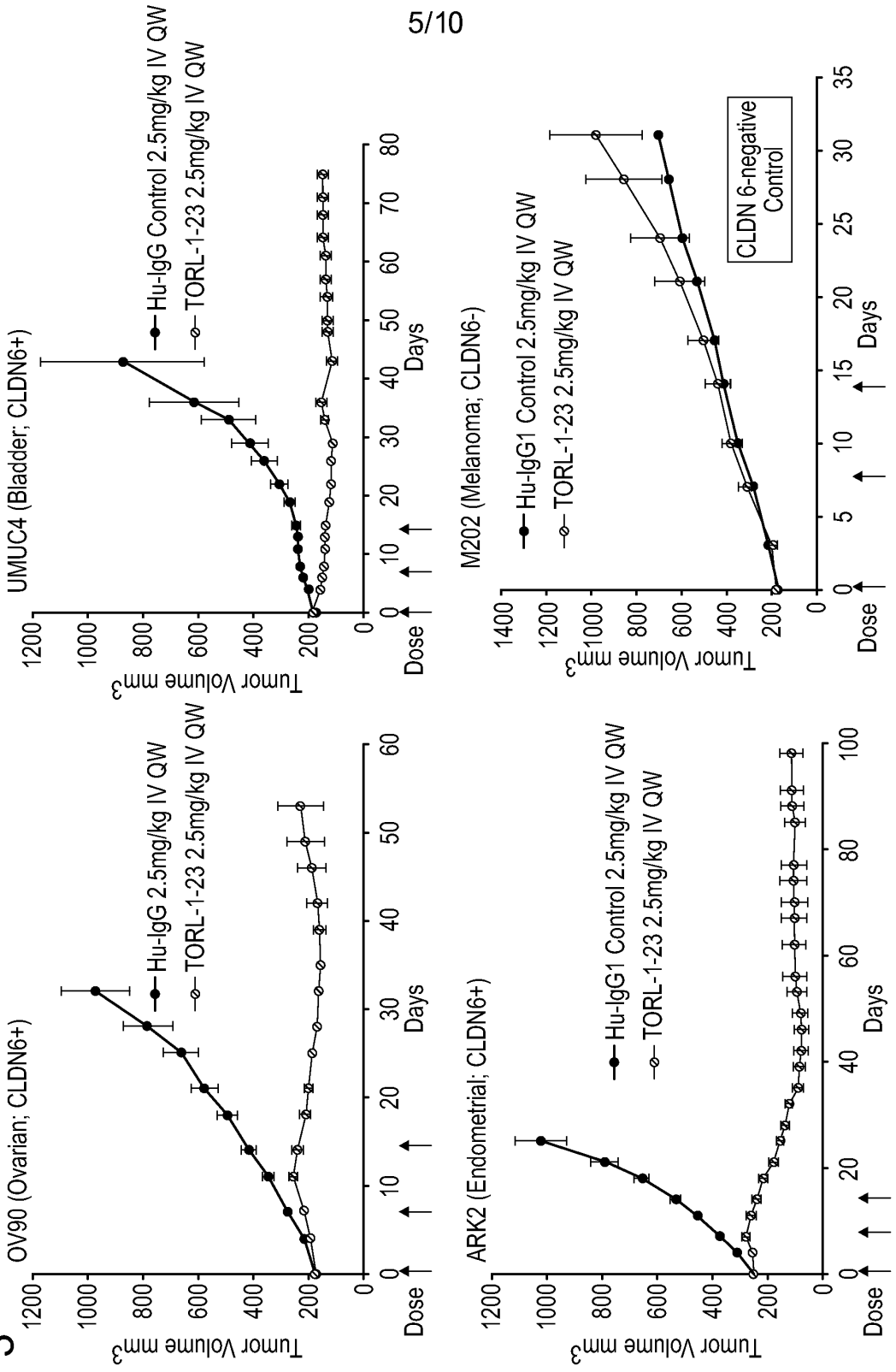

CDNL6+
CLDN6-
PD
PR
100
80
60
40
20
0
-20
-40
-60
-80
-100
% Change Tumor Size
Time (months)

FIG. 2B

FIG. 3


UMUC4 (Bladder; CLDN6+)
Hu-IgG Control 2.5mg/kg IV QW
TORL-1-23 2.5mg/kg IV QW
Tumor Volume mm3
Days
Dose
M202 (Melanoma; CLDN6-)
Hu-IgG1 Control 2.5mg/kg IV QW
TORL-1-23 2.5mg/kg IV QW
CLDN 6-negative Control
OV90 (Ovarian; CLDN6+)
Hu-IgG 2.5mg/kg IV QW
TORL-1-23 2.5mg/kg IV QW
ARK2 (Endometrial; CLDN6+)
Hu-IgG1 Control 2.5mg/kg IV QW
TORL-1-23 2.5mg/kg IV QW

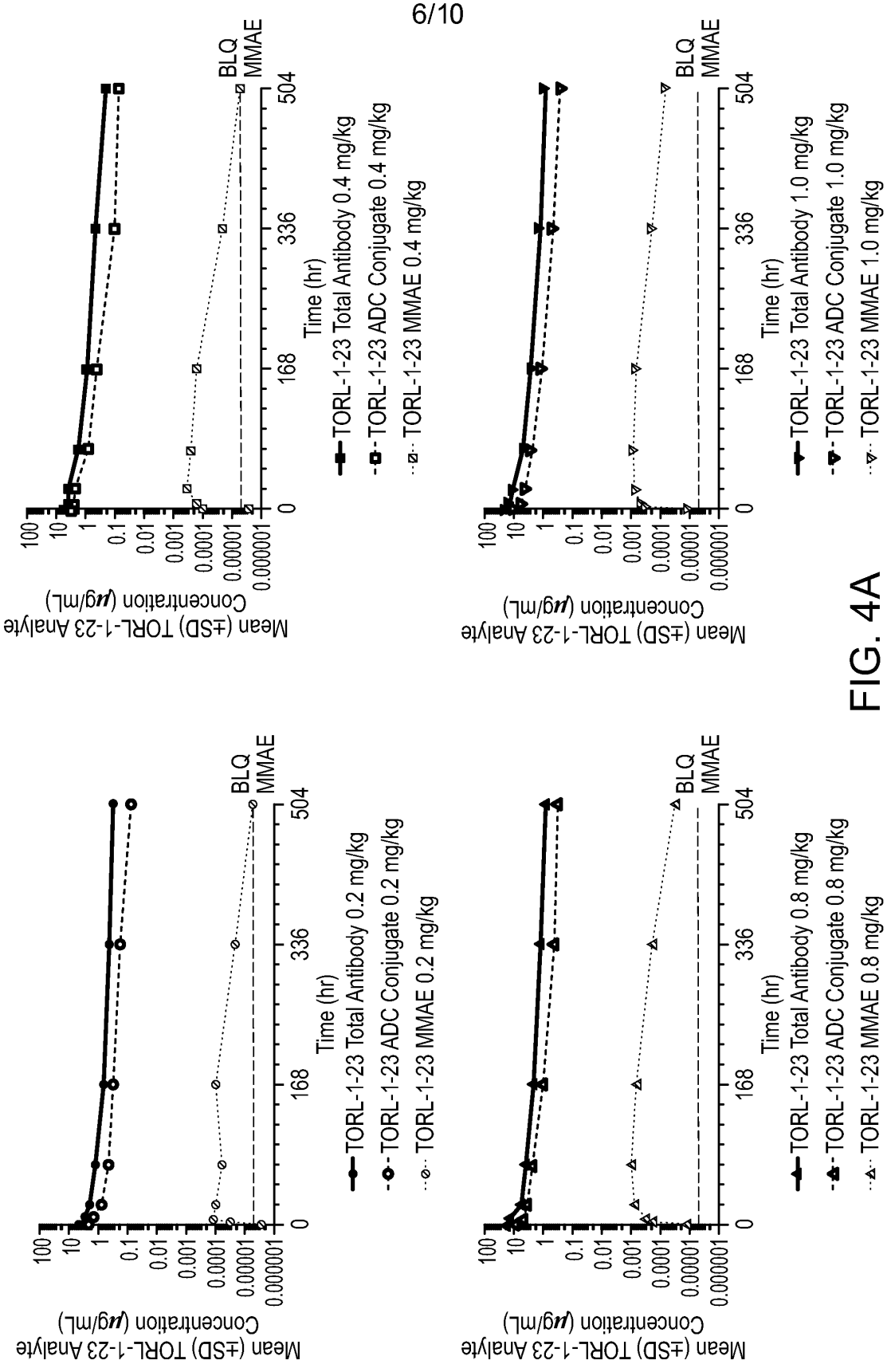

Mean (±SD) TORL-1-23 Analyte Concentration (μg/mL)
100
10
1
0.1
0.01
0.001
0.0001
0.00001
0.000001
0
168
336
504
Time (hr)
BLQ
MMAE
TORL-1-23 Total Antibody 0.4 mg/kg
TORL-1-23 ADC Conjugate 0.4 mg/kg
TORL-1-23 MMAE 0.4 mg/kg
TORL-1-23 Total Antibody 1.0 mg/kg
TORL-1-23 ADC Conjugate 1.0 mg/kg
TORL-1-23 MMAE 1.0 mg/kg
TORL-1-23 Total Antibody 0.2 mg/kg
TORL-1-23 ADC Conjugate 0.2 mg/kg
TORL-1-23 MMAE 0.2 mg/kg
TORL-1-23 Total Antibody 0.8 mg/kg
TORL-1-23 ADC Conjugate 0.8 mg/kg
TORL-1-23 MMAE 0.8 mg/kg

FIG. 4A

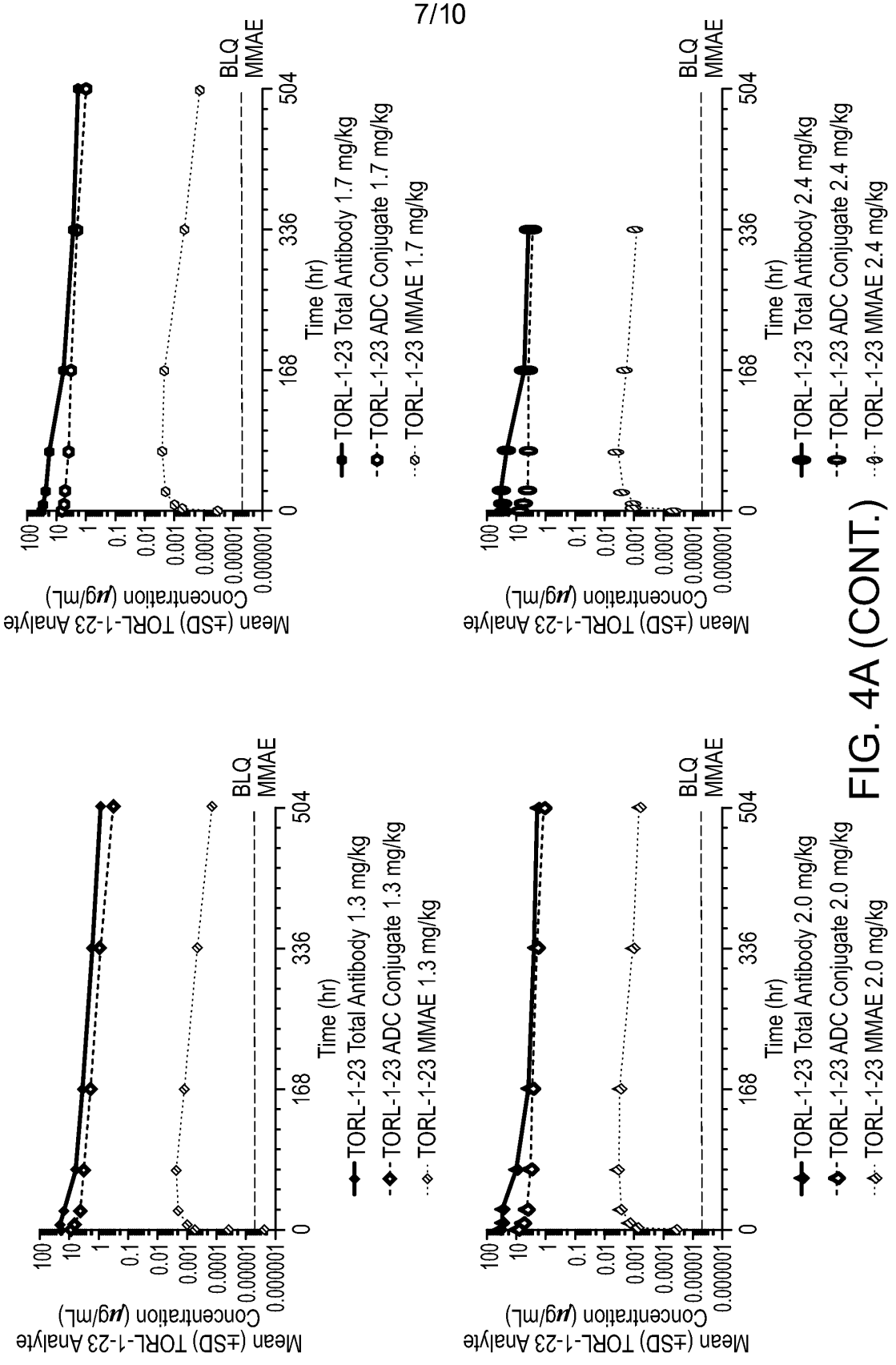

Mean (±SD) TORL-1-23 Analyte Concentration (μg/mL)
100
10
1
0.1
0.01
0.001
0.0001
0.00001
0.000001
0
168
336
504
Time (hr)
BLQ
MMAE
TORL-1-23 Total Antibody 1.7 mg/kg
TORL-1-23 ADC Conjugate 1.7 mg/kg
TORL-1-23 MMAE 1.7 mg/kg
TORL-1-23 Total Antibody 2.4 mg/kg
TORL-1-23 ADC Conjugate 2.4 mg/kg
TORL-1-23 MMAE 2.4 mg/kg
TORL-1-23 Total Antibody 1.3 mg/kg
TORL-1-23 ADC Conjugate 1.3 mg/kg
TORL-1-23 MMAE 1.3 mg/kg
TORL-1-23 Total Antibody 2.0 mg/kg
TORL-1-23 ADC Conjugate 2.0 mg/kg
TORL-1-23 MMAE 2.0 mg/kg

FIG. 4A (CONT.)

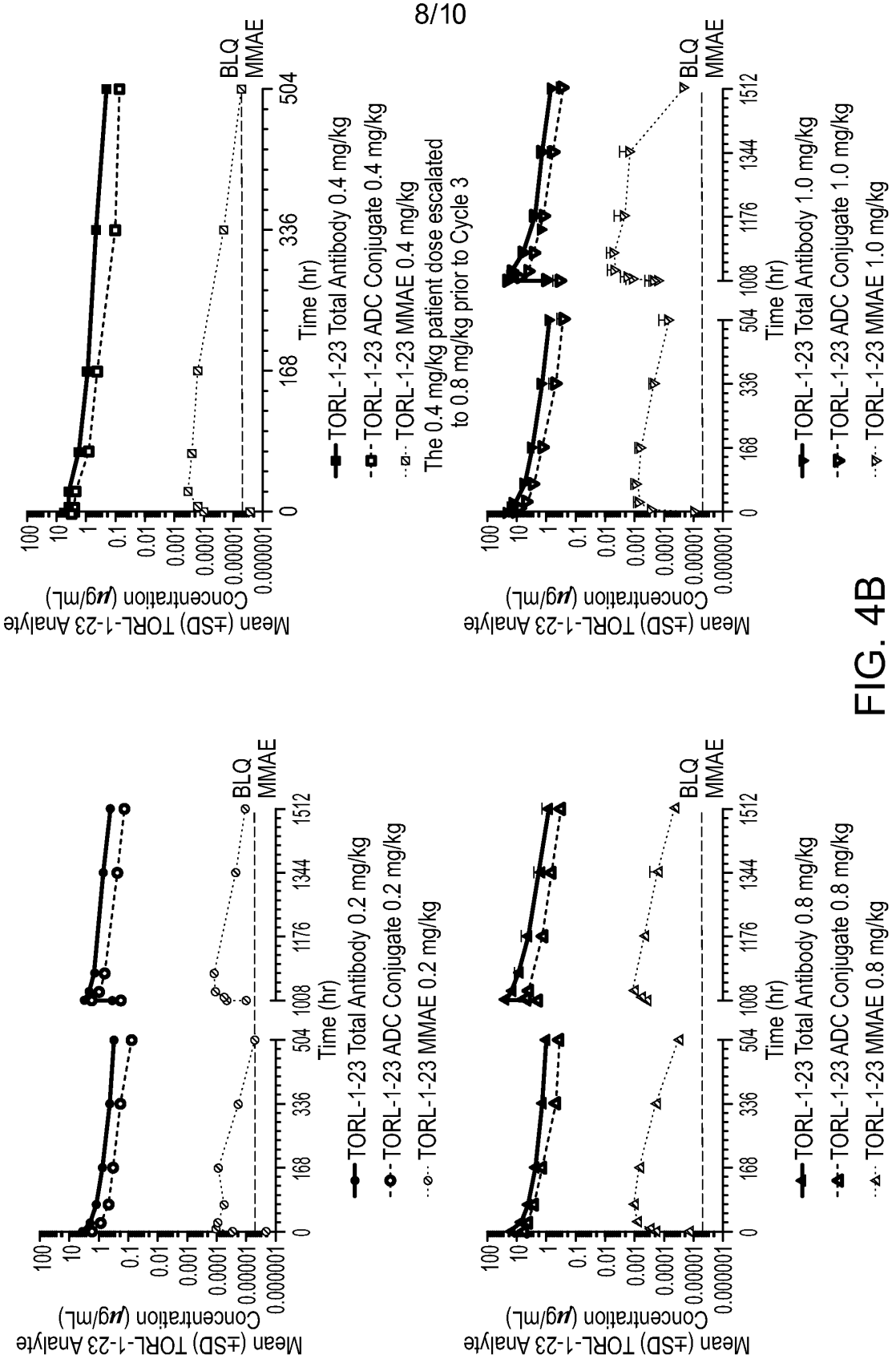

Mean (±SD) TORL-1-23 Analyte Concentration (μg/mL)
100
10
1
0.1
0.01
0.001
0.0001
0.00001
0.000001
0
168
336
504
1008
1176
1344
1512
Time (hr)
BLQ
MMAE
TORL-1-23 Total Antibody 0.4 mg/kg
TORL-1-23 ADC Conjugate 0.4 mg/kg
TORL-1-23 MMAE 0.4 mg/kg
The 0.4 mg/kg patient dose escalated to 0.8 mg/kg prior to Cycle 3
TORL-1-23 Total Antibody 1.0 mg/kg
TORL-1-23 ADC Conjugate 1.0 mg/kg
TORL-1-23 MMAE 1.0 mg/kg
TORL-1-23 Total Antibody 0.2 mg/kg
TORL-1-23 ADC Conjugate 0.2 mg/kg
TORL-1-23 MMAE 0.2 mg/kg
TORL-1-23 Total Antibody 0.8 mg/kg
TORL-1-23 ADC Conjugate 0.8 mg/kg
TORL-1-23 MMAE 0.8 mg/kg

FIG. 4B

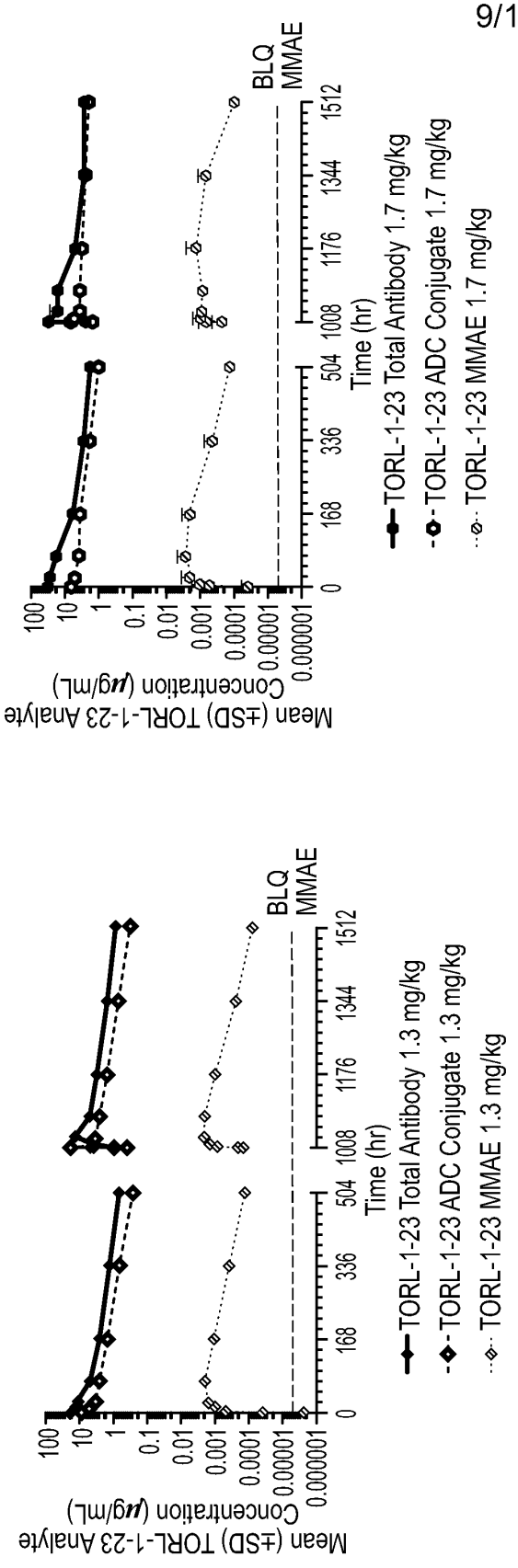
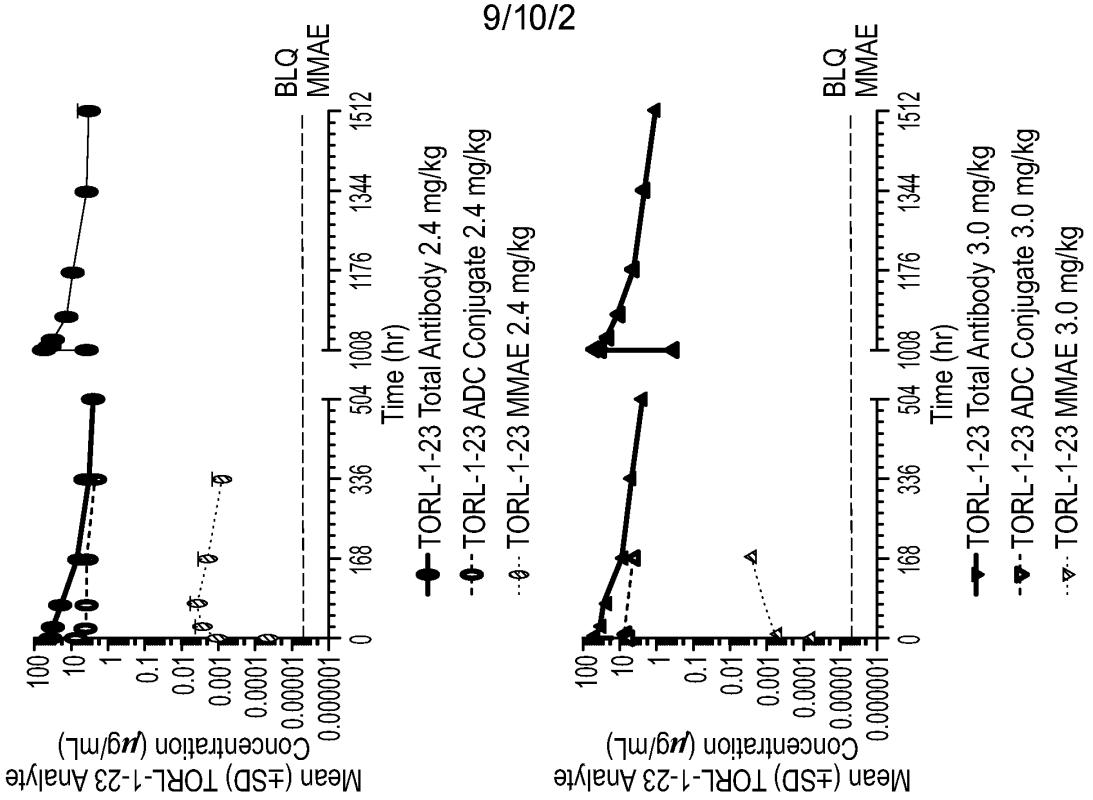

Mean (±SD) TORL-1-23 Analyte Concentration (μg/mL)
100
10
1
0.1
0.01
0.001
0.0001
0.00001
0.000001
0
168
336
504
1008
1176
1344
1512
Time (hr)
BLQ
MMAE
TORL-1-23 Total Antibody 1.3 mg/kg
TORL-1-23 ADC Conjugate 1.3 mg/kg
TORL-1-23 MMAE 1.3 mg/kg
Mean (±SD) TORL-1-23 Analyte Concentration (μg/mL)
100
10
1
0.1
0.01
0.001
0.0001
0.00001
0.000001
0
168
336
504
1008
1176
1344
1512
Time (hr)
BLQ
MMAE
TORL-1-23 Total Antibody 1.7 mg/kg
TORL-1-23 ADC Conjugate 1.7 mg/kg
TORL-1-23 MMAE 1.7 mg/kg

FIG. 4B (CONT.)


Mean (±SD) TORL-1-23 Analyte Concentration (μg/mL)
100
10
1
0.1
0.01
0.001
0.0001
0.00001
0.000001
0
168
336
504
1008
1176
1344
1512
Time (hr)
BLQ
MMAE
TORL-1-23 Total Antibody 2.4 mg/kg
TORL-1-23 ADC Conjugate 2.4 mg/kg
TORL-1-23 MMAE 2.4 mg/kg
TORL-1-23 Total Antibody 3.0 mg/kg
TORL-1-23 ADC Conjugate 3.0 mg/kg
TORL-1-23 MMAE 3.0 mg/kg

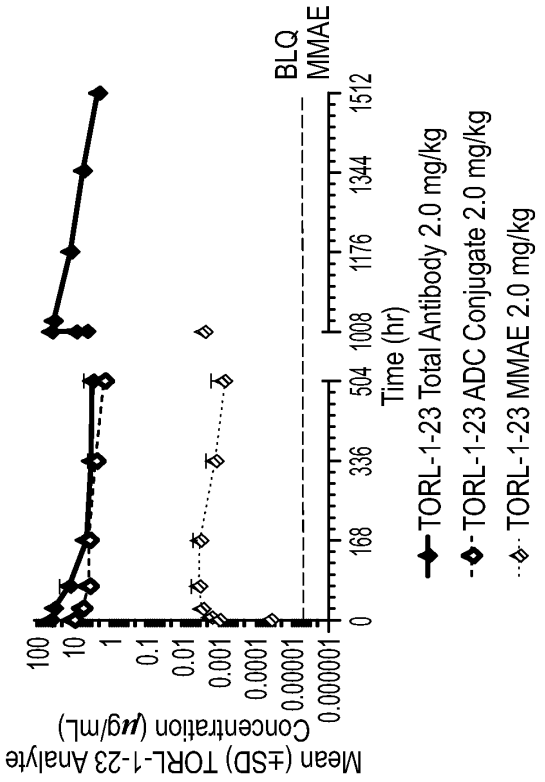

Mean (±SD) TORL-1-23 Analyte Concentration (μg/mL)
100
10
1
0.1
0.01
0.001
0.0001
0.00001
0.000001
0
168
336
504
1008
1176
1344
1512
Time (hr)
BLQ
MMAE
TORL-1-23 Total Antibody 2.0 mg/kg
TORL-1-23 ADC Conjugate 2.0 mg/kg
TORL-1-23 MMAE 2.0 mg/kg

FIG. 4B (CONT.)

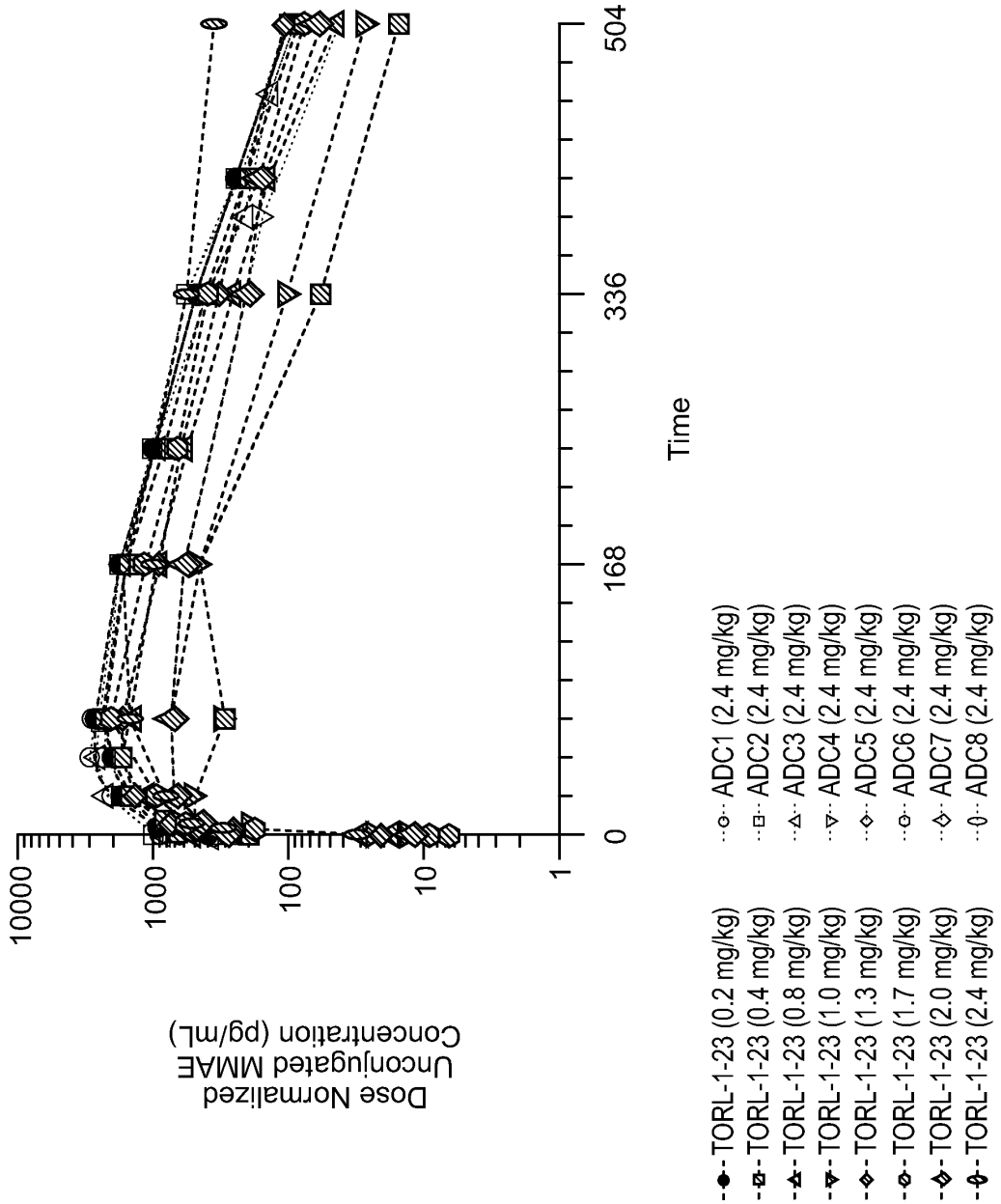

Dose Normalized
Unconjugated MMAE
Concentration (pg/mL)
10000
1000
100
10
1
0
168
336
504
Time
TORL-1-23 (0.2 mg/kg)
TORL-1-23 (0.4 mg/kg)
TORL-1-23 (0.8 mg/kg)
TORL-1-23 (1.0 mg/kg)
TORL-1-23 (1.3 mg/kg)
TORL-1-23 (1.7 mg/kg)
TORL-1-23 (2.0 mg/kg)
TORL-1-23 (2.4 mg/kg)
ADC1 (2.4 mg/kg)
ADC2 (2.4 mg/kg)
ADC3 (2.4 mg/kg)
ADC4 (2.4 mg/kg)
ADC5 (2.4 mg/kg)
ADC6 (2.4 mg/kg)
ADC7 (2.4 mg/kg)
ADC8 (2.4 mg/kg)

FIG. 5