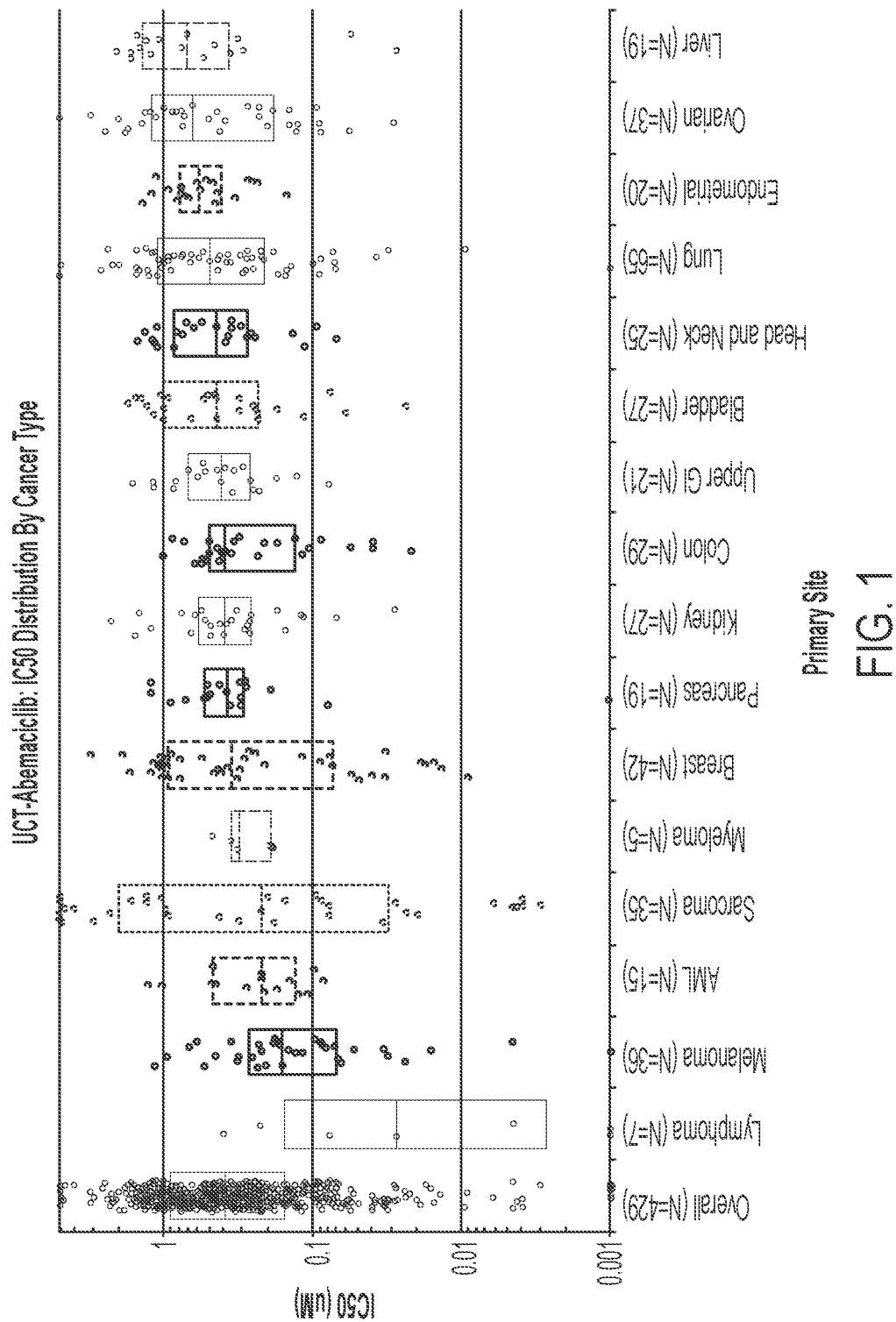




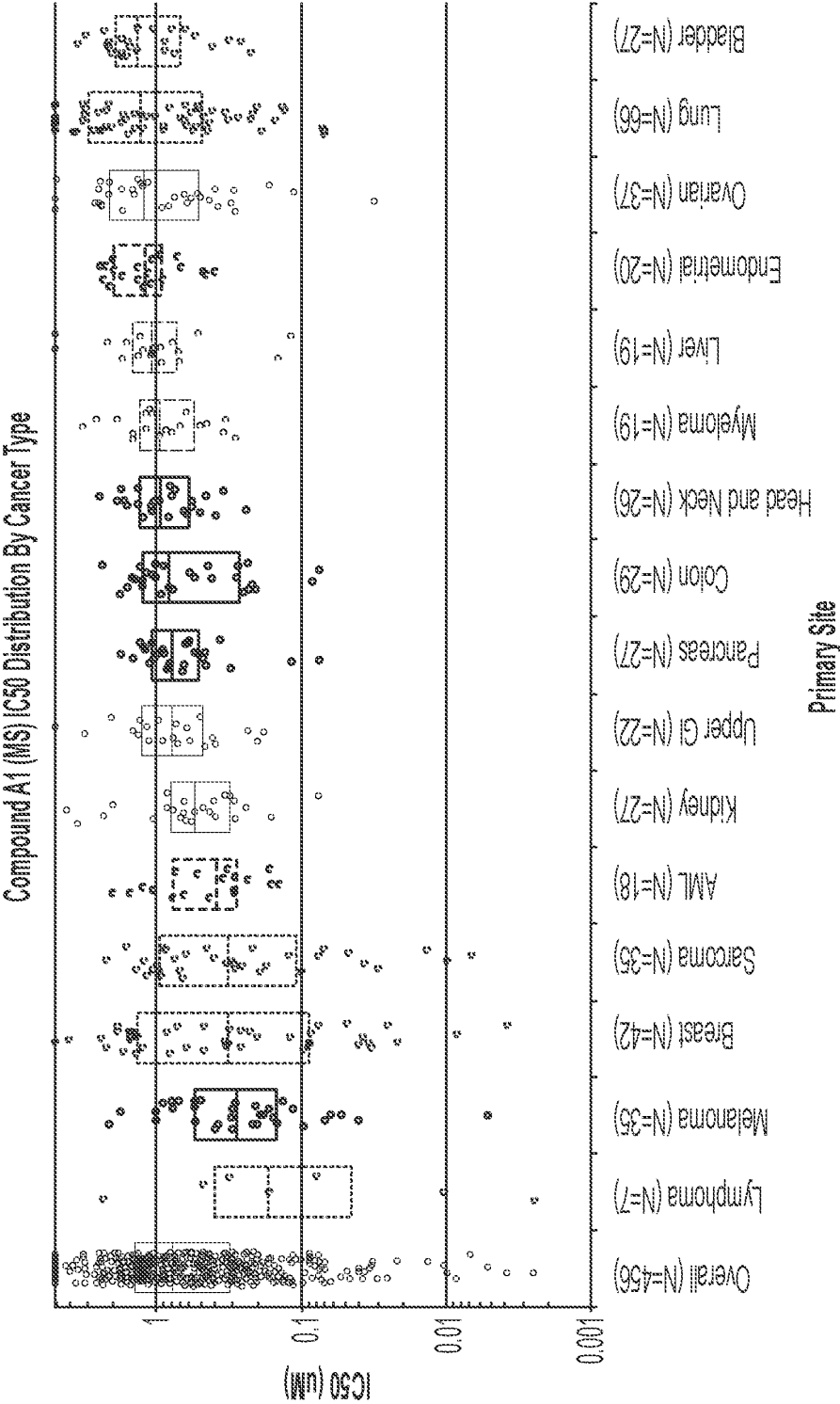


FIG. 2

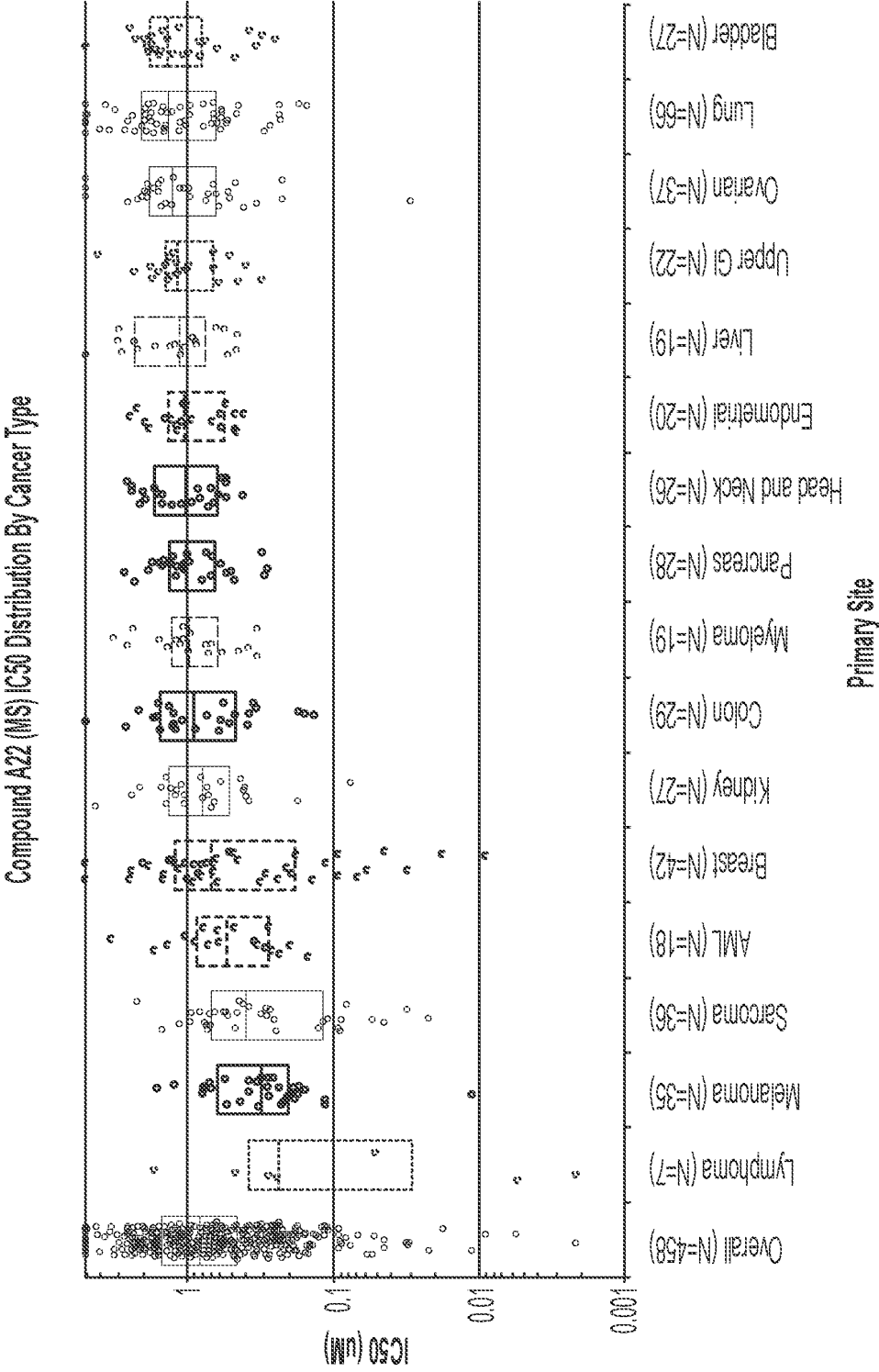


FIG. 3

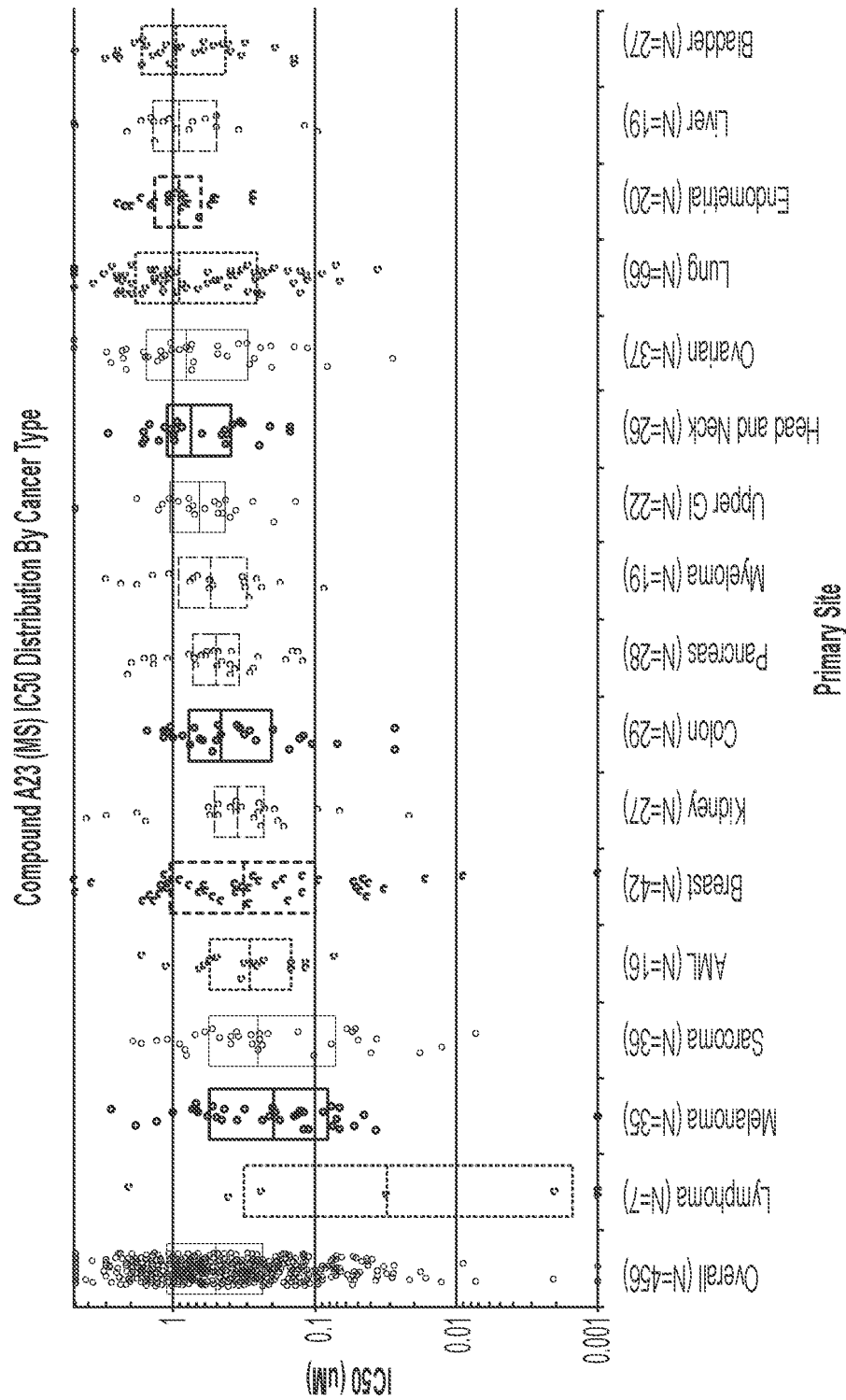


FIG. 4

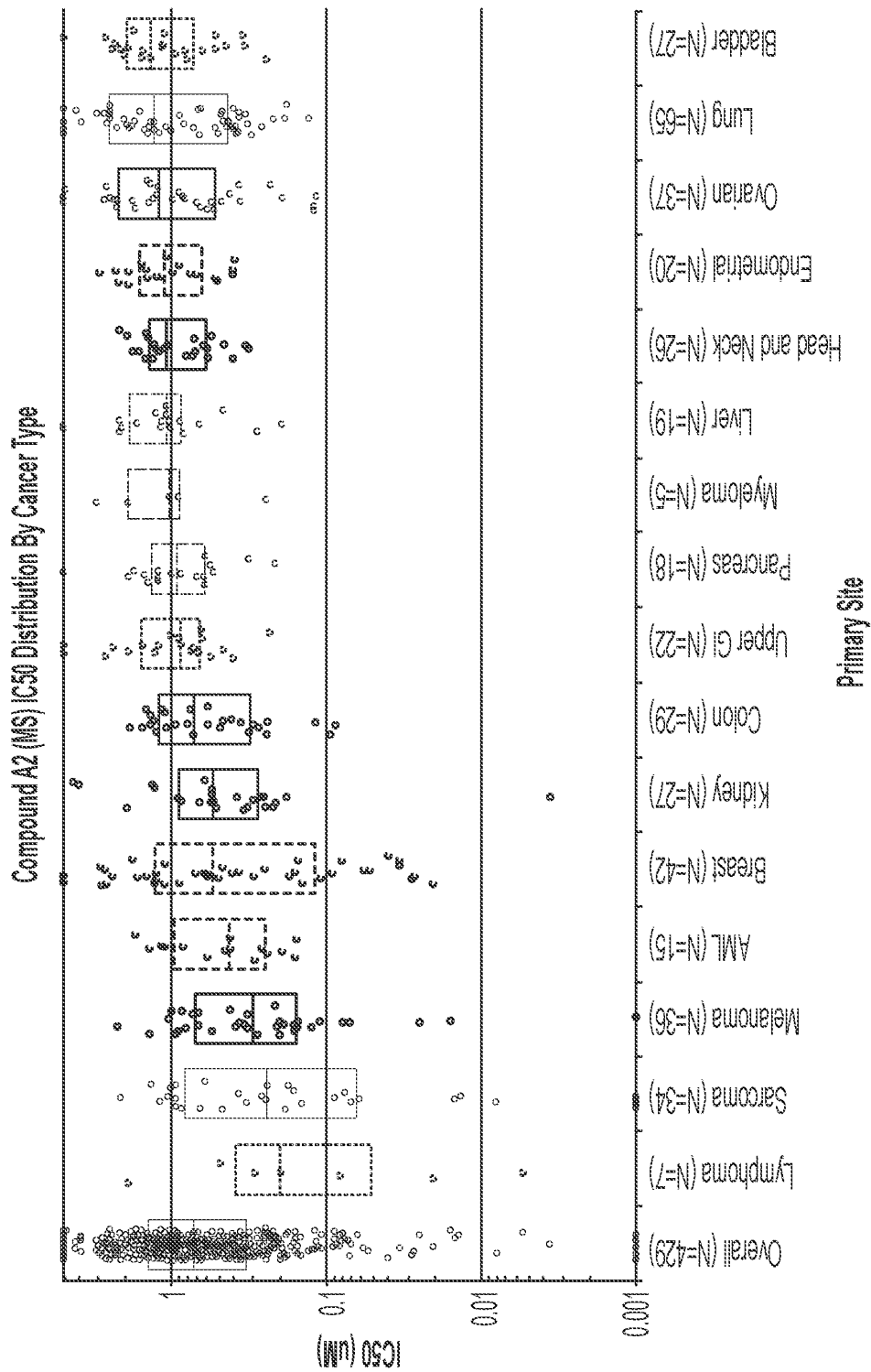


FIG. 5

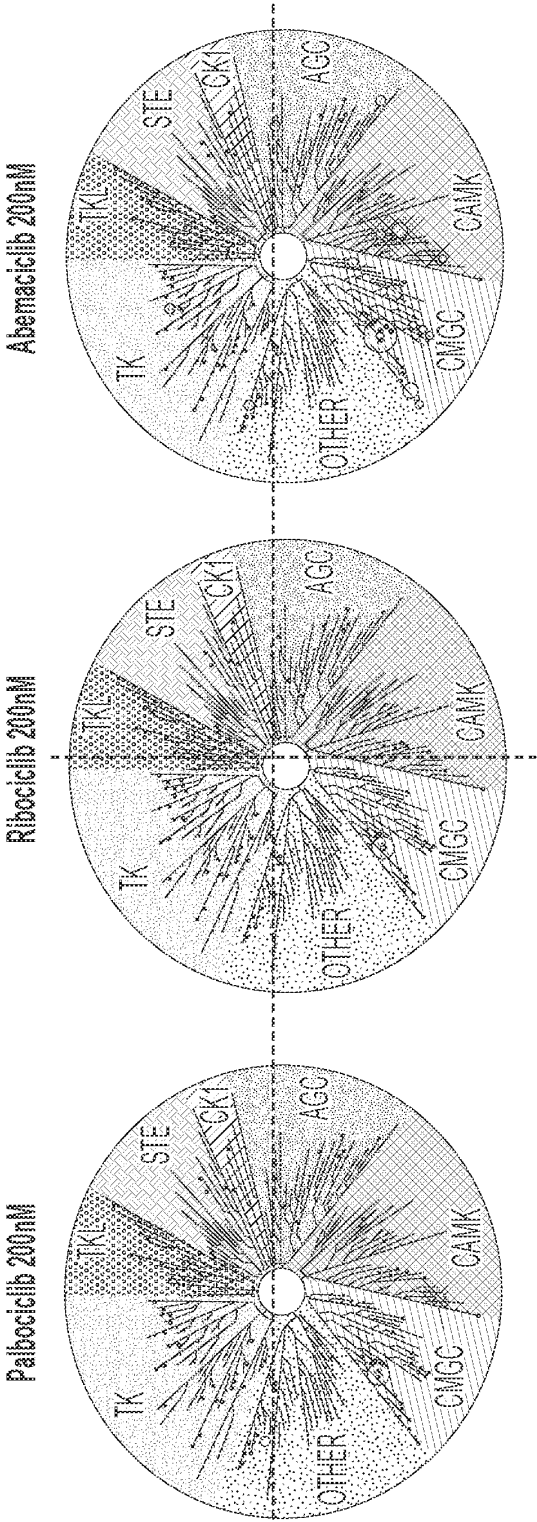


FIG. 6

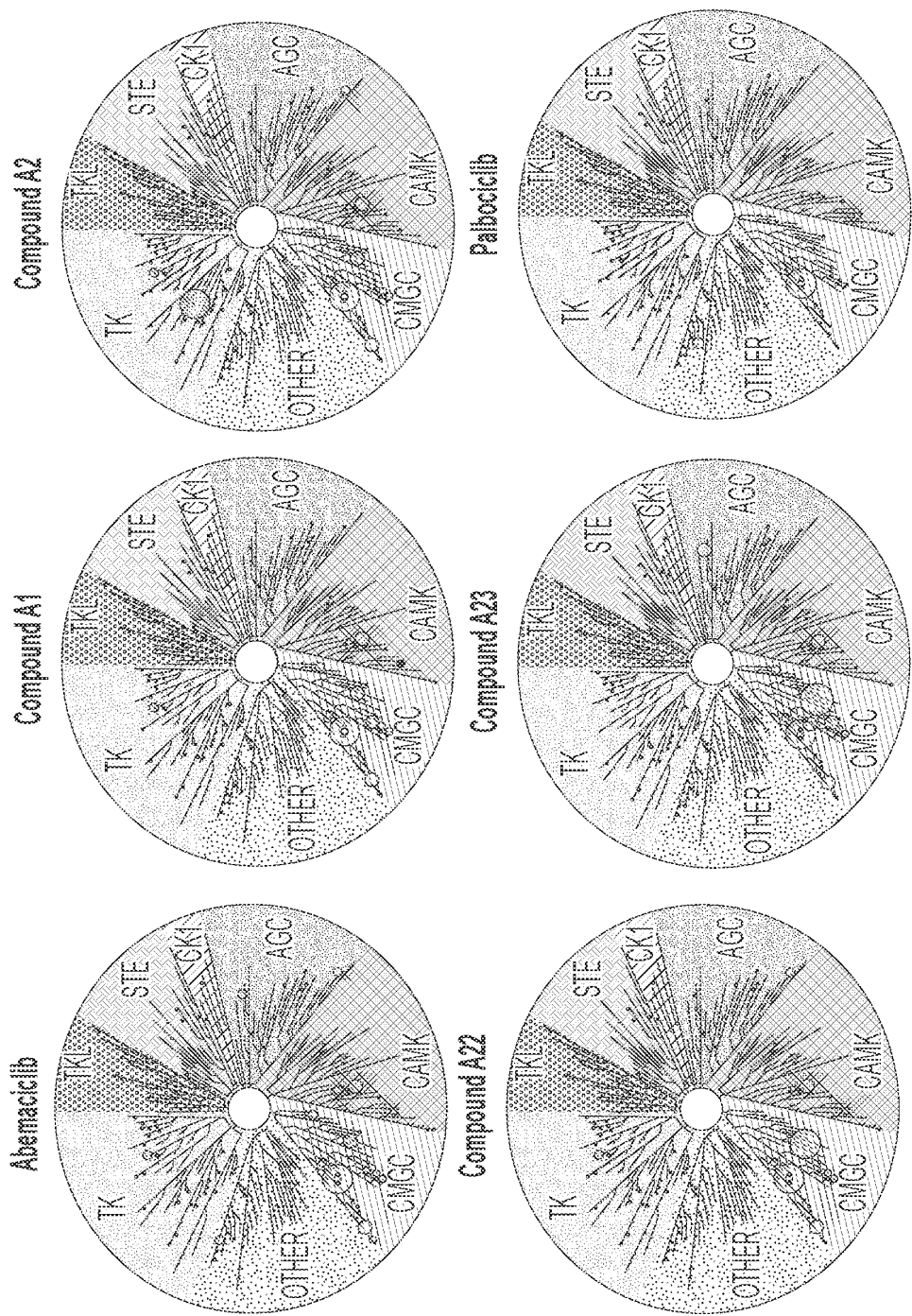


FIG. 7

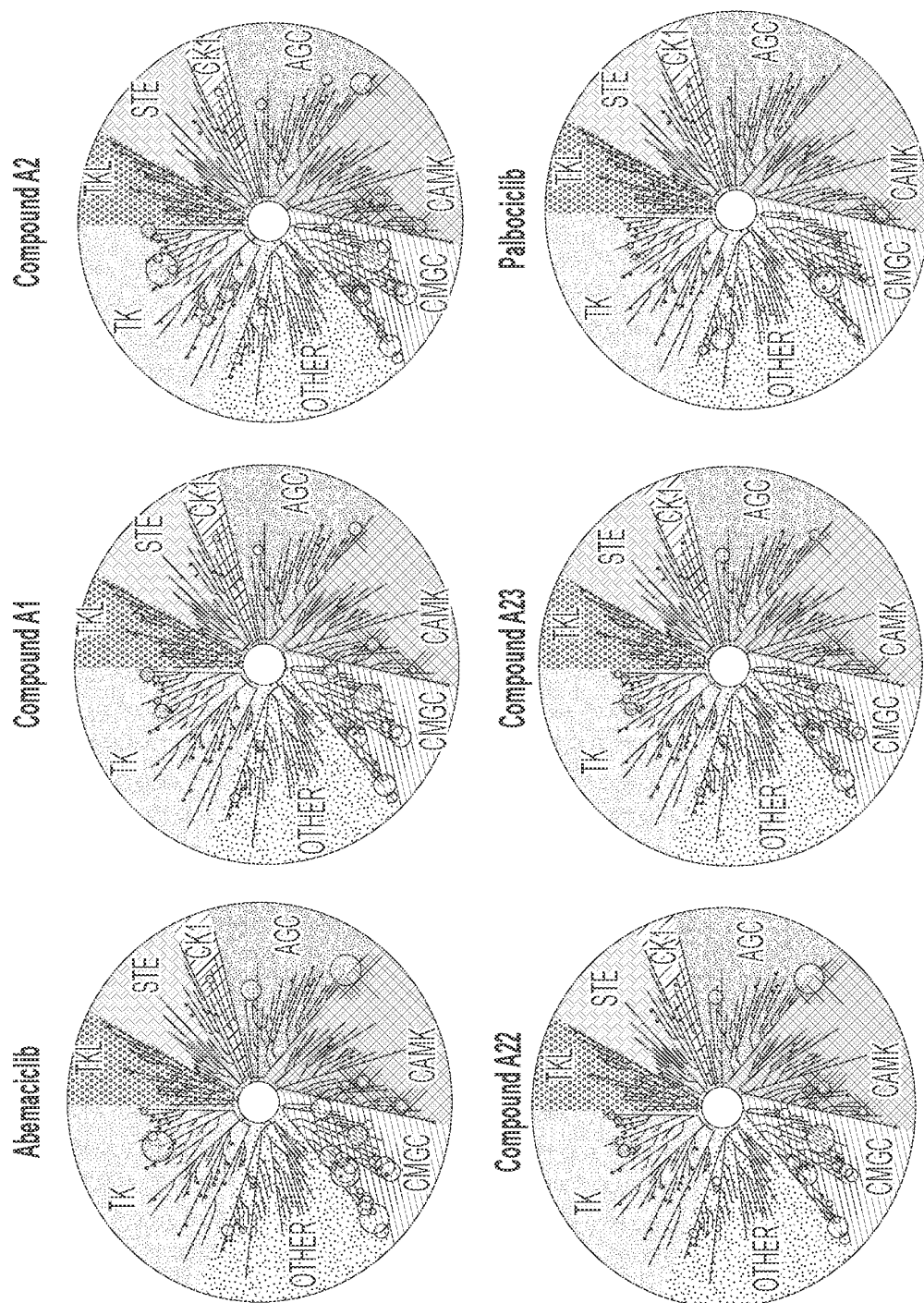


FIG. 8

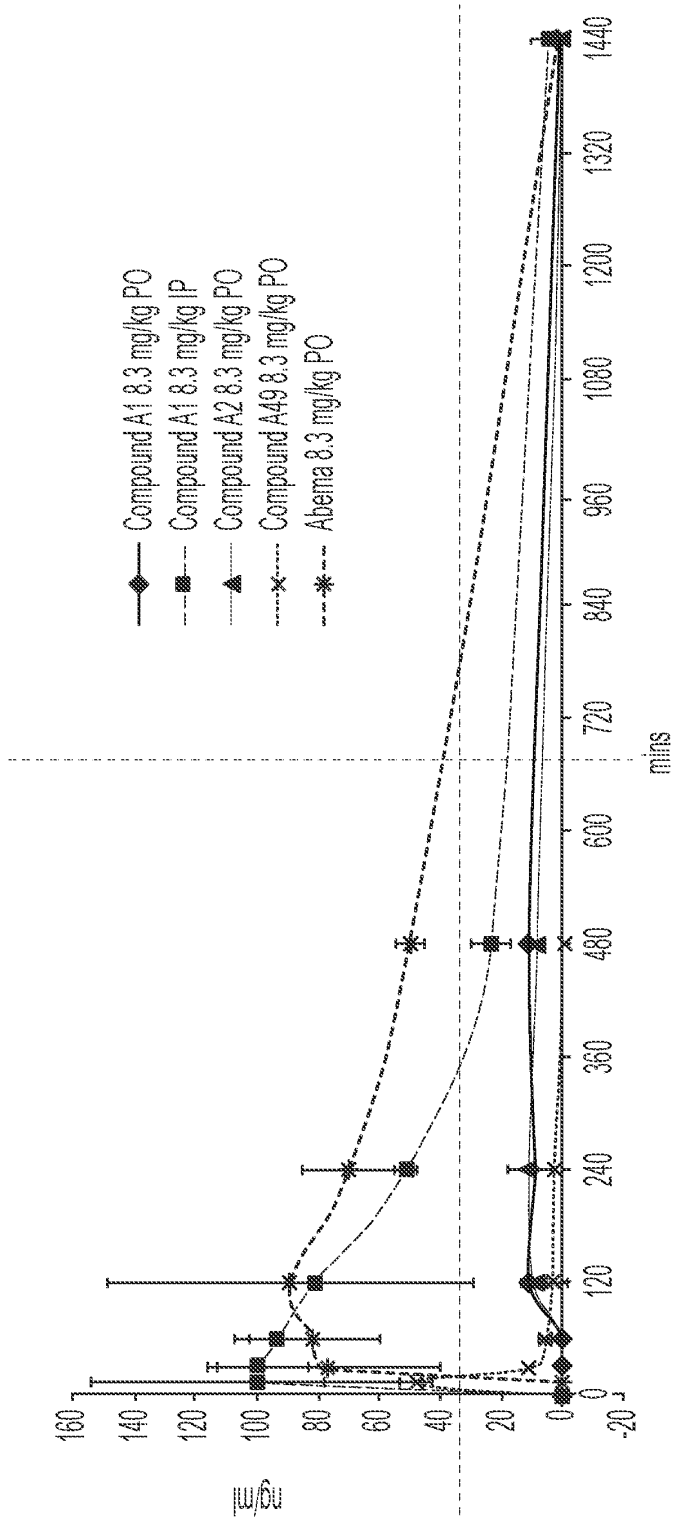


FIG. 9

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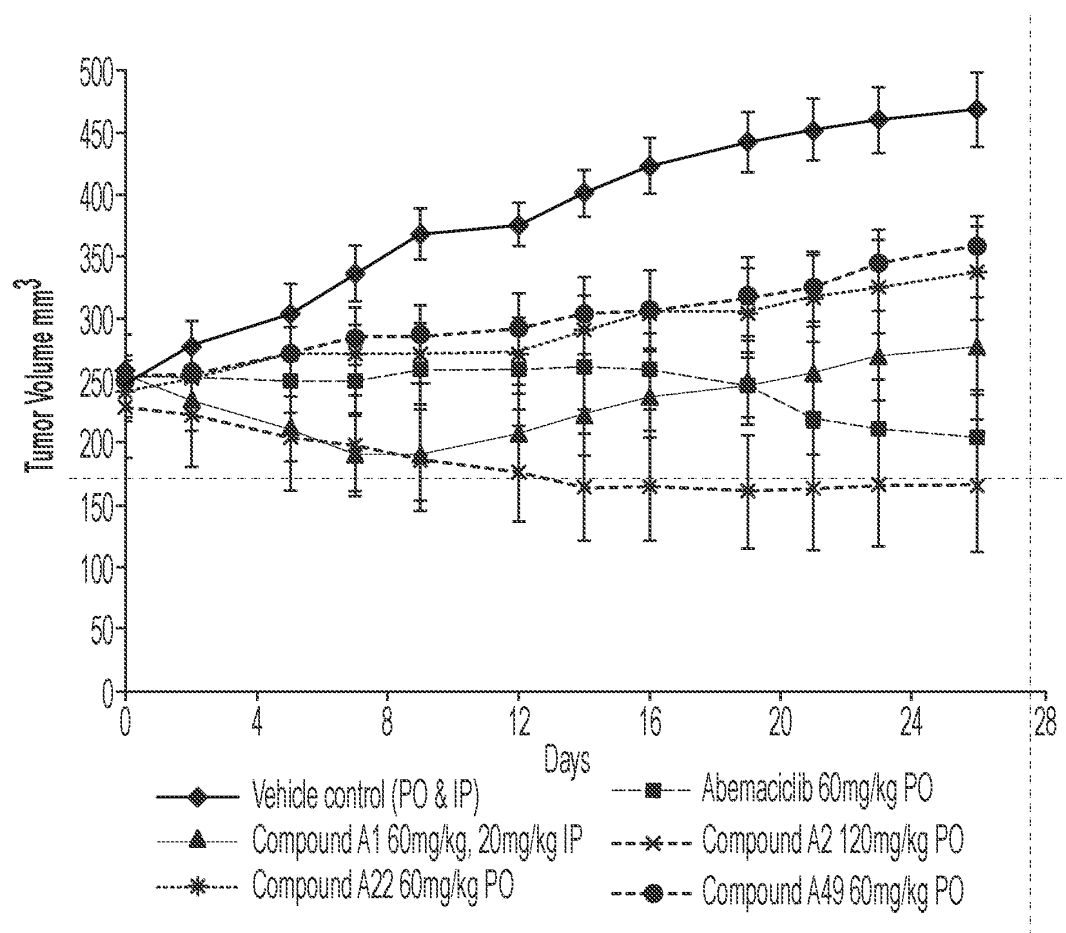


FIG. 10

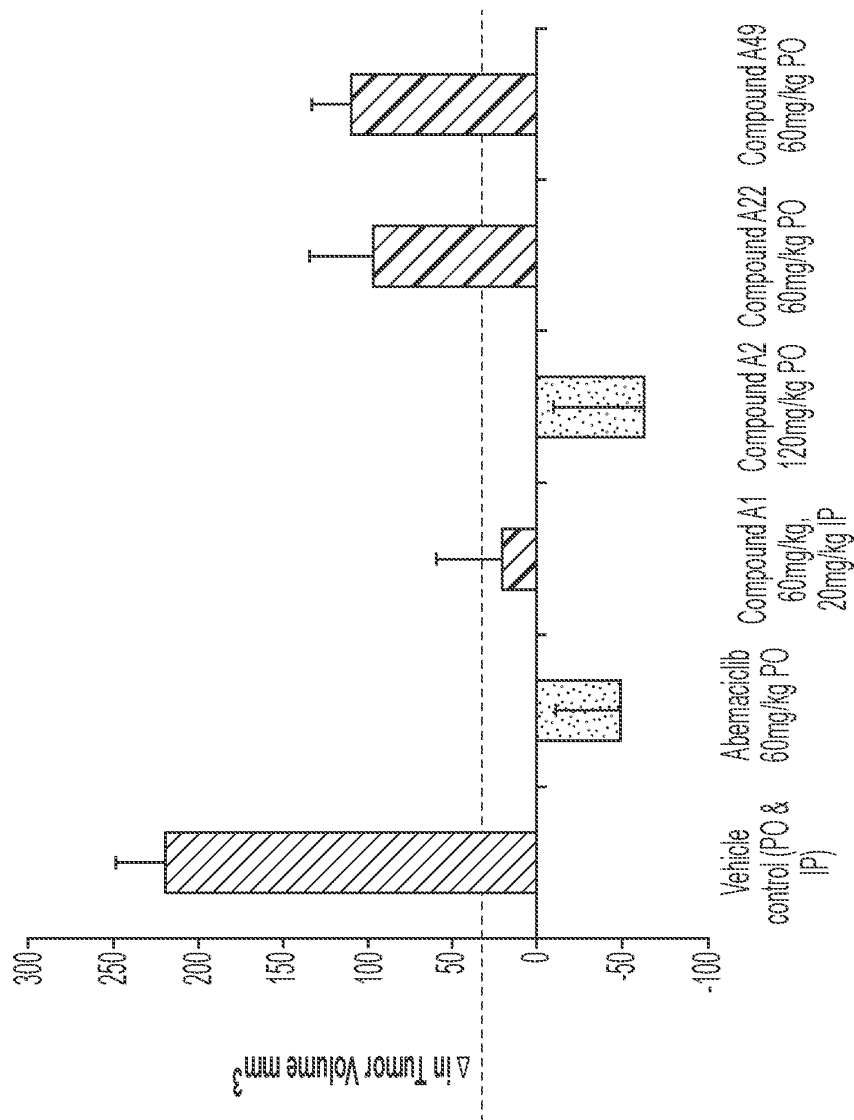


FIG. 11

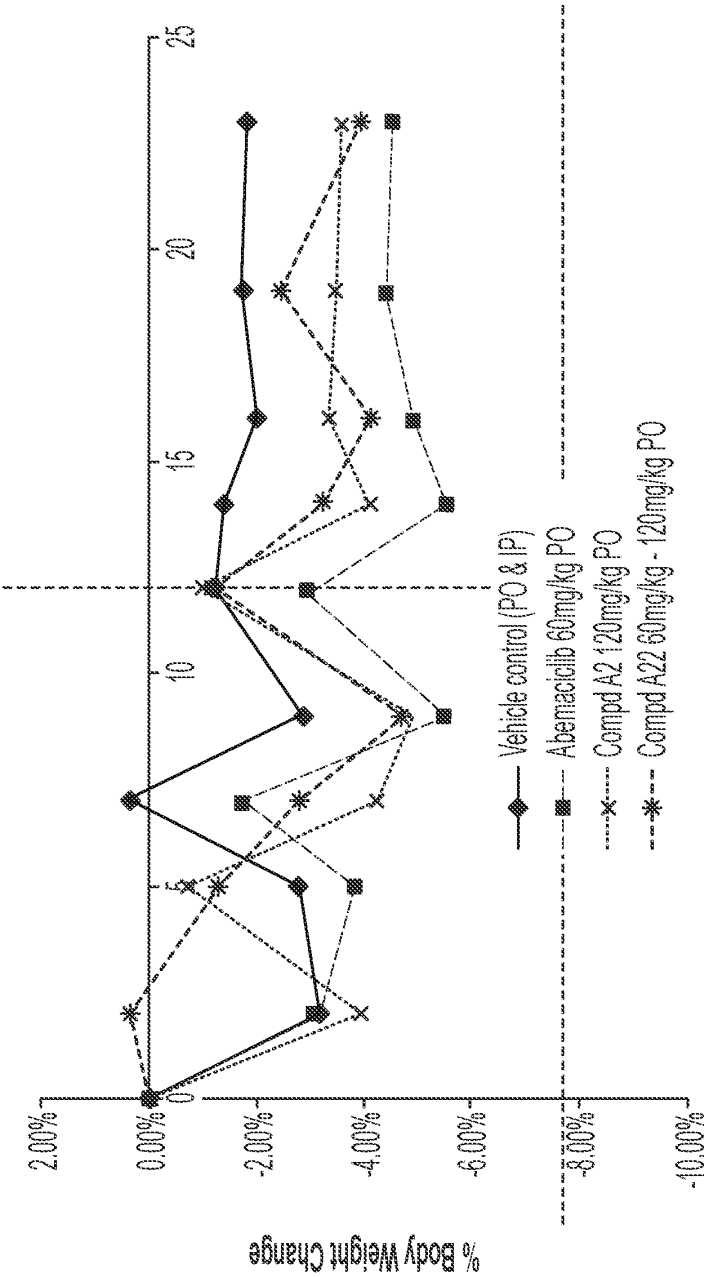


FIG. 12A

Group	0	2	5	7	9	12	14	16	19	23
Vehicle control (PO & IP)	0.00%	-3.20%	-2.77%	0.36%	-2.88%	-1.15%	-1.40%	-1.97%	-1.71%	-1.83%
Abemaciclib 60mg/kg PO	0.00%	-3.08%	-3.83%	-1.71%	-5.46%	-2.92%	-5.52%	-4.87%	-4.39%	-4.50%
Compound A2 120mg/kg PO	0.00%	-3.87%	-0.82%	-4.22%	-4.89%	-0.98%	-4.03%	-3.34%	-3.47%	-3.60%
Compound A22 60mg/kg - 120mg/kg PO	0.00%	0.34%	-1.31%	-2.81%	-4.71%	-1.24%	-3.22%	-4.12%	-2.46%	-3.91%

FIG. 12B

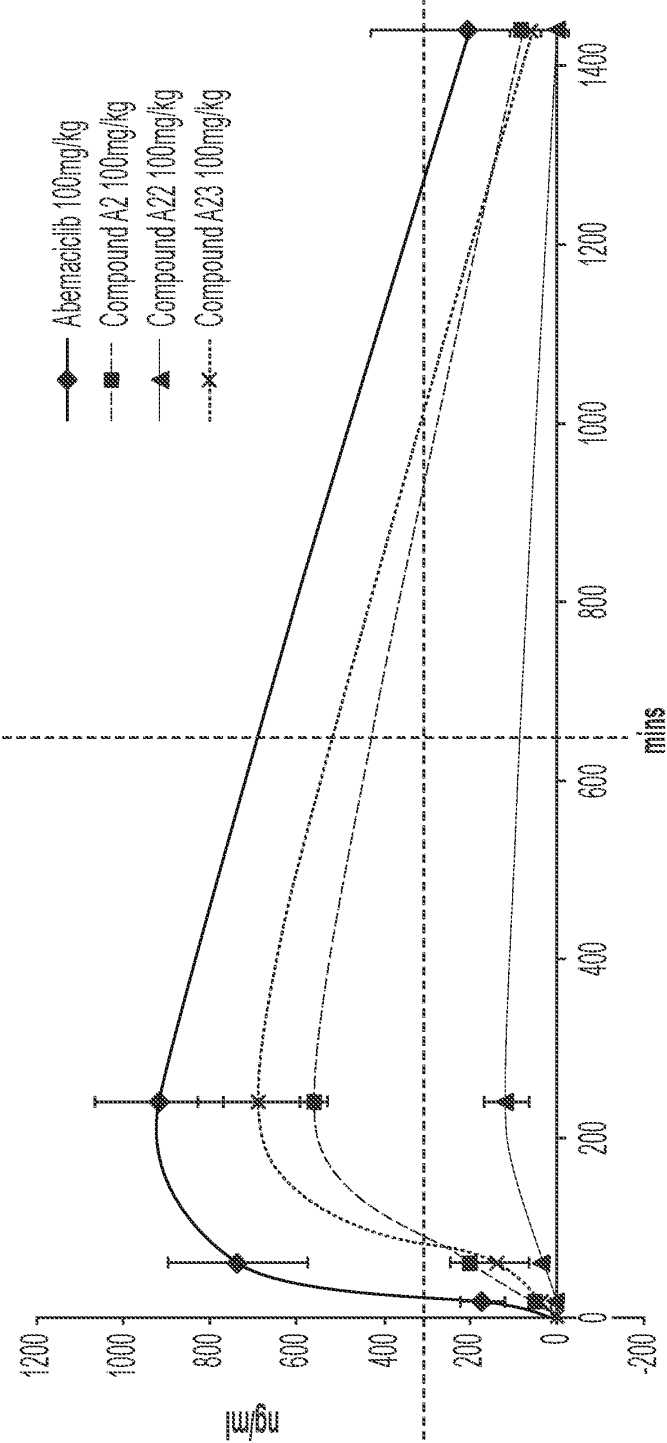


FIG. 13

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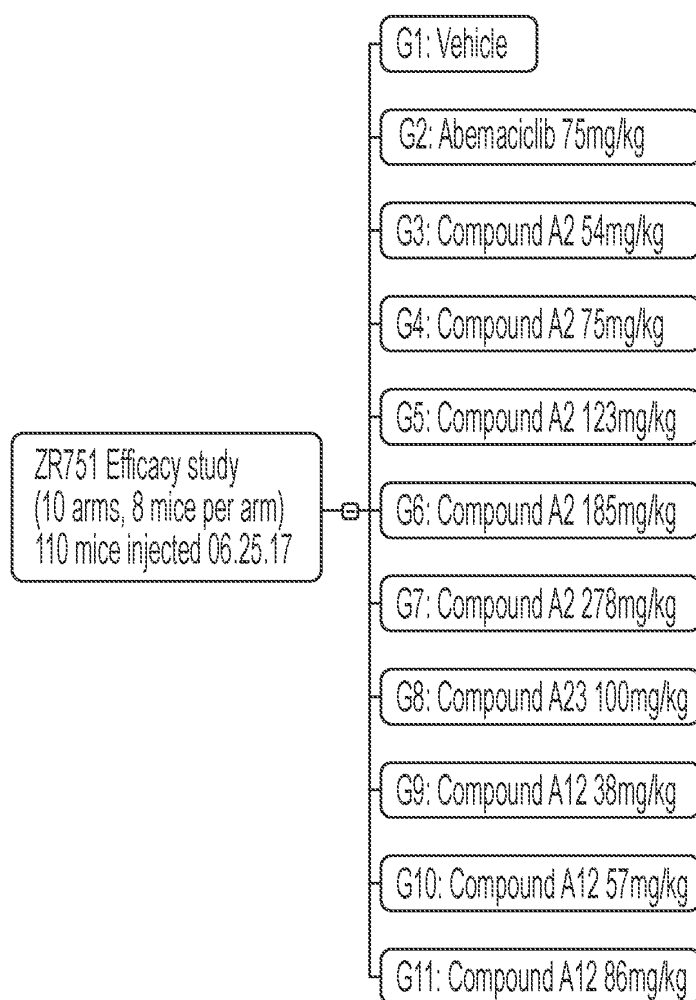


FIG. 14

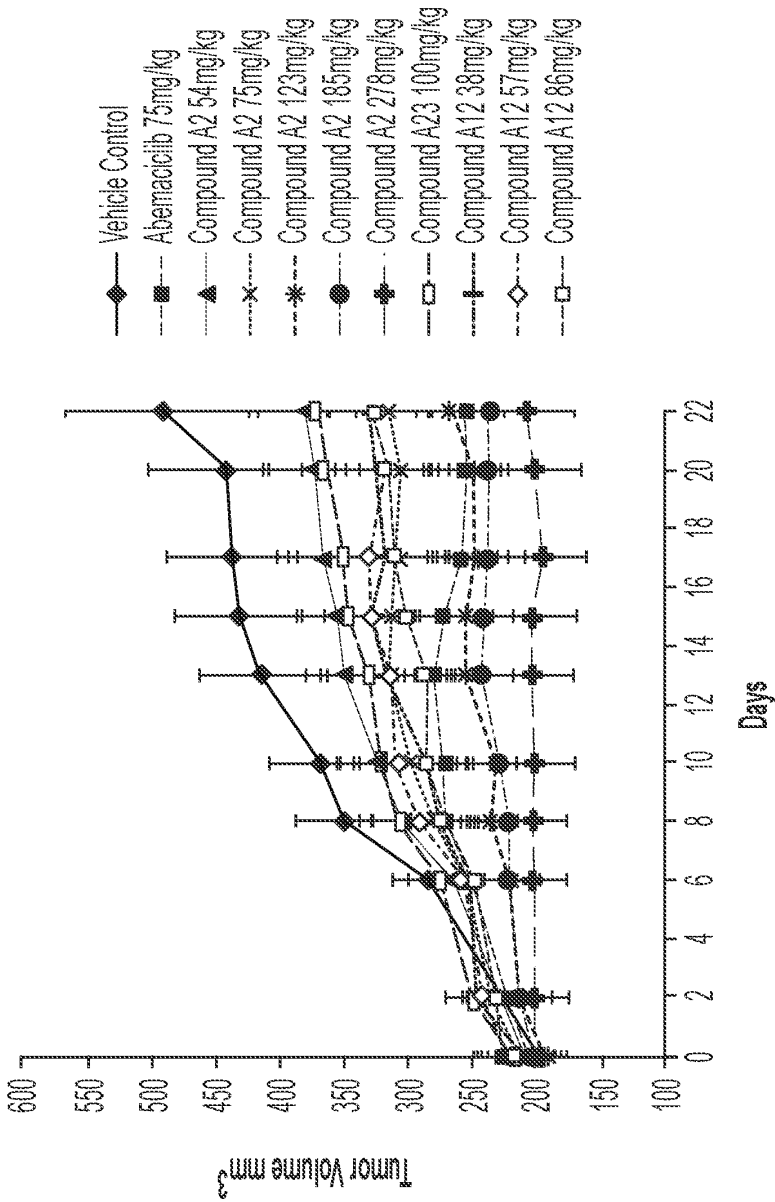


FIG. 15

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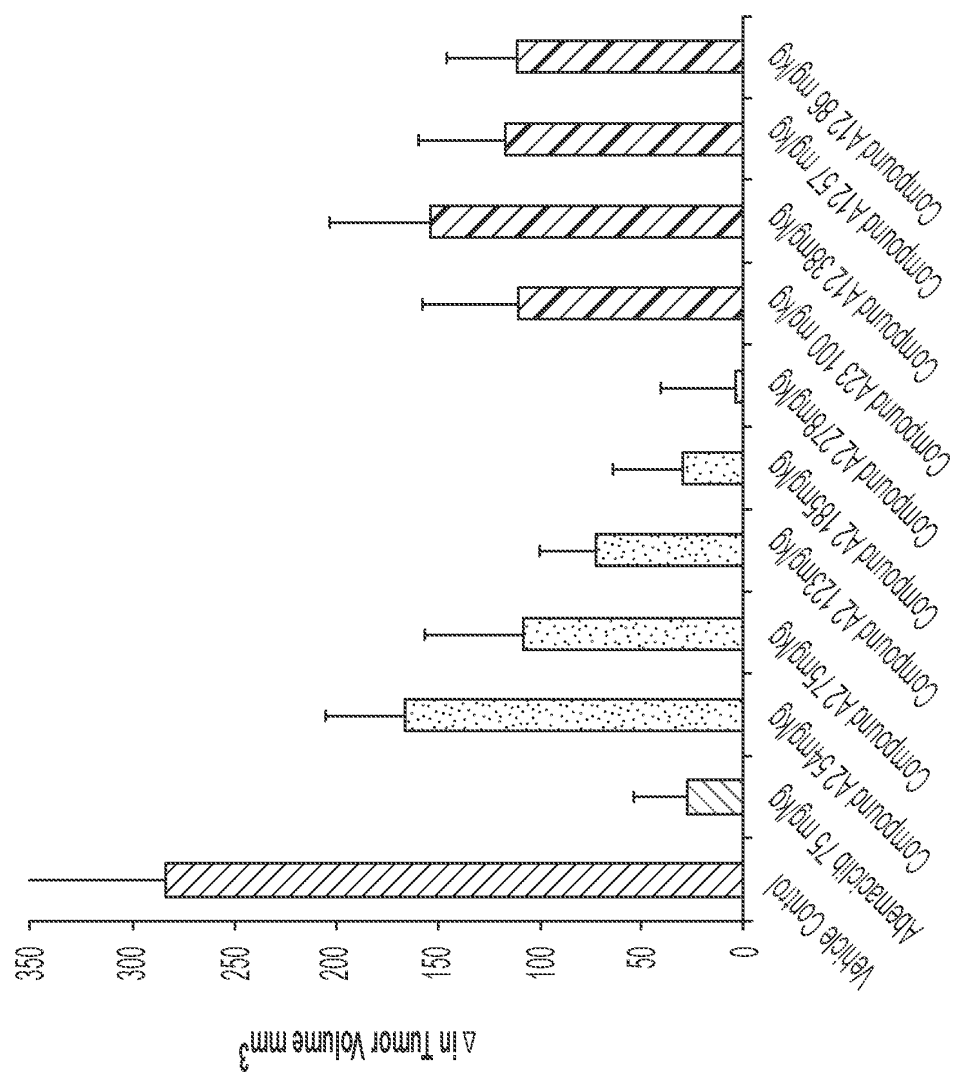


FIG. 16

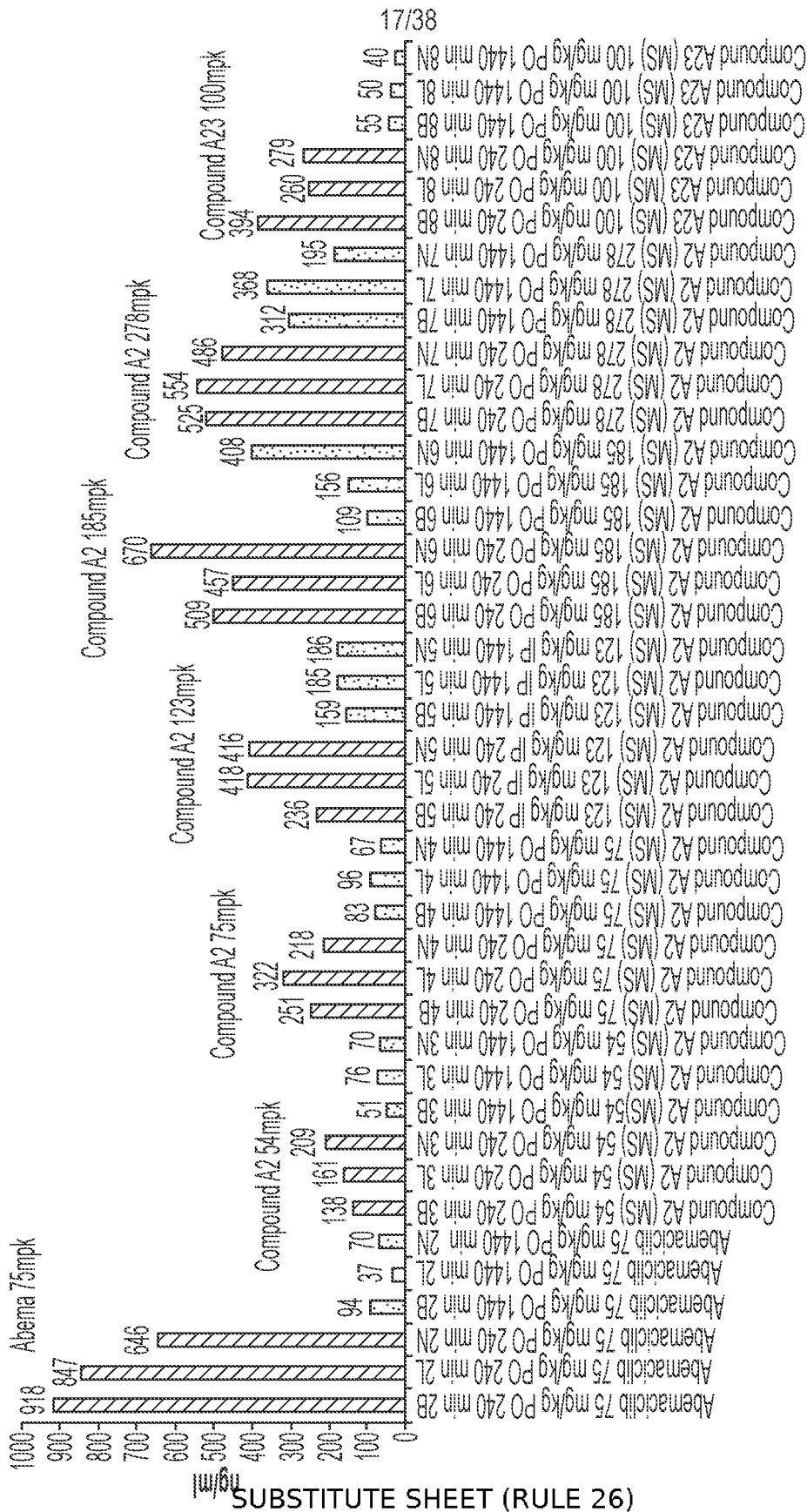
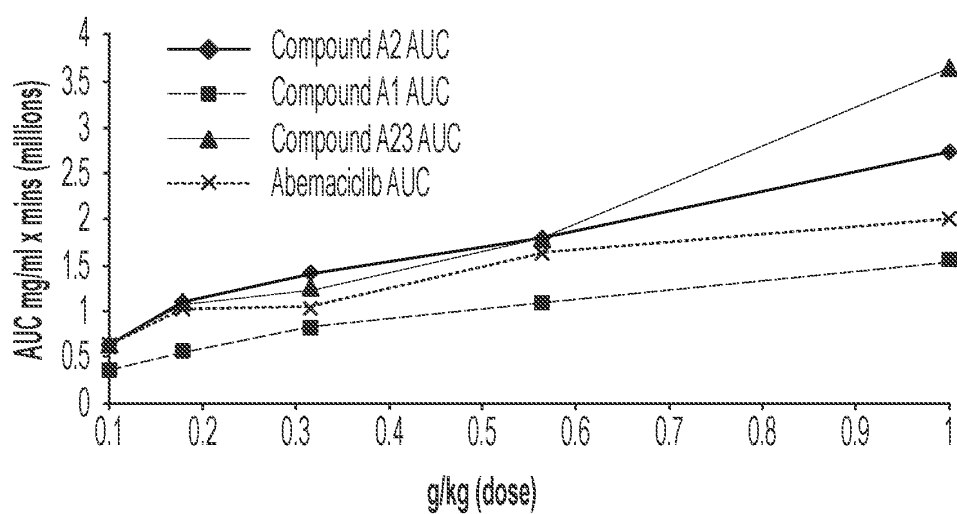


FIG. 17

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Dose	Compound A2 AUC	Compound A1 AUC	Compound A23 AUC	Abemaciclib AUC	ng/ml x mins (millions)
100	0.63	0.35	0.64	0.64	
178	1.10	0.55	1.08	1.02	
316	1.41	0.82	1.24	1.05	
564	1.79	1.09	1.80	1.63	
1000	2.72	1.54	3.68	2.01	

FIG. 18A

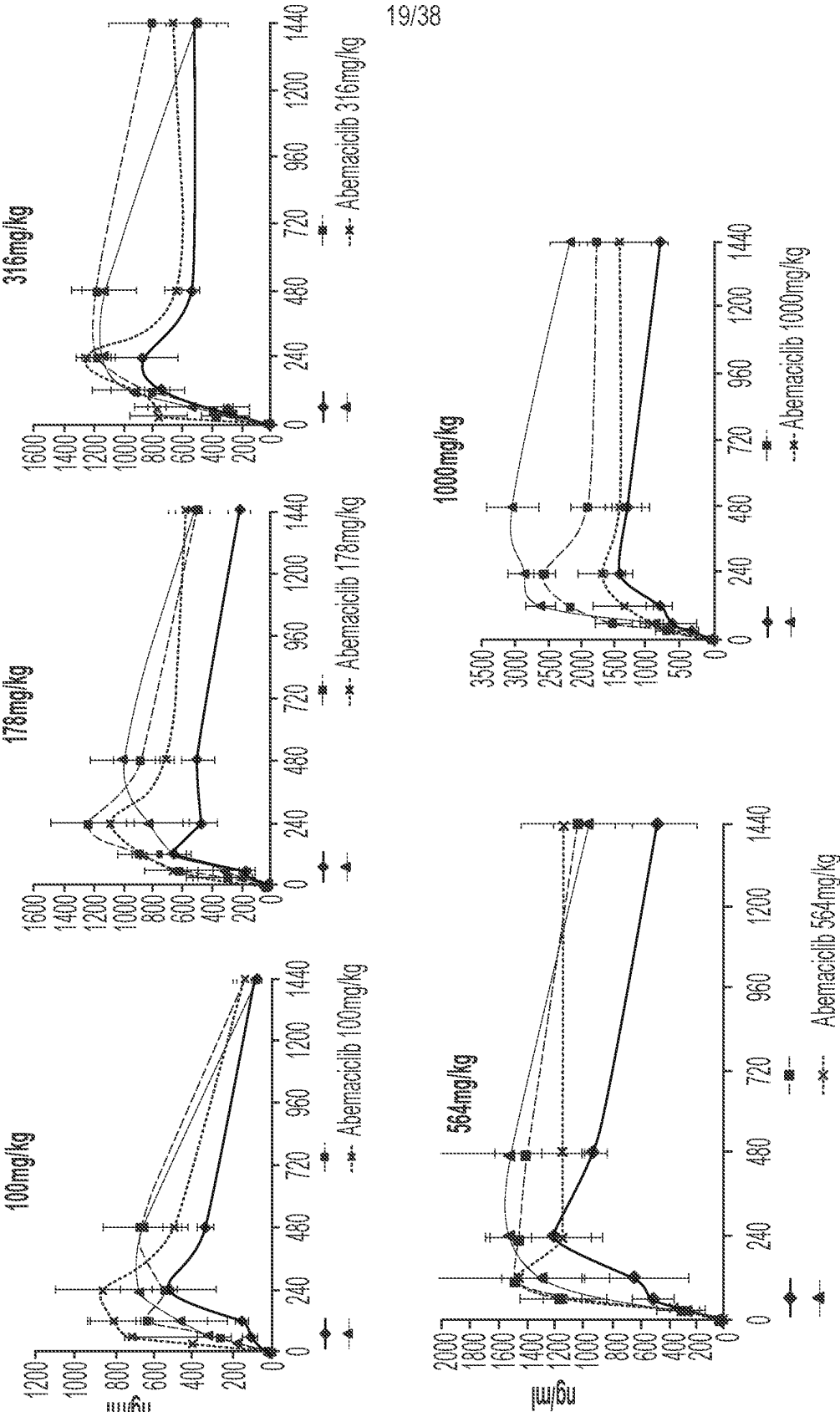


FIG. 18B

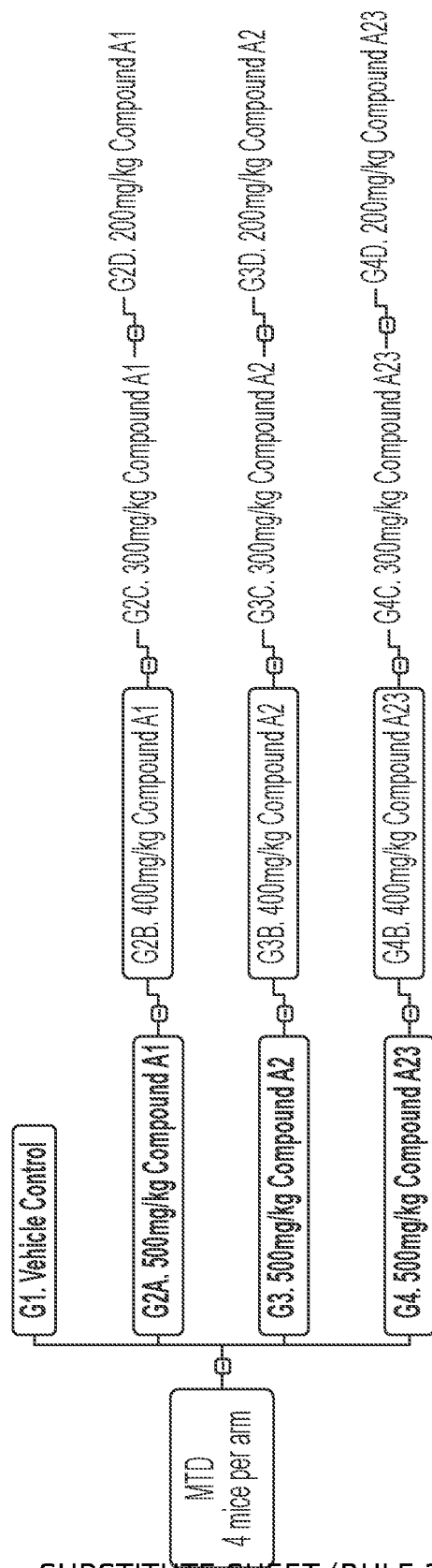


FIG. 19

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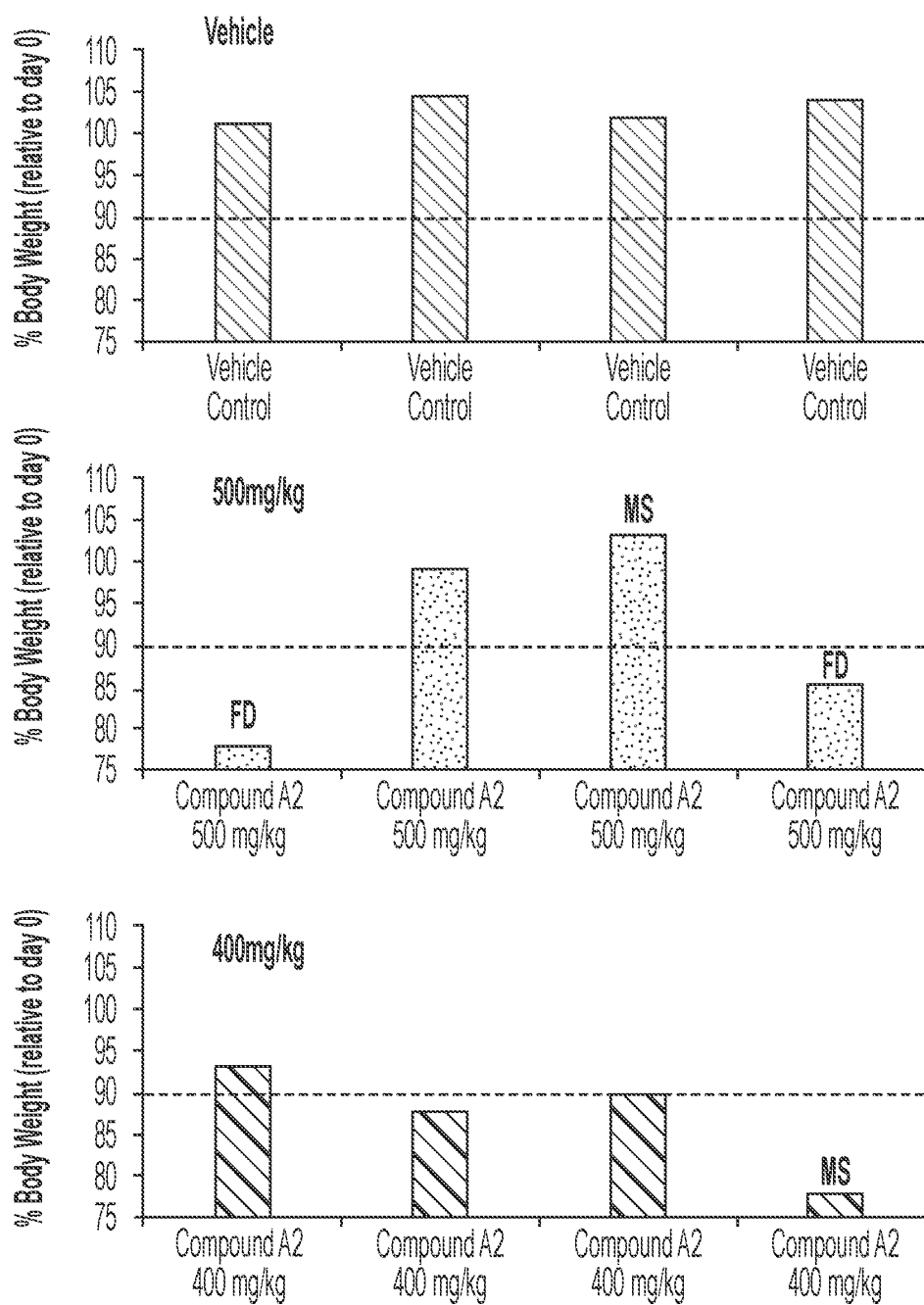


FIG. 20

SUBSTITUTE SHEET (RULE 26)

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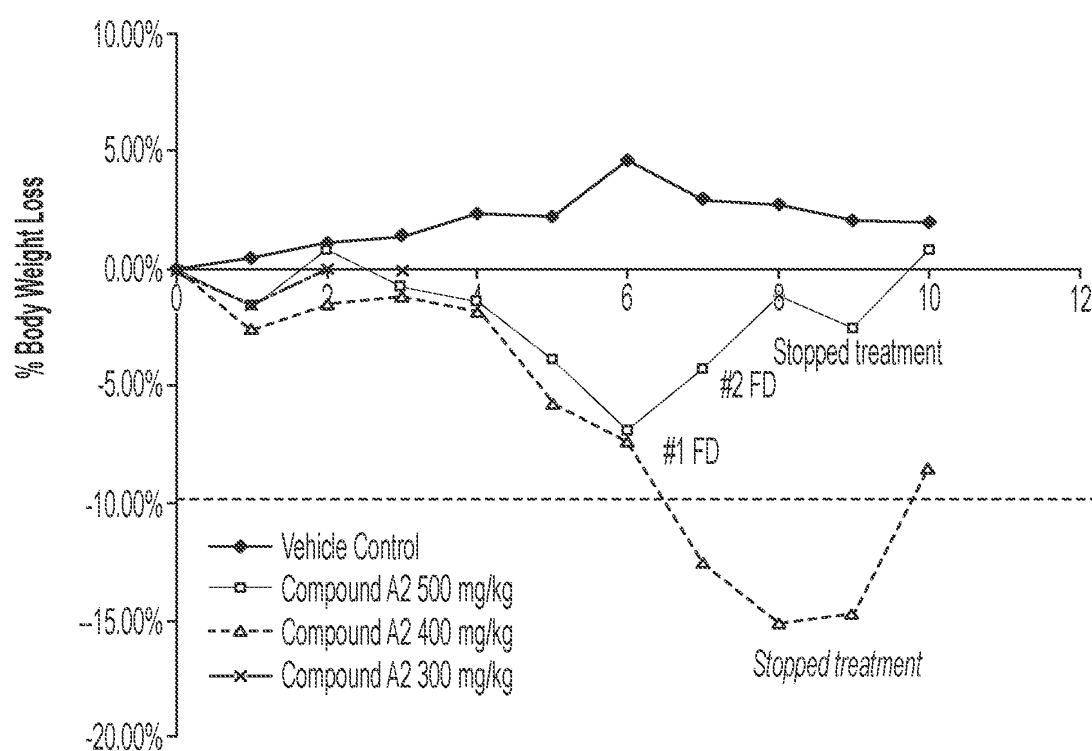


FIG. 21

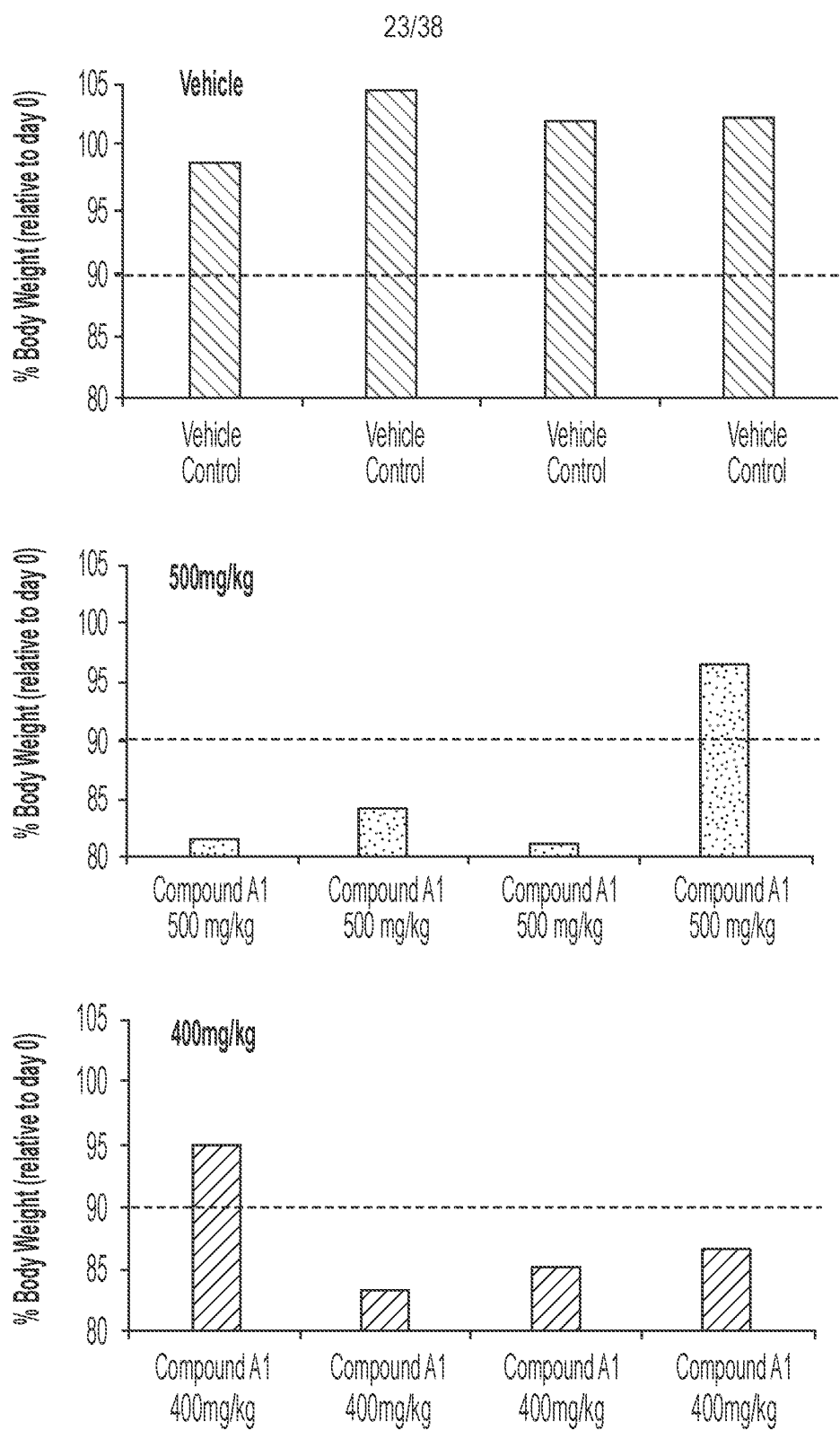


FIG. 22

SUBSTITUTE SHEET (RULE 26)

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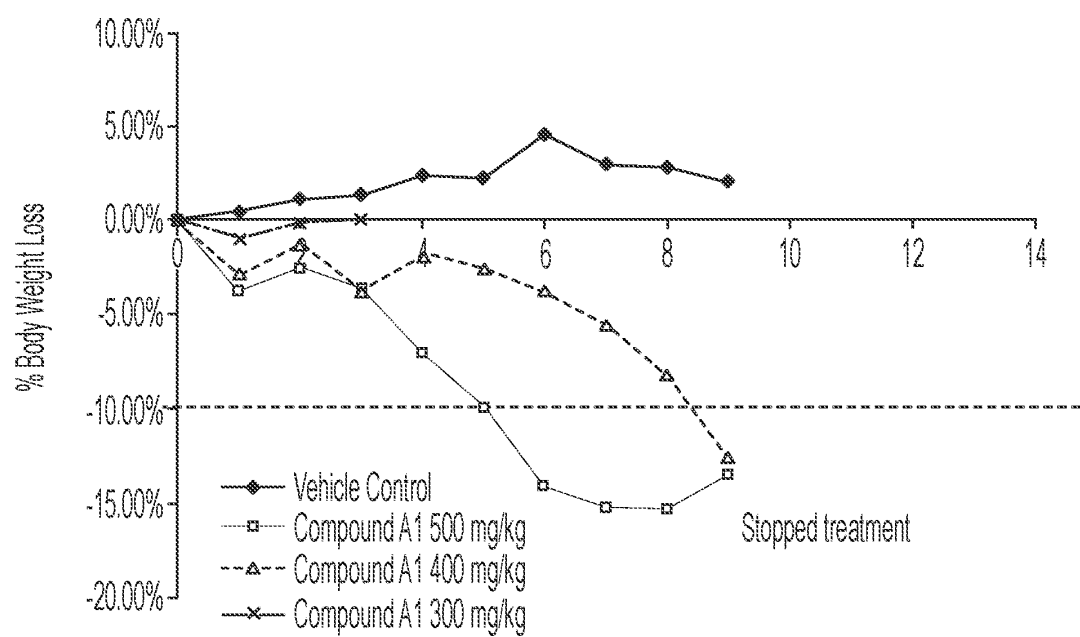


FIG. 23

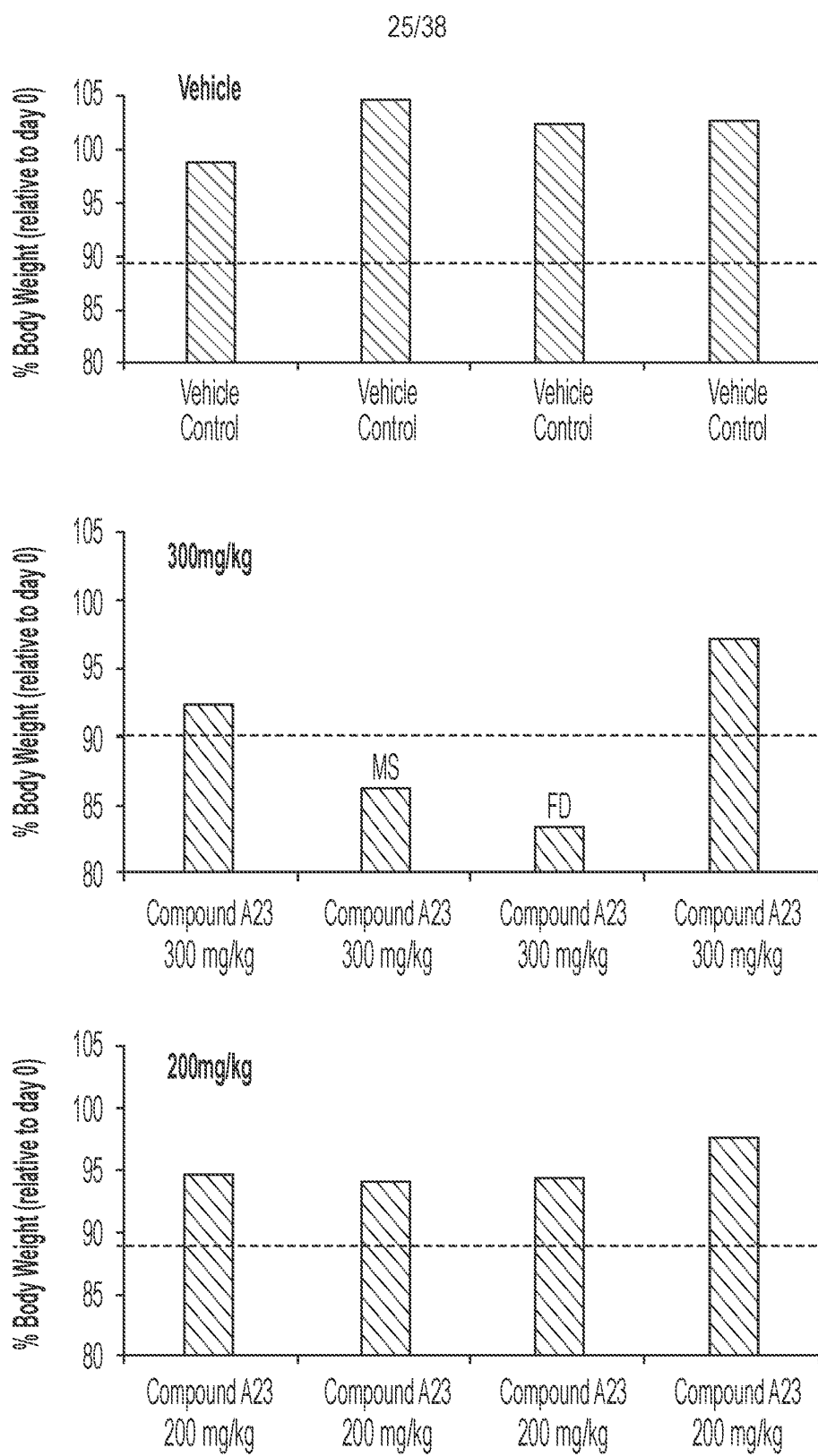


FIG. 24

SUBSTITUTE SHEET (RULE 26)

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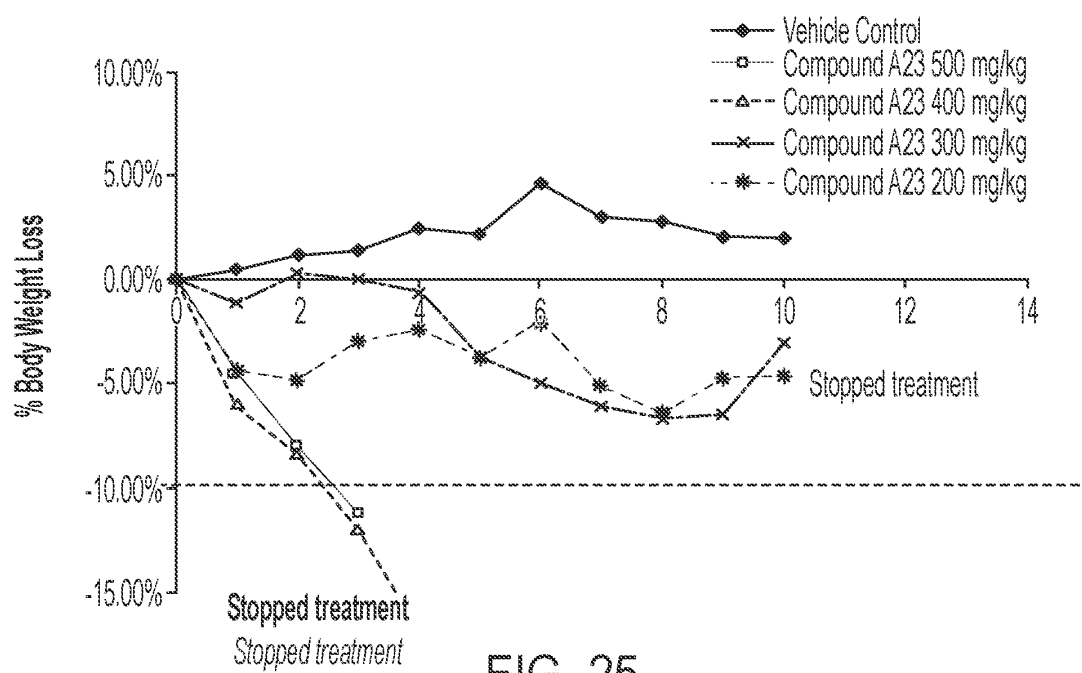


FIG. 25

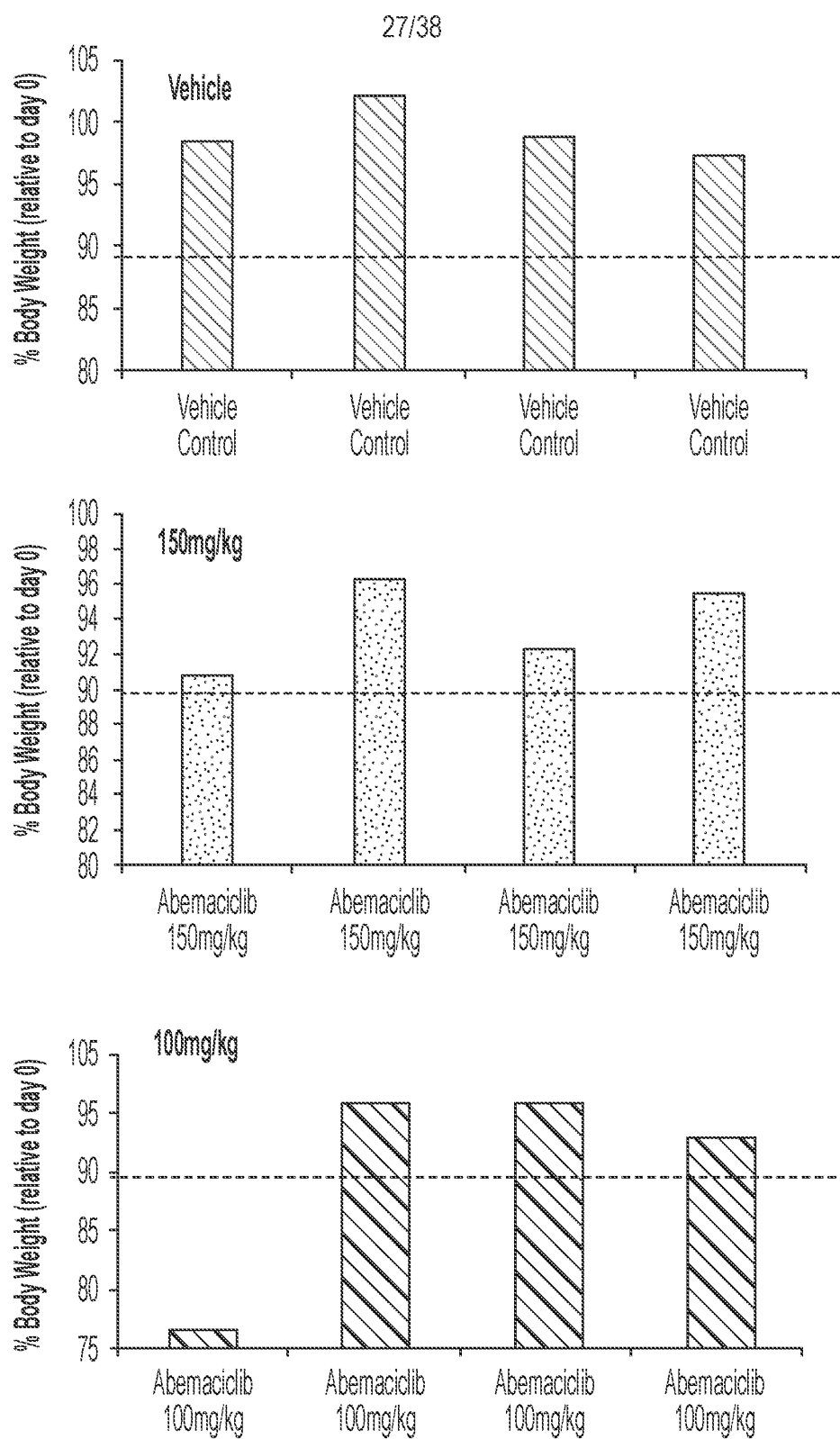


FIG. 26
SUBSTITUTE SHEET (RULE 26)

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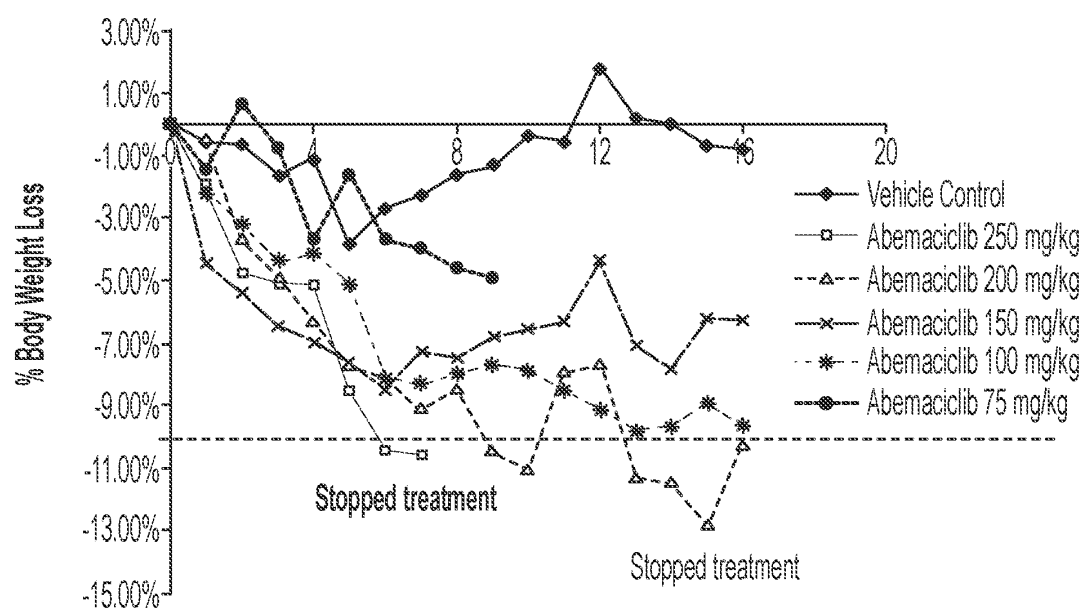


FIG. 27

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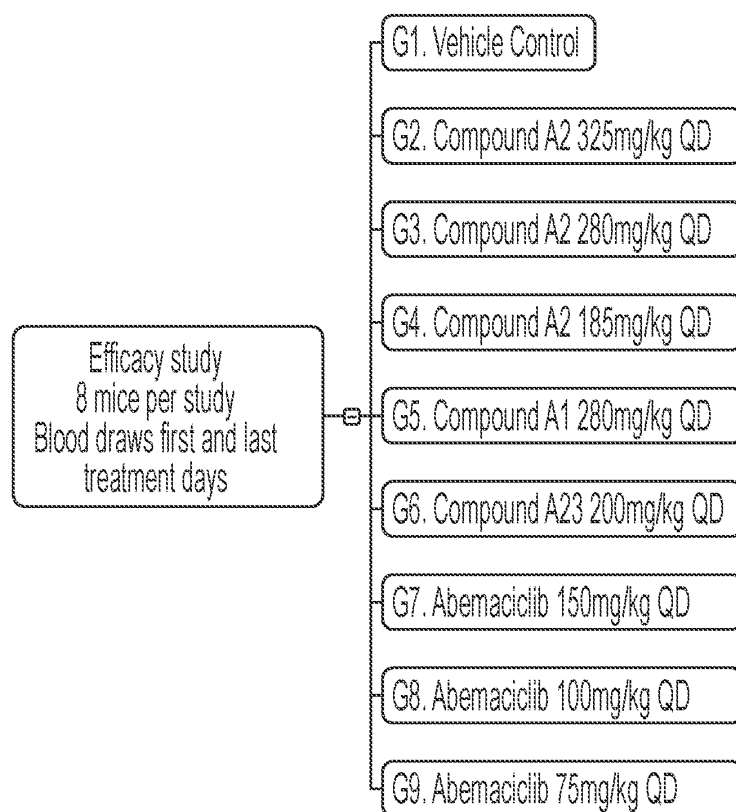


FIG. 28

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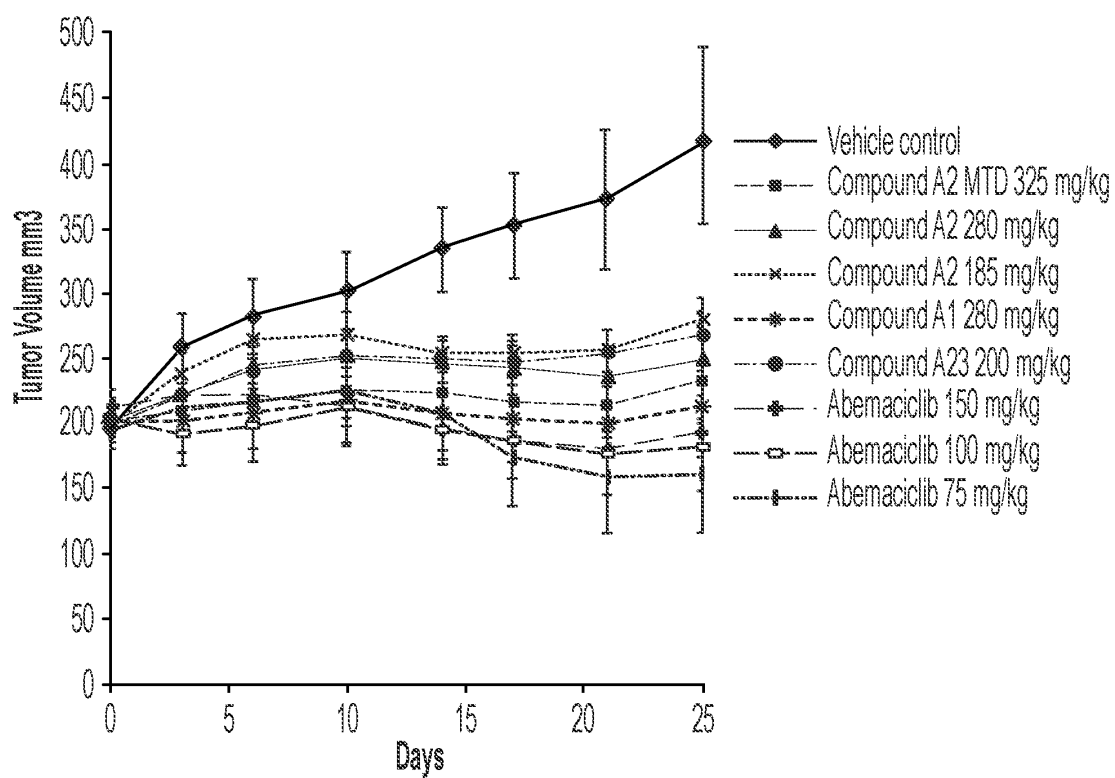


FIG. 29

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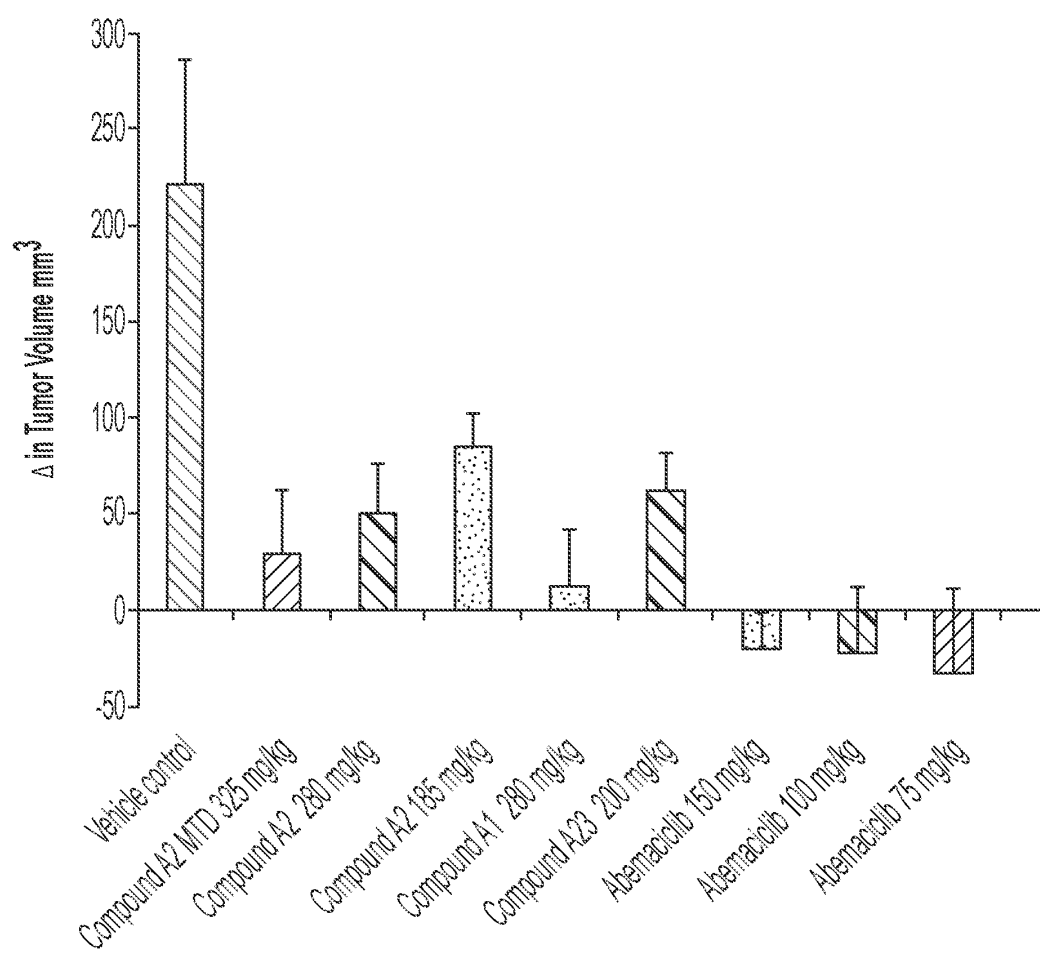


FIG. 30

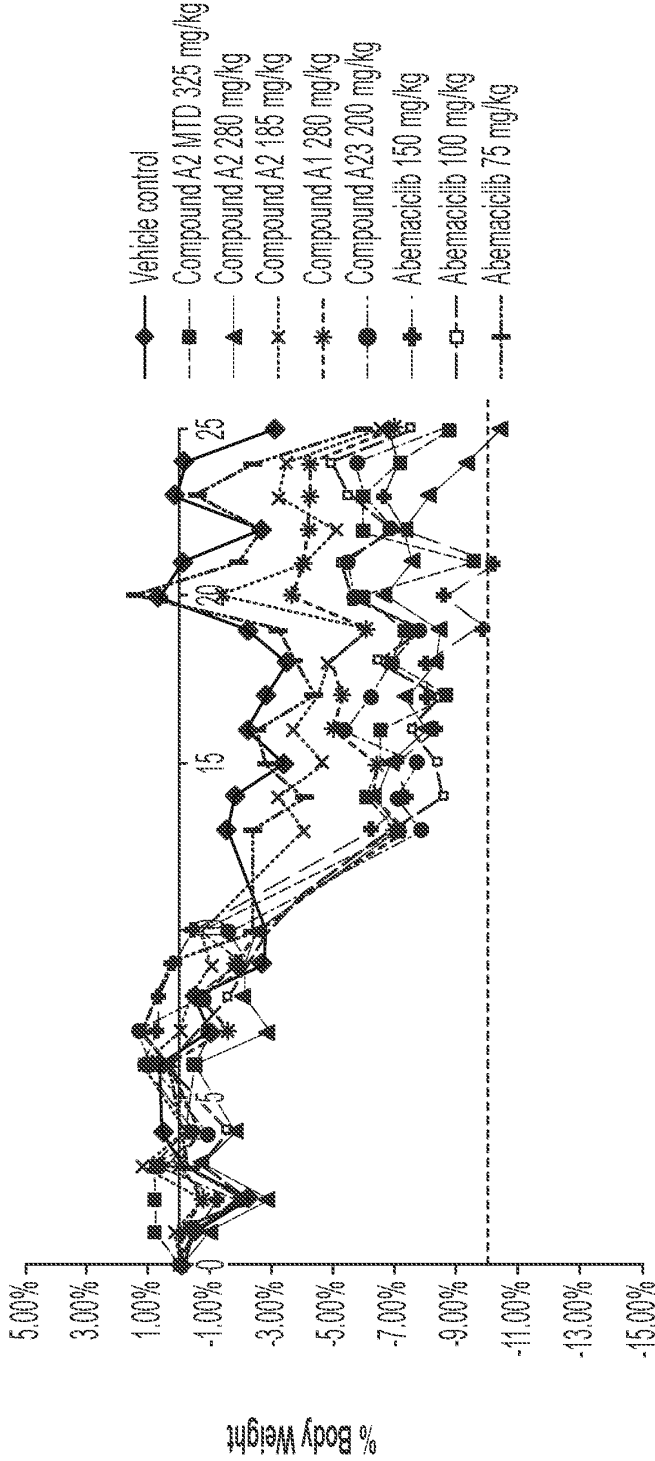


FIG. 31

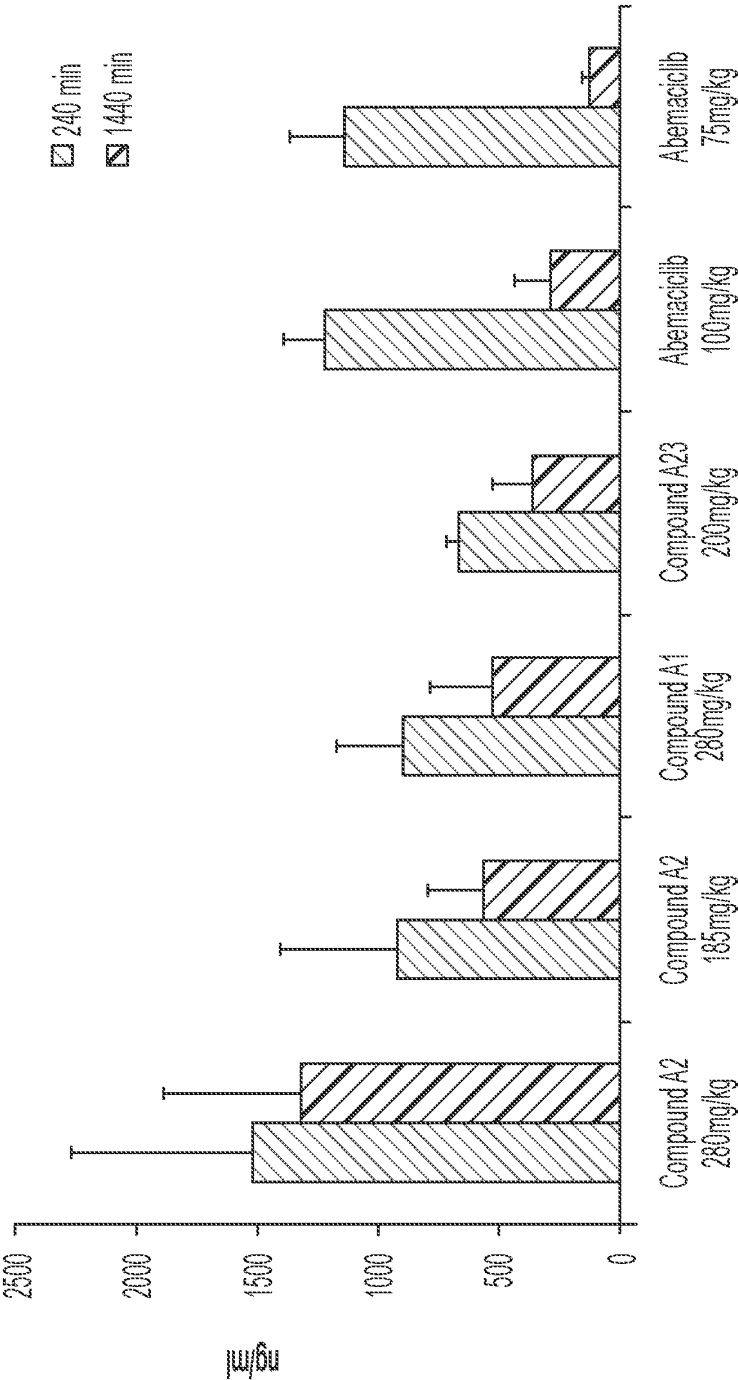


FIG. 32

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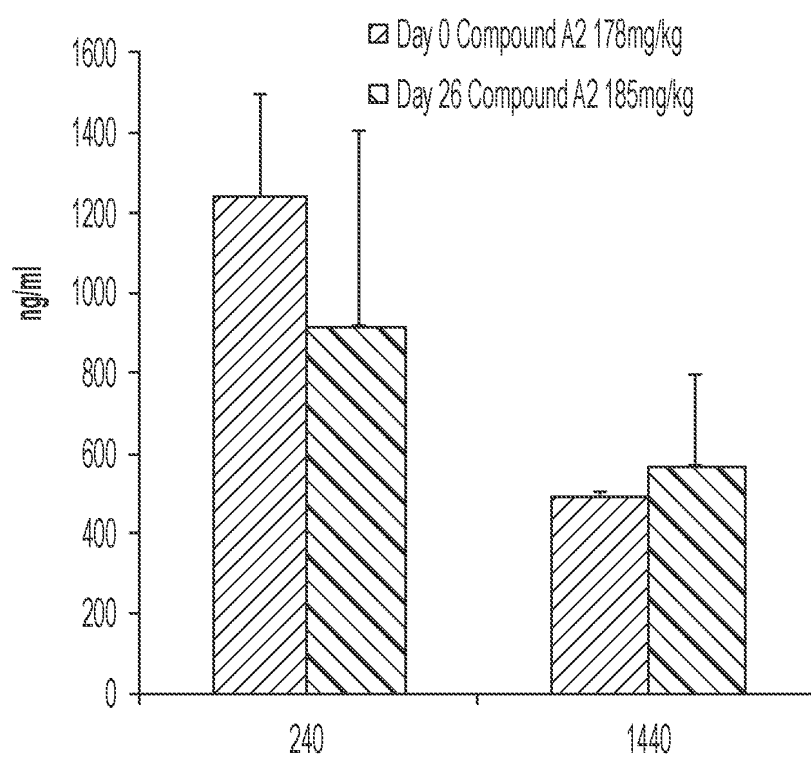


FIG. 33

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Group	Dates/Study Days												
	7/18/2017	7/20/2017	7/24/2017	7/26/2017	7/28/2017	7/31/2017	8/2/2017	8/4/2017	8/7/2017	8/9/2017	8/11/2017		
	0	2	6	8	10	13	15	17	20	22	24		
Vehicle	0.00%	-0.34%	1.86%	-0.33%	-0.32%	3.72%	-0.12%	1.83%	1.09%	0.07%	2.51%		
Abemaciclib 75 mg/kg	0.00%	-2.77%	-1.94%	-2.43%	-1.70%	-0.49%	-3.47%	-3.59%	-2.52%	-4.40%	-2.01%		
Compound A2 54mg/kg	0.00%	-2.01%	-0.30%	-3.22%	-2.69%	-0.15%	-4.54%	-5.90%	-1.00%	-4.89%	-3.32%		
Compound A2 75mg/kg	0.00%	-0.80%	-1.18%	-1.84%	-2.09%	0.06%	-2.85%	-1.19%	-1.07%	-3.93%	-1.96%		
Compound A2 123mg/kg	0.00%	-4.07%	-4.06%	-5.24%	-5.60%	-3.01%	-5.29%	-6.00%	-4.34%	-7.22%	-6.06%		
Compound A2 185mg/kg	0.00%	-1.81%	-0.67%	-3.15%	-2.36%	-1.91%	-4.14%	-5.18%	-4.23%	-6.83%	-6.29%		
Compound A2 278mg/kg	0.00%	-3.39%	-4.22%	-5.26%	-4.89%	-5.27%	-7.12%	-6.53%	-6.30%	-8.57%	-8.68%		
Compound A23 100mg/kg	0.00%	-0.84%	-0.59%	-2.53%	-2.52%	0.38%	-3.36%	-1.16%	0.43%	-3.62%	-3.07%		
Compound A12 38mg/kg	0.00%	-3.53%	-0.47%	-0.74%	-0.39%	2.42%	-1.36%	0.36%	1.31%	-2.25%	0.91%		
Compound A12 57mg/kg	0.00%	-2.83%	-1.74%	-2.35%	-2.14%	-0.05%	-2.81%	-0.78%	-0.13%	-3.23%	-0.50%		
Compound A12 86mg/kg	0.00%	0.54%	1.37%	1.24%	0.53%	3.93%	1.44%	1.74%	3.33%	0.02%	2.90%		

FIG. 34

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Compound	0	1	2	3	4	6	7	8	9	10	13
Vehicle Control	0.00%	-0.51%	-2.21%	-0.04%	0.58%	0.58%	-1.00%	-0.38%	-2.69%	-2.83%	-1.56%
A2 MTD 325mg/kg	0.00%	0.76%	0.76%	0.76%	-0.21%	-0.49%	-1.09%	-0.40%	-2.68%	-0.84%	-7.02%
A2 280mg/kg	0.00%	-0.98%	-2.87%	-0.69%	-1.85%	-0.46%	-2.93%	-2.05%	-2.12%	-2.36%	-7.06%
A2 185mg/kg	0.00%	0.14%	-0.80%	1.16%	-0.31%	1.11%	-0.05%	-0.35%	-1.03%	-0.74%	-4.07%
A1 280mg/kg	0.00%	-0.08%	-0.66%	-0.04%	-0.68%	-0.40%	-1.57%	-0.51%	-1.84%	-2.48%	-7.35%
A23 200mg/kg	0.00%	-0.40%	-1.87%	0.81%	-0.92%	0.92%	1.30%	-0.81%	-2.18%	-1.62%	-7.95%
Abemaciclib 150 mg/kg	0.00%	-0.57%	-1.24%	0.47%	-0.51%	1.20%	0.74%	0.64%	0.23%	-0.44%	-6.20%
Abemaciclib 100 mg/kg	0.00%	0.00%	-2.21%	-0.78%	-1.59%	0.46%	-0.77%	-1.56%	-2.08%	-2.94%	-6.80%
Abemaciclib 75 mg/kg	0.00%	-0.39%	-2.25%	-0.32%	-0.12%	0.27%	1.20%	0.71%	0.29%	-2.40%	-2.35%

FIG. 35A

Compound	14	15	16	17	18	19	20	21	22	23	24	25
Vehicle Control	-1.80%	-3.38%	-2.19%	-2.81%	-3.49%	-2.20%	0.66%	-0.04%	-2.67%	0.09%	-0.17%	-3.05%
A2 MTD 325mg/kg	-6.03%	-6.54%	-6.57%	-8.78%	-6.93%	-7.33%	-5.96%	-9.58%	-5.98%	-6.02%	-7.17%	-8.78%
A2 280mg/kg	-6.41%	-6.73%	-8.00%	-7.32%	-8.34%	-8.45%	-6.65%	-7.54%	-7.21%	-8.11%	-9.27%	-10.39%
A2 185mg/kg	-3.22%	-4.63%	-3.71%	-4.54%	-4.84%	-5.97%	-1.25%	-4.05%	-5.12%	-3.23%	-3.44%	-6.47%
A1 280mg/kg	-6.26%	-6.39%	-4.98%	-5.27%	-4.95%	-6.10%	-3.63%	-3.98%	-4.12%	-4.21%	-4.20%	-6.96%
A23 200mg/kg	-7.11%	-7.69%	-5.40%	-6.23%	-6.81%	-7.76%	-5.71%	-5.50%	-6.88%	-6.00%	-5.73%	-8.74%
Abemaciclib 150 mg/kg	-7.37%	-7.03%	-8.30%	-8.03%	-8.03%	-9.79%	-8.58%	-10.20%	-7.43%	-6.60%	-7.14%	-6.75%
Abemaciclib 100 mg/kg	-8.58%	-8.37%	-7.52%	-8.40%	-6.50%	-7.91%	-5.62%	-5.28%	-6.86%	-5.47%	-4.95%	-7.55%
Abemaciclib 75 mg/kg	-4.05%	-2.81%	-2.58%	-4.39%	-3.79%	-3.26%	1.41%	-2.00%	-2.74%	-0.49%	-2.47%	5.98%

FIG. 35B

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Xenograft Model/Group	No. Mice	Mean tumor volume mm ³			T/C	Max Mean % Body Weight Loss	No. of mice to drop below 10% BWL	Deaths	
		Day 1	Day last	Change = DLast-D1				TR	NTR
ZR751 Vehicle Control	8	196	417	221	-	3.5	0	0	0
325mg/kg Compound A2	8	203	234	30	0.561	9.6	5	2	0
280mg/kg Compound A2	8	198	249	51	0.597	10.4	5	0	0
185mg/kg Compound A2	8	197	281	85	0.675	6.5	0	0	0
280mg/kg Compound A1	8	200	213	13	0.510	7.0	2	0	0
200mg/kg Compound A23	8	206	267	62	0.642	8.7	3	1	0
150mg/kg Abemaciclib	8	212	193	-19	0.464	10.2	6	2	0
100mg/kg Abemaciclib	8	203	182	-21	0.436	8.6	3	0	0
75mg/kg Abemaciclib	8	194	161	-33	0.386	6.0	1	0	0

T/C: ratio of final tumor volume in treated vs control
 TR: Treatment Related, NTR: Non-Treatment related

FIG. 36

SUBSTITUTE SHEET (RULE 26)

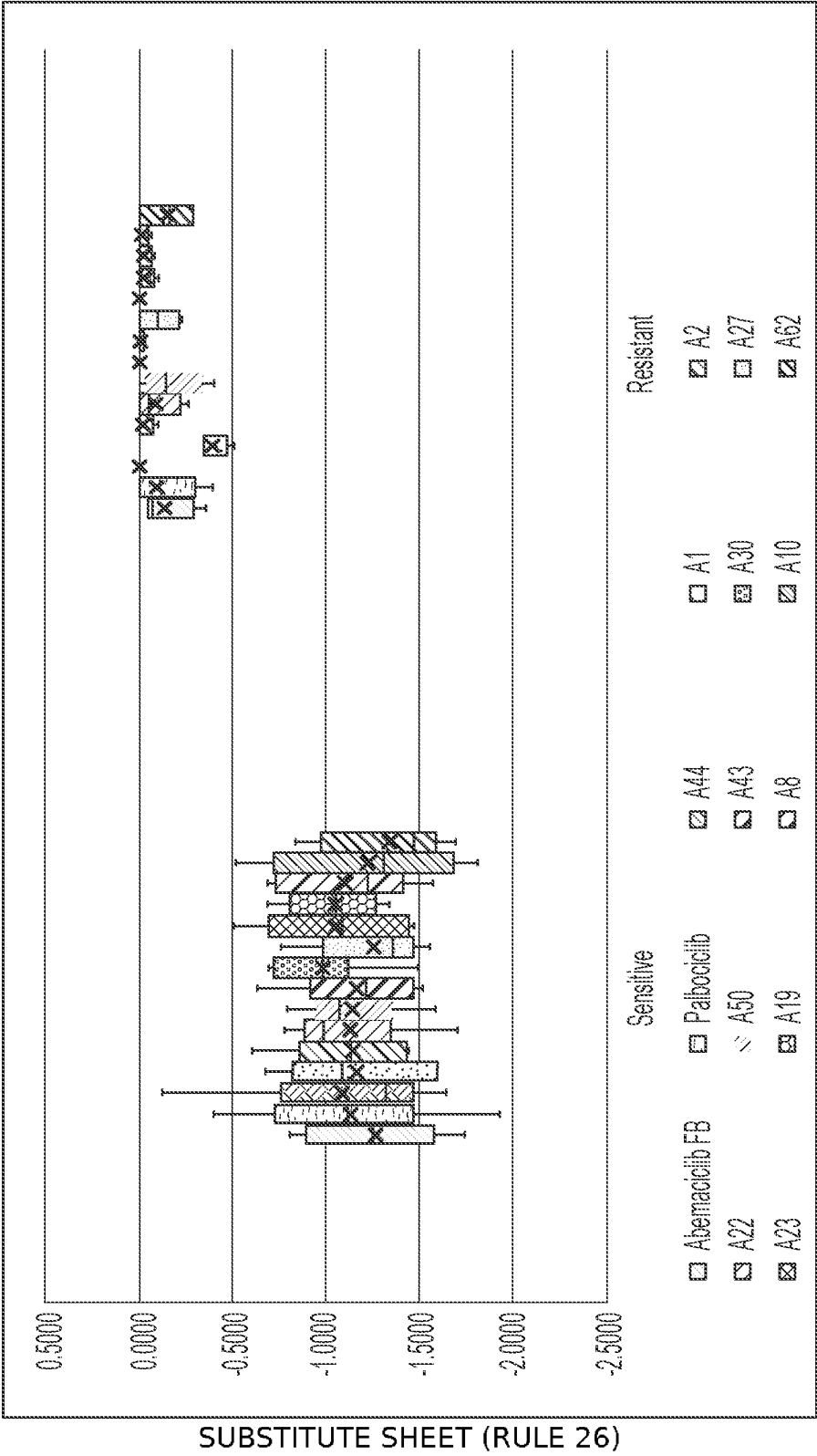


FIG. 37