

Inhibitor	Biochemical IC ₅₀ (nM)									
		CDK 2 /E	CDK 2 /A	CDK 1 /B	CDK 3 /E	CDK4 /D1	CDK 5 /p35	CDK6 /D3	CDK 7 /H	CDK 9 /T
Cmpd. I	IC ₅₀ (nM)	0.6	2.4	30	13	133	18	338	1000	73
	<i>Fold vs CDK2/ E</i>	1	4	55	23	241	32	615	1818	132
PF- 0710409 1	IC ₅₀ (nM)	2.4	7	66	ND	>1000	ND	>1000	ND	363
	<i>Fold vs CDK2/ E</i>	1	4	28	ND	>1000	ND	>1000	ND	152

FIG. 1A

Compound I Residence Time (min)				
	CDK2/E	CDK2/A2	CDK1/B	CDK9/T
Minutes	17	17	0.055	0.21
<i>Fold decrease vs CDK2/E</i>	1	1	309	81

FIG. 1B

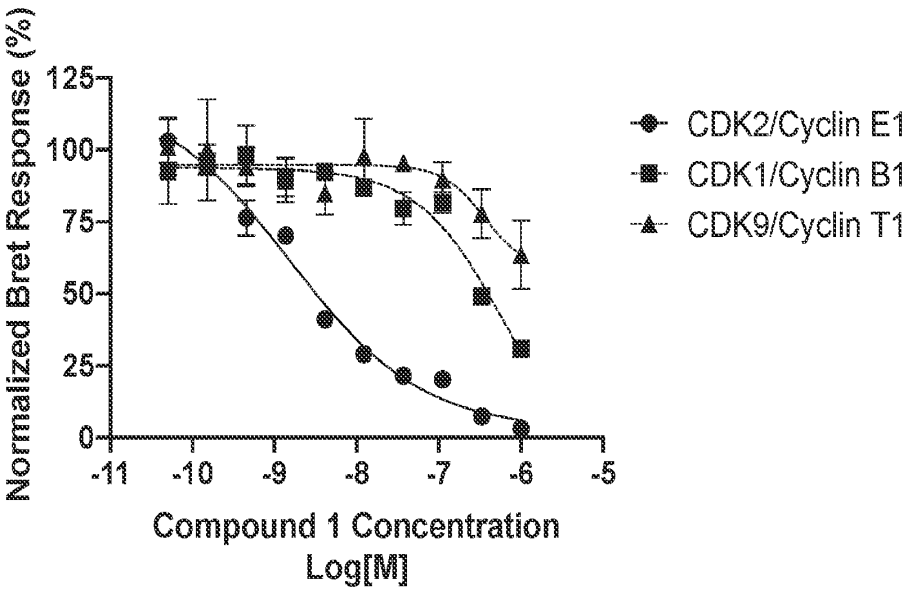


FIG. 2A

Compound I Intracellular IC ₅₀			
	CDK2/E	CDK1/B	CDK9/T
IC ₅₀ (nM)	1.5	629	>1000
Fold vs CDK2/E	1	419	>667

FIG. 2B

Compound I Intracellular NanoBRET IC ₅₀				
	CDK2/E	CDK1/B	CDK2/A	CDK9/T
IC ₅₀ (nM)	2.3	374	71.3	2950
Fold vs CDK2/E	1	163	31	1282

FIG. 2C

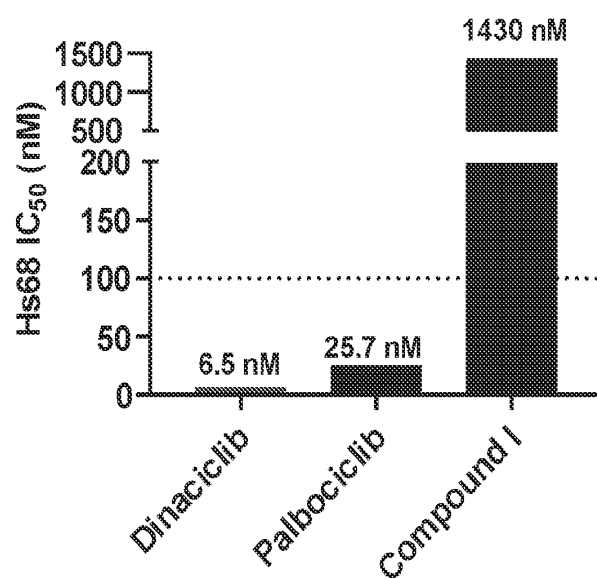


FIG. 3

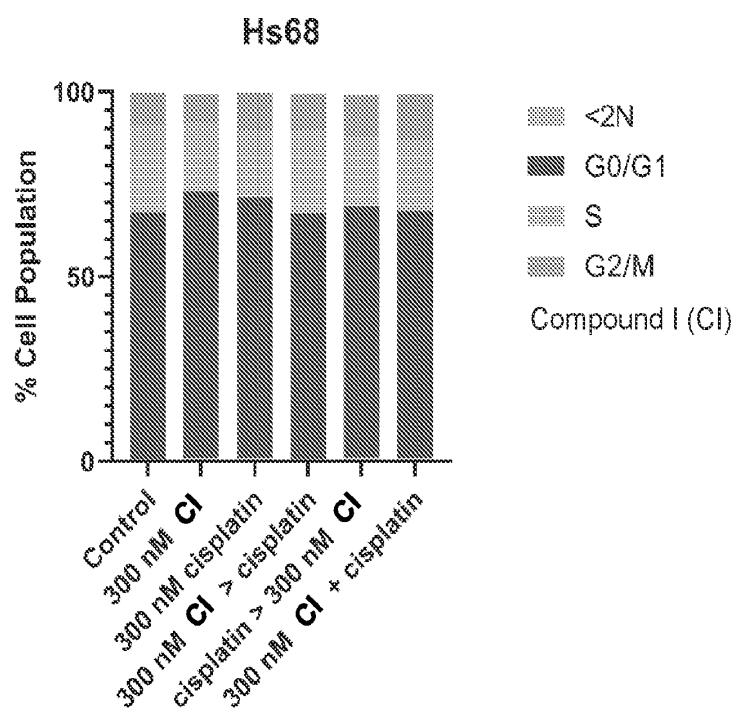


FIG. 4

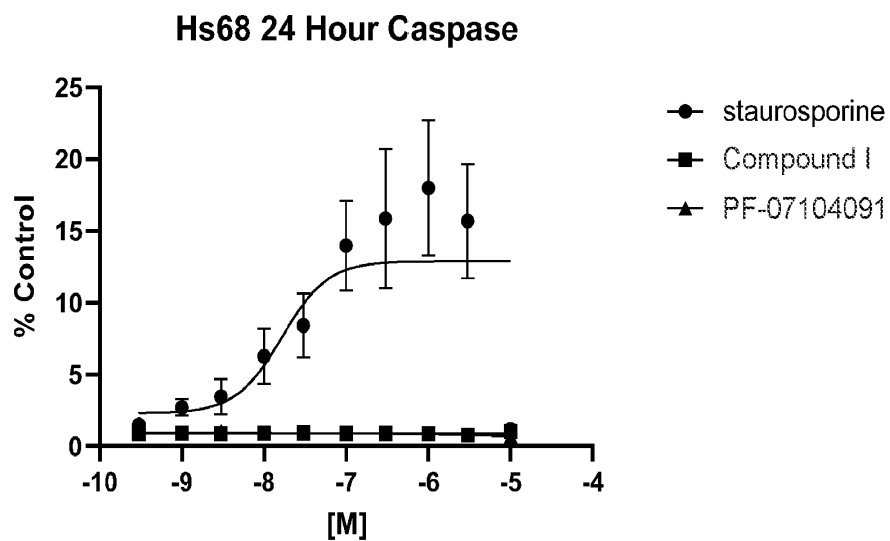


FIG. 5A

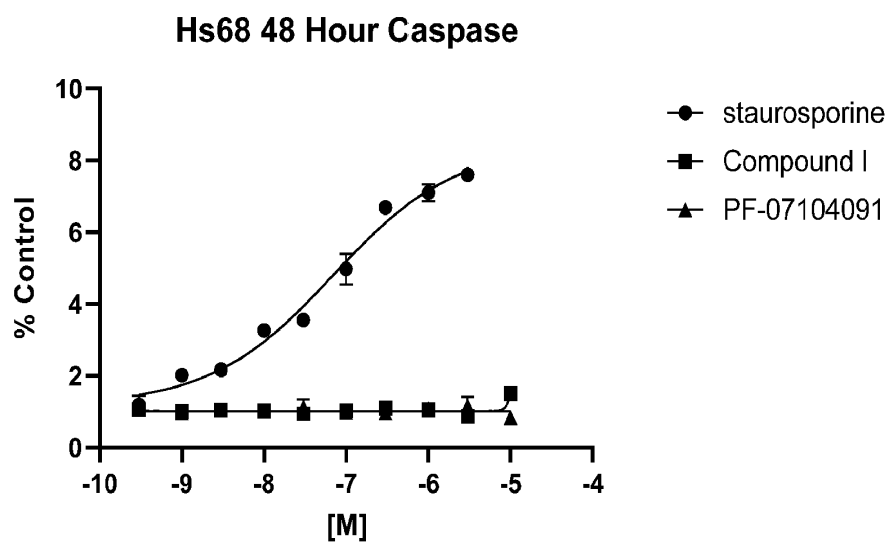


FIG. 5B

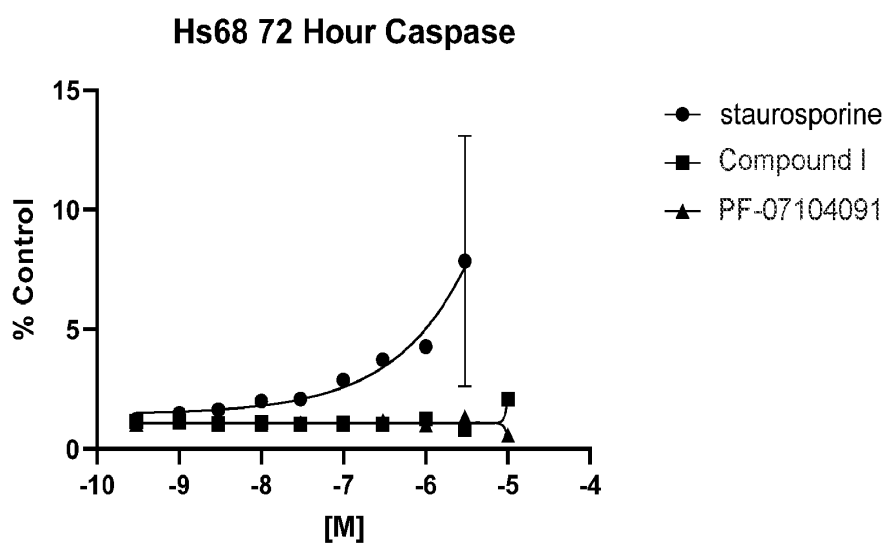


FIG. 5C

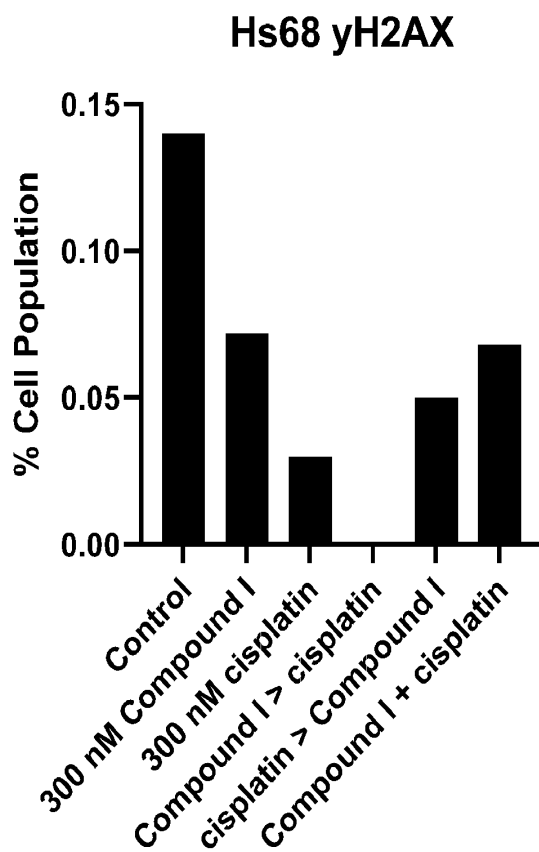


FIG. 6

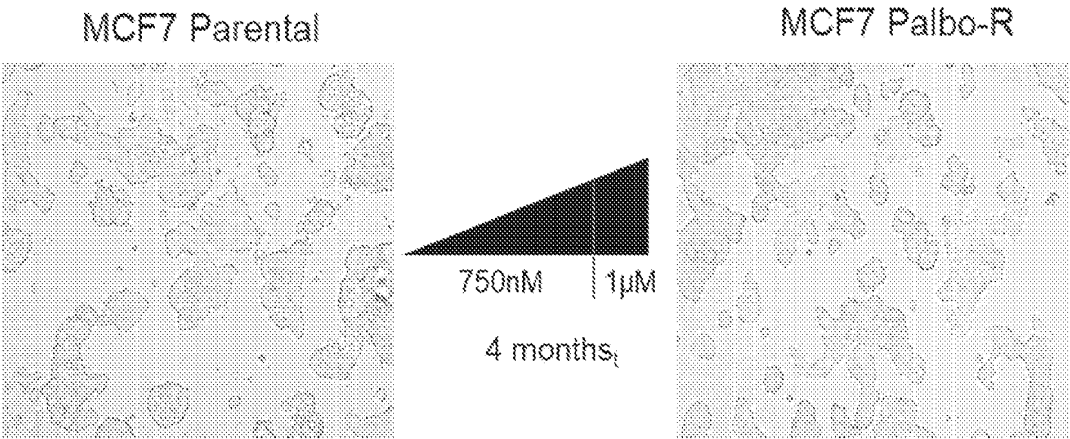


FIG. 7A

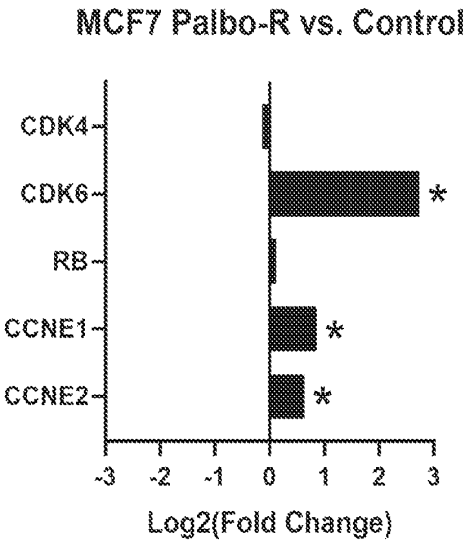


FIG. 7B

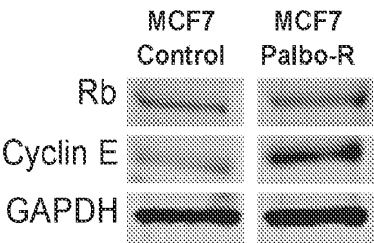


FIG. 7C

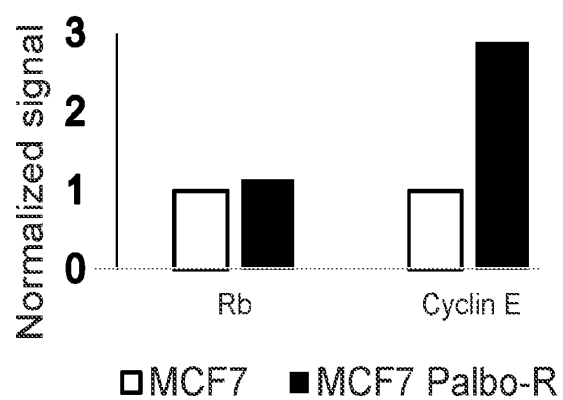


FIG. 7D

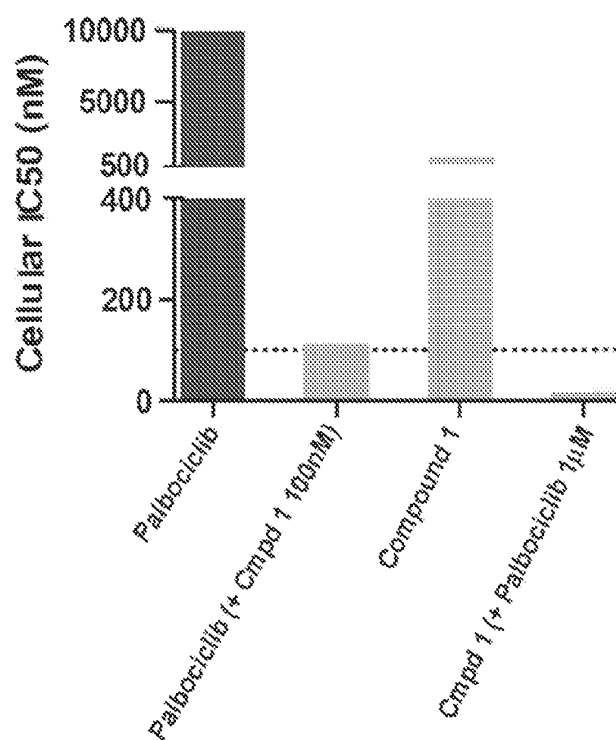


FIG. 8A

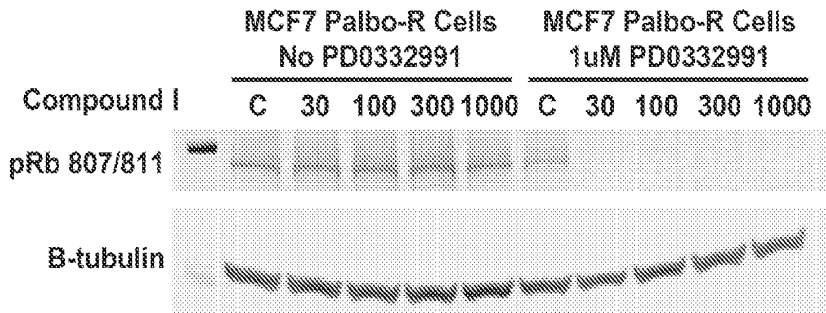


FIG. 8B

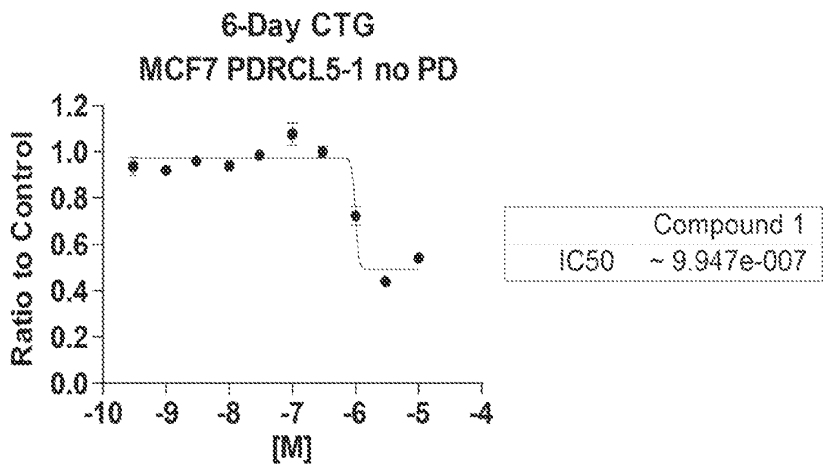


FIG. 8C

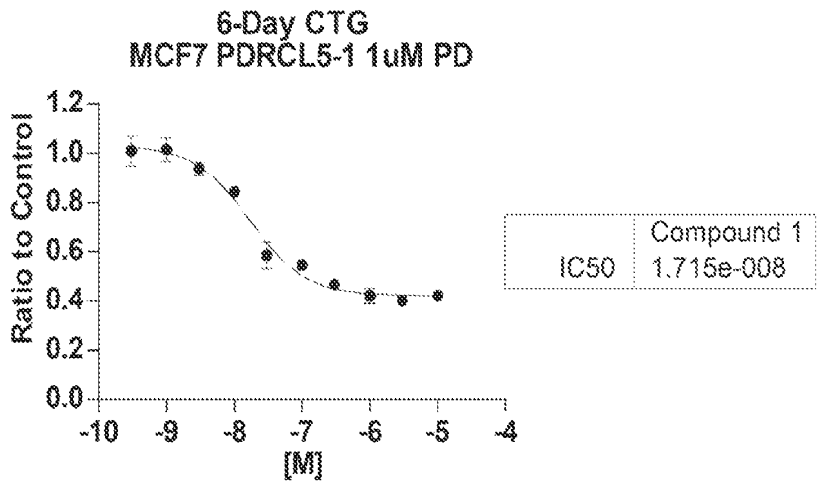


FIG. 8D

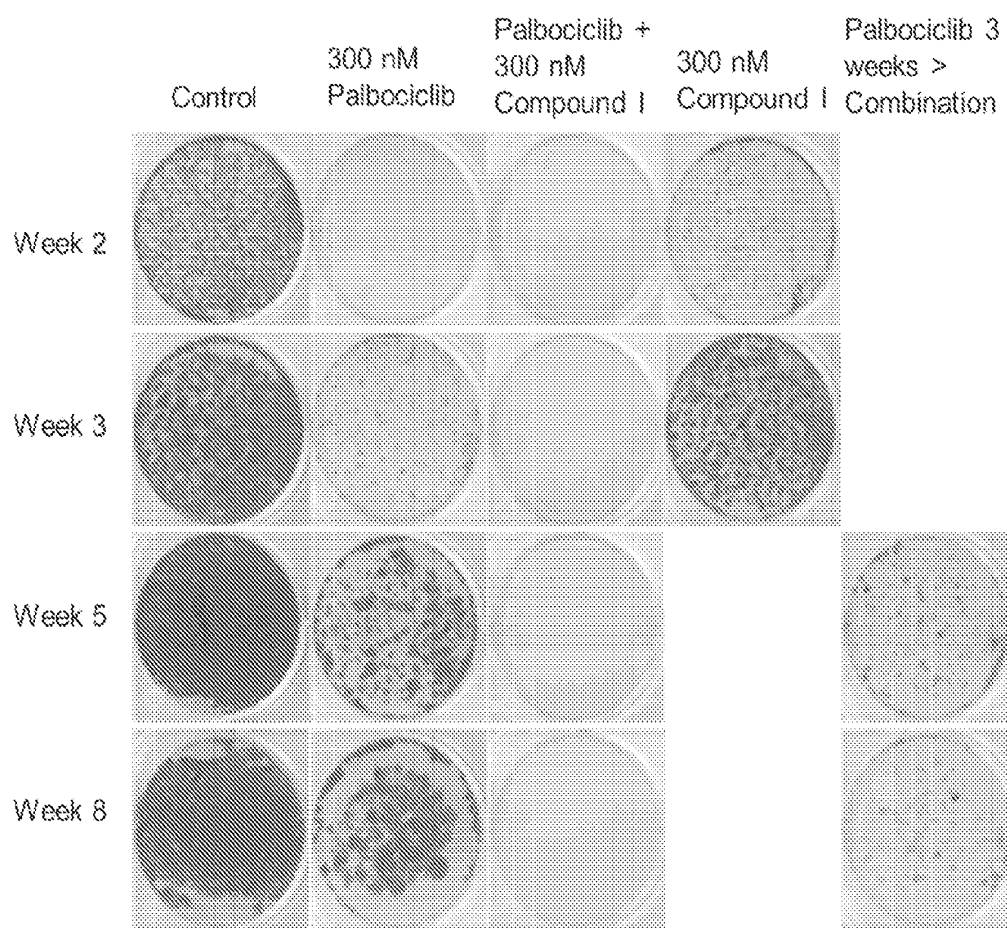


FIG. 8E

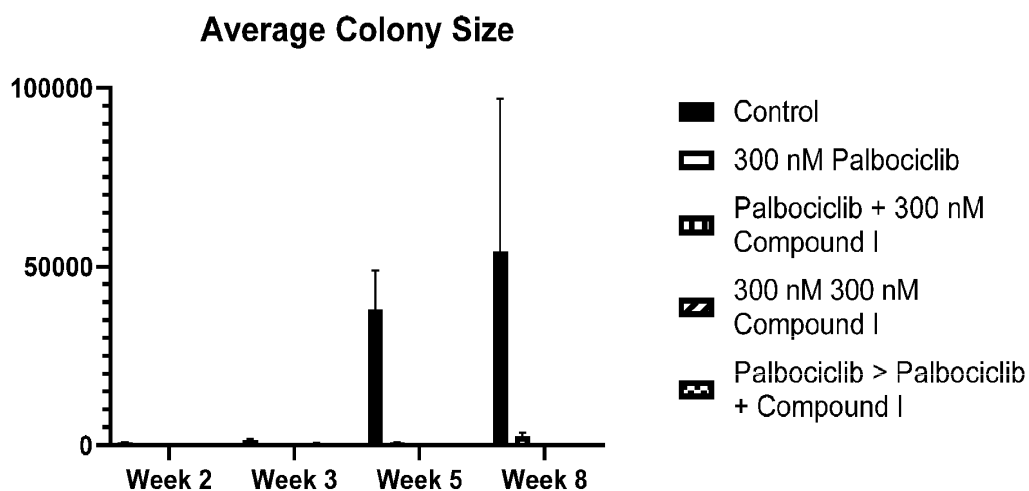


FIG. 8F

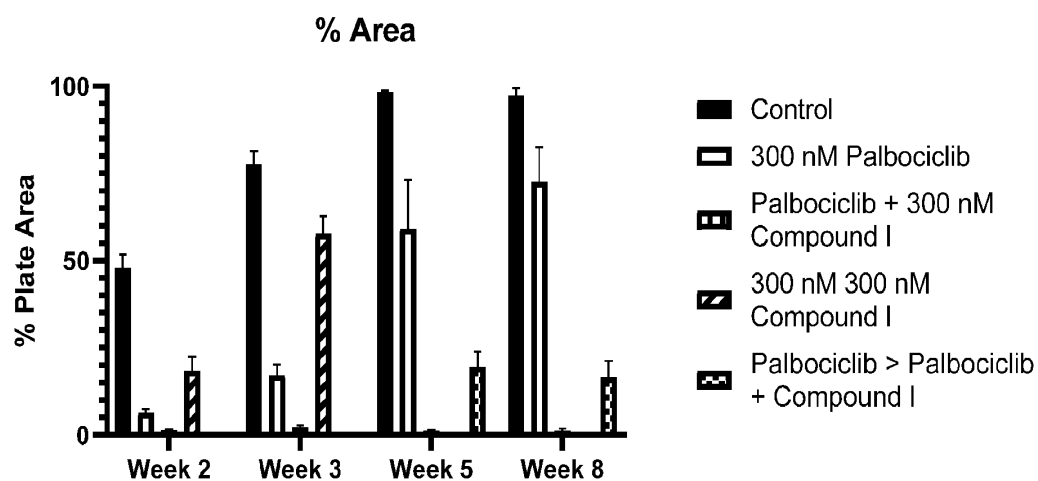


FIG. 8G

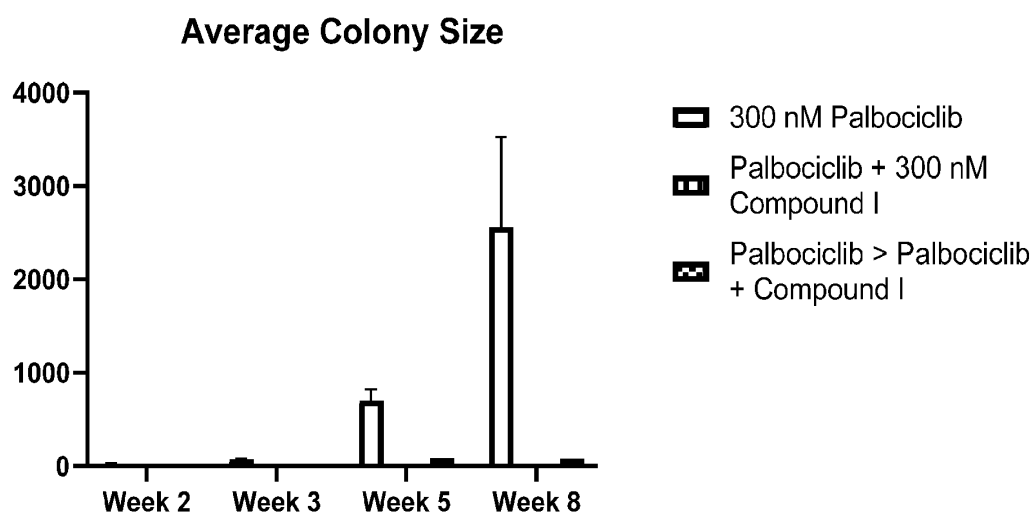


FIG. 8H

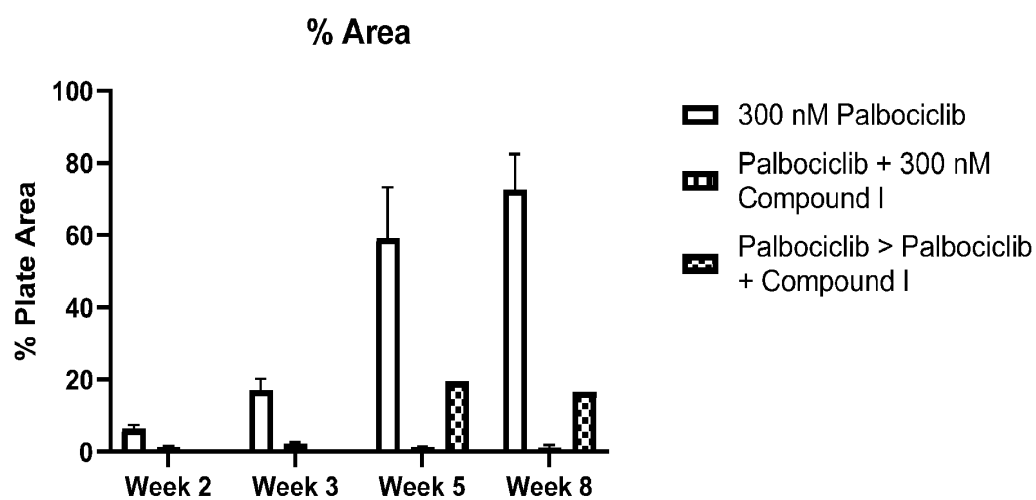


FIG. 8I

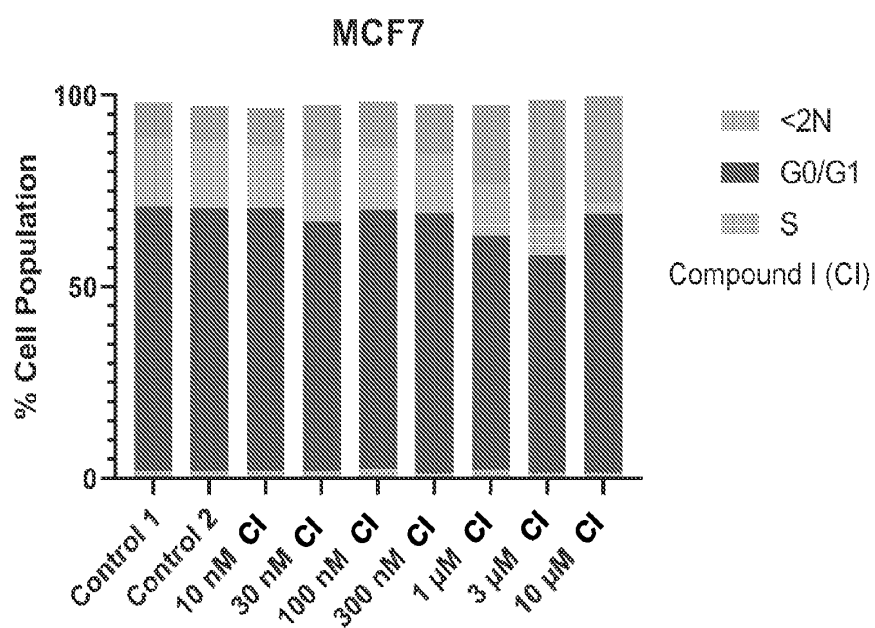


FIG. 9A

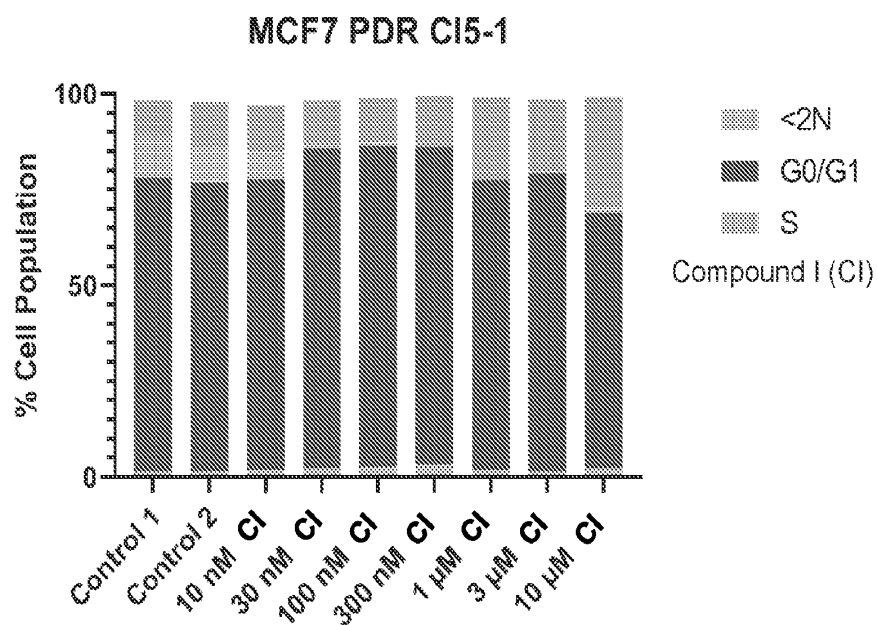


FIG. 9B

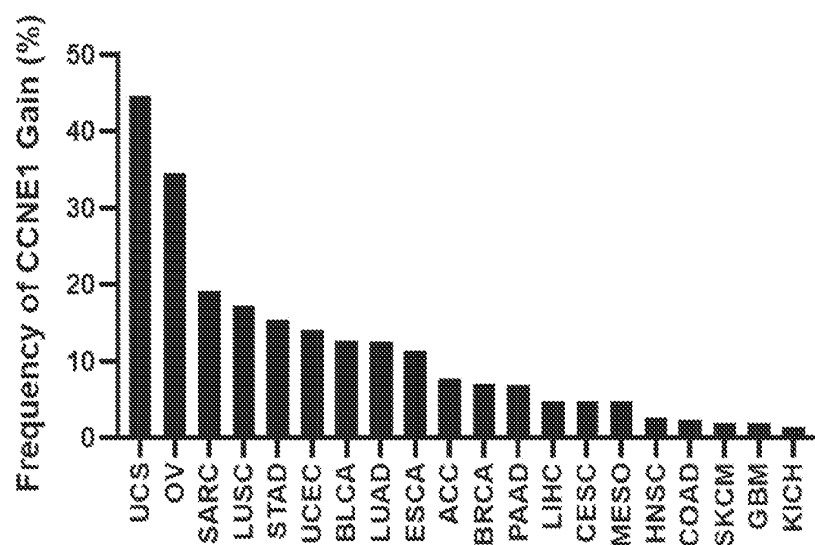


FIG. 10A

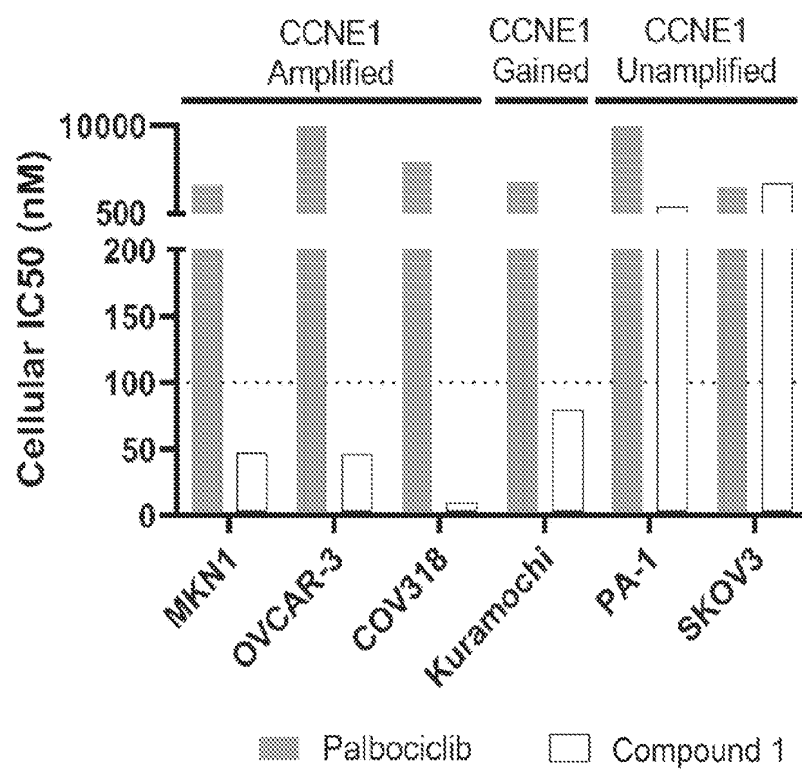


FIG. 10B

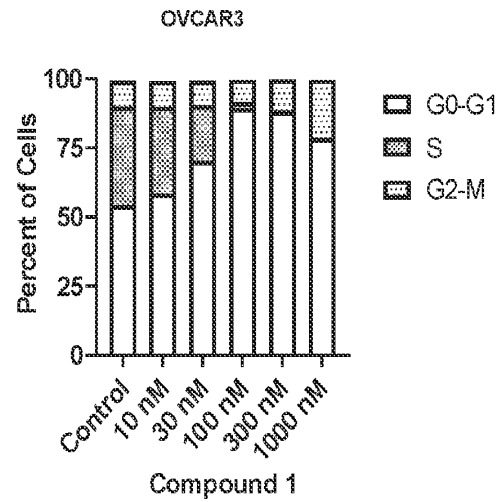


FIG. 10C

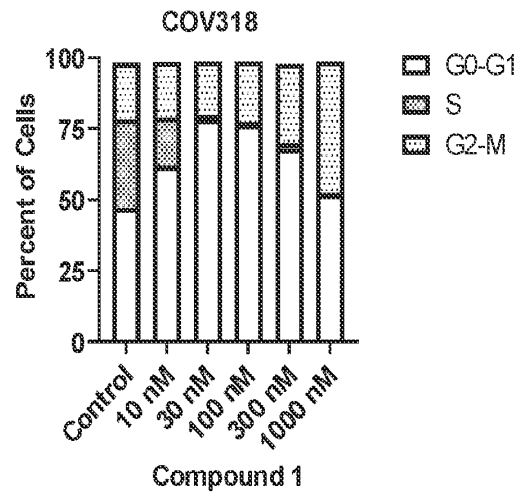
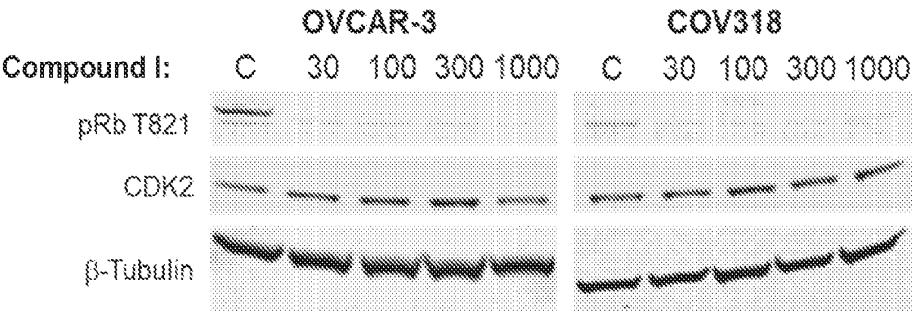


FIG. 10D



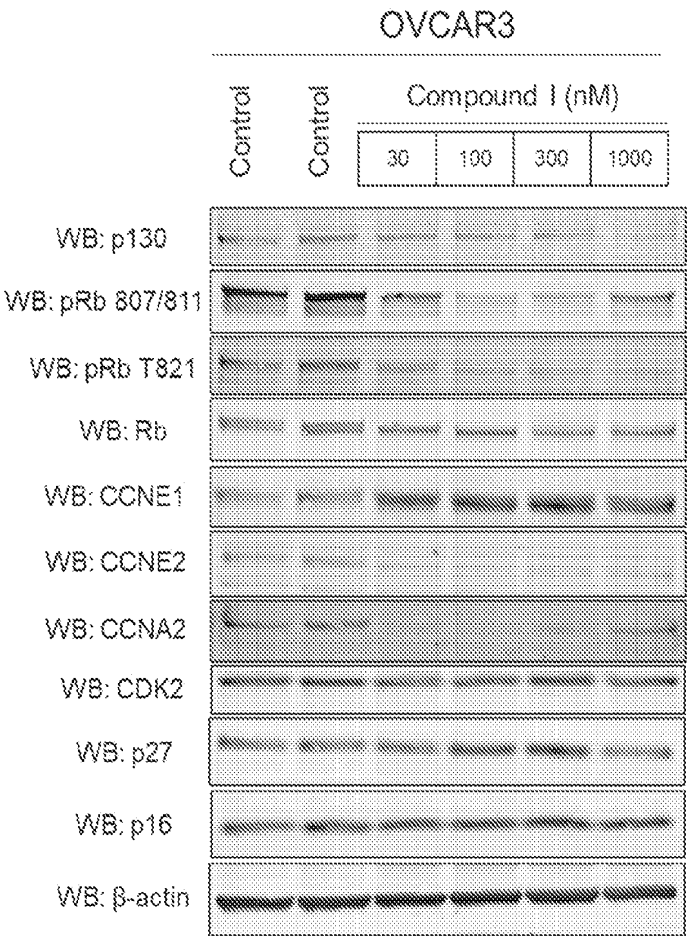


FIG. 11A

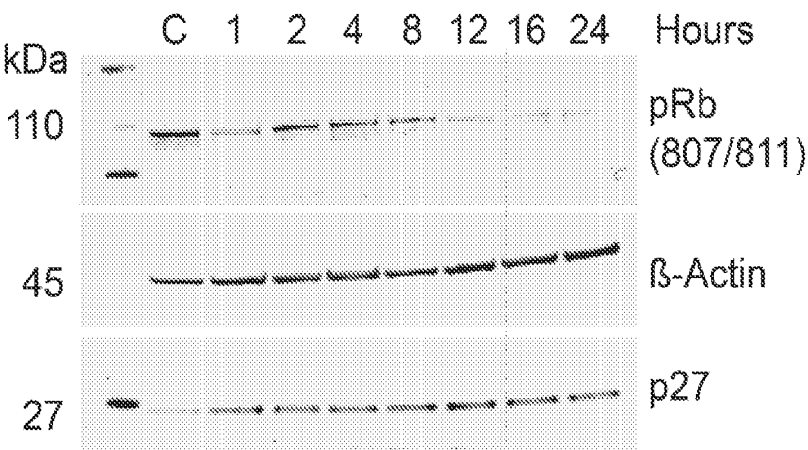


FIG. 11B

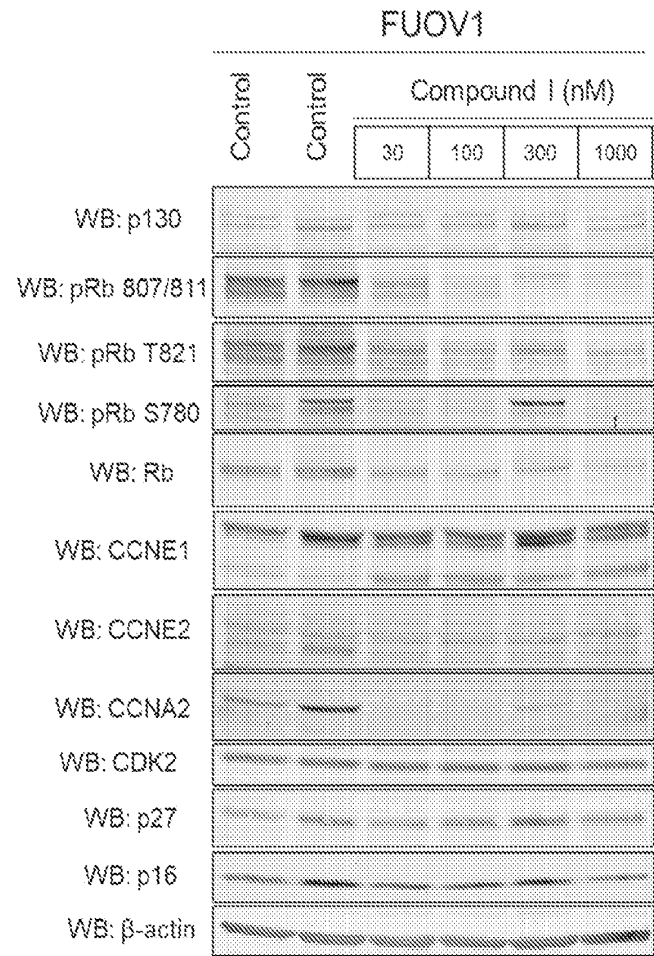


FIG. 12A

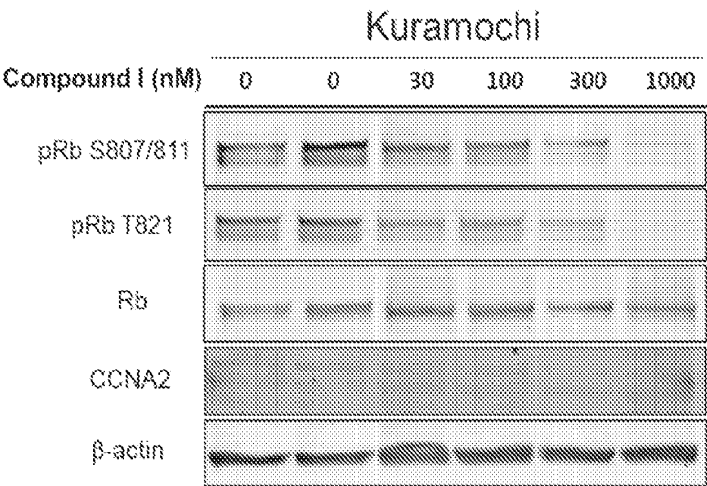


FIG. 12B

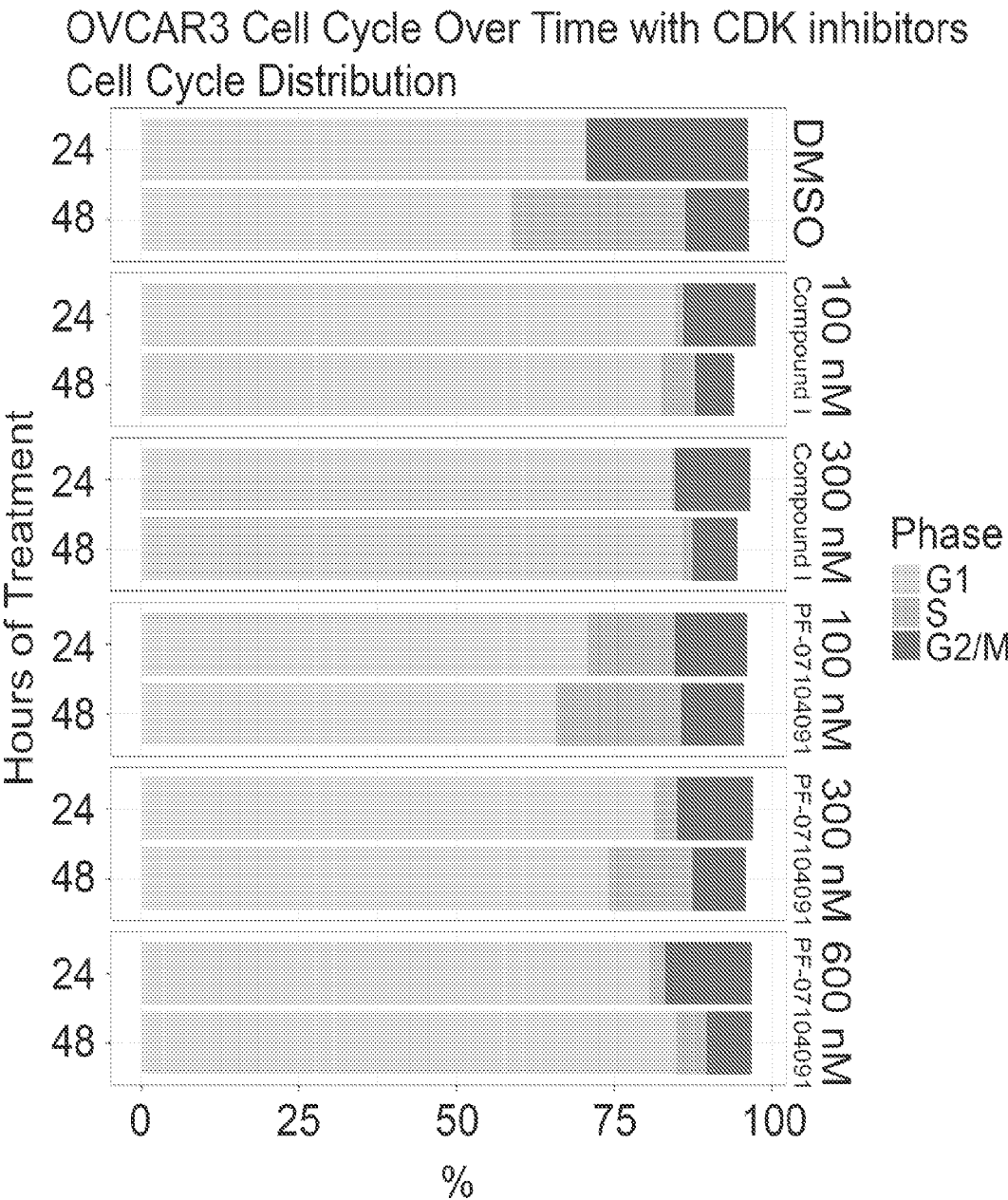


FIG. 13A

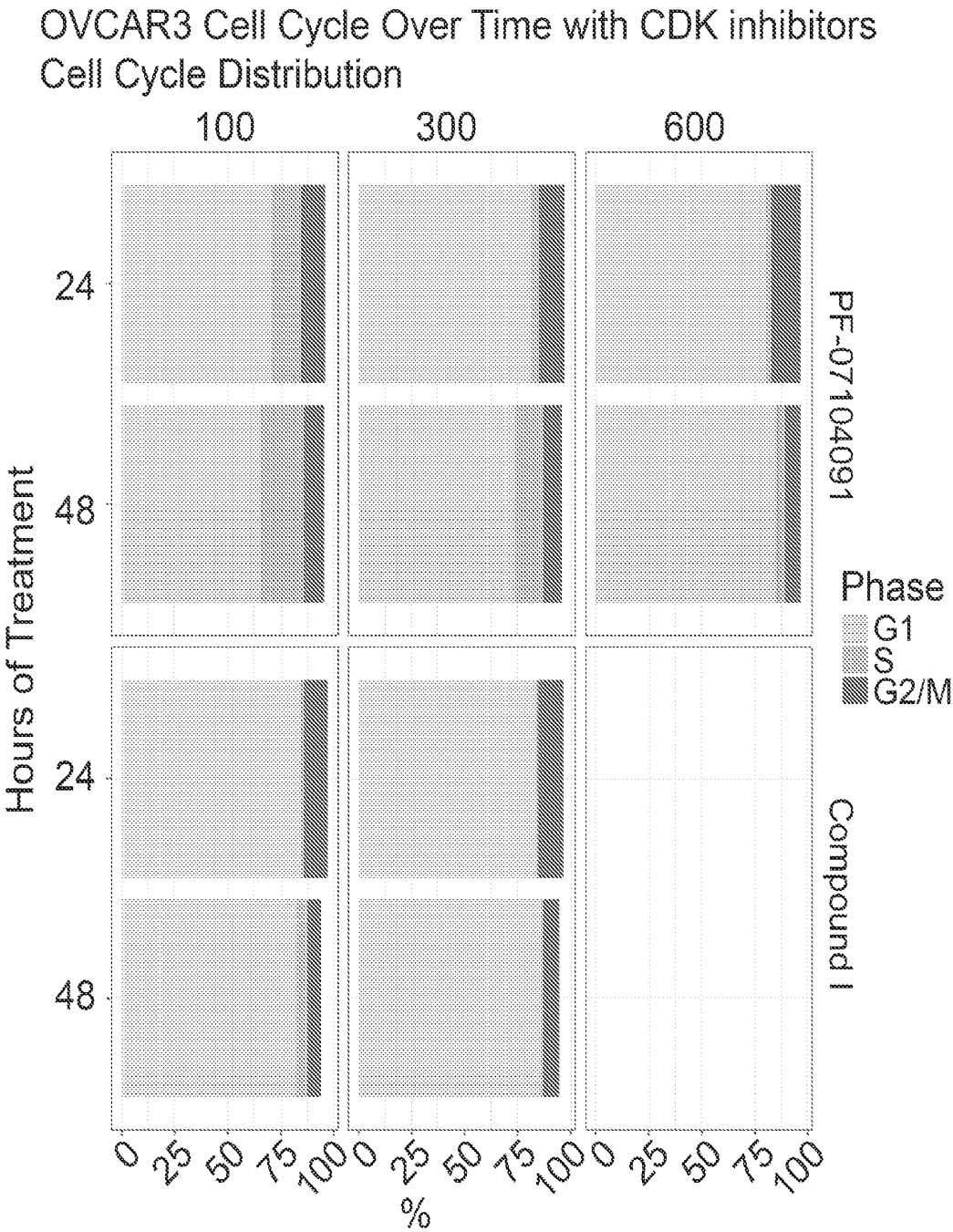


FIG. 13B

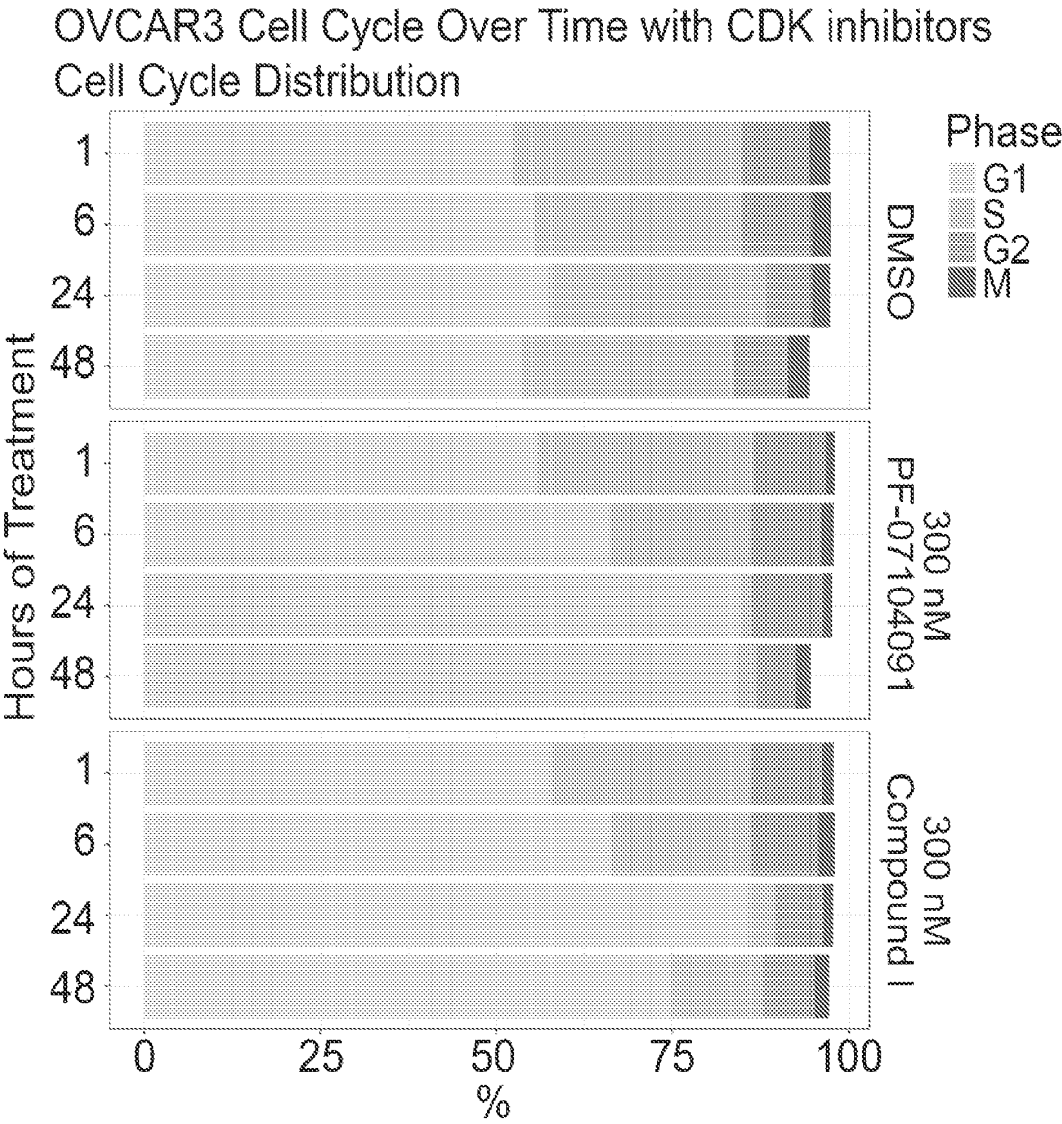


FIG. 13C

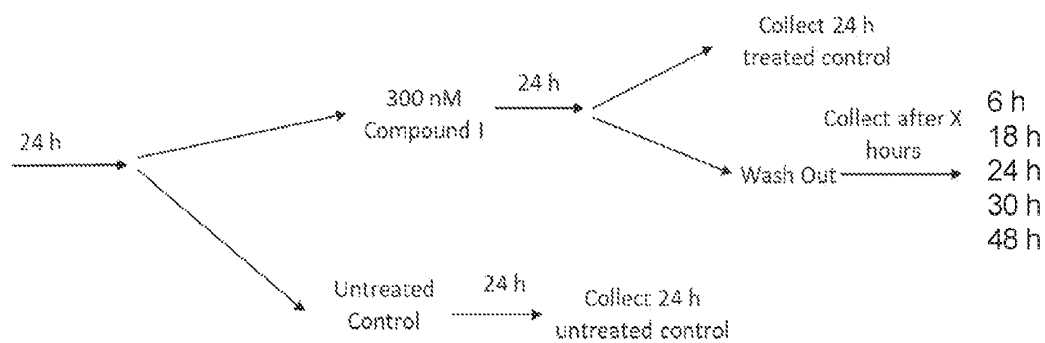


FIG. 14A

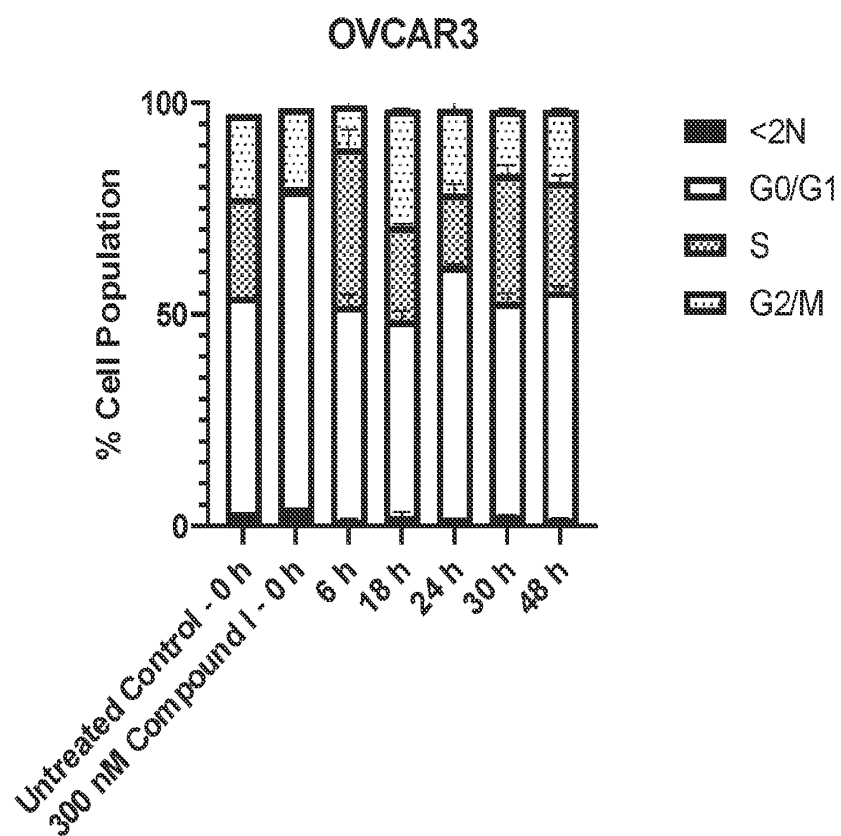
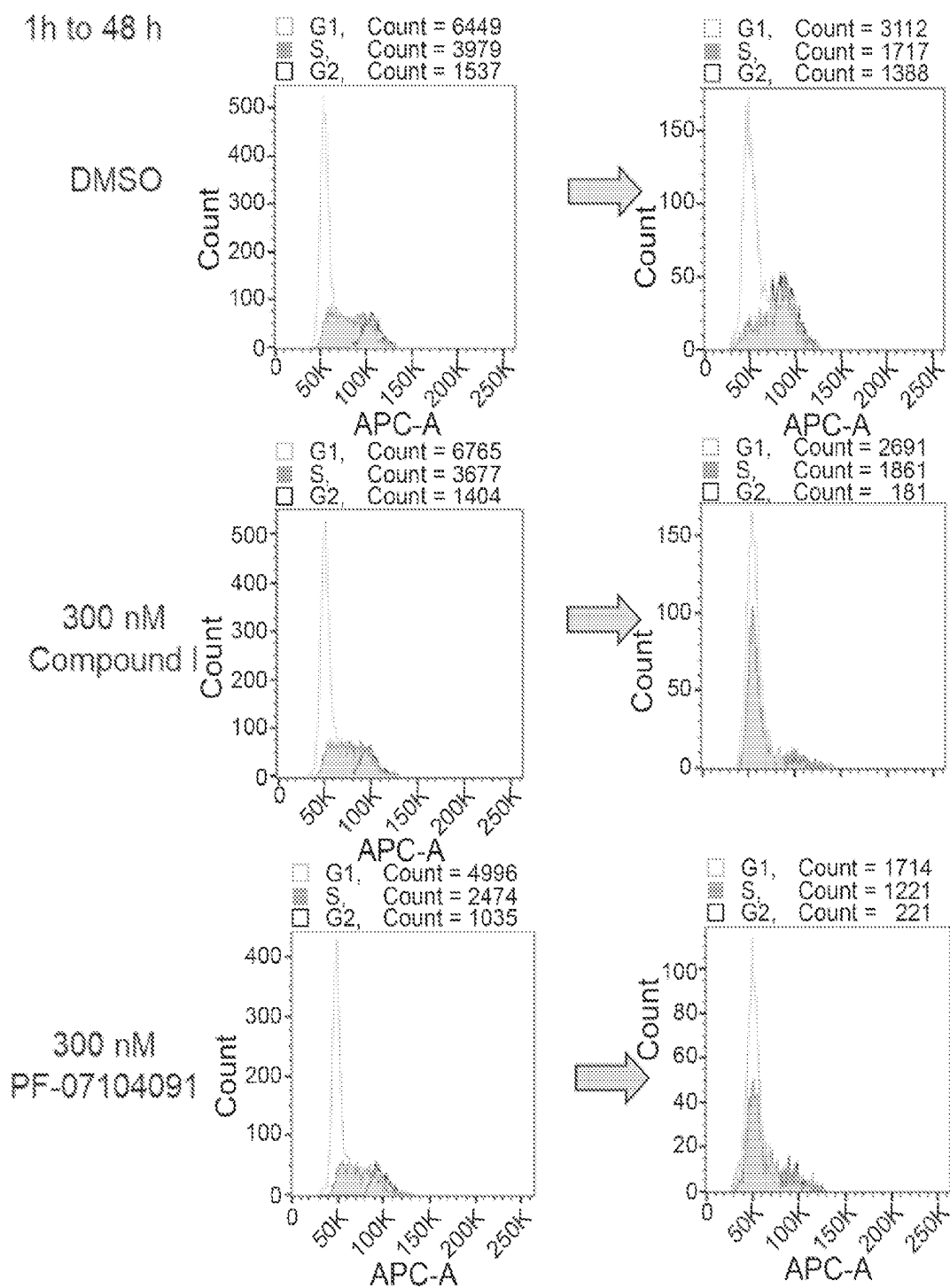


FIG. 14B



6h to 48 h

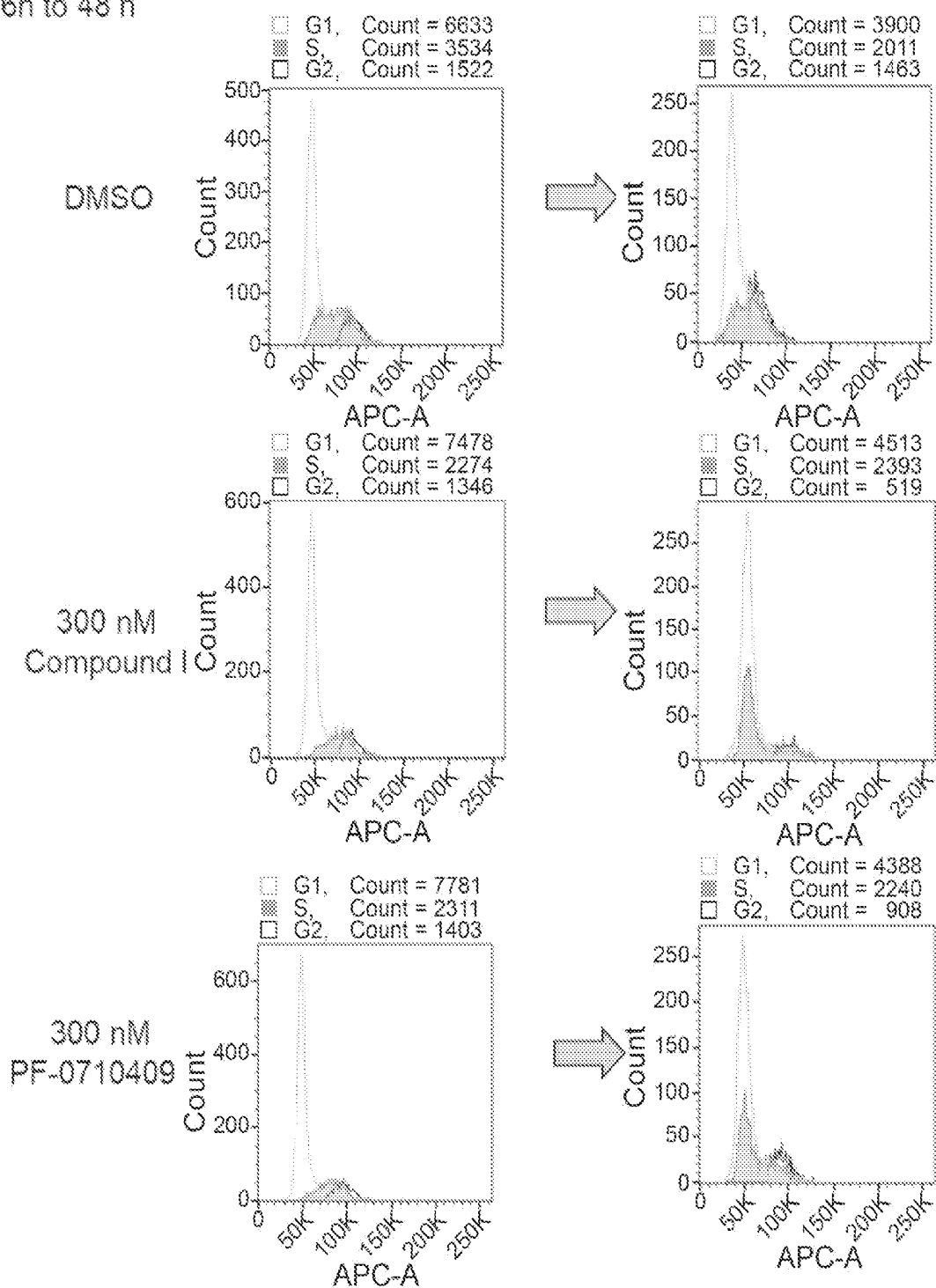


FIG. 15B

24h to 48 h

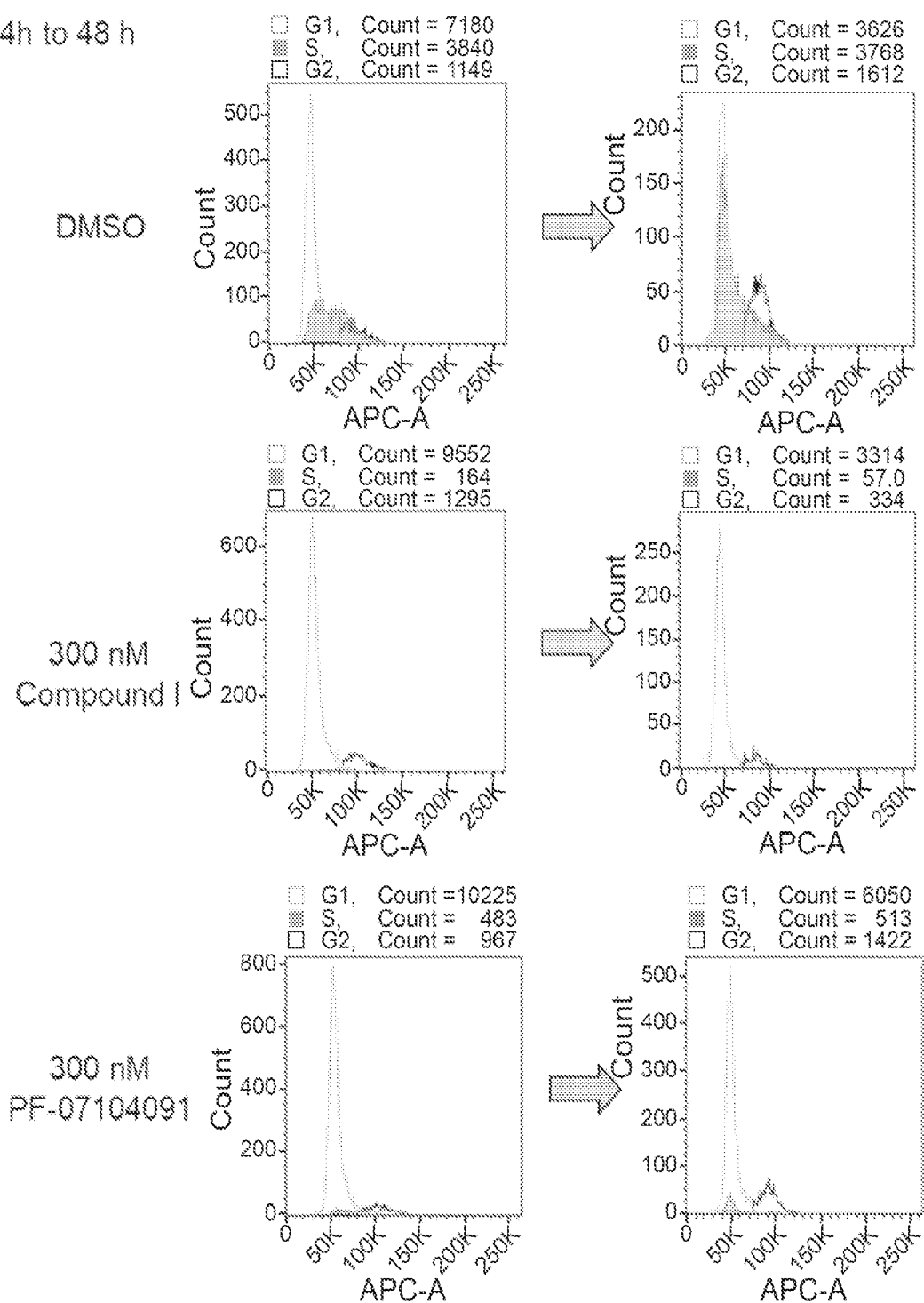


FIG. 15C

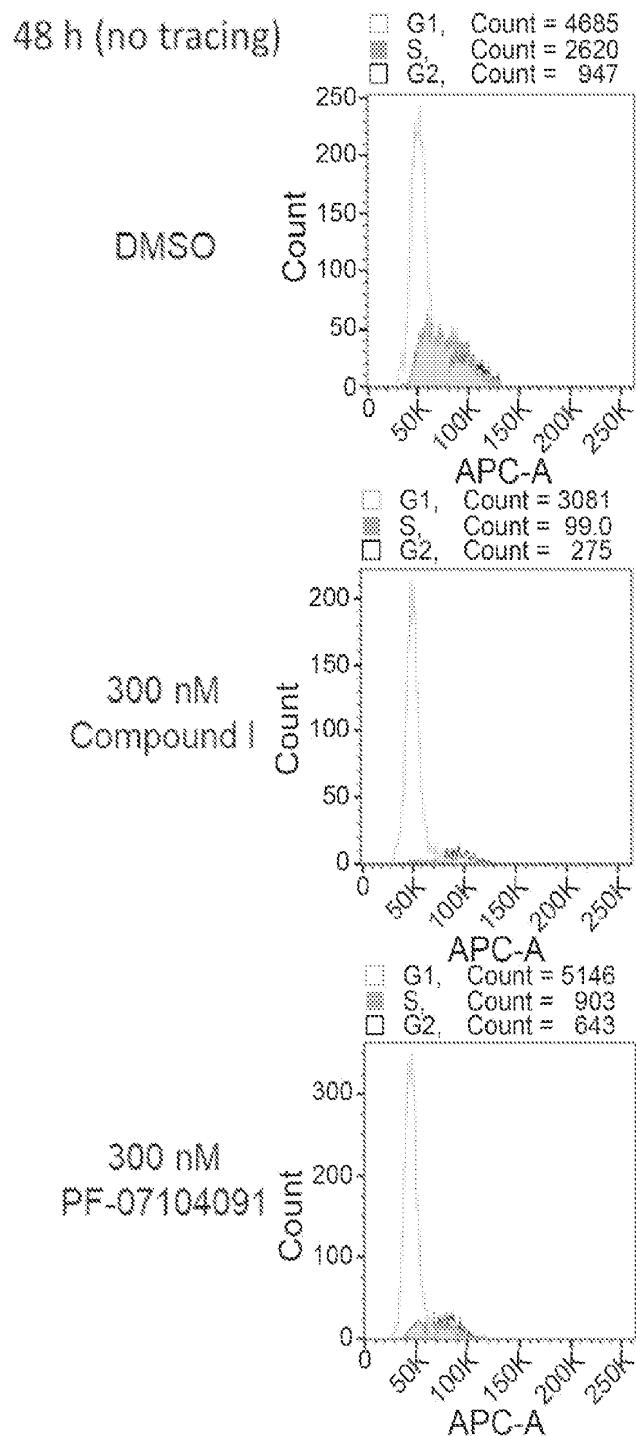


FIG. 15D

Kuramochi 6-Day CTG

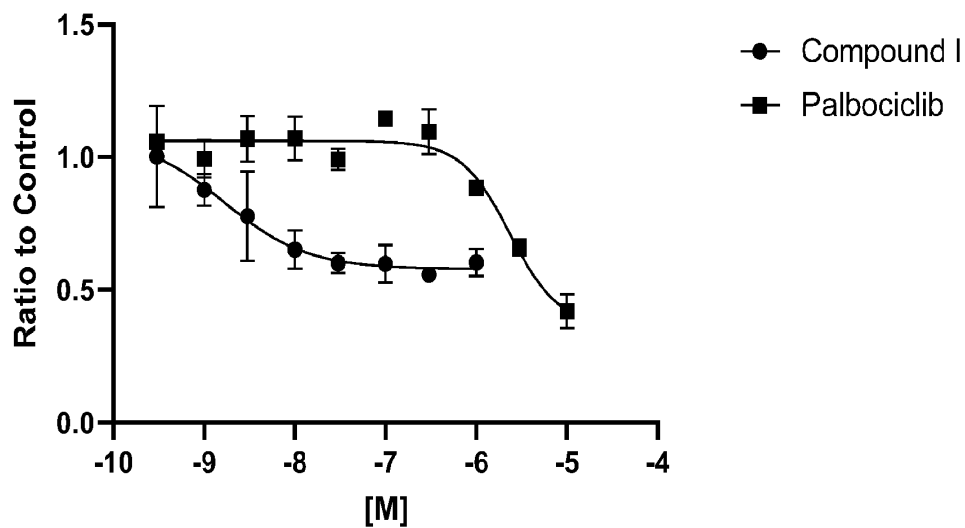


FIG. 16A

FUOV1 6-Day CTG

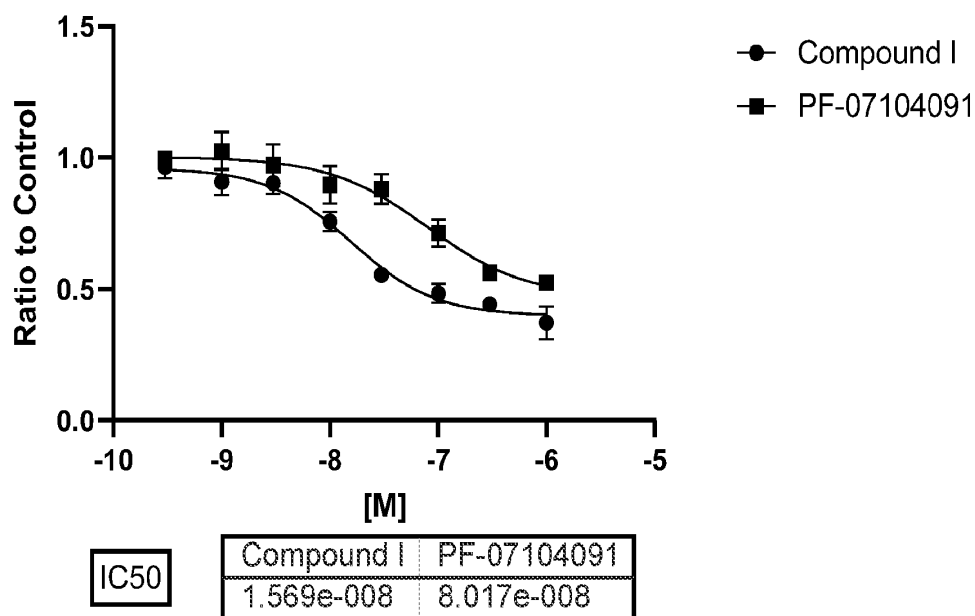


FIG. 16B

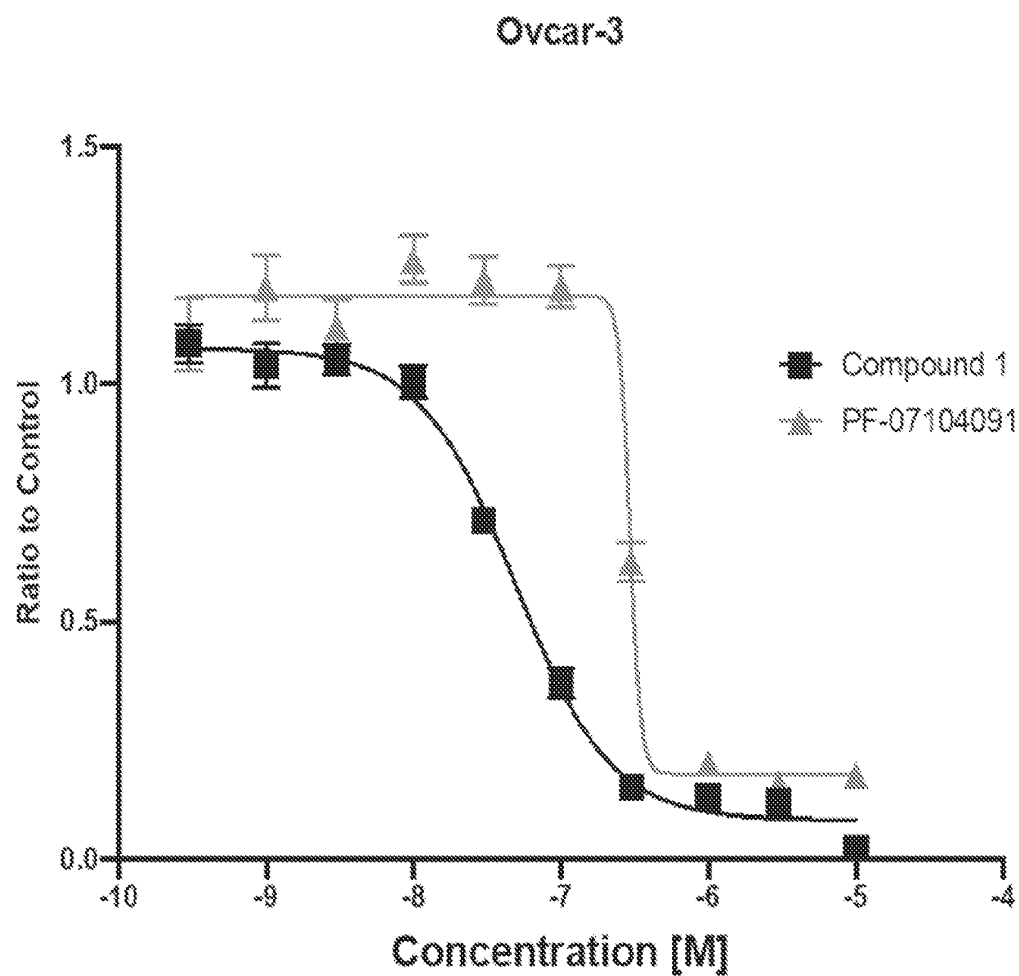


FIG. 17A

Comparison of Compound 1 and PF-07104091	
	OVCAR-1 IC ₅₀
Compound 1	49 nM
PF-07104091	294 nM

FIG. 17B

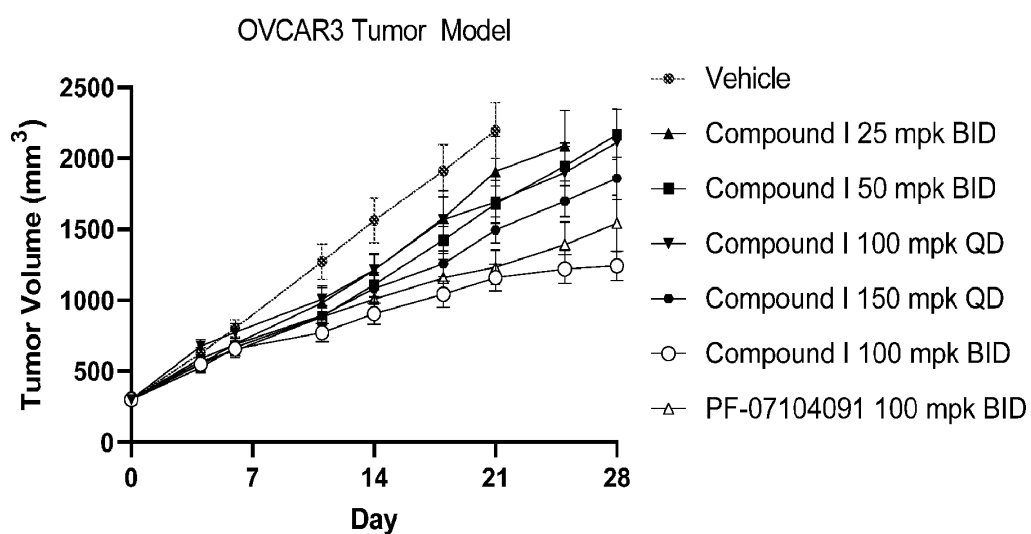


FIG. 18

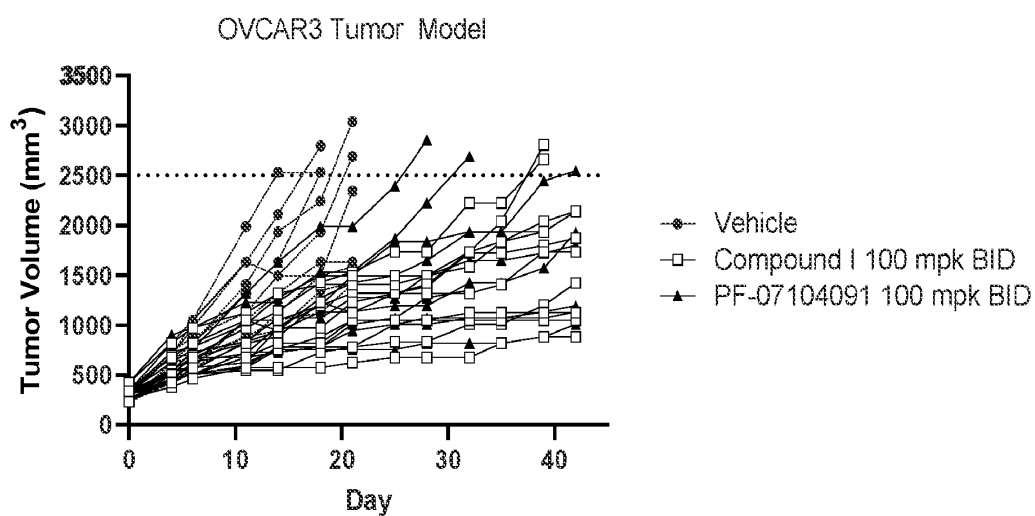


FIG. 19A

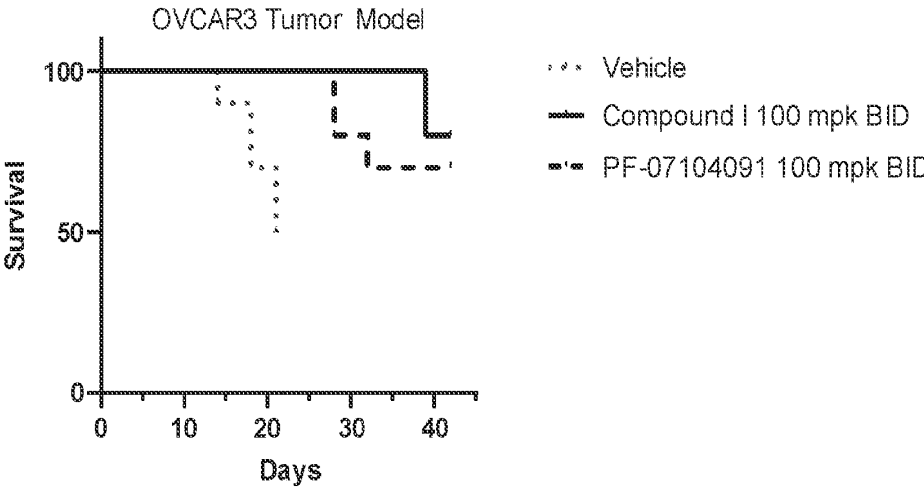


FIG. 19B

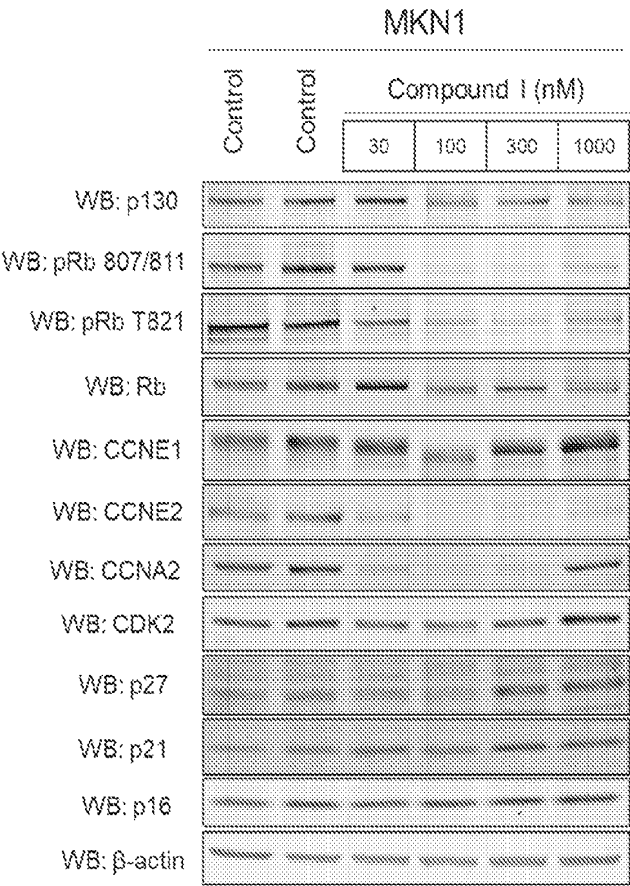
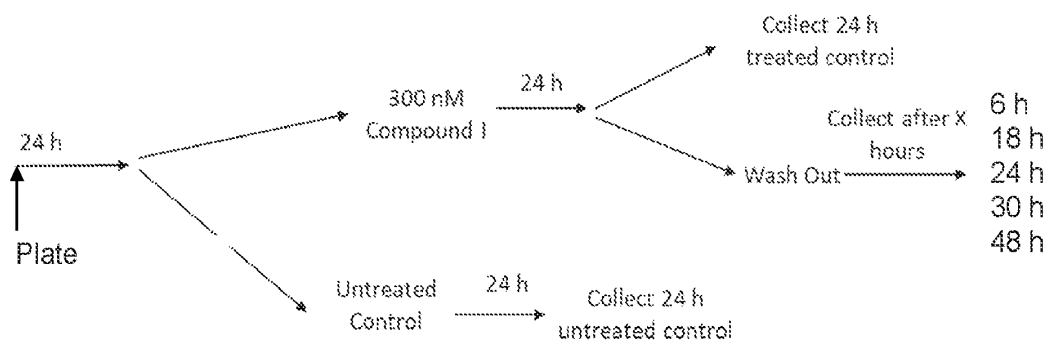


FIG. 20



Compound I (CI)

FIG. 21A

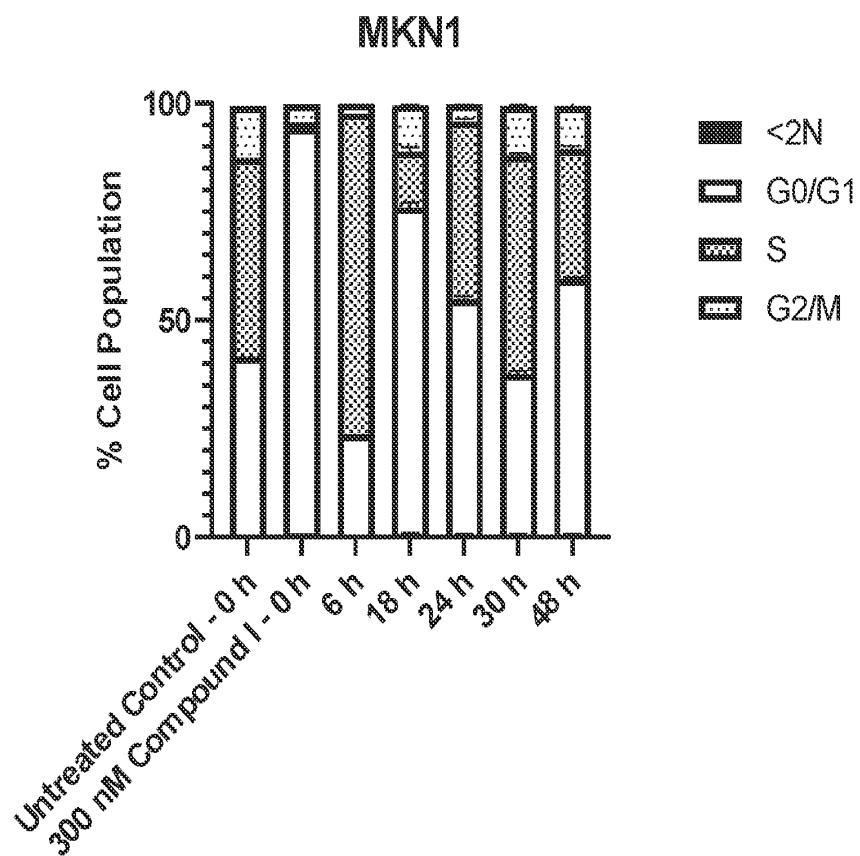


FIG. 21B

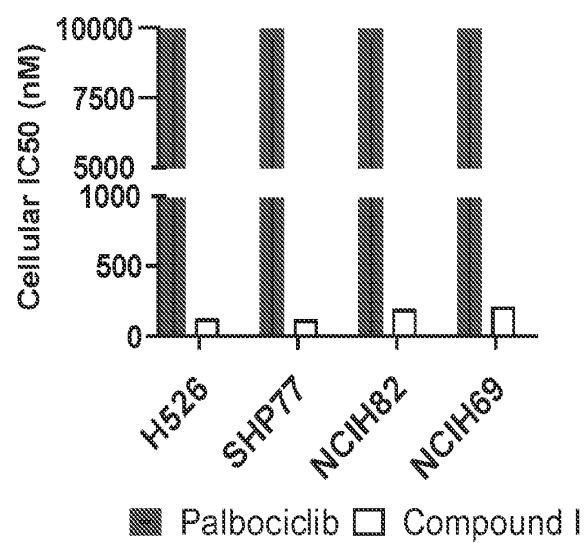


FIG. 22A

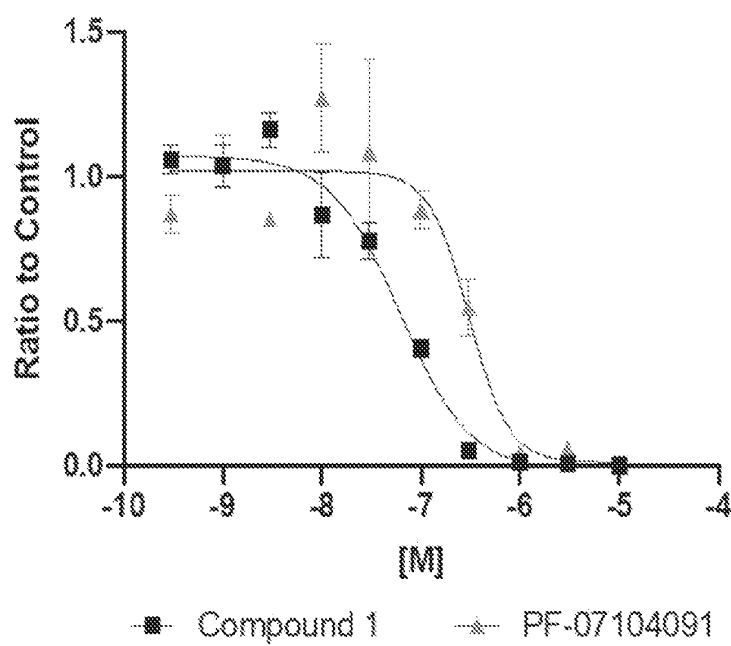
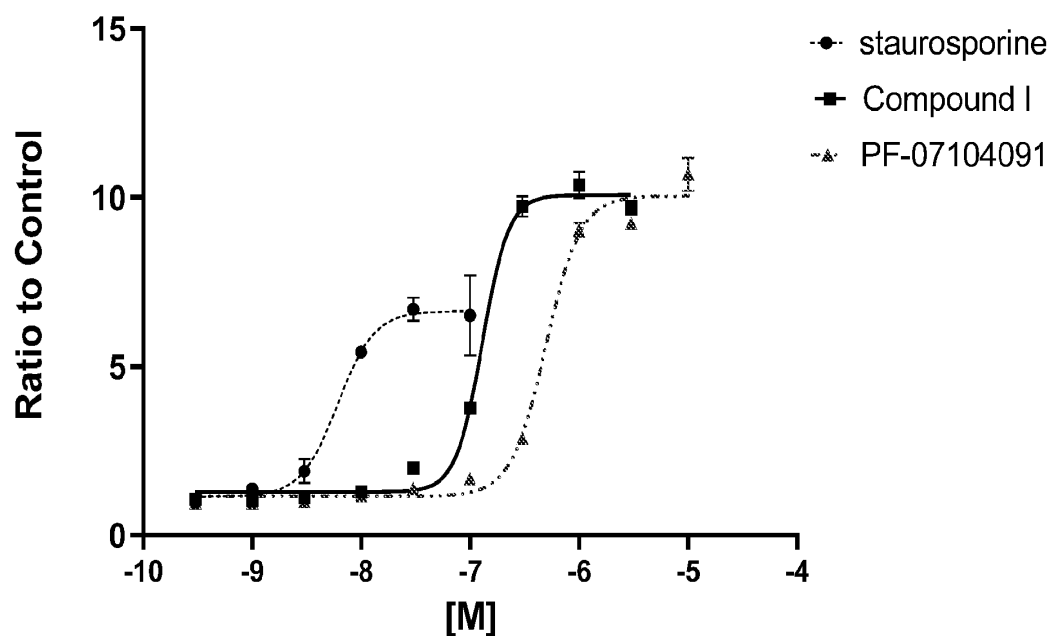


FIG. 22B

H526 IC ₅₀ (nM)	
Compound 1	62nM
PF-07104091	305 nM

FIG. 22C

H526 72hr Caspase 3/7 Activation



	staurosporine	Compound I	PF-07104091
IC50	6.094e-009	1.290e-007	4.950e-007

FIG. 22D

H526 Caspase IC ₅₀ (nM)	
Staurosporine	6 nM
Compound I	129 nM
PF-07104091	495 nM

FIG. 22E

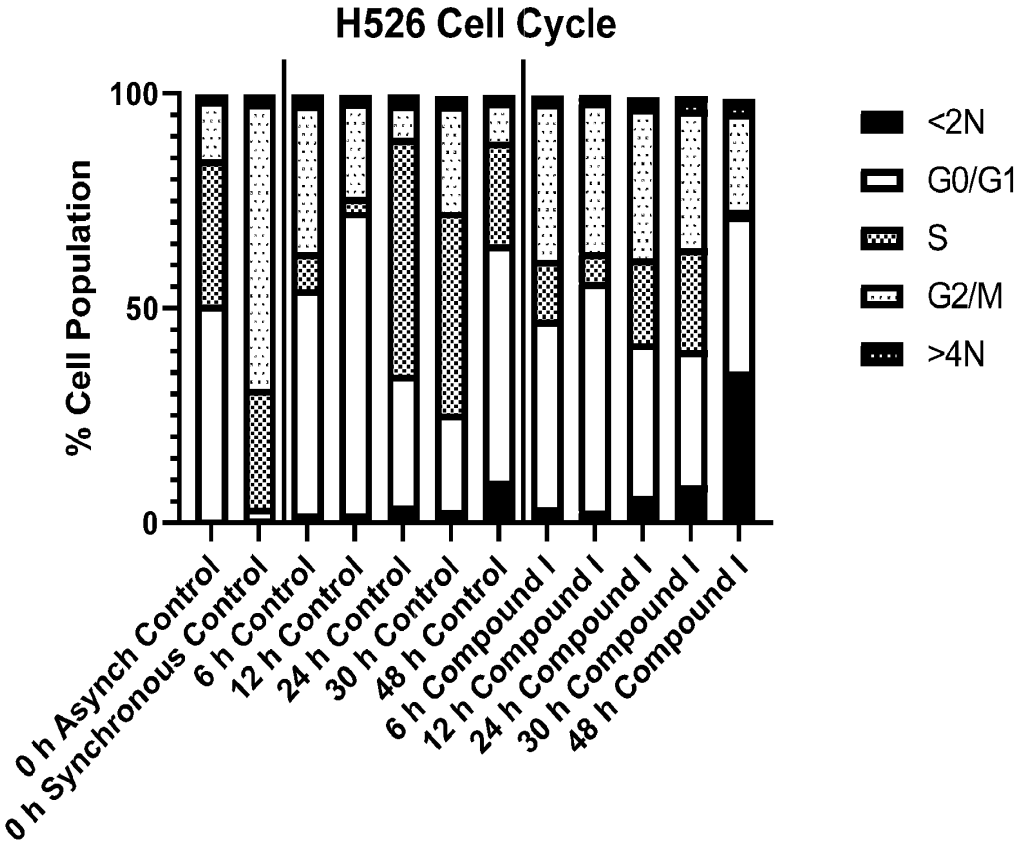


FIG. 23

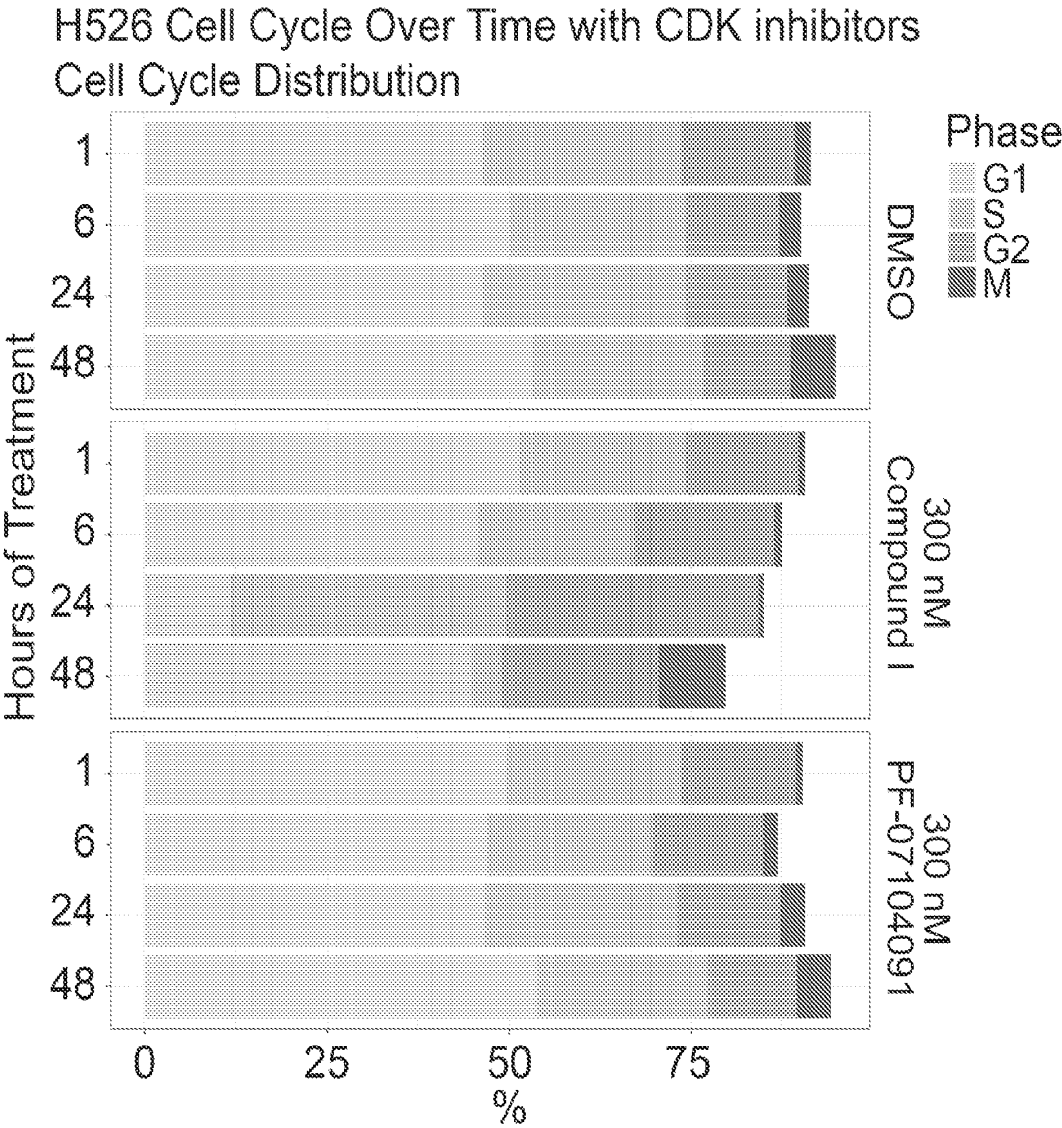


FIG. 24

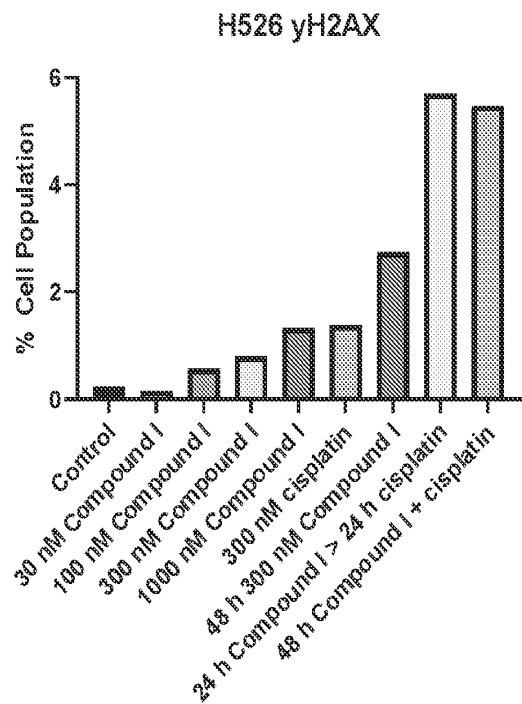


FIG. 25

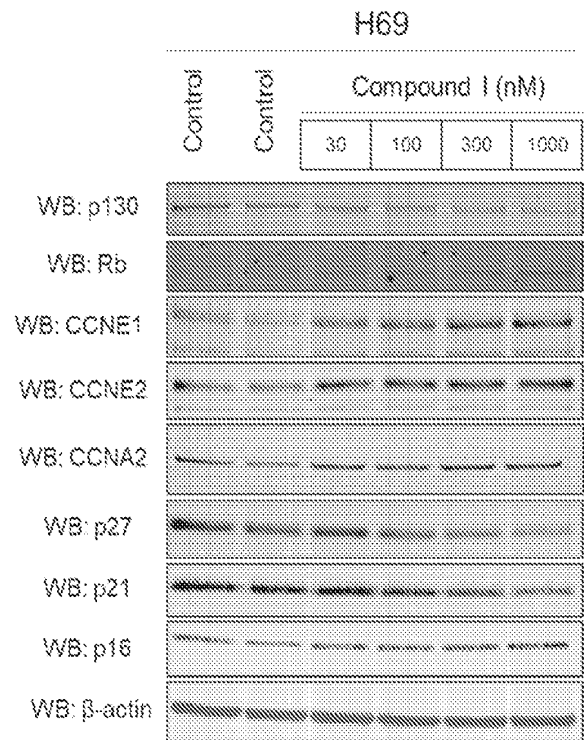


FIG. 26A

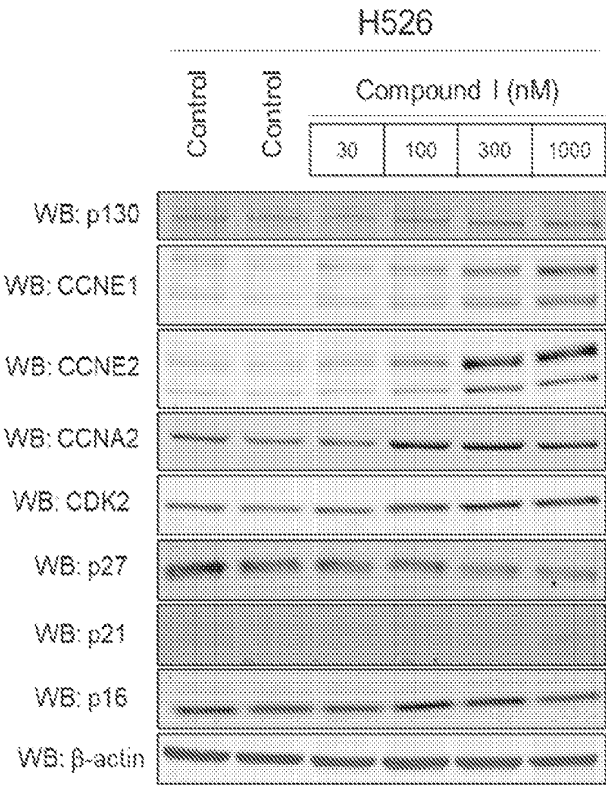


FIG. 26B

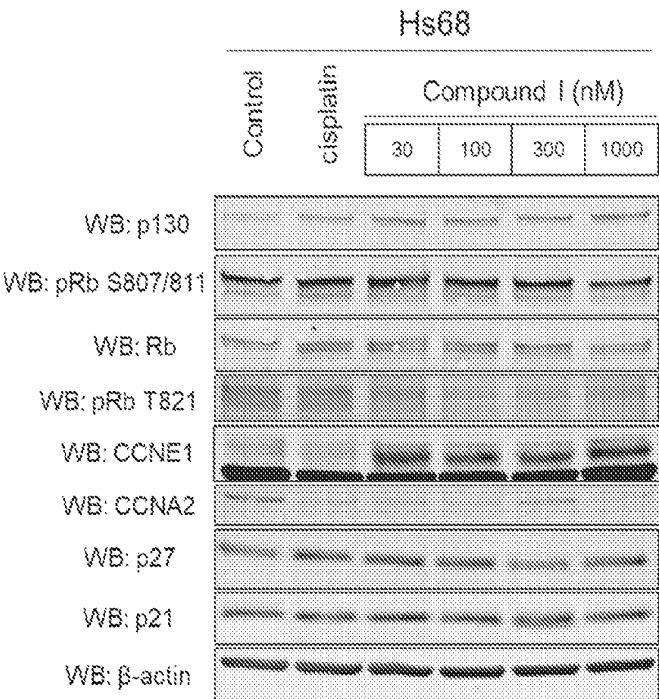
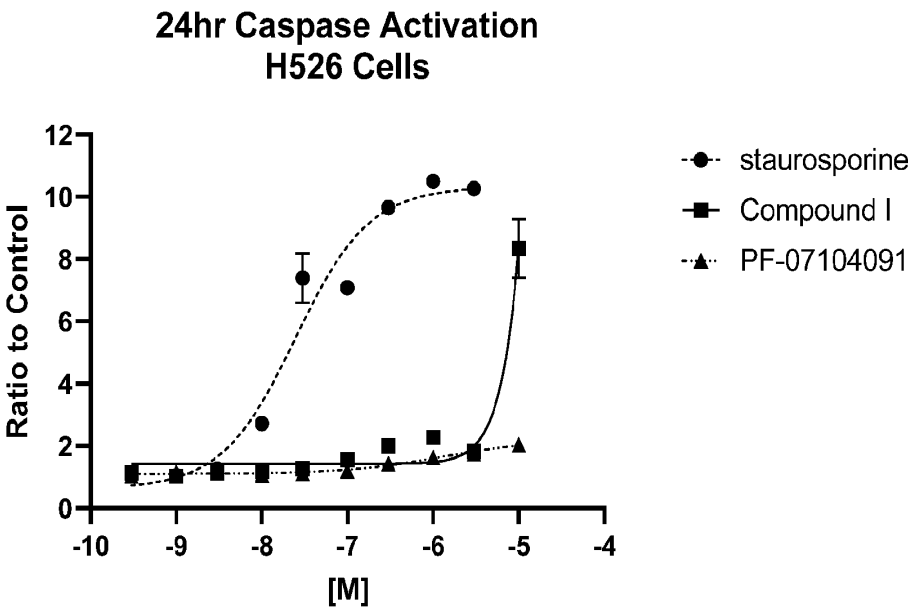
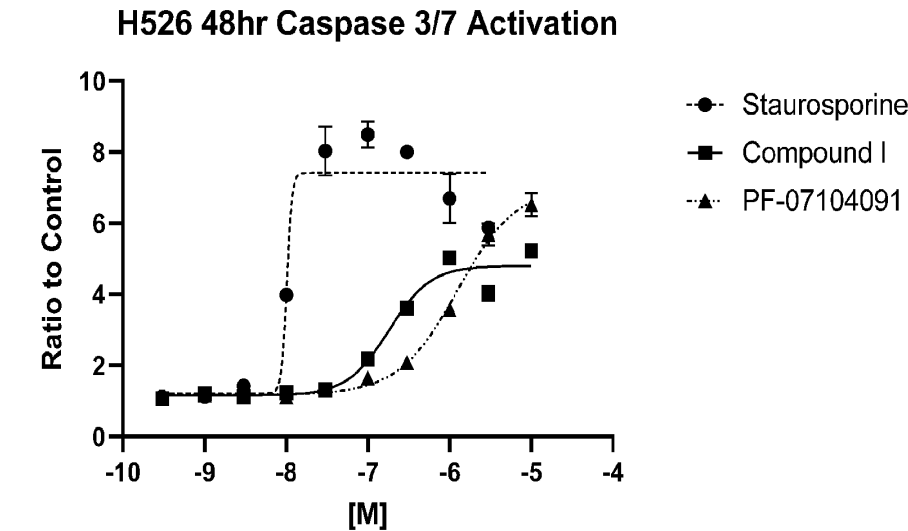


FIG. 26C



	staurosporine	Compound I	PF-07104091
IC50	2.481e-008	~ 0.0001657	1.287e-006

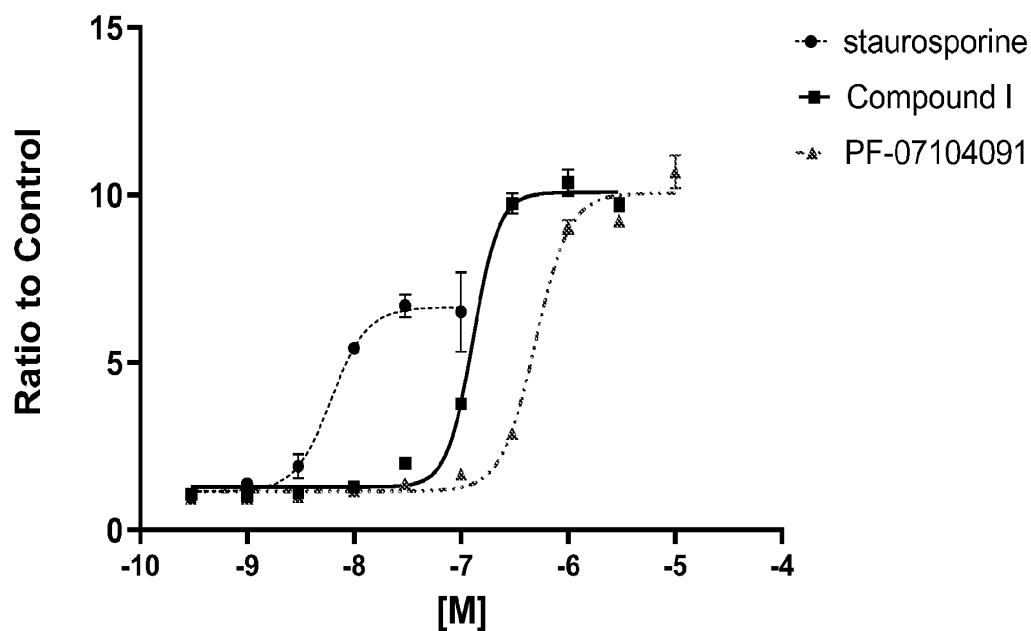
FIG. 27A



	Staurosporine	Compound I	PF-07104091
IC50	~ 1.016e-008	1.809e-007	1.251e-006

FIG. 27B

H526 72hr Caspase 3/7 Activation



	staurosporine	Compound I	PF-07104091
IC50	6.094e-009	1.290e-007	4.950e-007

FIG. 27C

H526 Caspase IC ₅₀ (nM)	
Staurosporine	6 nM
Compound 1	129 nM
PF-07104091	495 nM

FIG. 27D

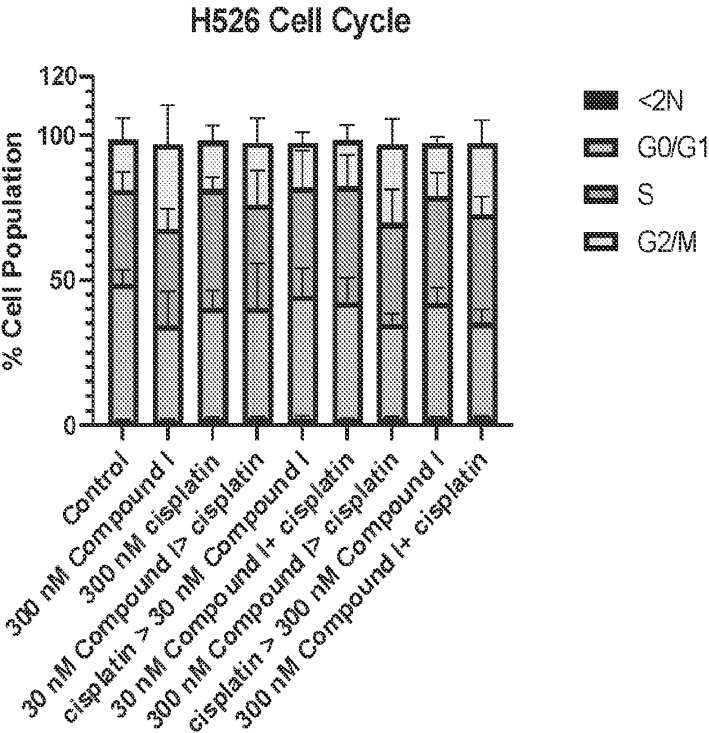
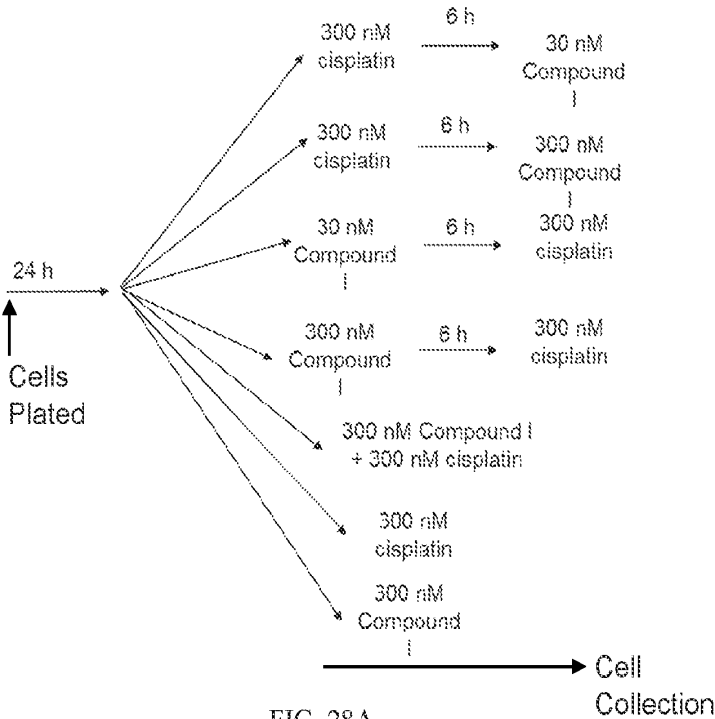


FIG. 28B

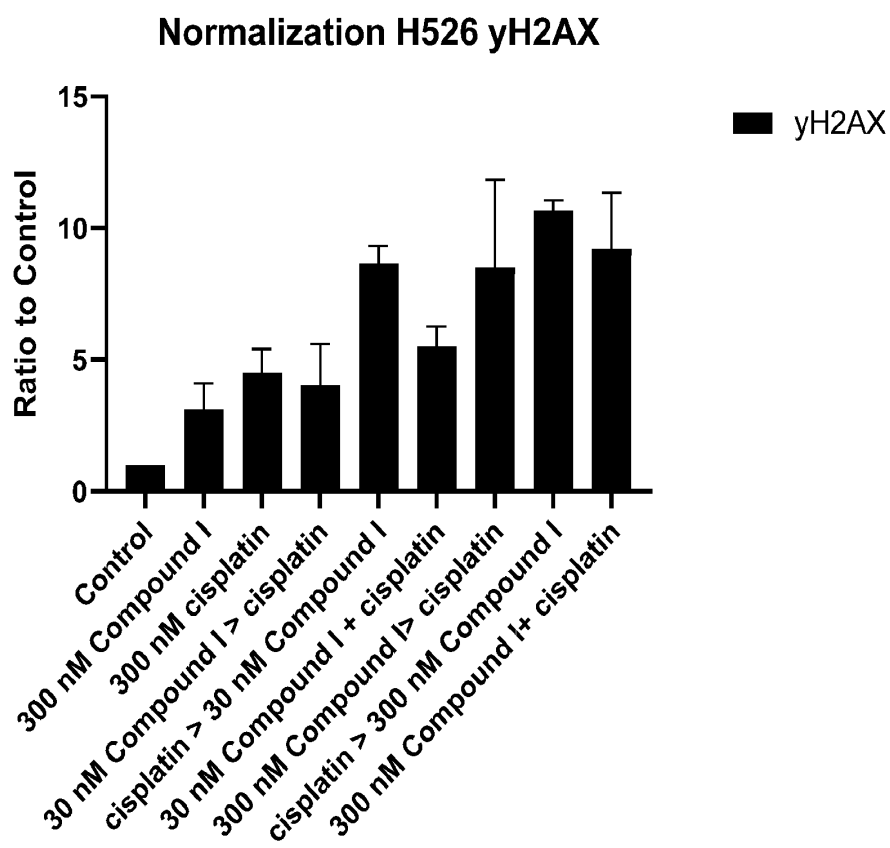


FIG. 28C

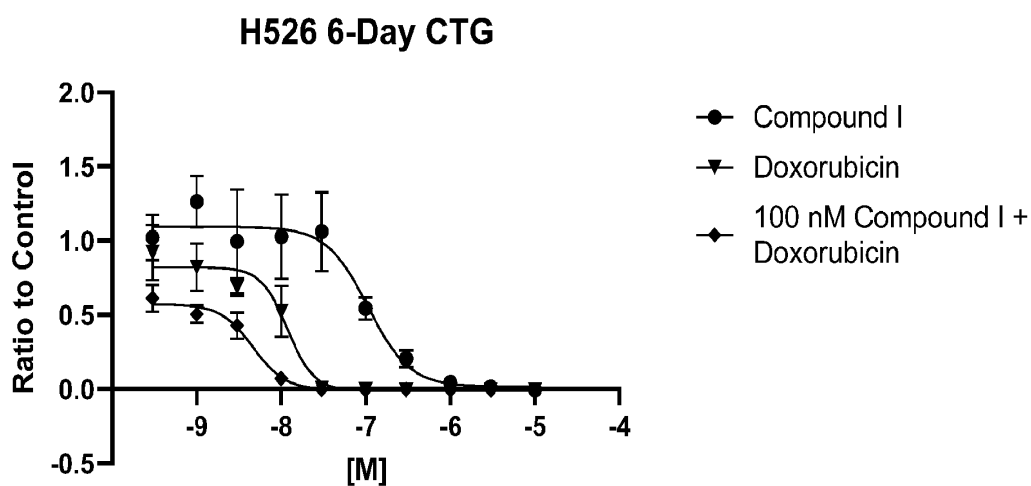


FIG. 29A

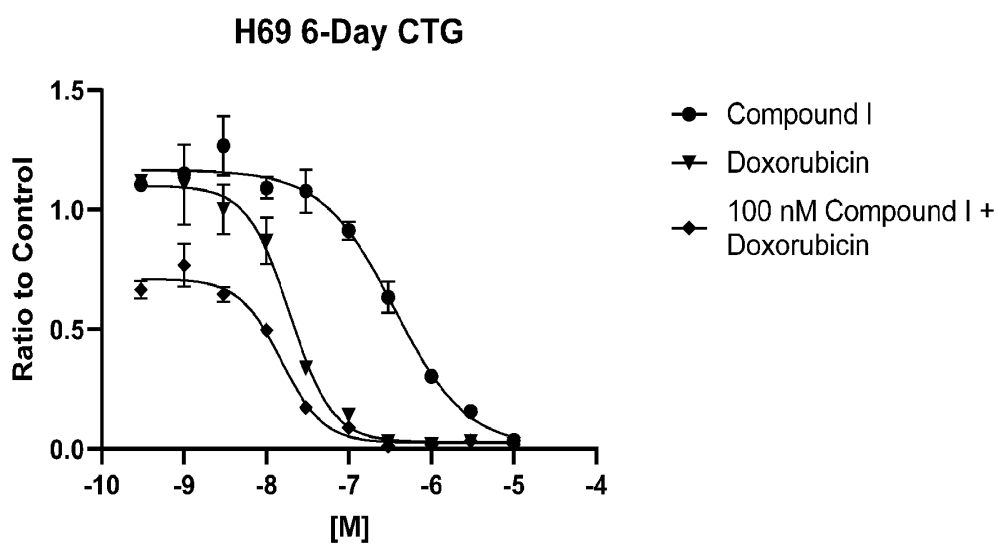


FIG. 29B

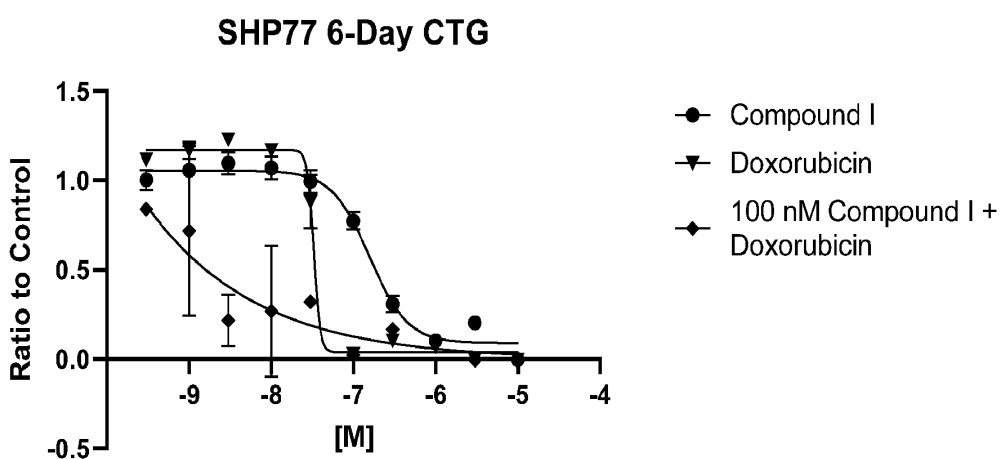


FIG. 29C

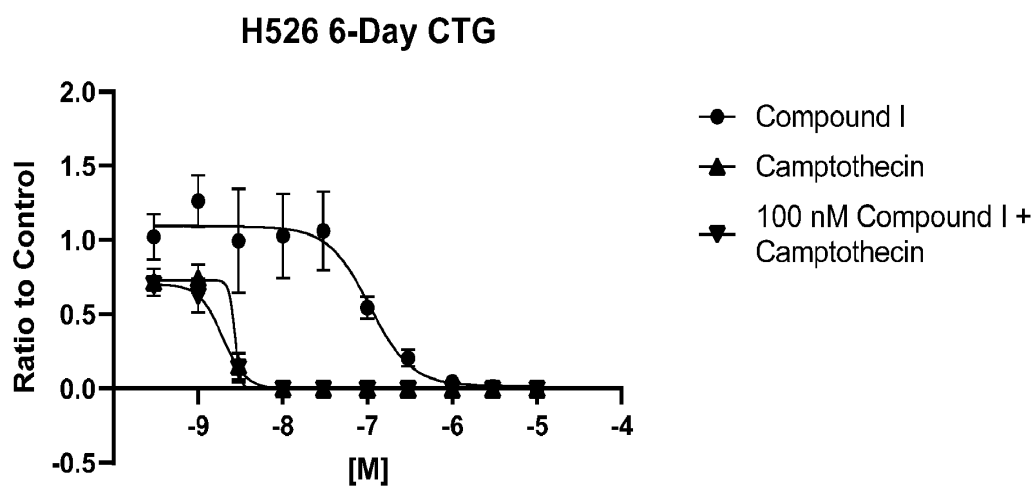


FIG. 30A

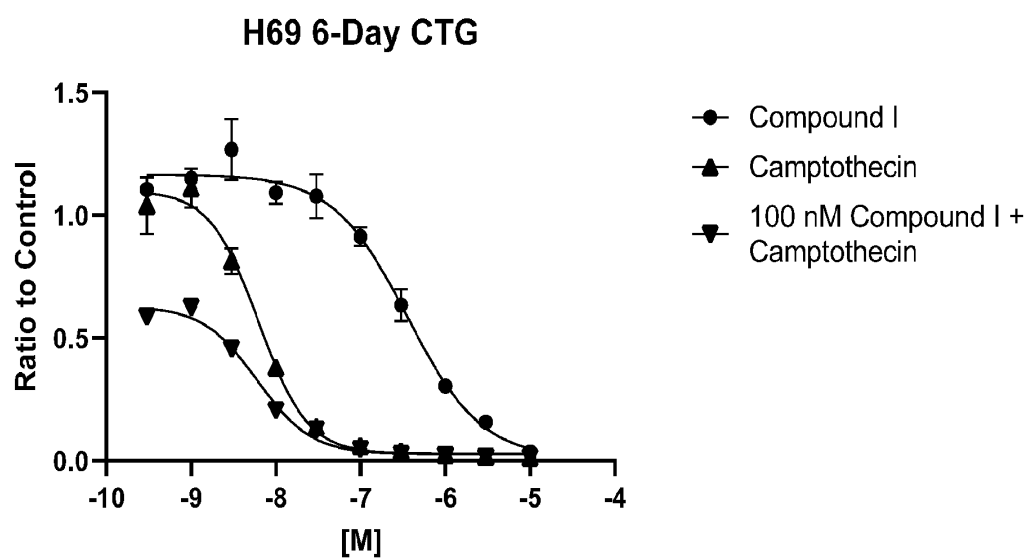


FIG. 30B

SHP77 6-Day CTG

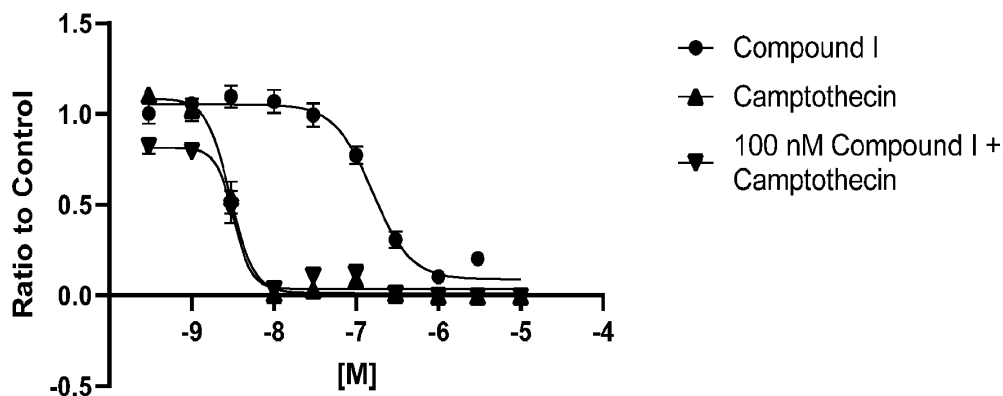
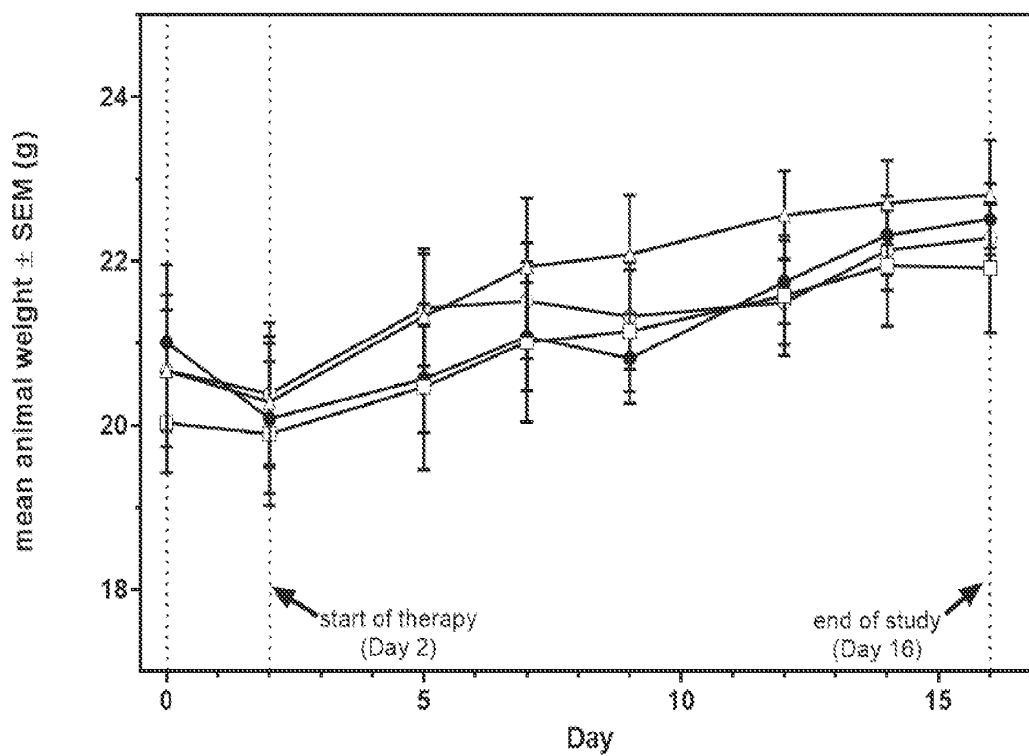


FIG. 30C



- ◆ Group 1: Vehicle (10 ml/kg) 2x daily p.o. (n=6/6)
- ◇ Group 2: Compound I conc.1 (75 mg/kg, 10 ml/kg) 2x daily p.o. (n=6/6)
- Group 3: Compound I conc.2 (100 mg/kg, 10 ml/kg) 2x daily p.o. (n=6/6)
- △ Group 4: Compound I conc.3 (150 mg/kg, 10 ml/kg) 2x daily p.o. (n=6/6)

FIG. 31A

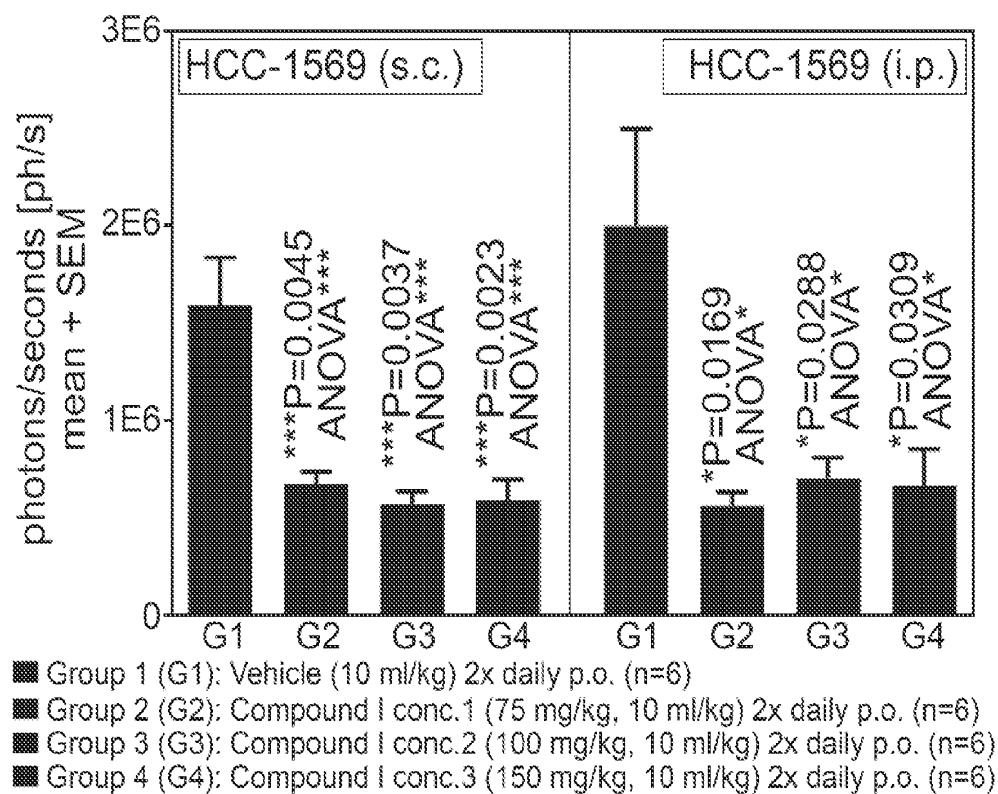


FIG. 31B

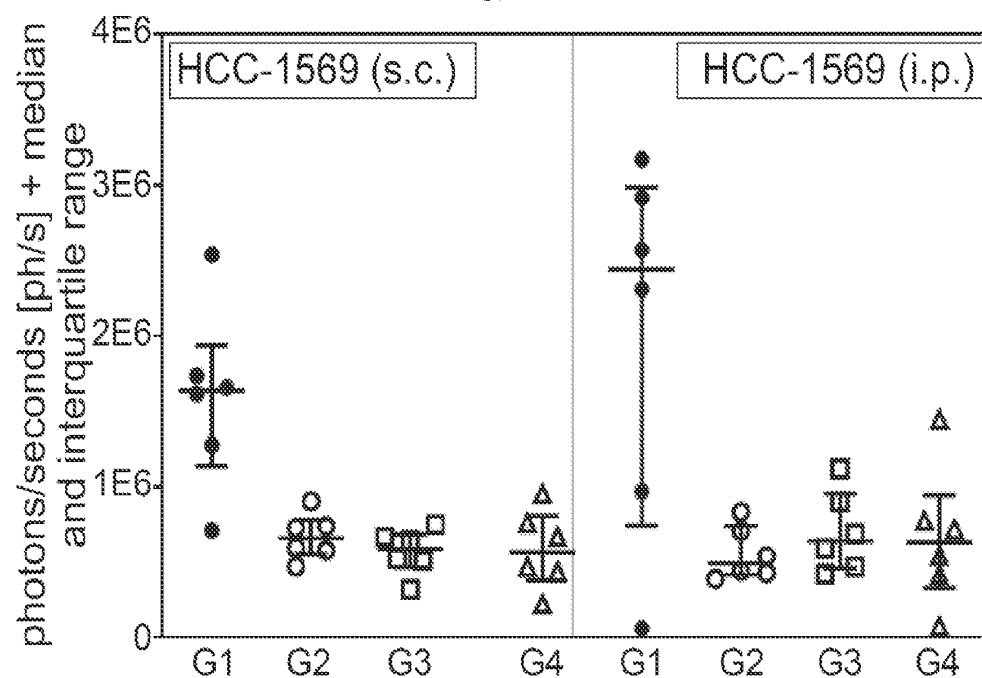


FIG. 31C

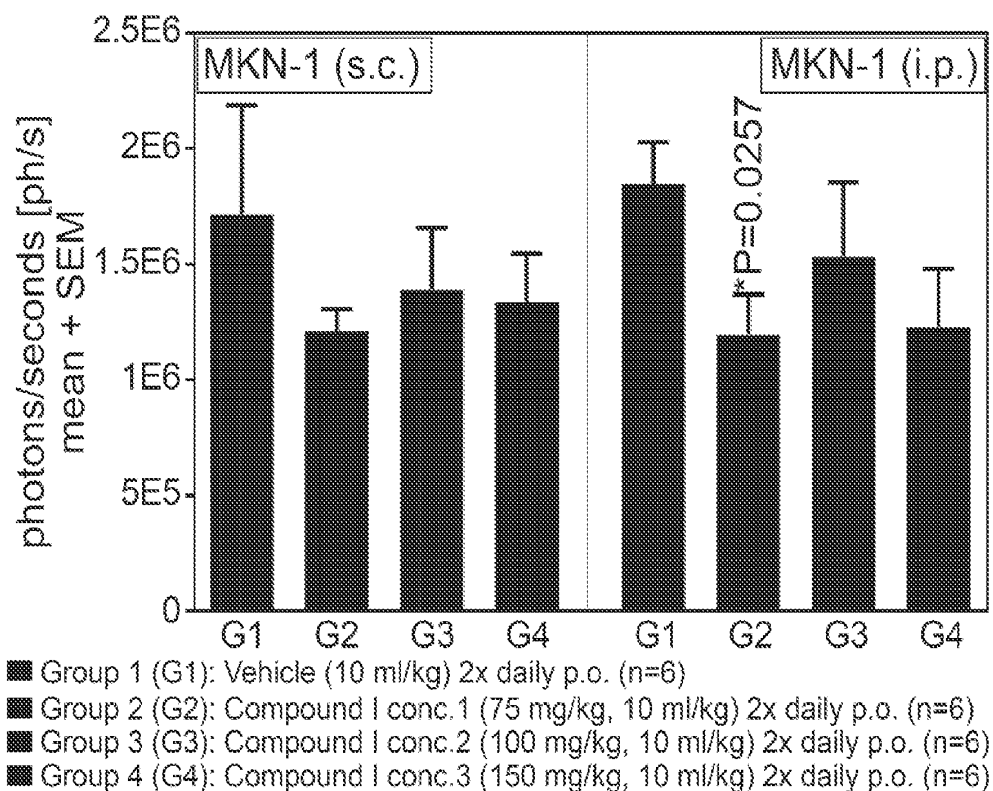


FIG. 31D

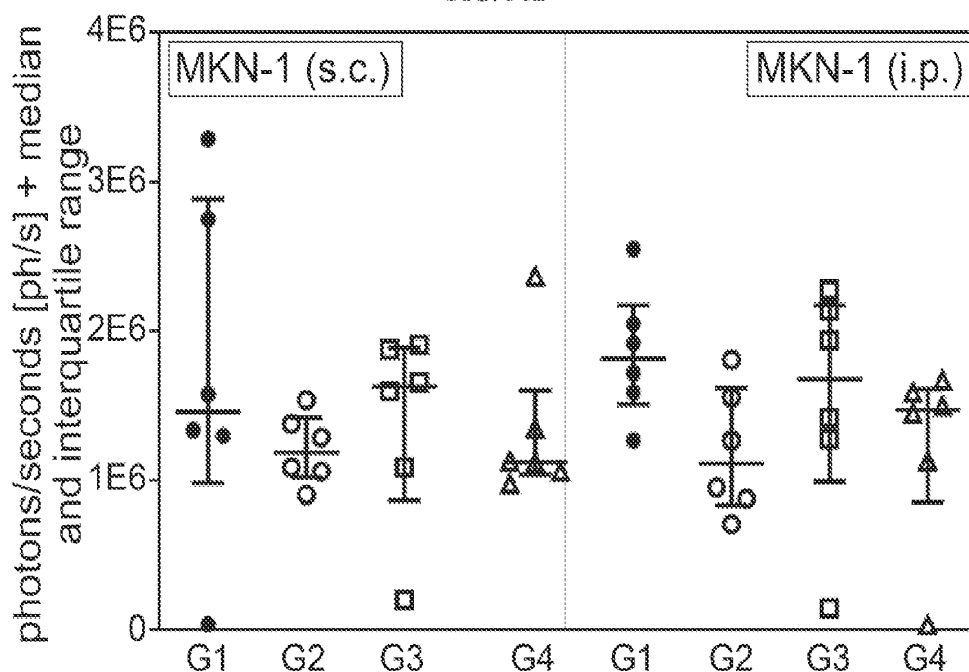


FIG. 31E

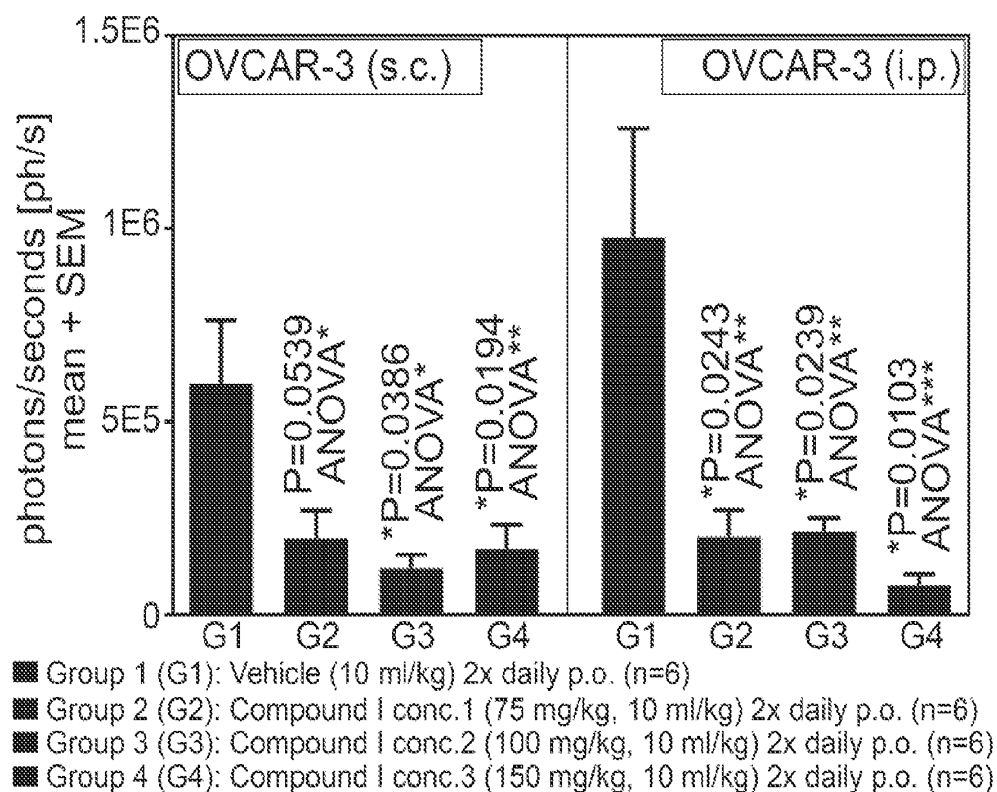


FIG. 31F

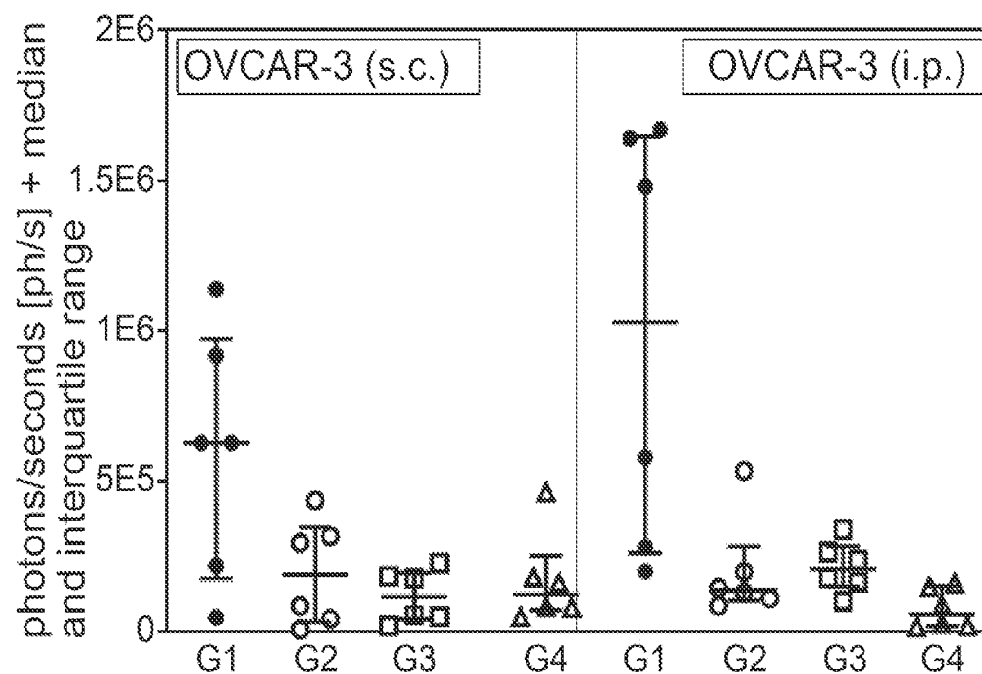


FIG. 31G

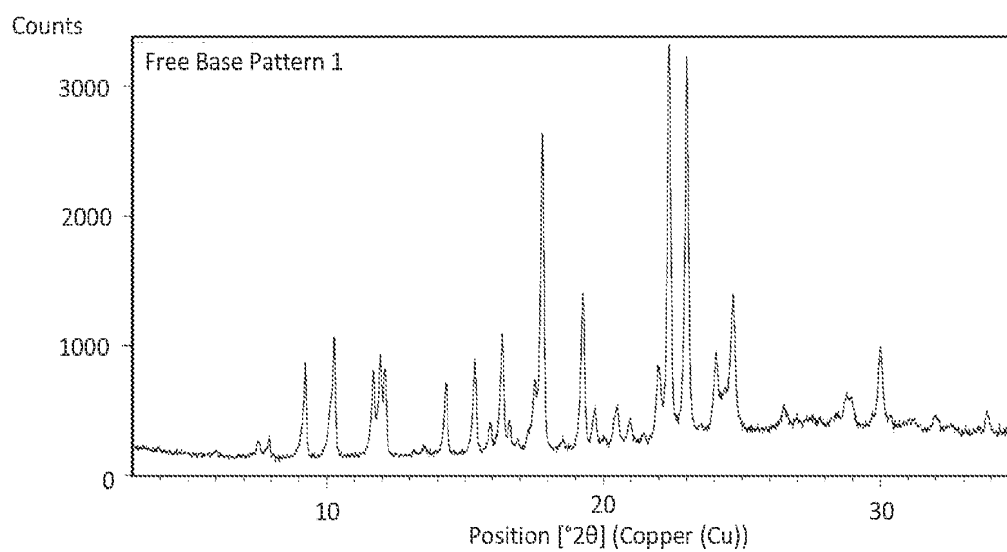


FIG. 32

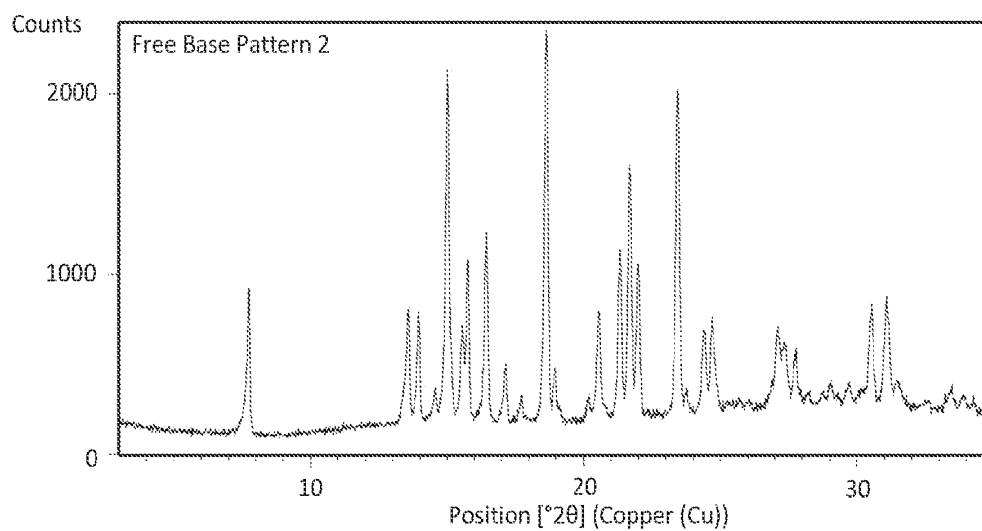


FIG. 33

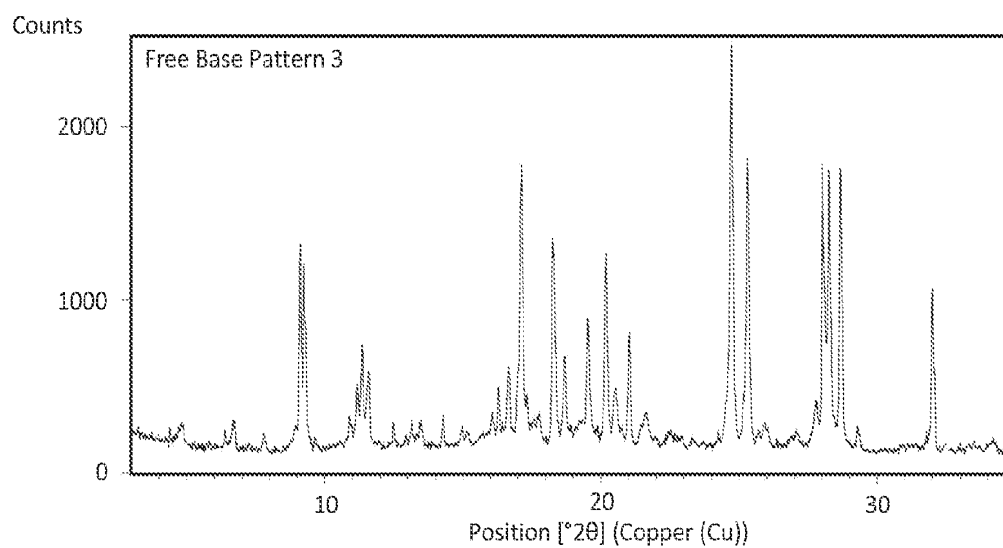


FIG. 34

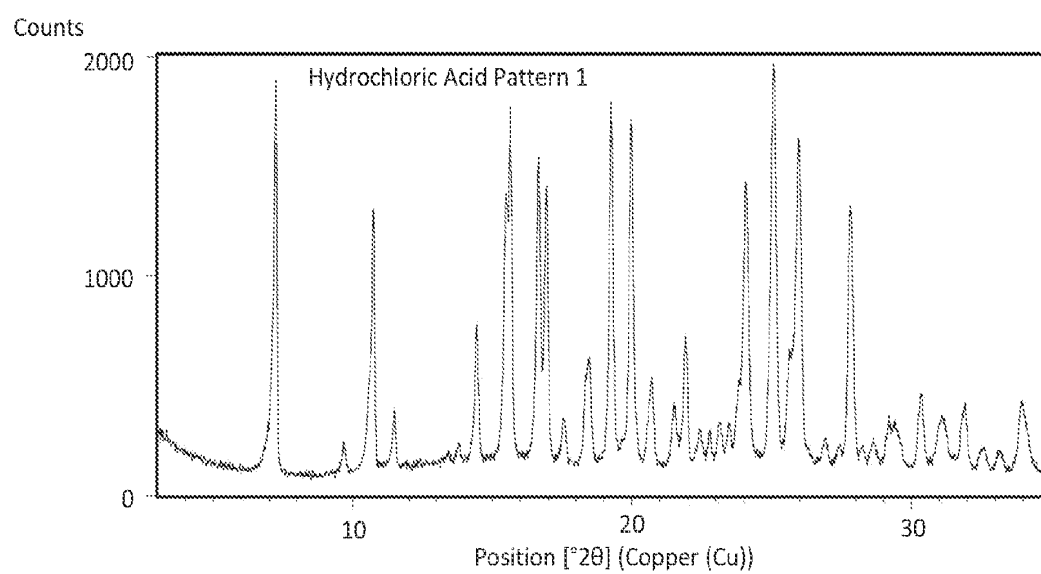


FIG. 35

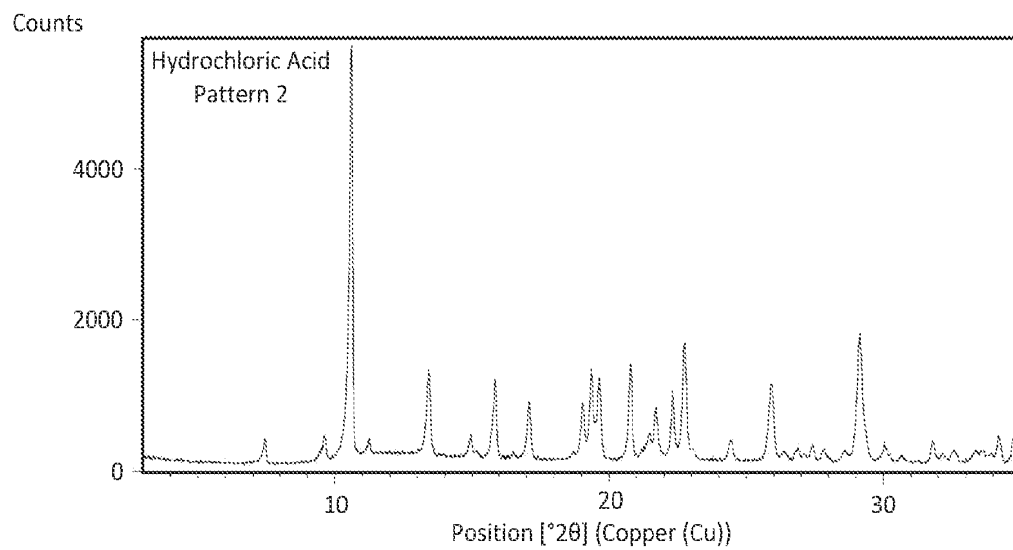


FIG. 36

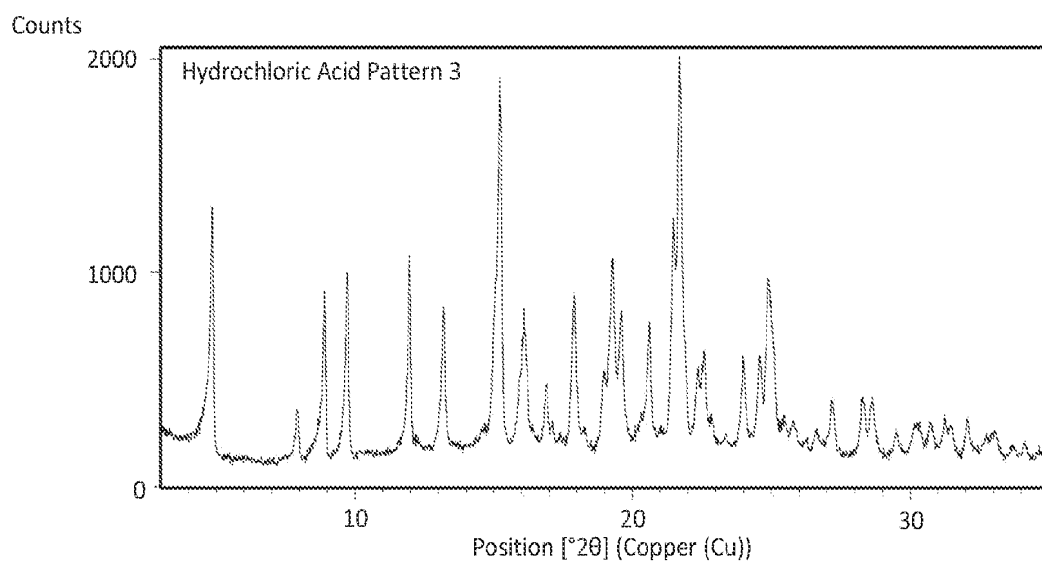


FIG. 37

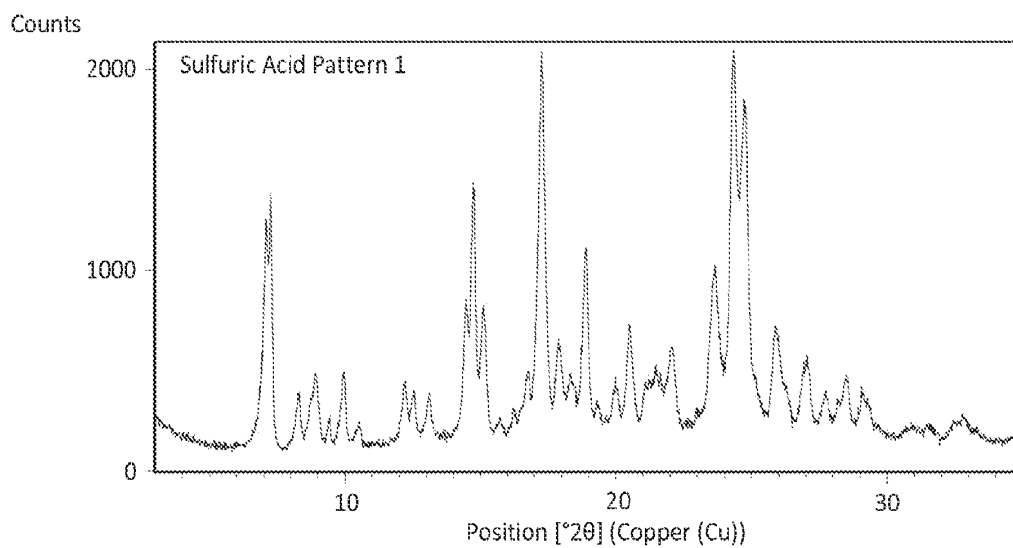


FIG. 38

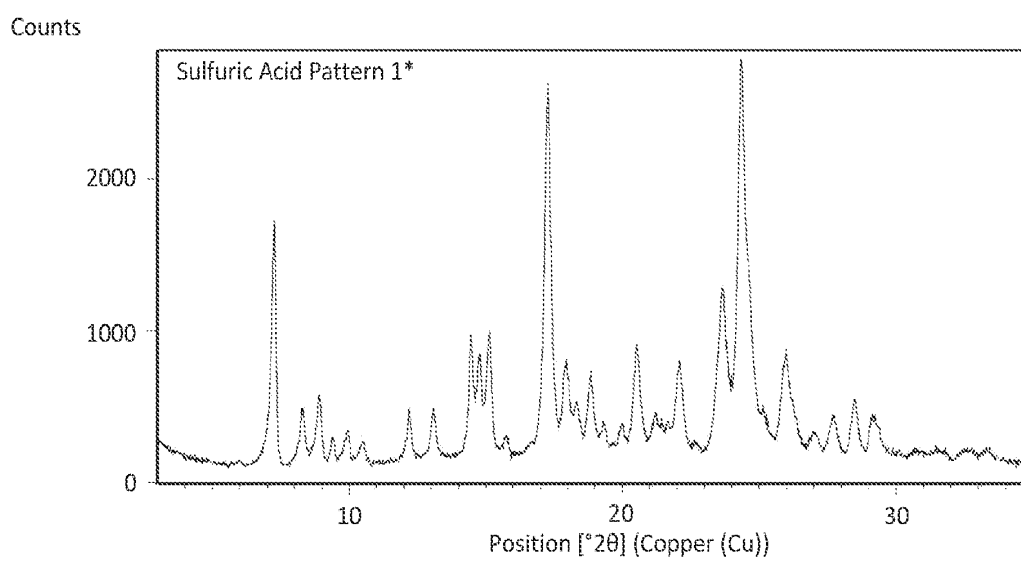


FIG. 39

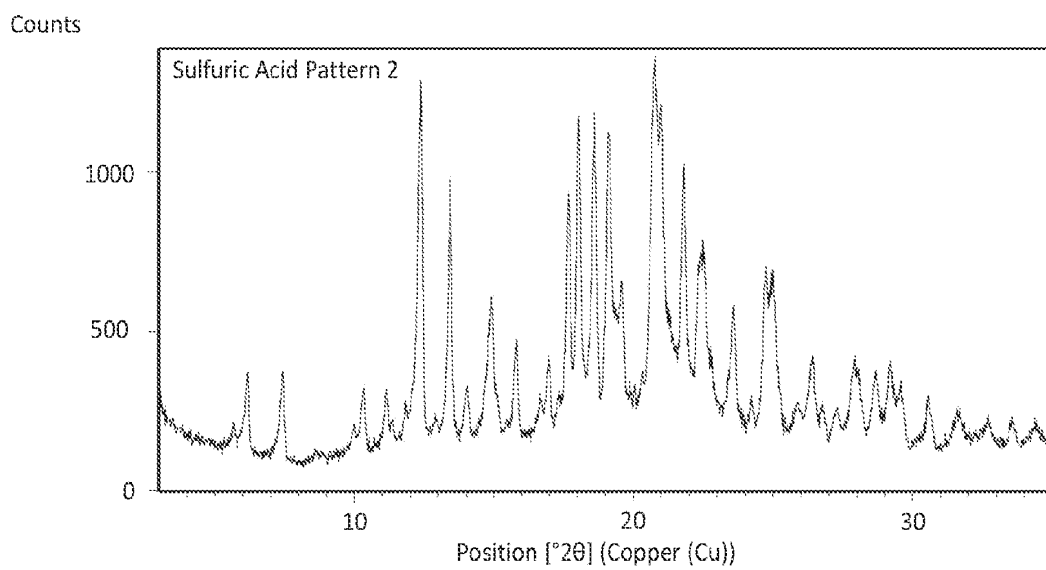


FIG. 40

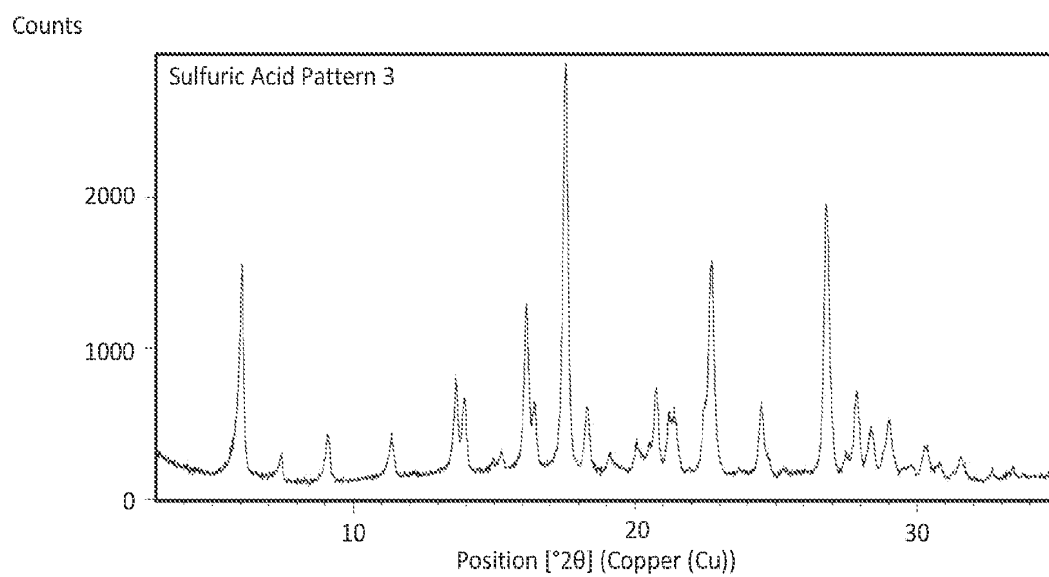


FIG. 41

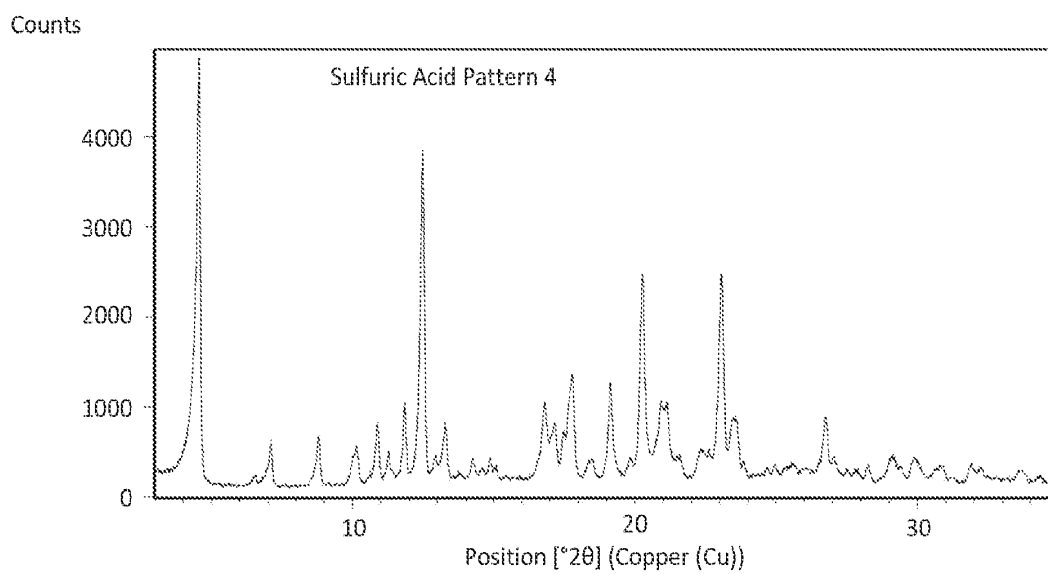


FIG. 42

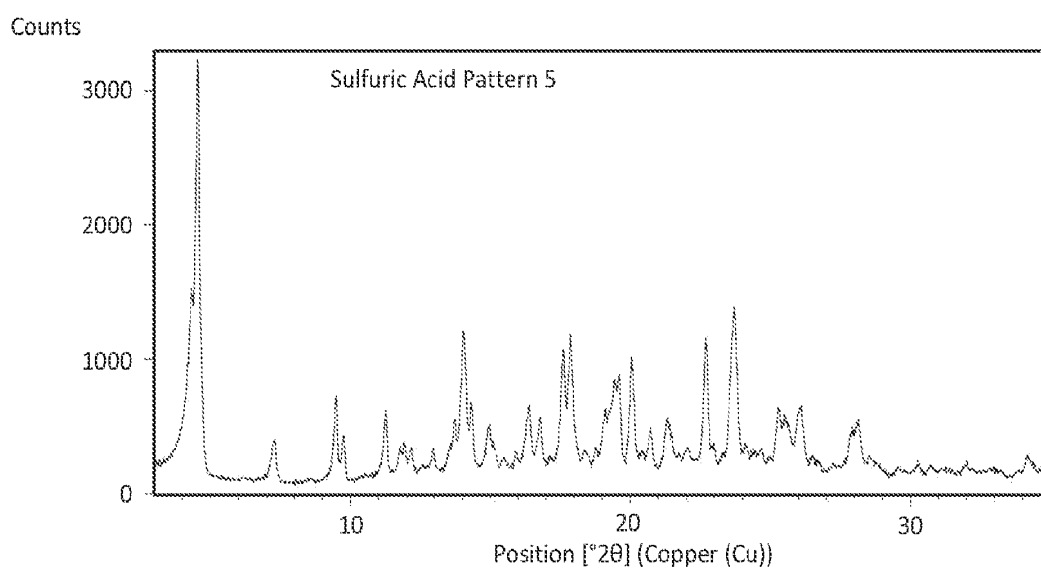


FIG. 43

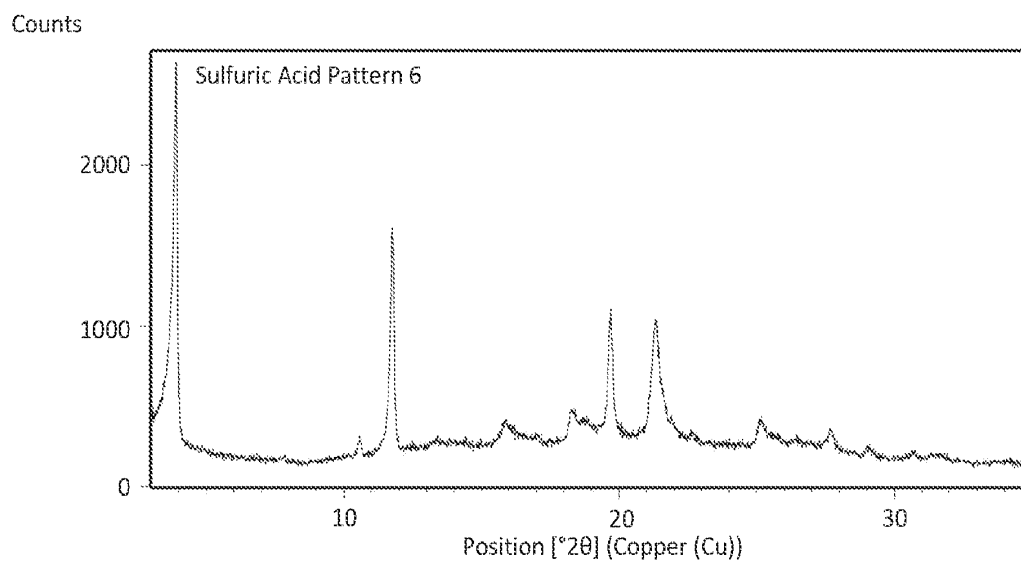


FIG. 44

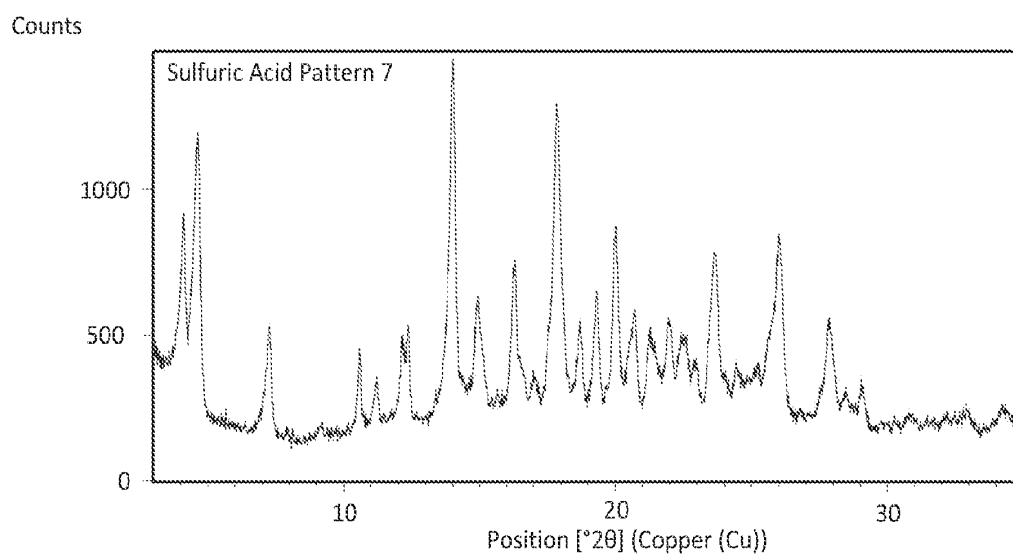


FIG. 45

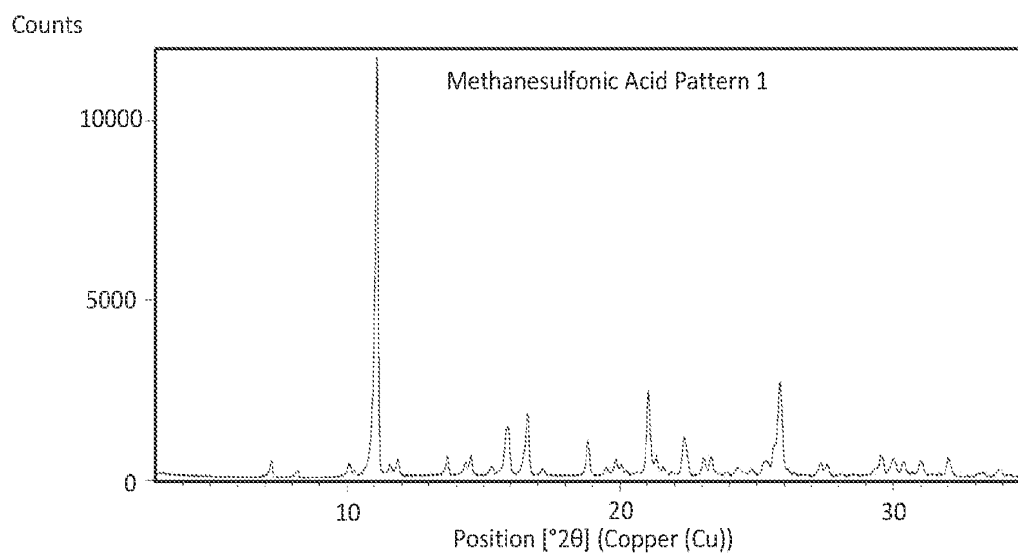


FIG. 46

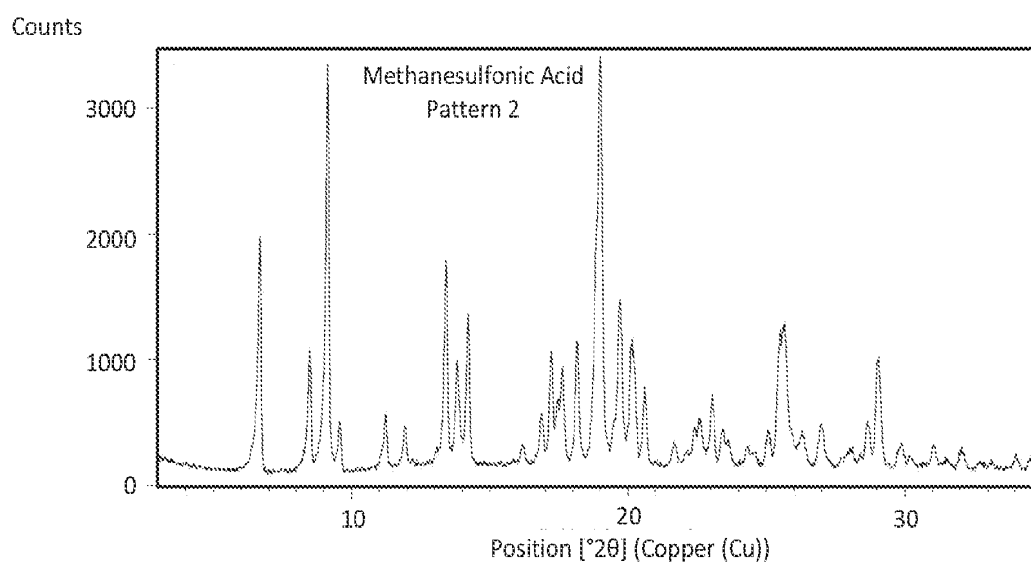


FIG. 47

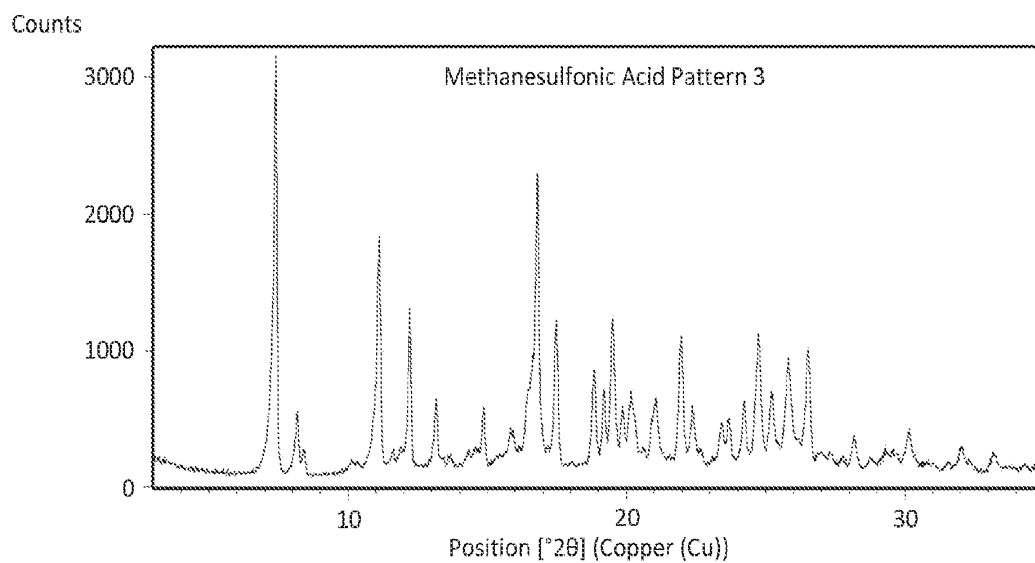


FIG. 48

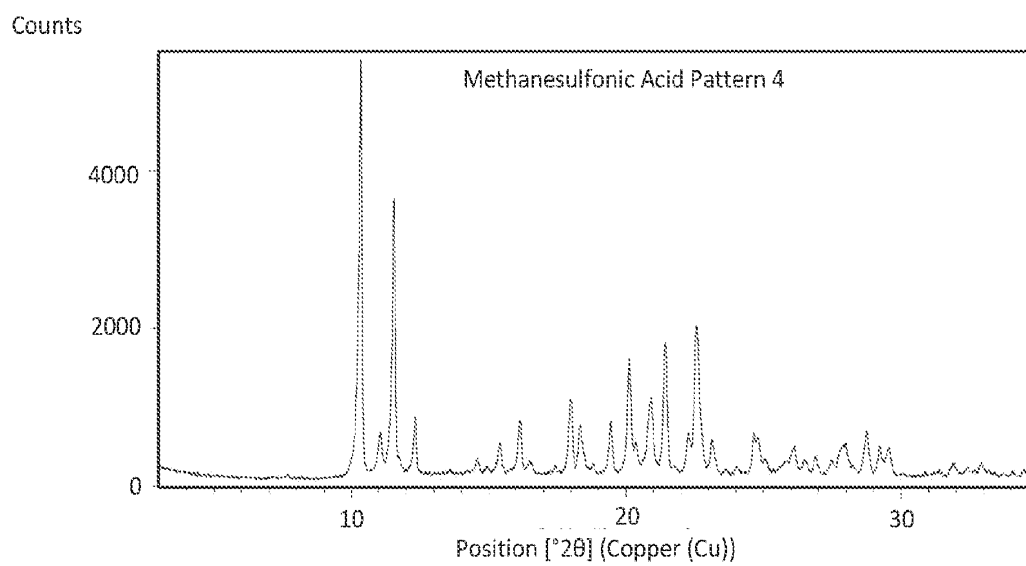


FIG. 49

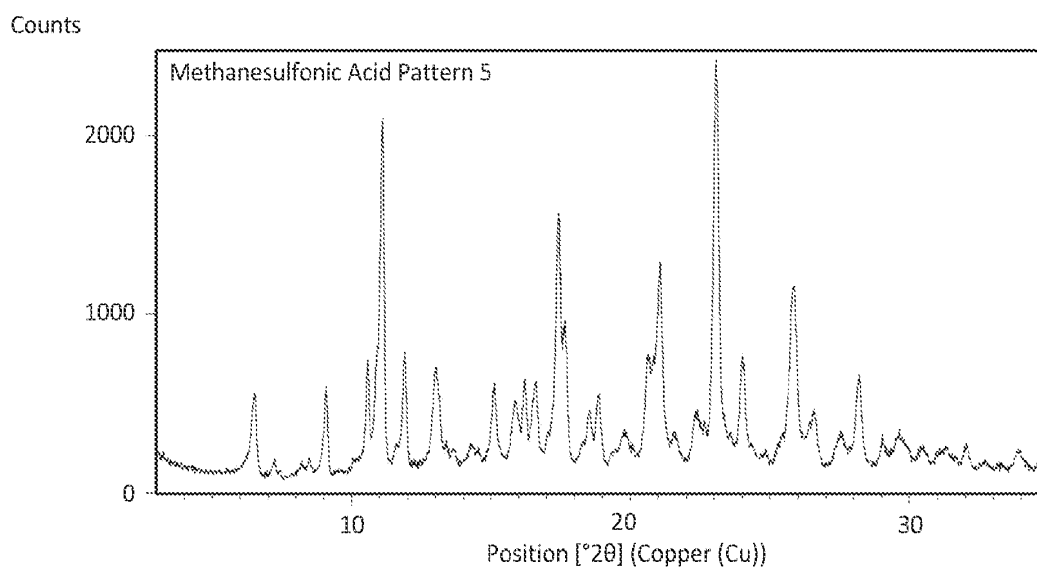


FIG. 50

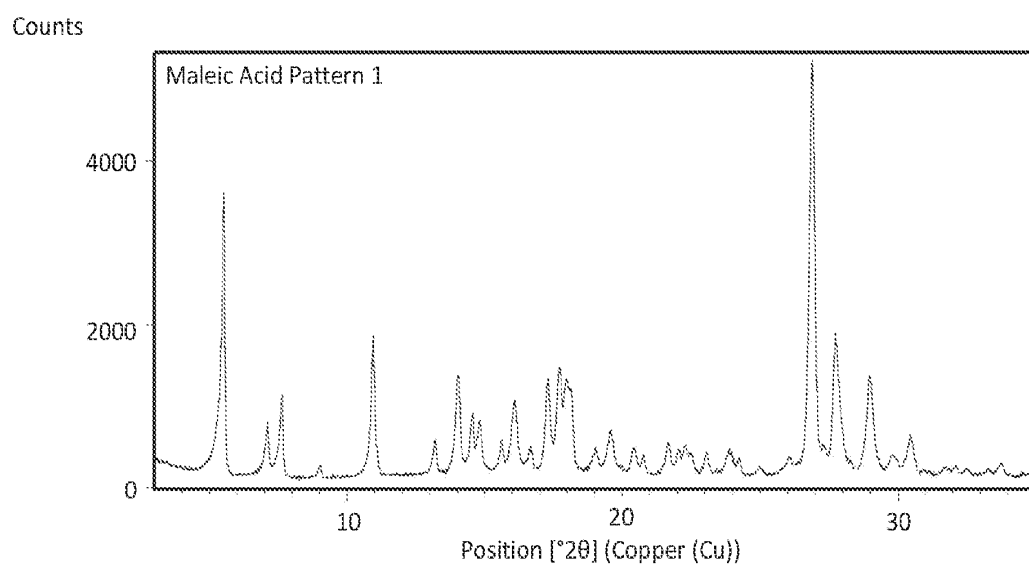


FIG. 51

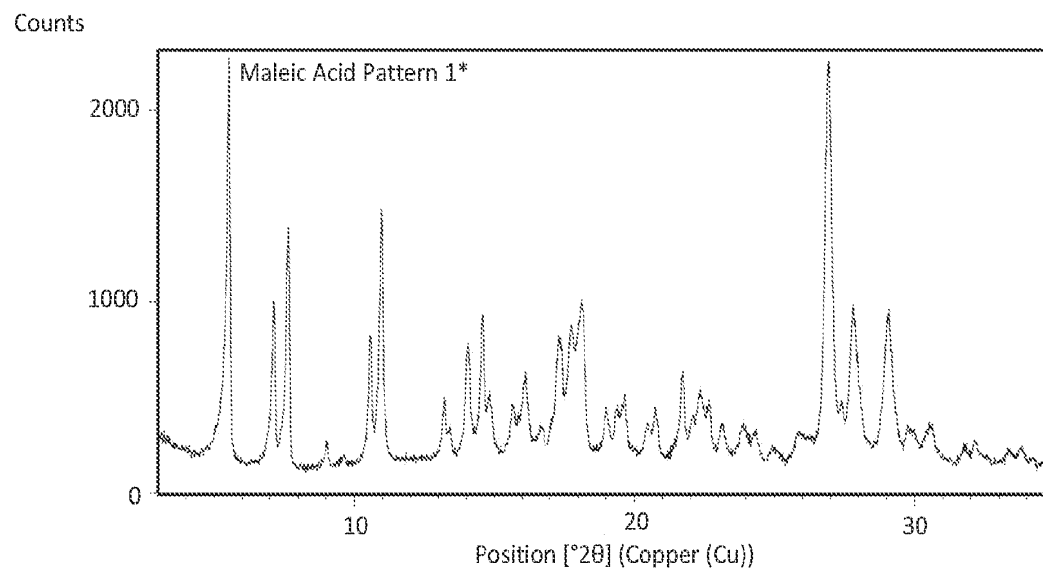


FIG. 52

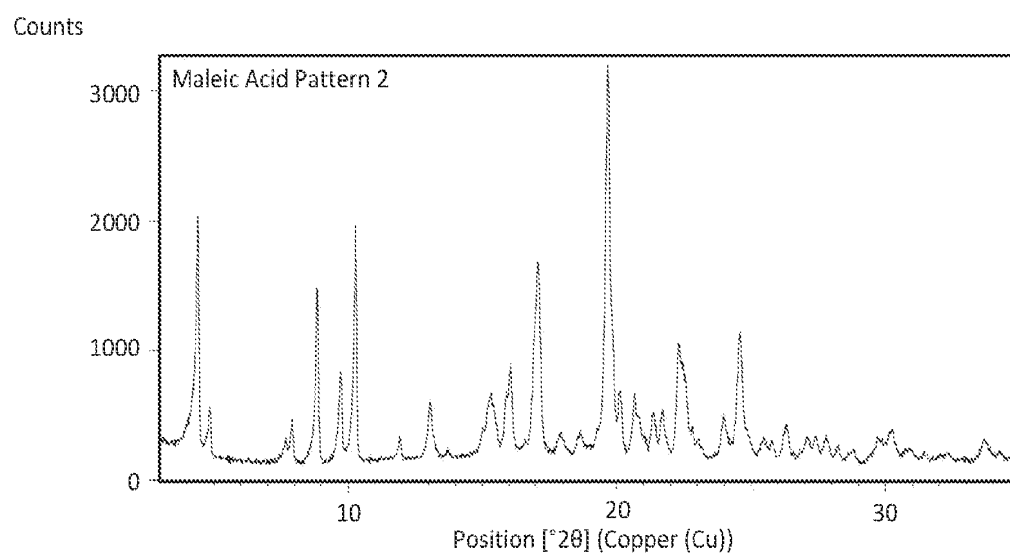


FIG. 53

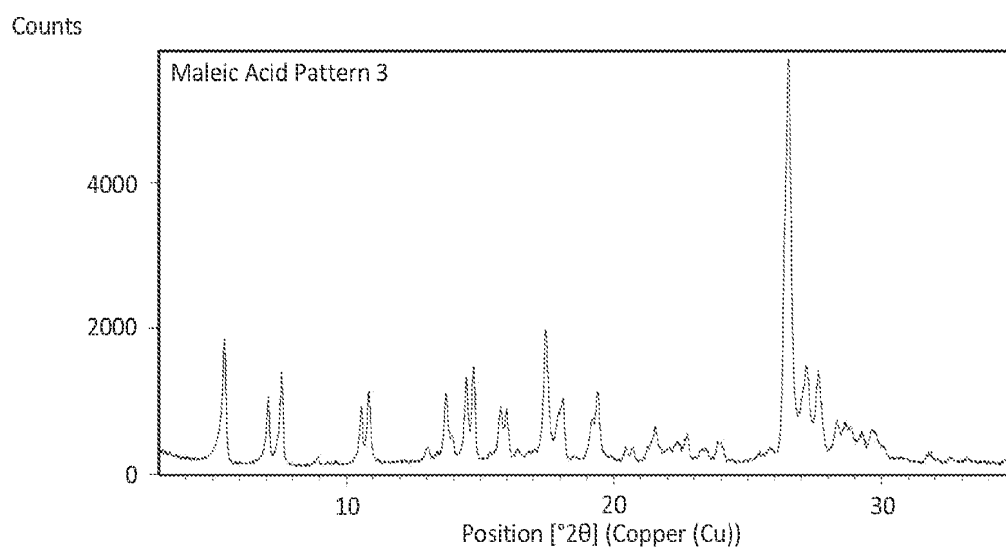


FIG. 54

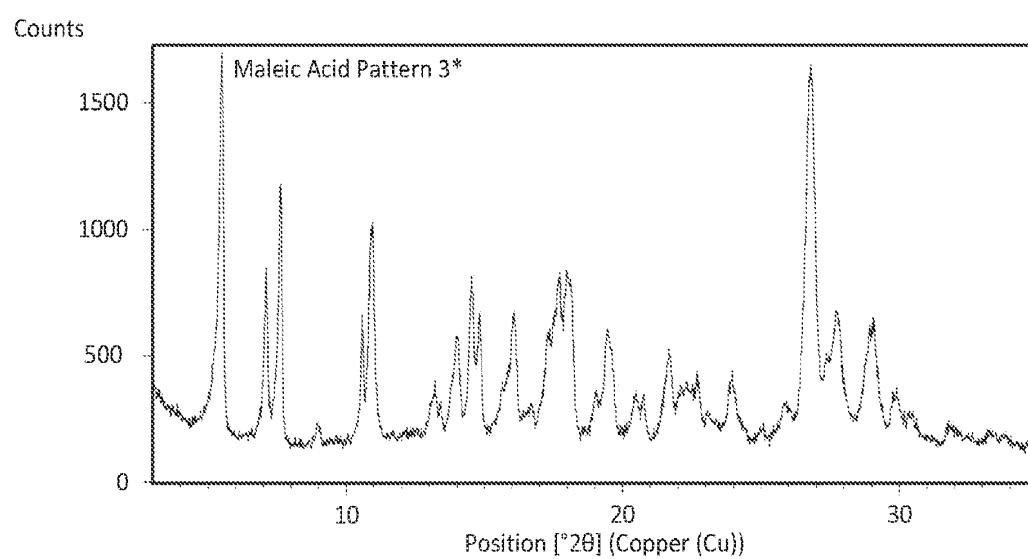


FIG. 55

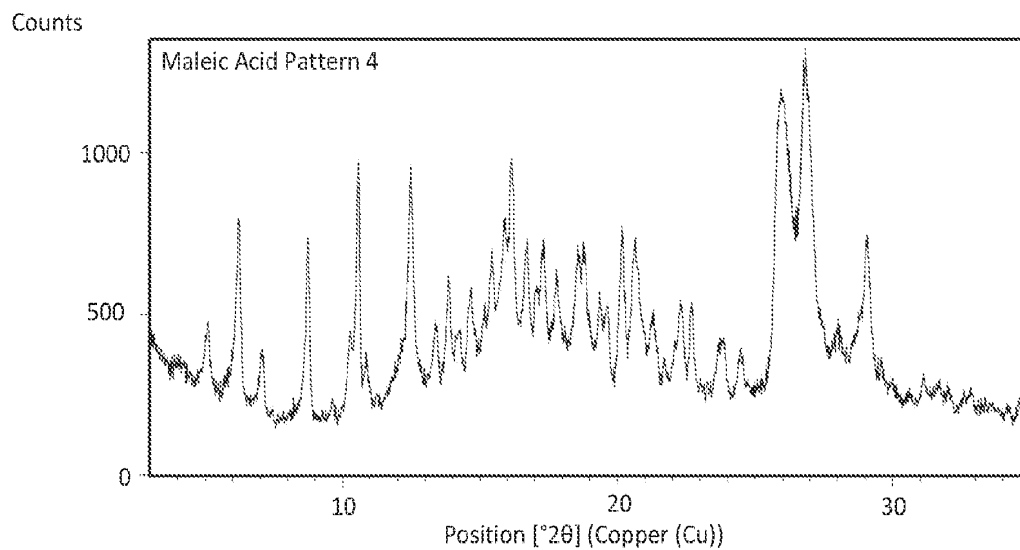


FIG. 56

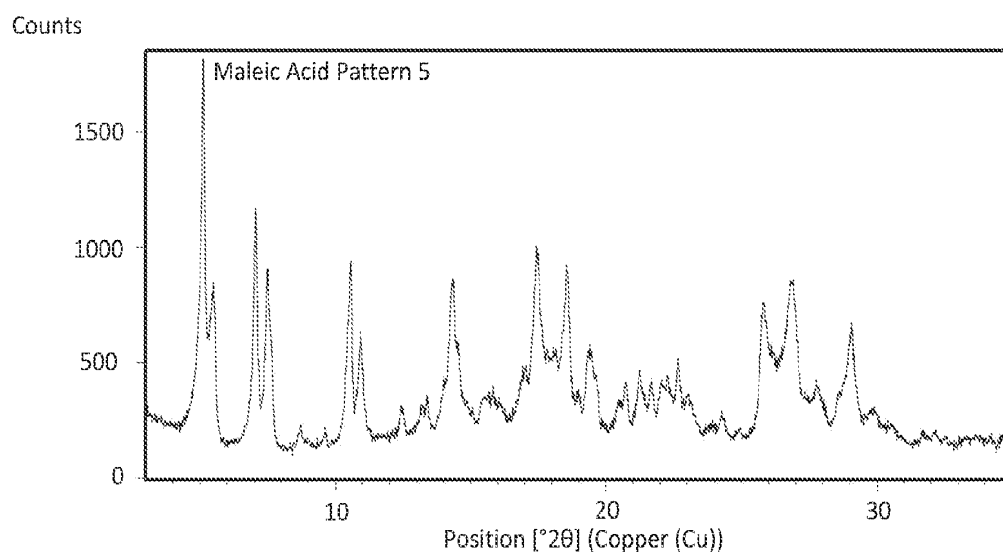


FIG. 57

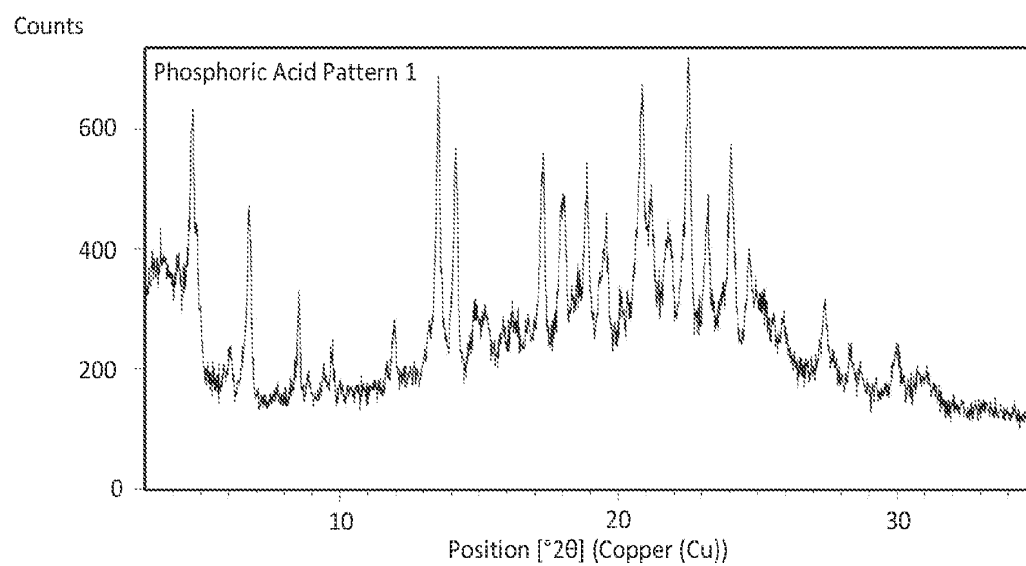


FIG. 58

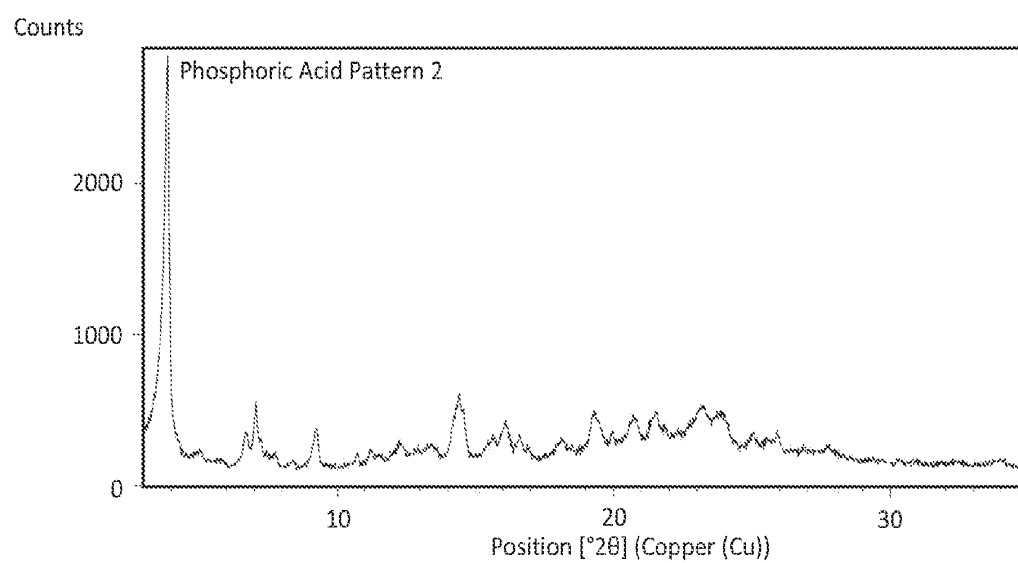


FIG. 59

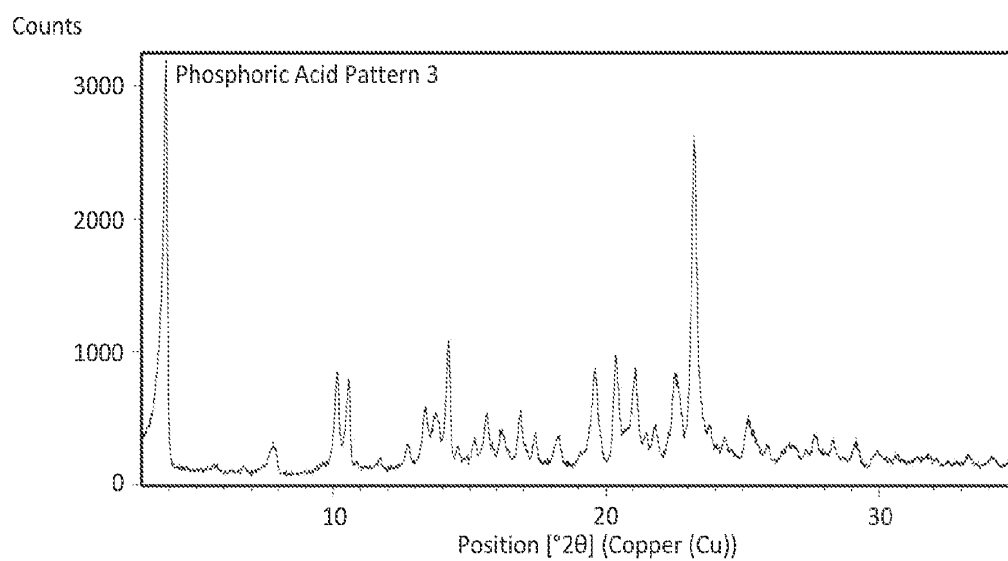


FIG. 60

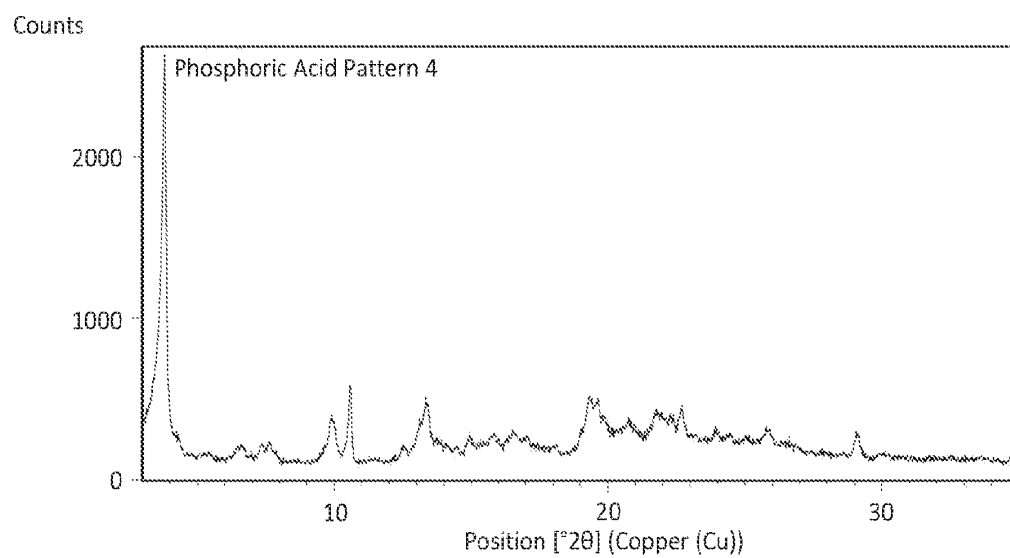


FIG. 61

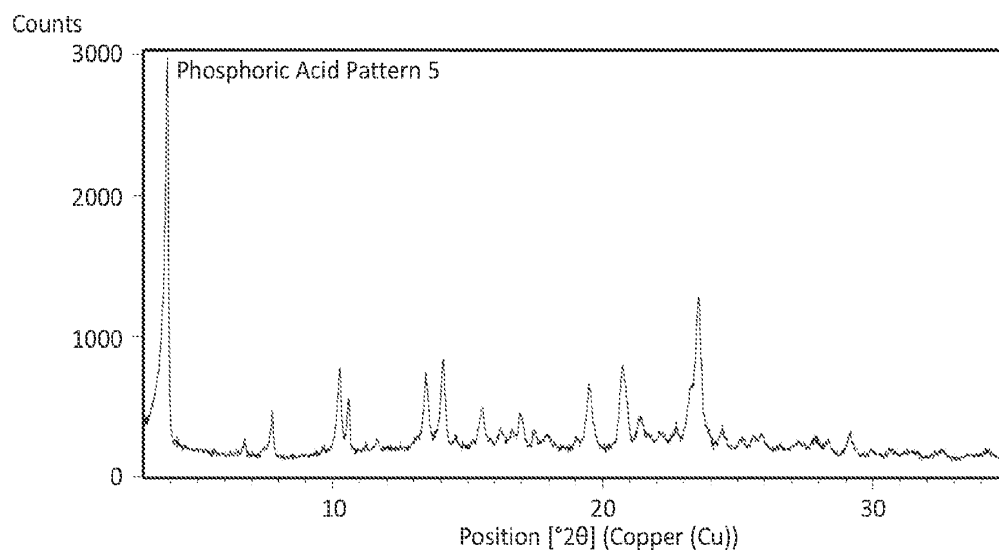


FIG. 62

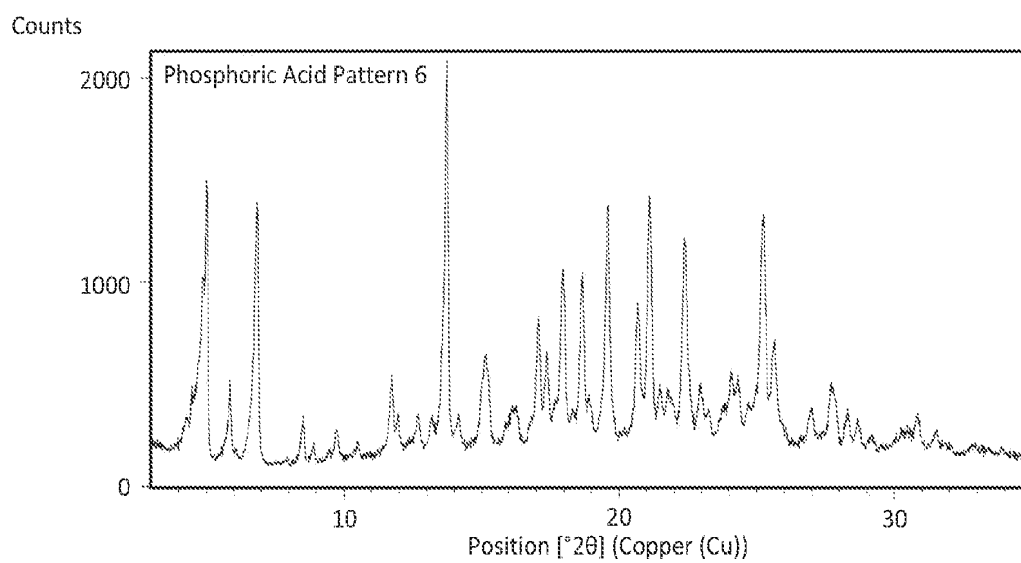


FIG. 63

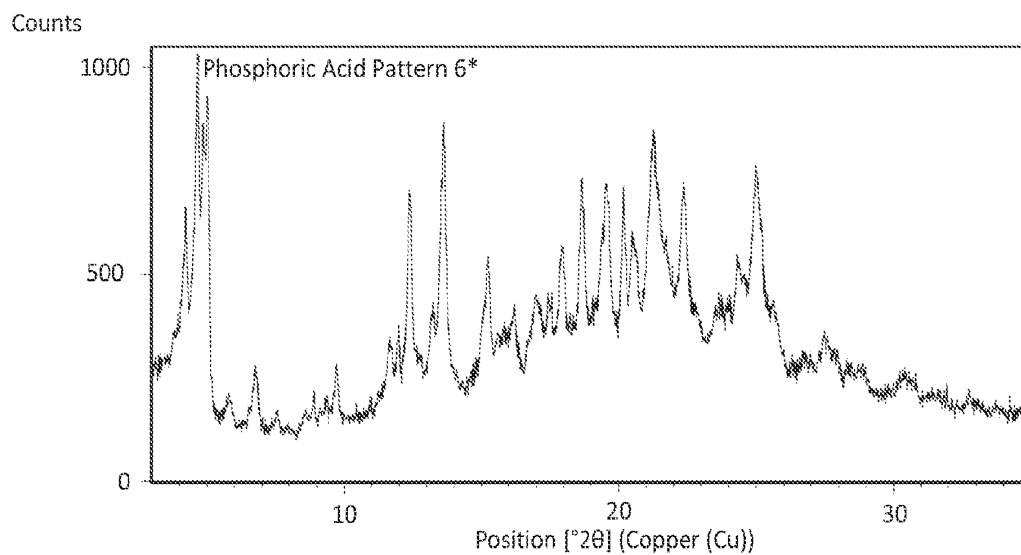


FIG. 64

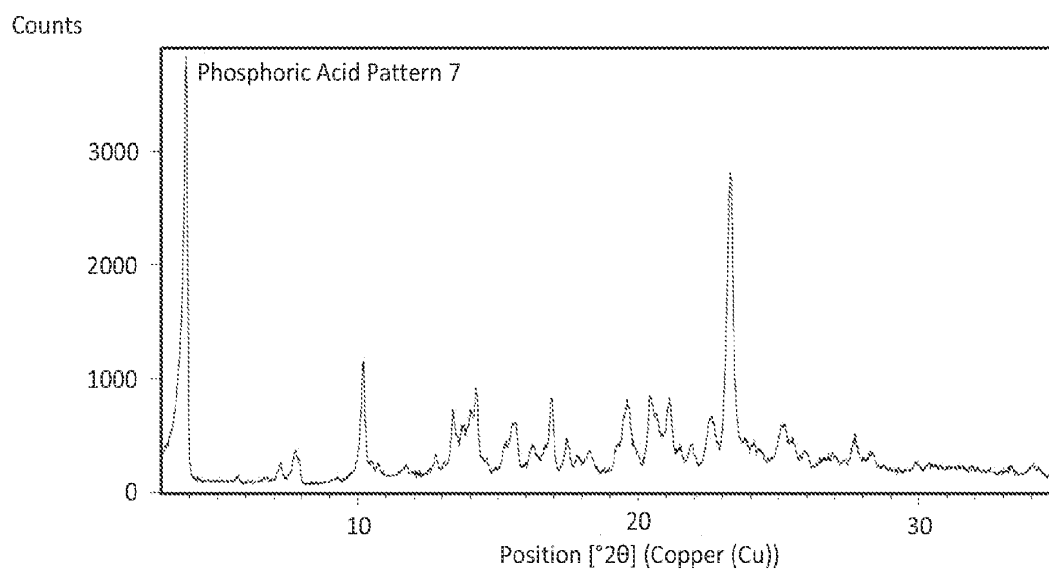


FIG. 65

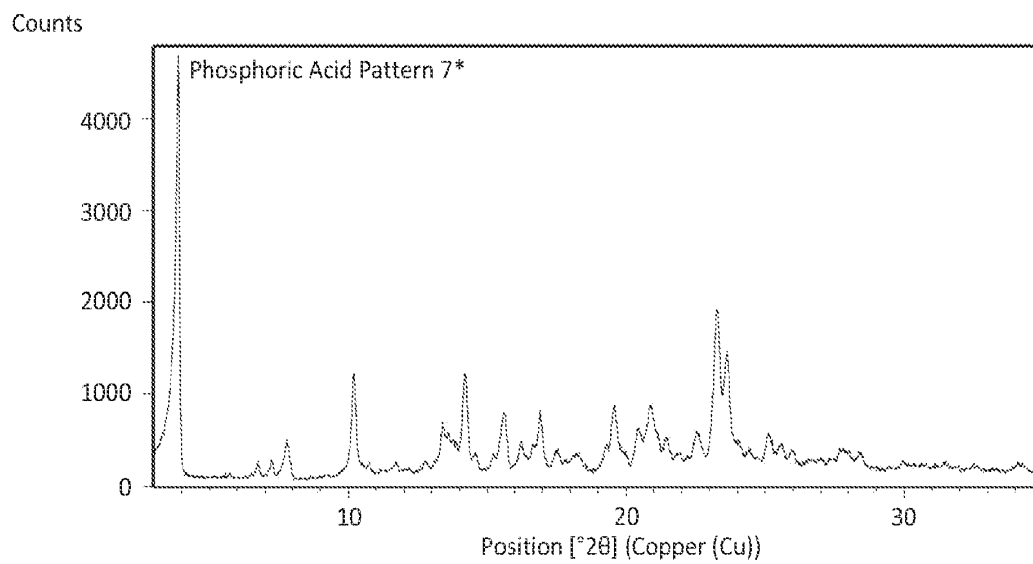


FIG. 66

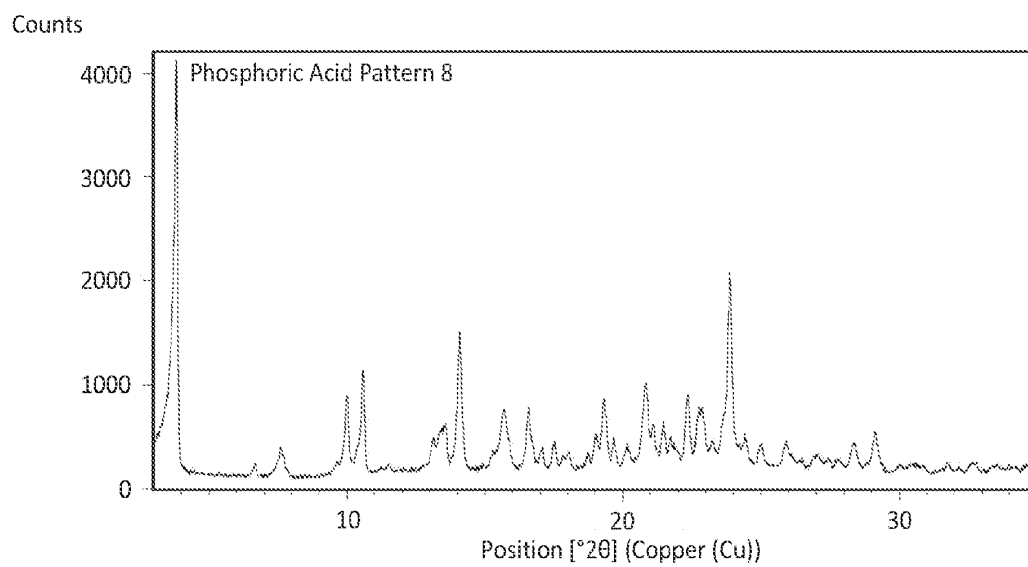


FIG. 67

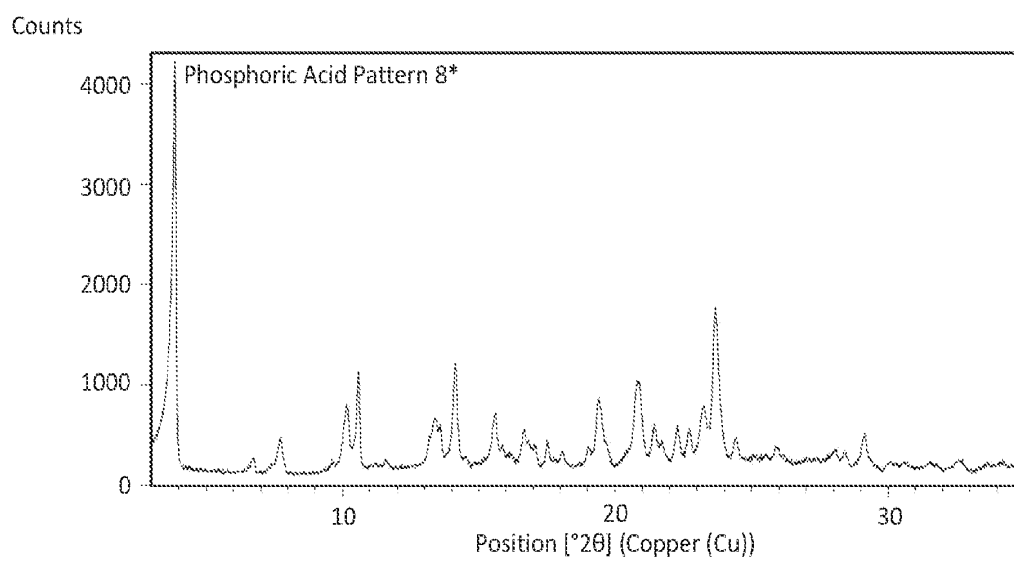


FIG. 68

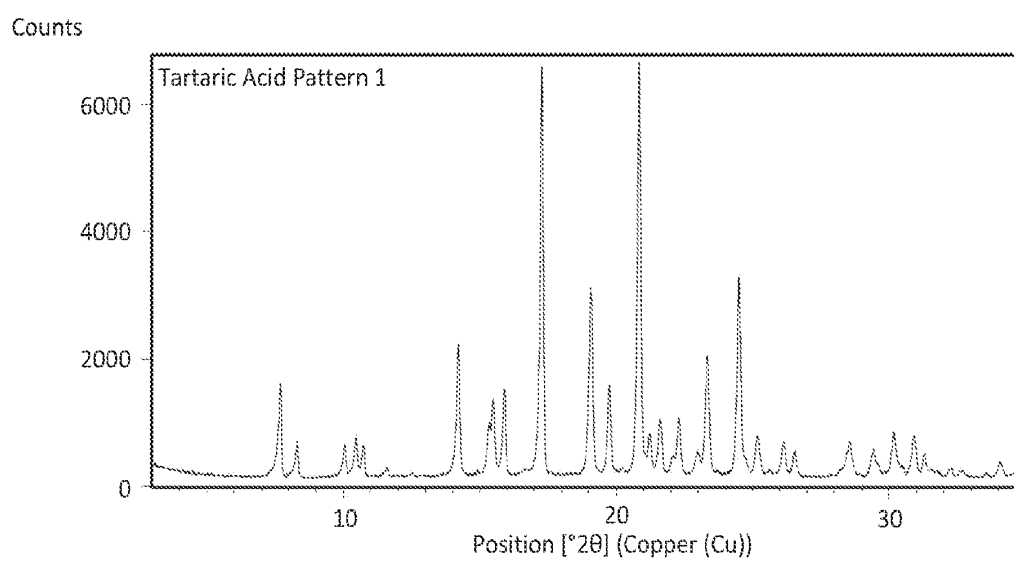


FIG. 69

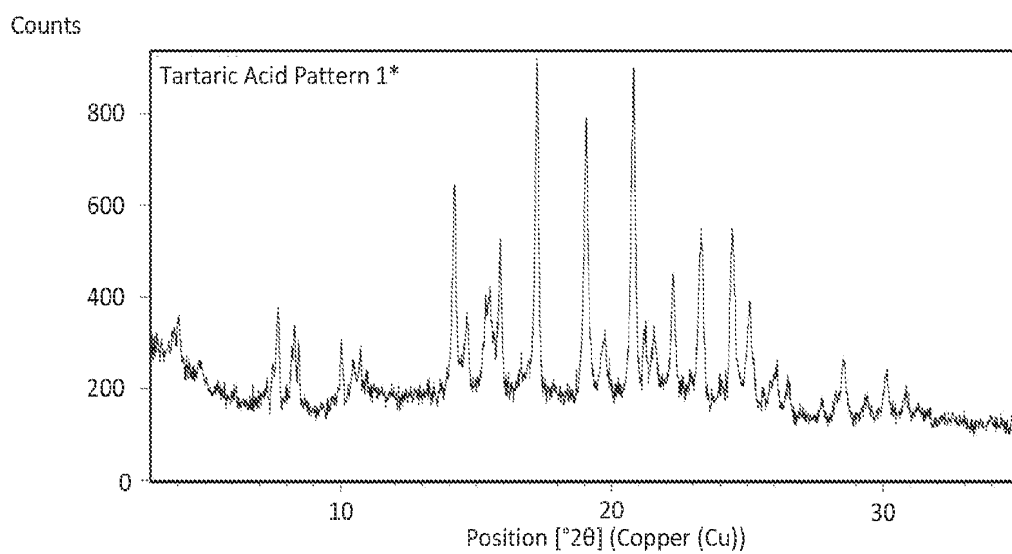


FIG. 70

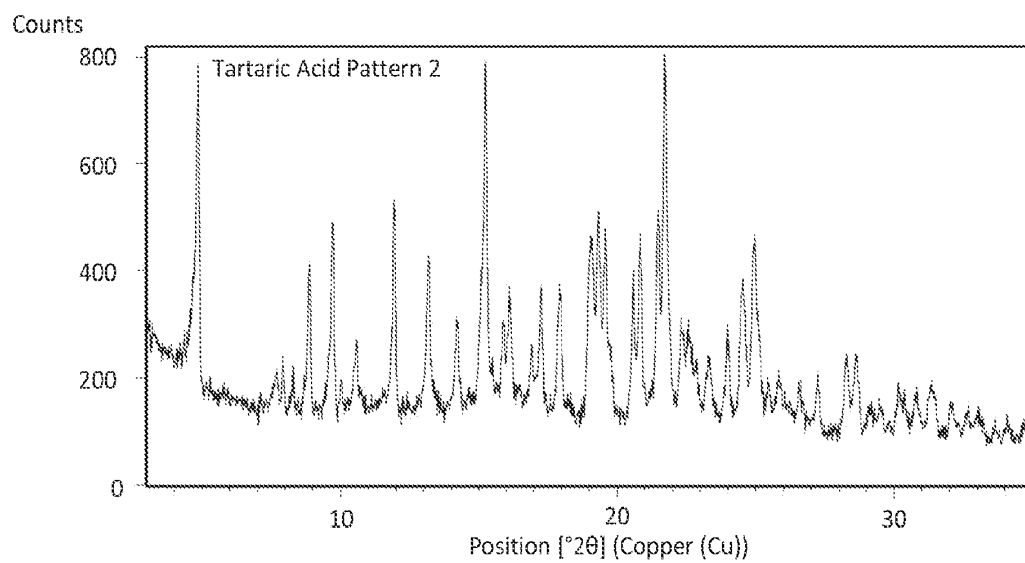


FIG. 71

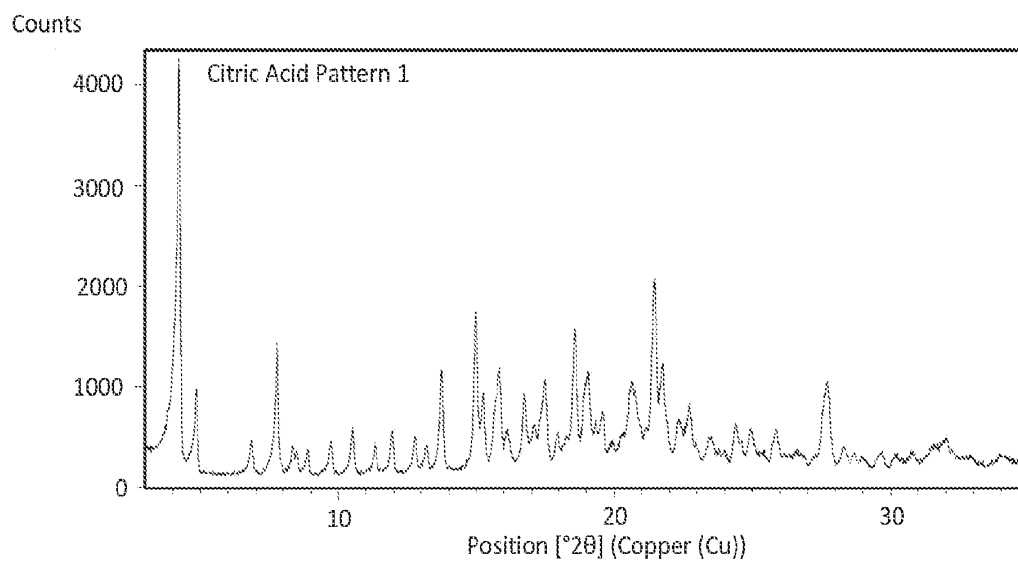


FIG. 72

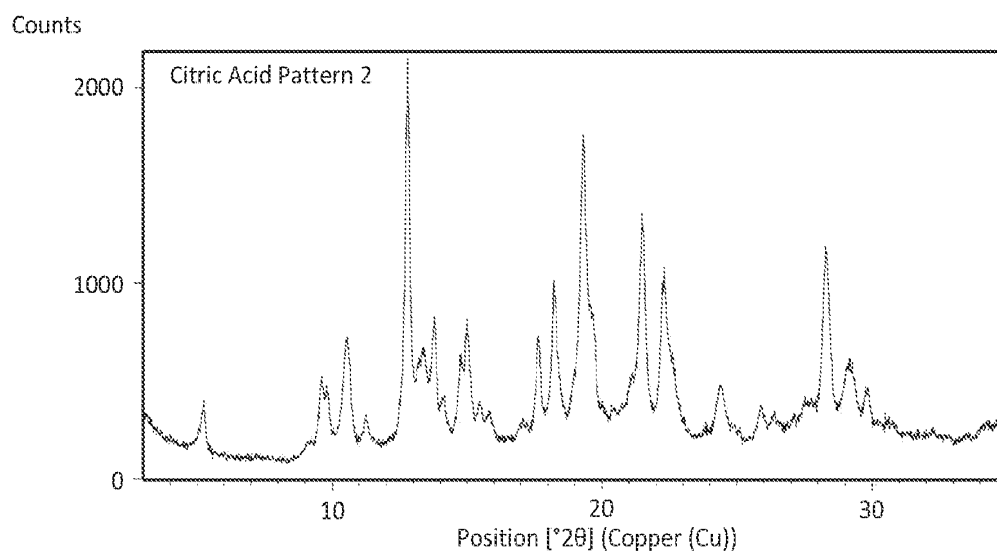


FIG. 73

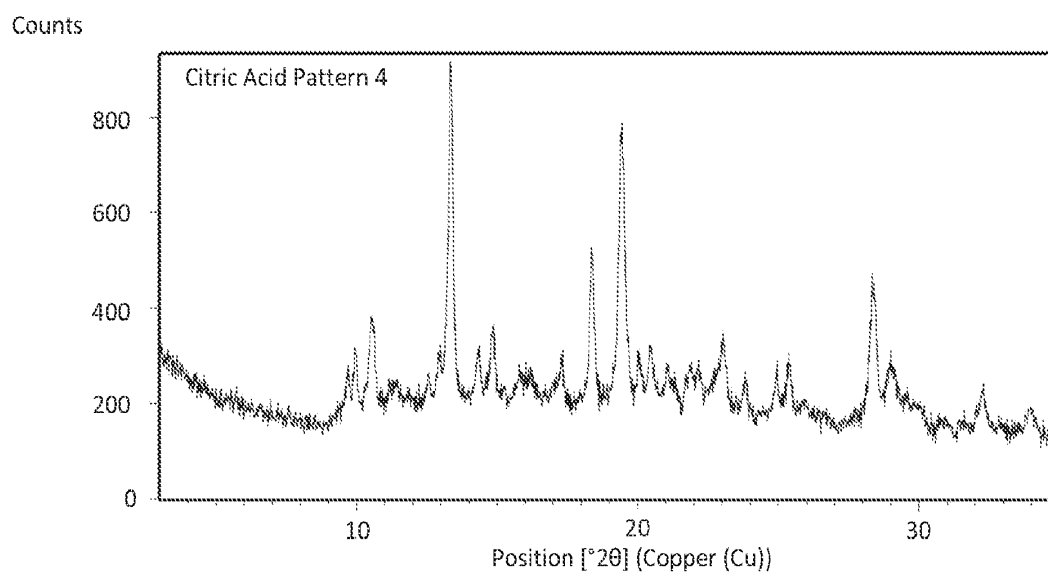


FIG. 74

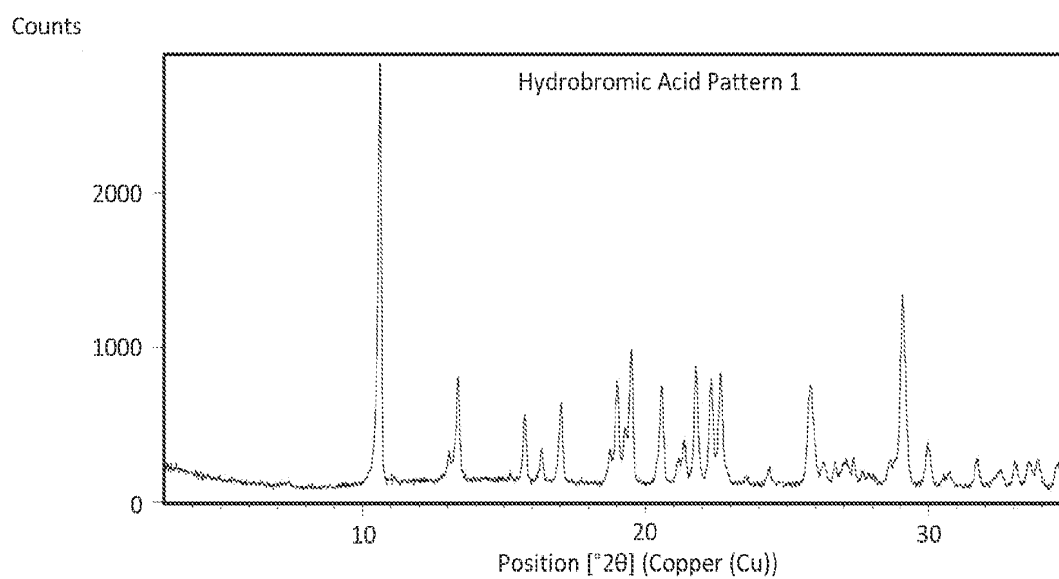


FIG. 75

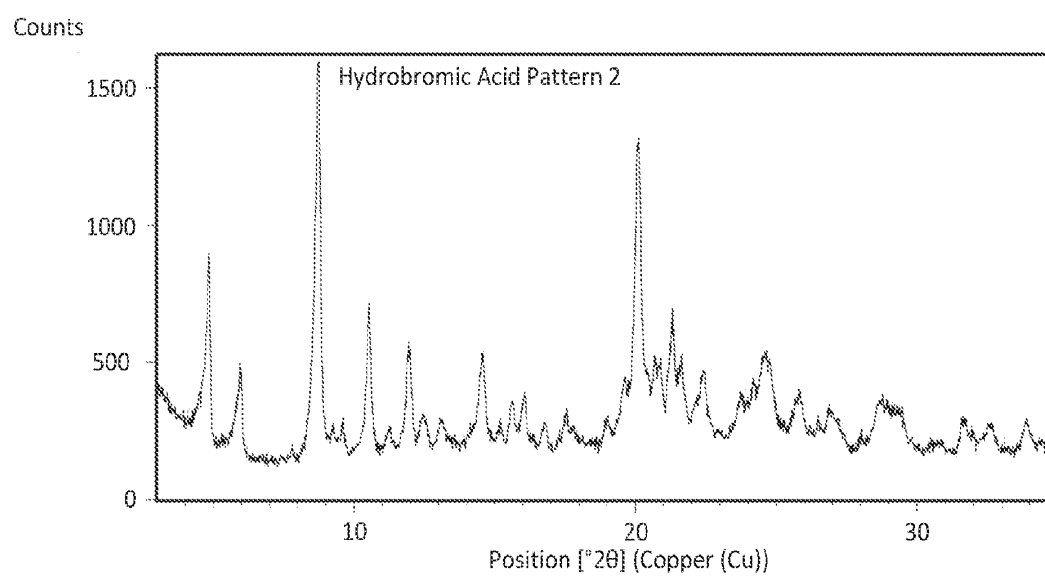


FIG. 76

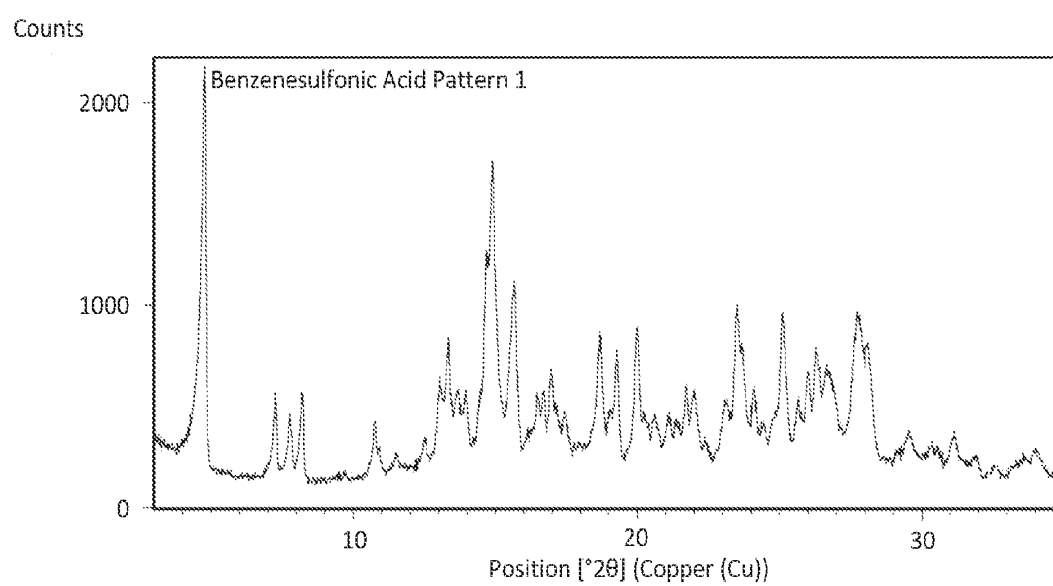


FIG. 77

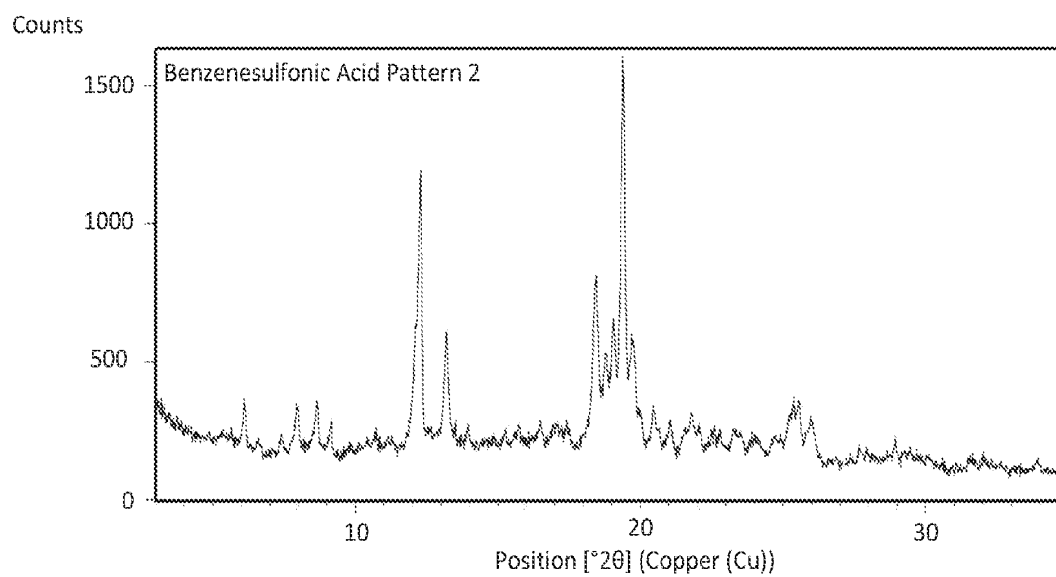


FIG. 78

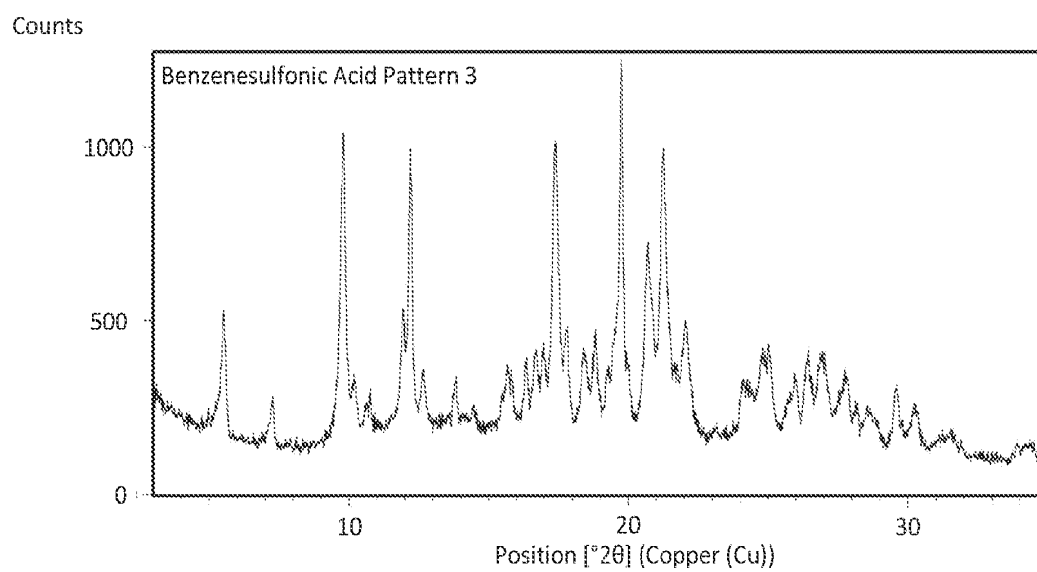


FIG. 79

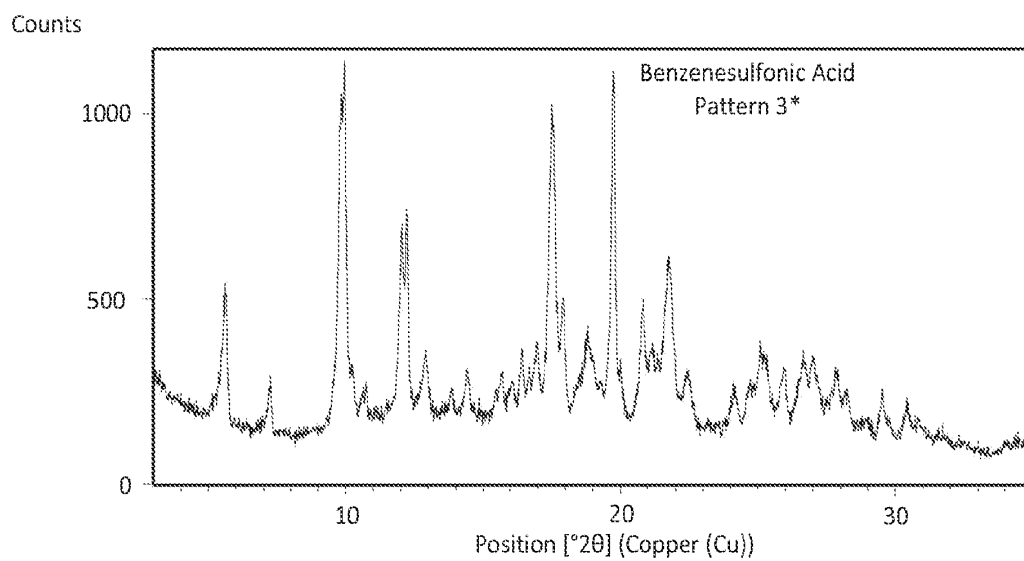


FIG. 80

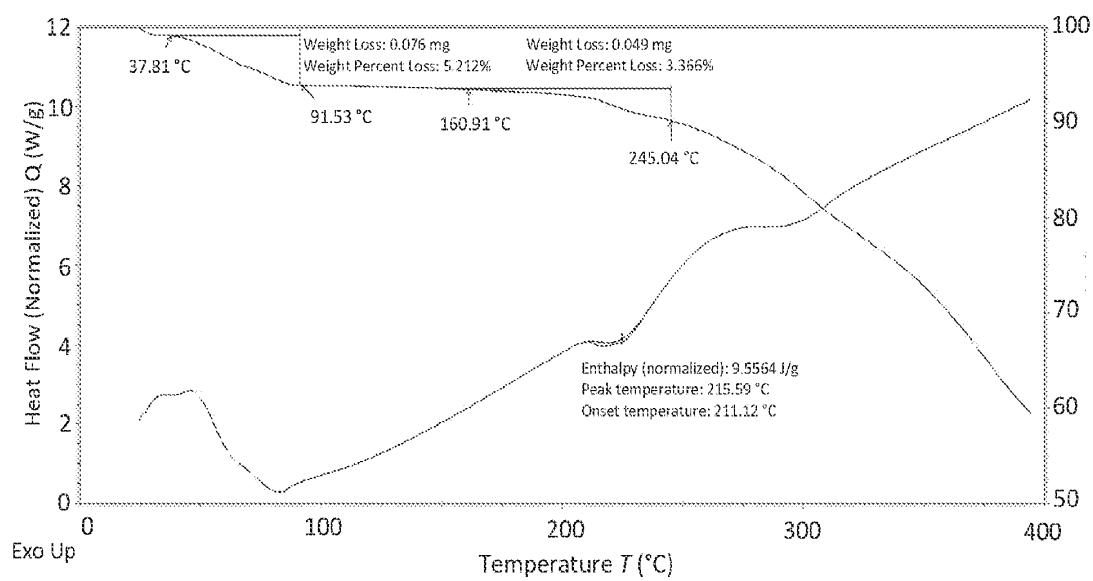


FIG. 81

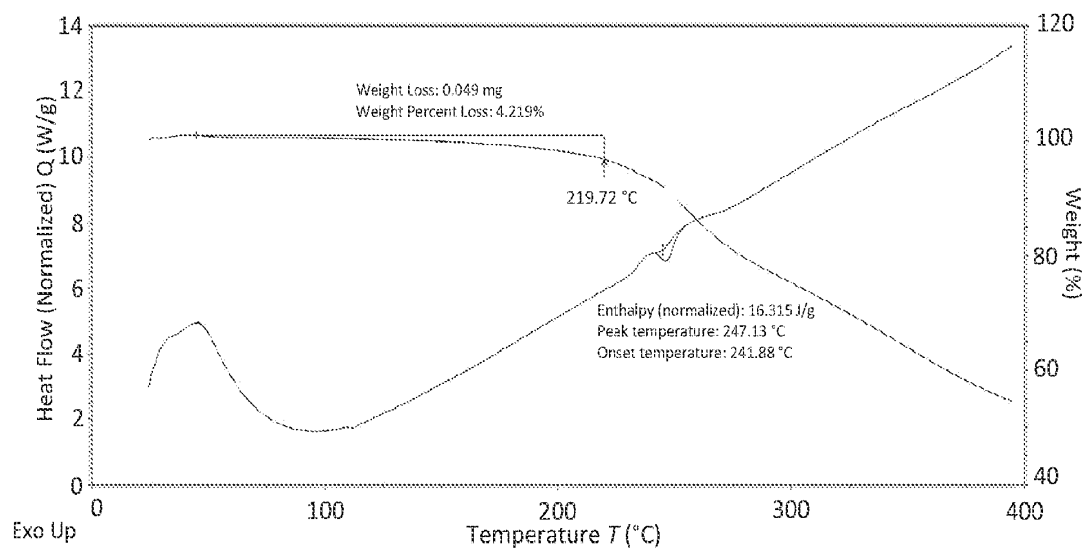


FIG. 82

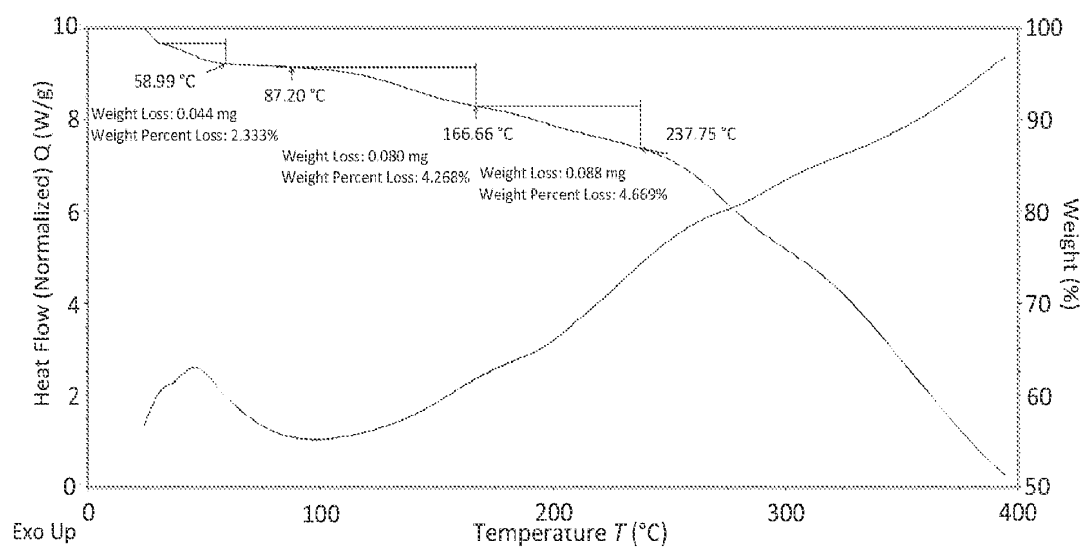


FIG. 83

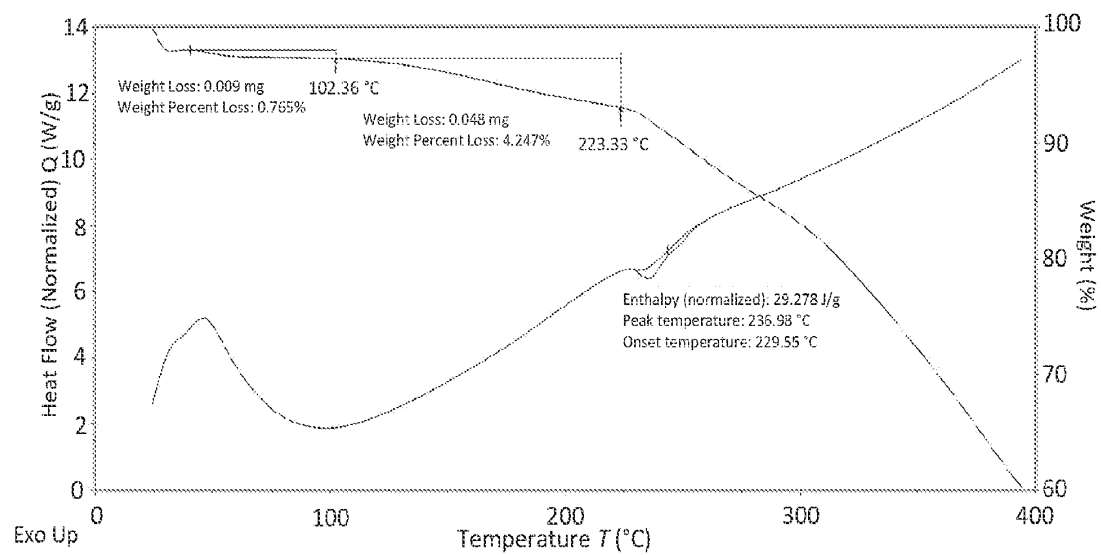


FIG. 84

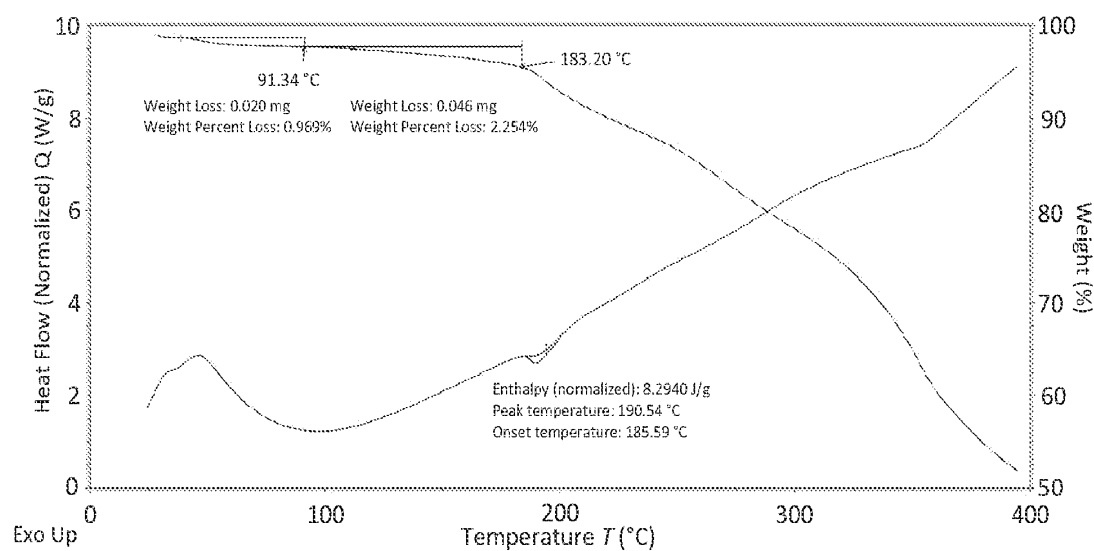


FIG. 85

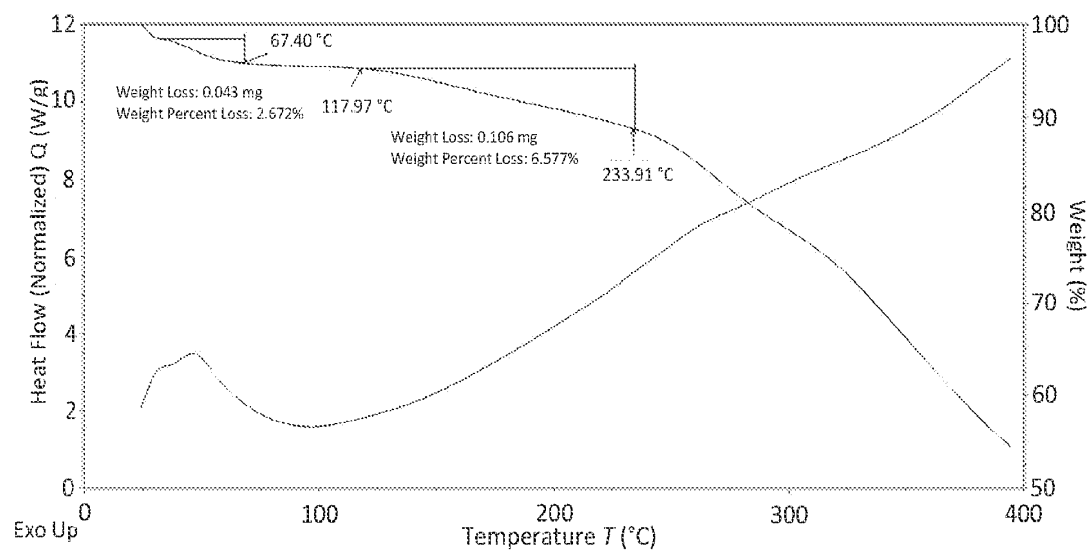


FIG. 86

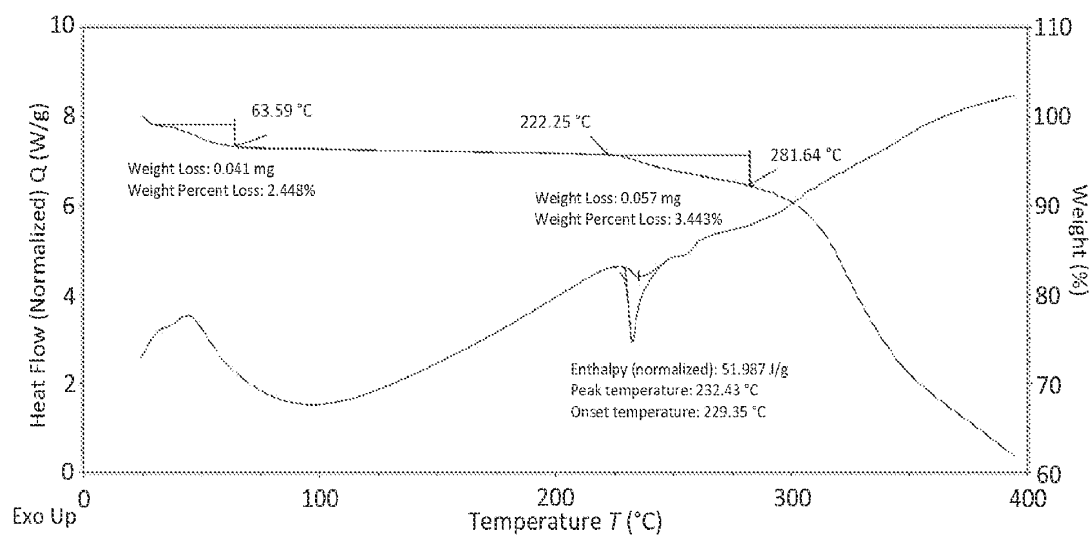


FIG. 87

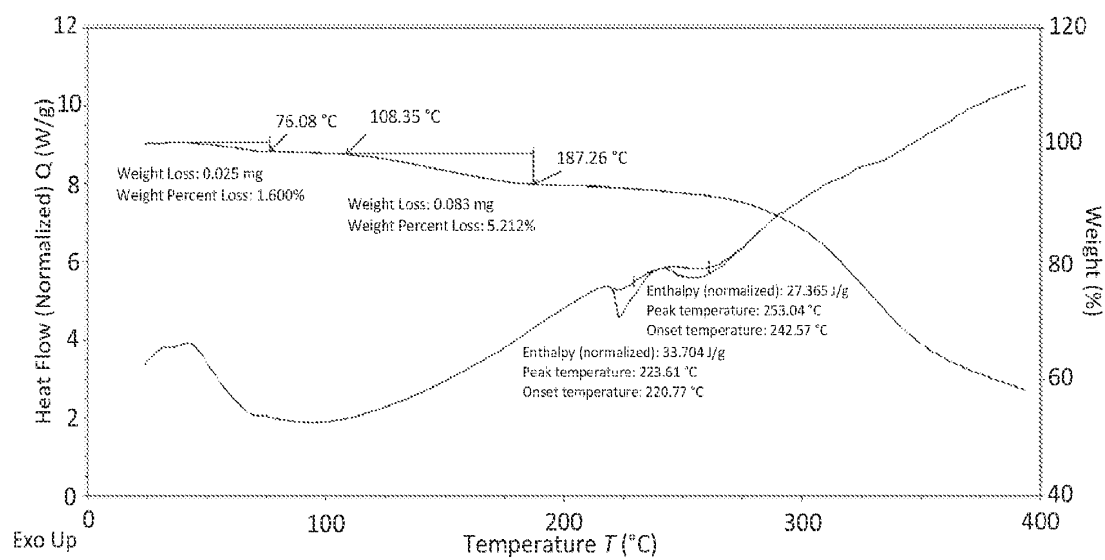


FIG. 88

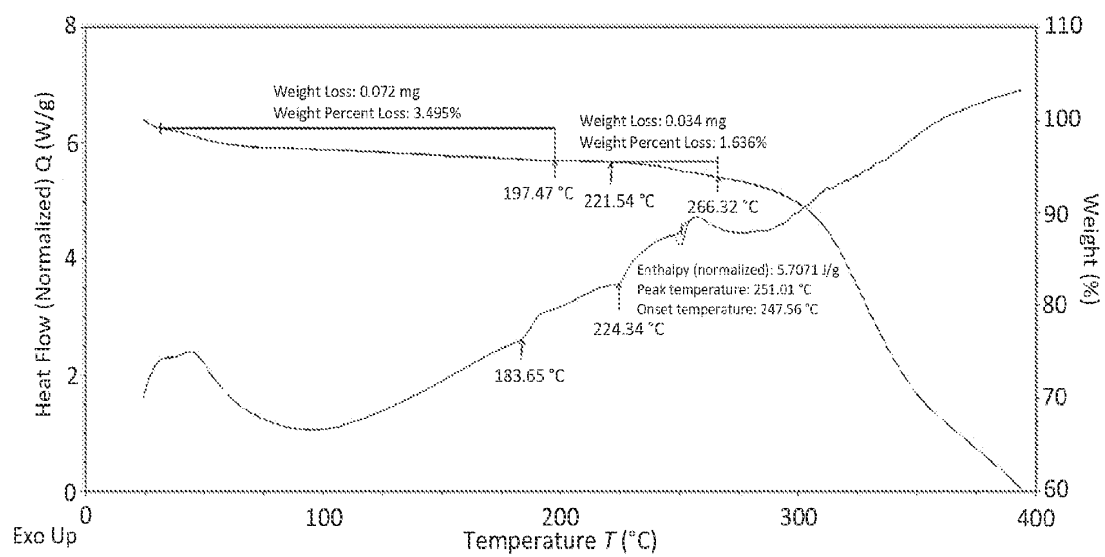


FIG. 89

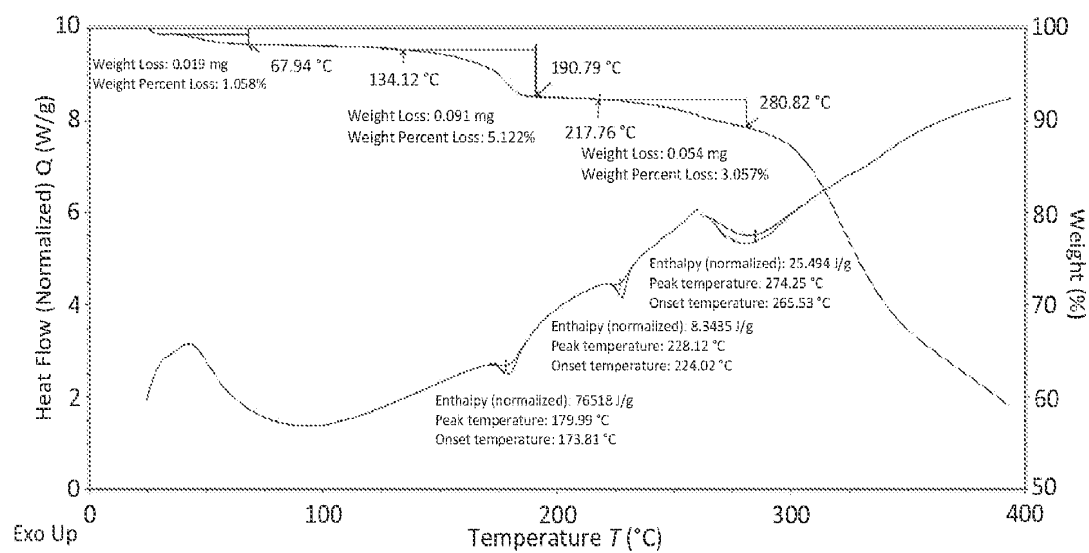


FIG. 90

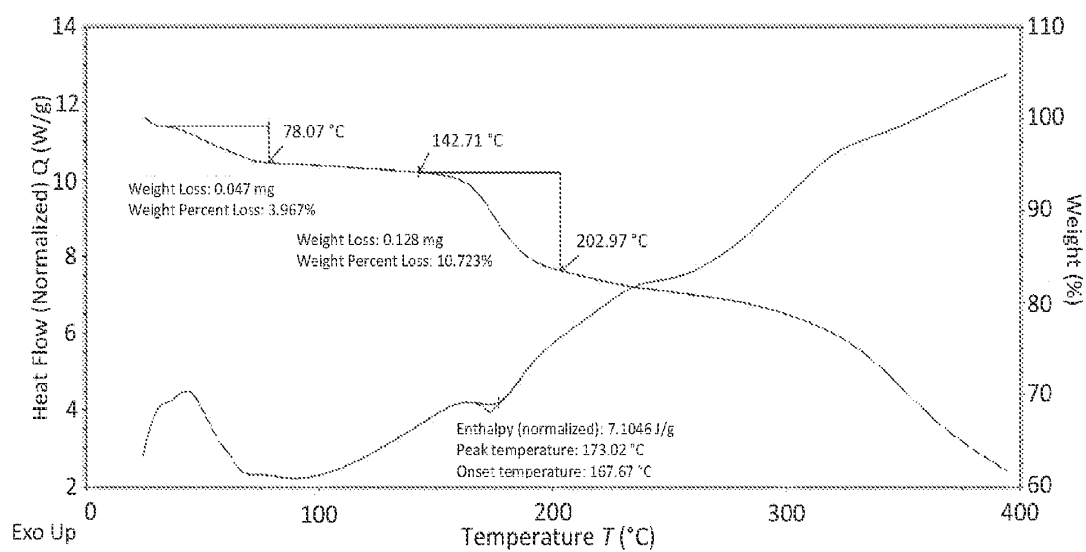


FIG. 91

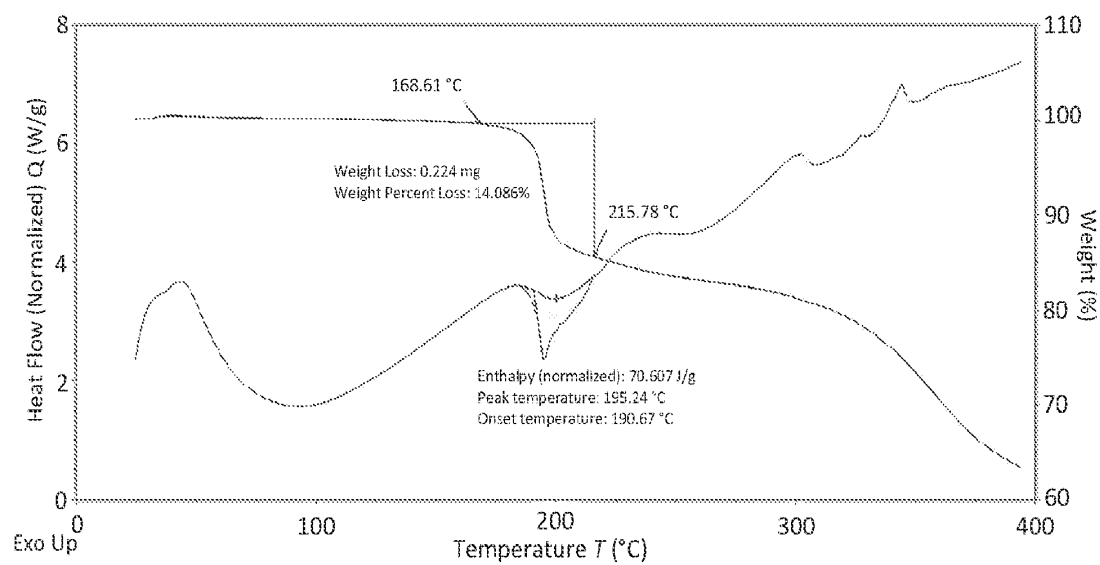


FIG. 92

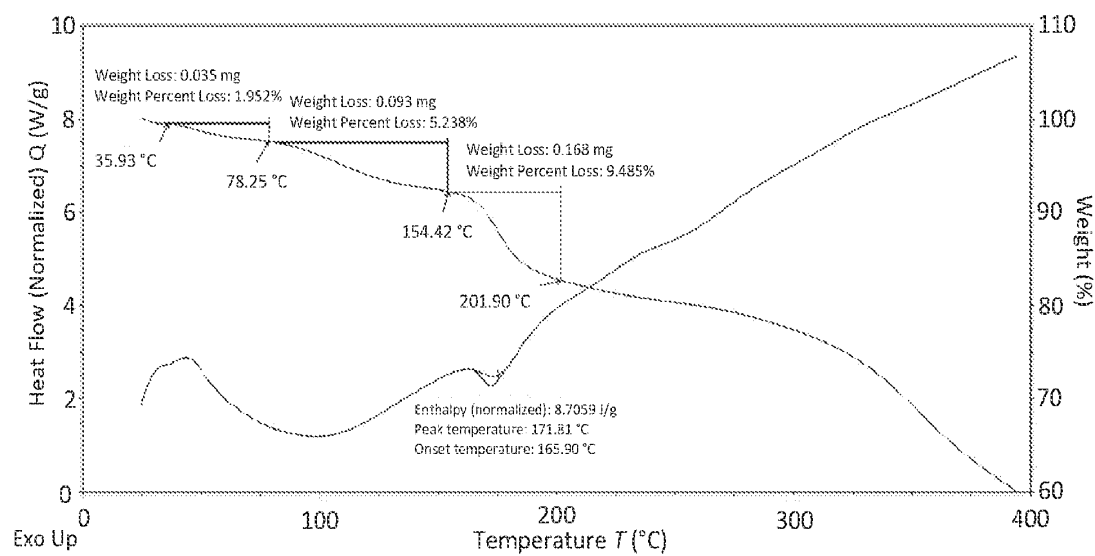


FIG. 93

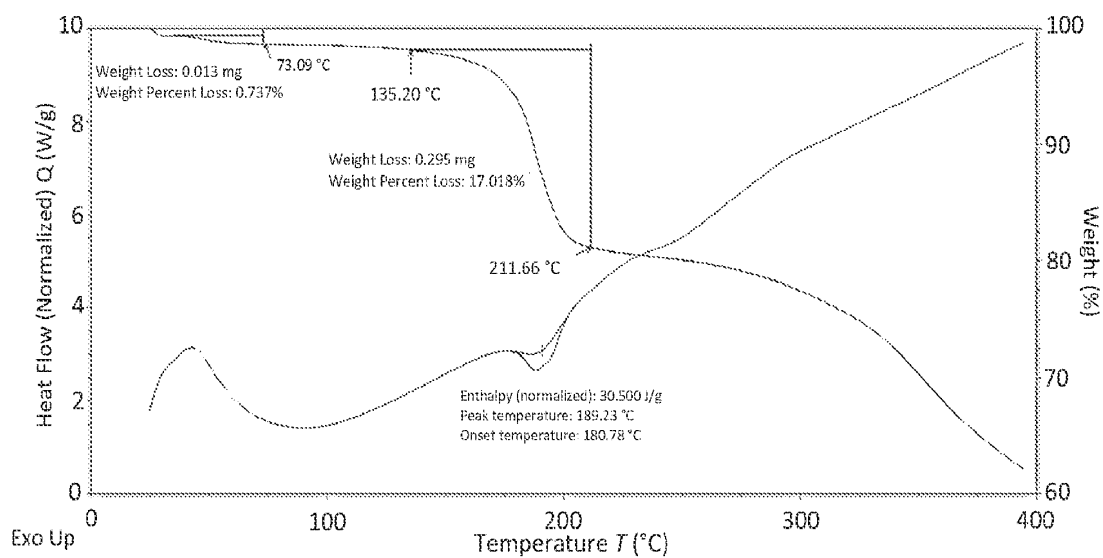


FIG. 94

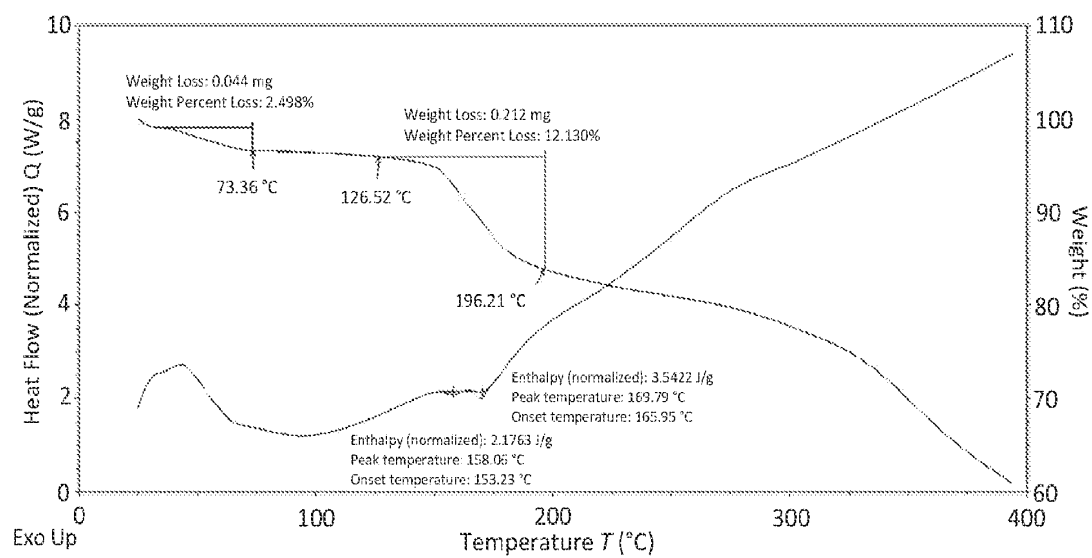


FIG. 95

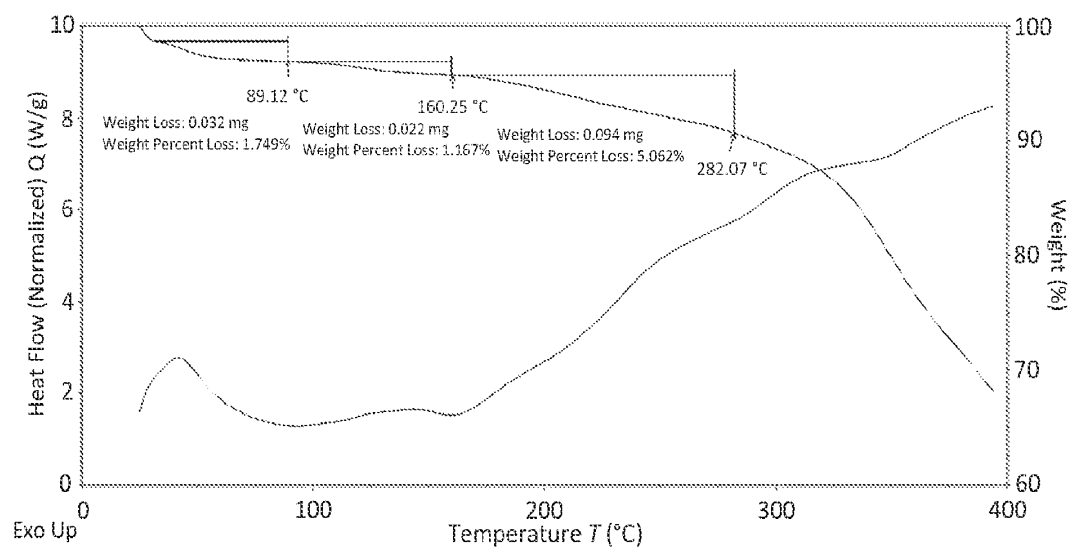


FIG. 96

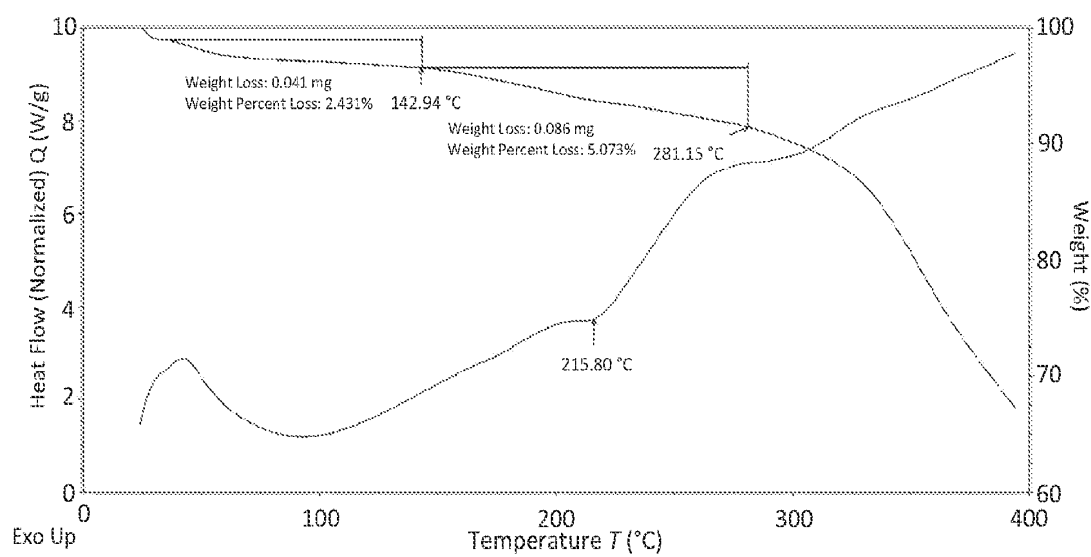


FIG. 97

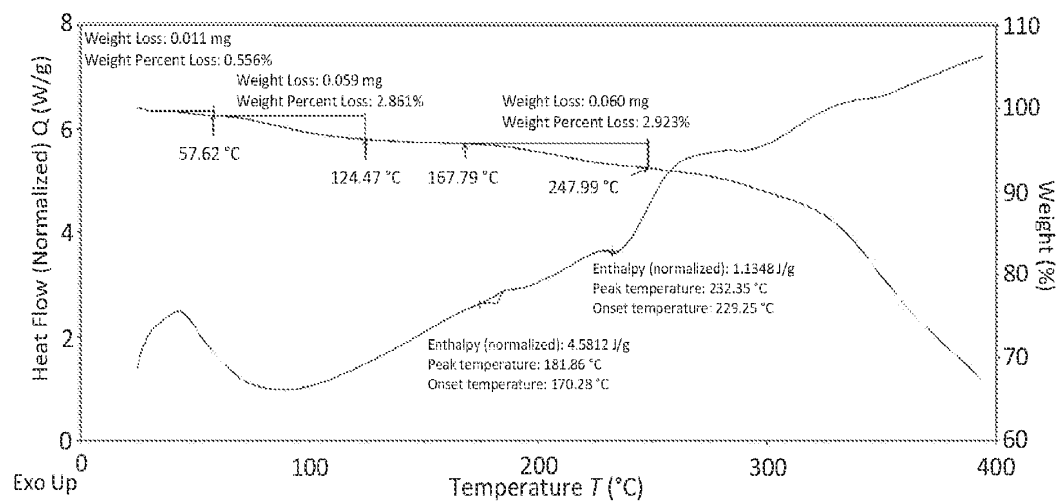


FIG. 98

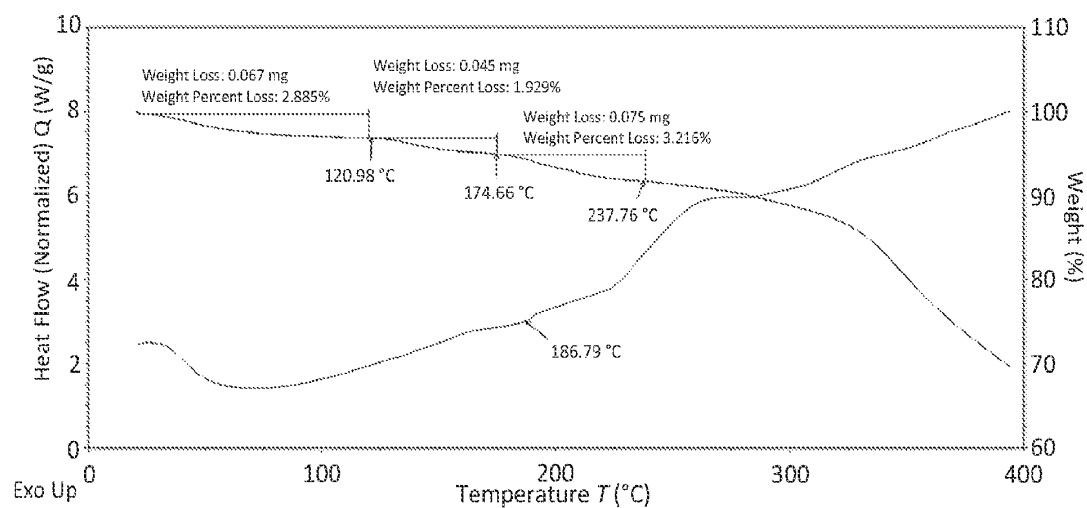


FIG. 99

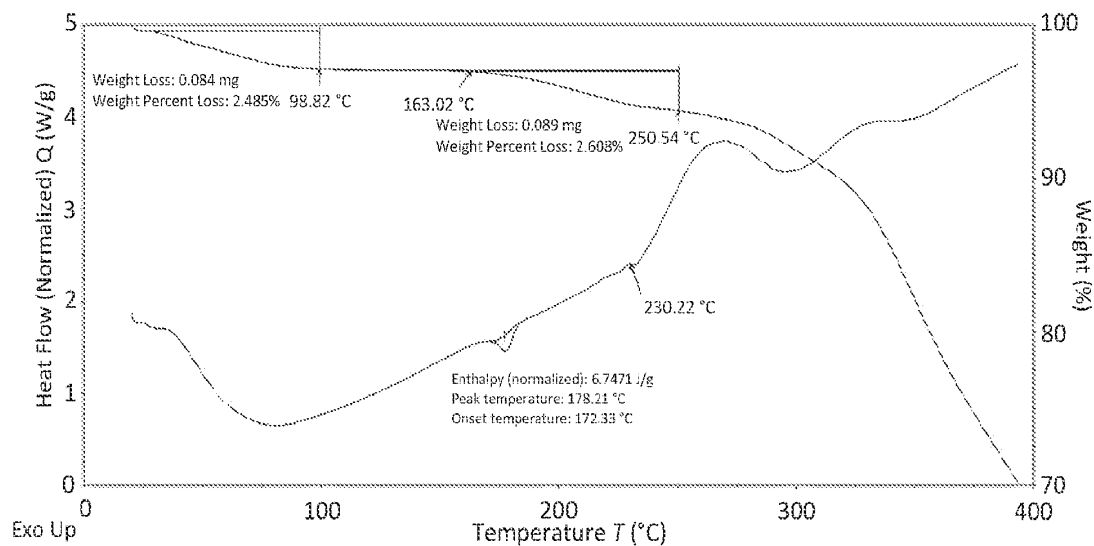


FIG. 100

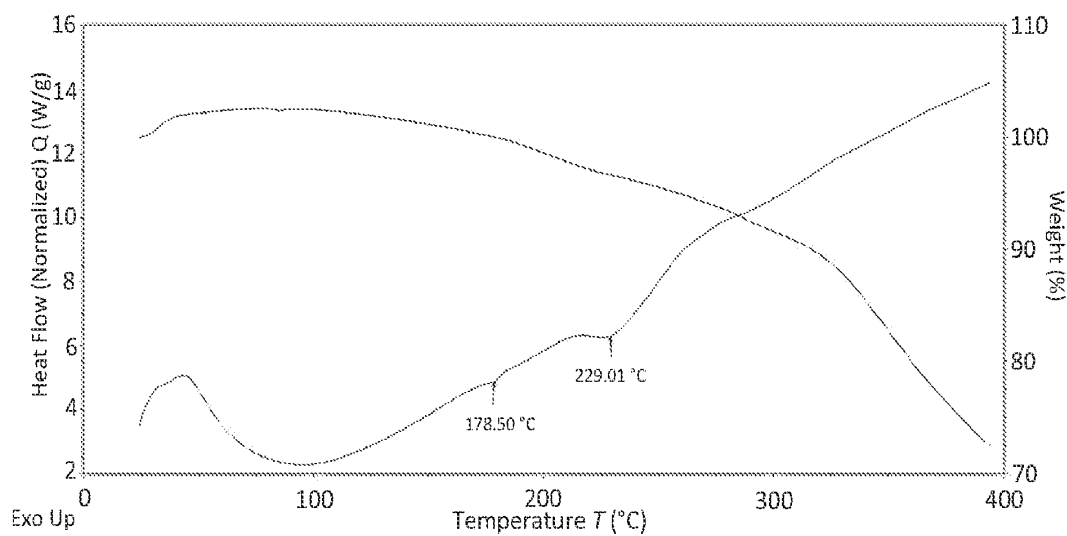


FIG. 101

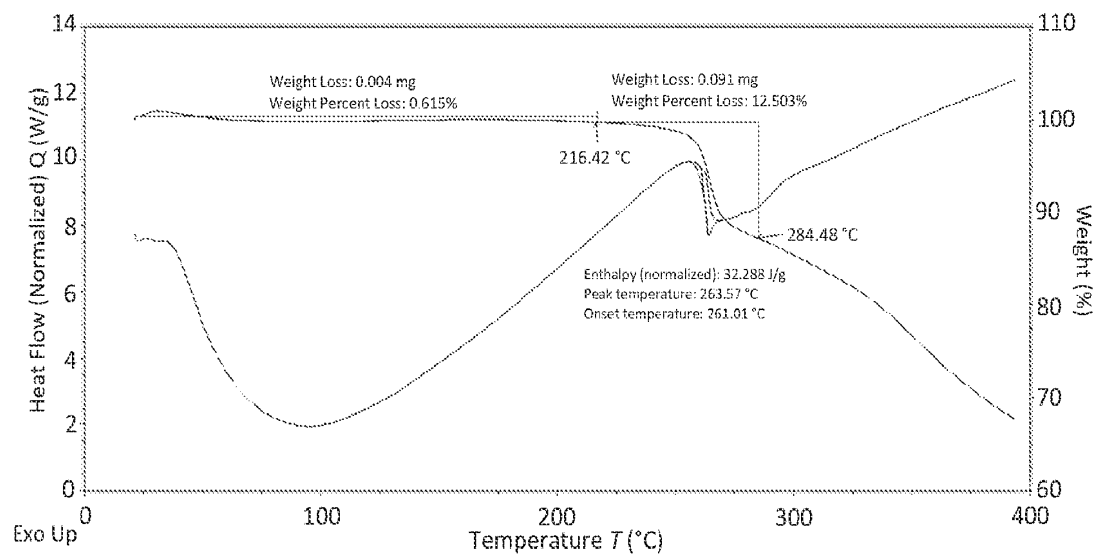


FIG. 102

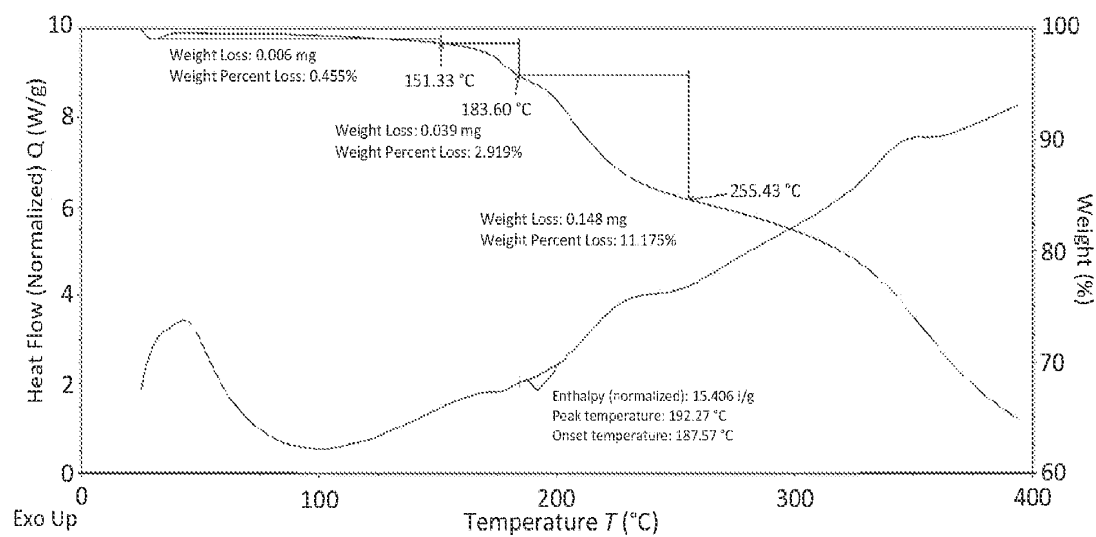


FIG. 103

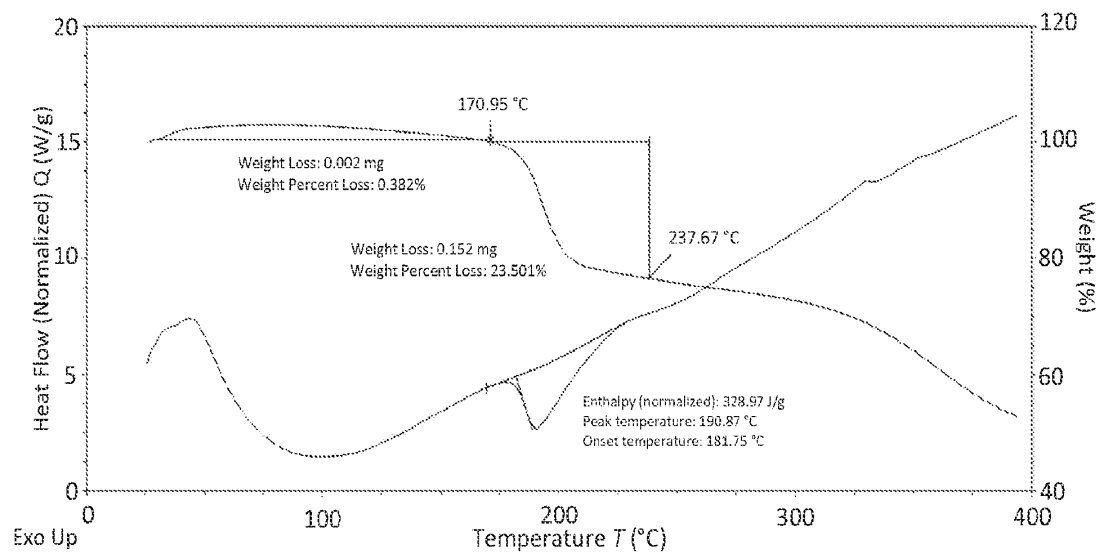


FIG. 104

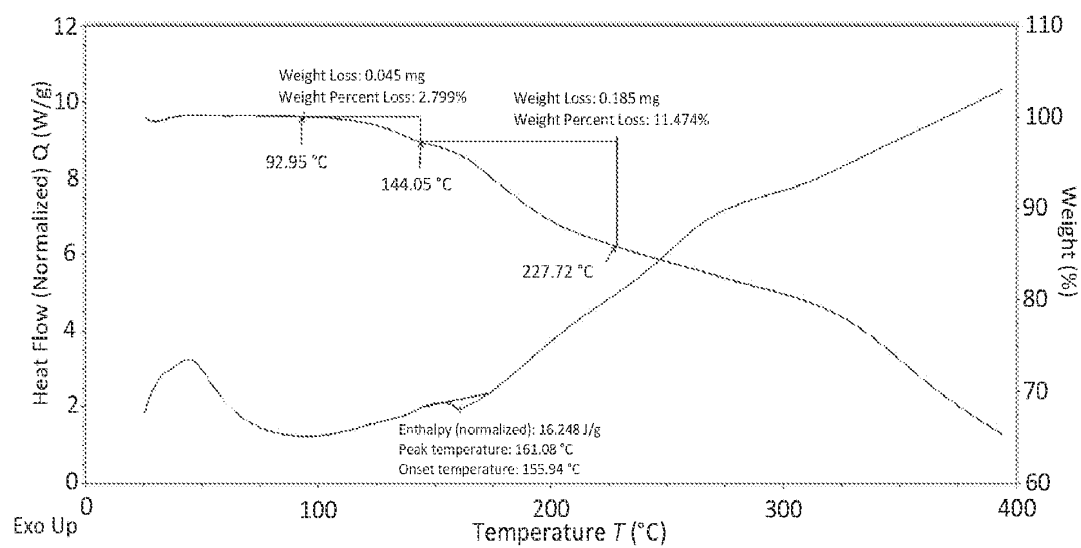


FIG. 105

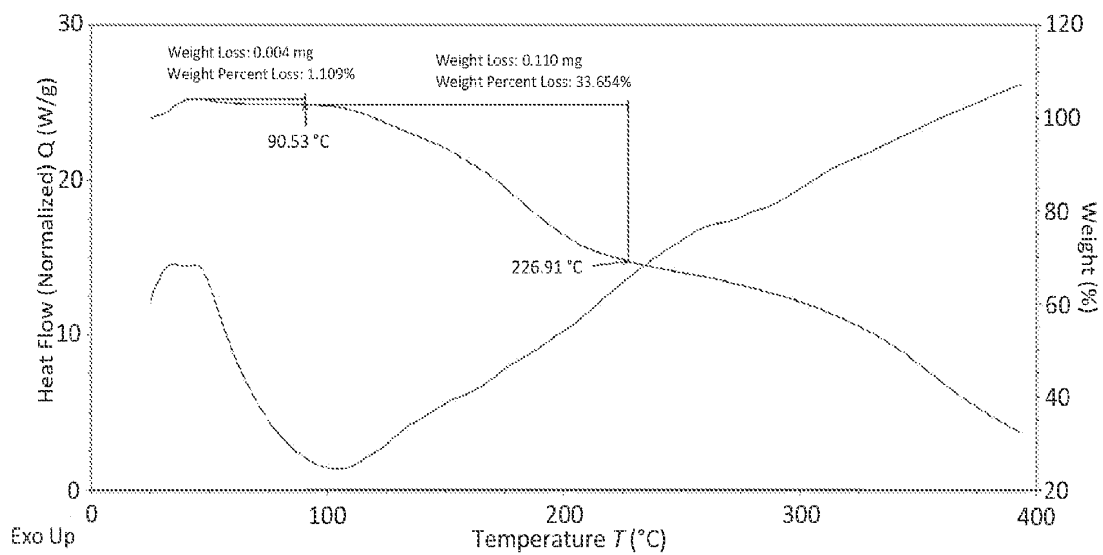


FIG. 106

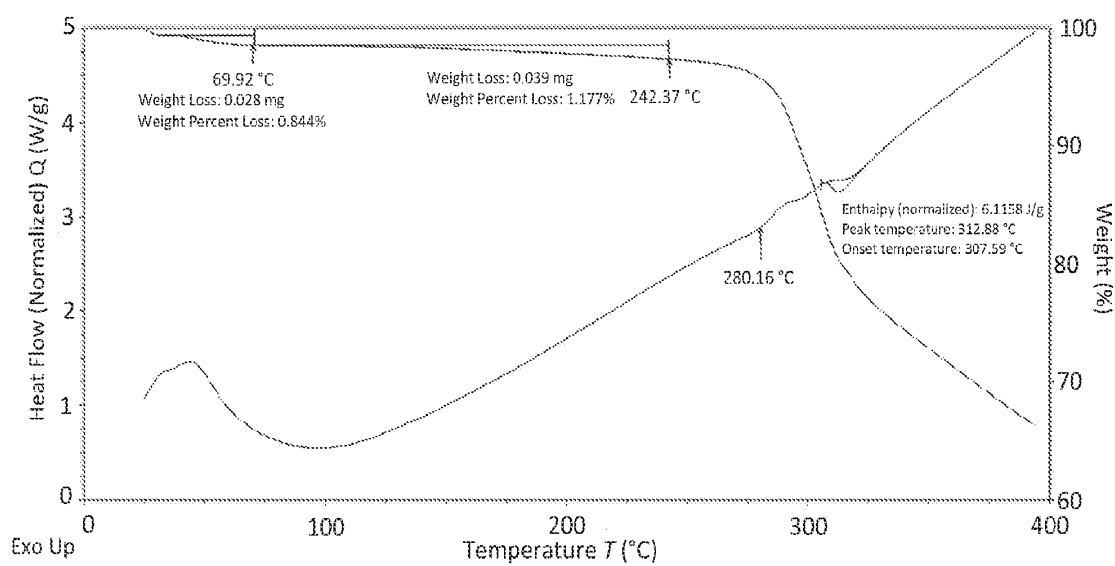


FIG. 107

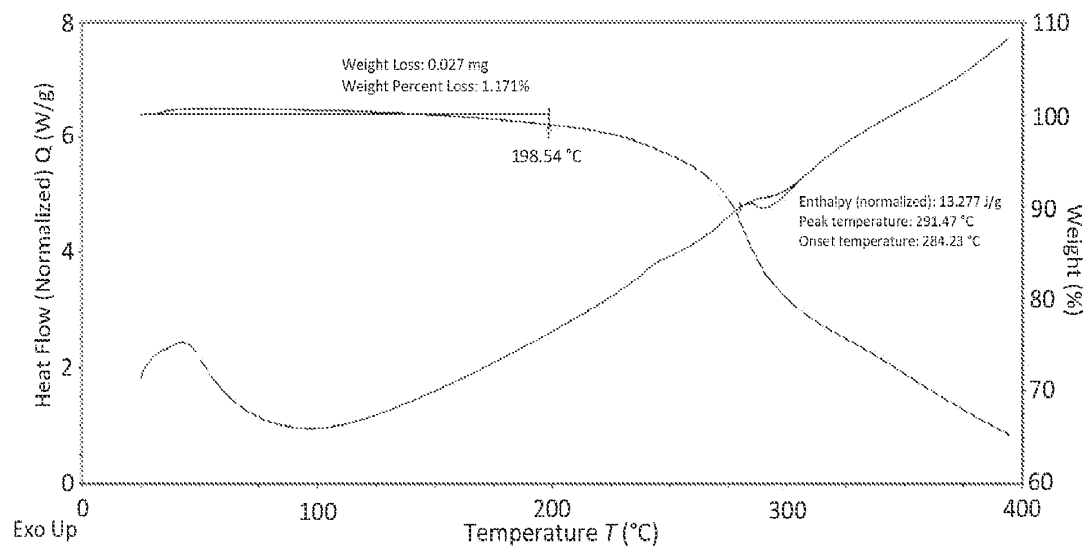


FIG. 108

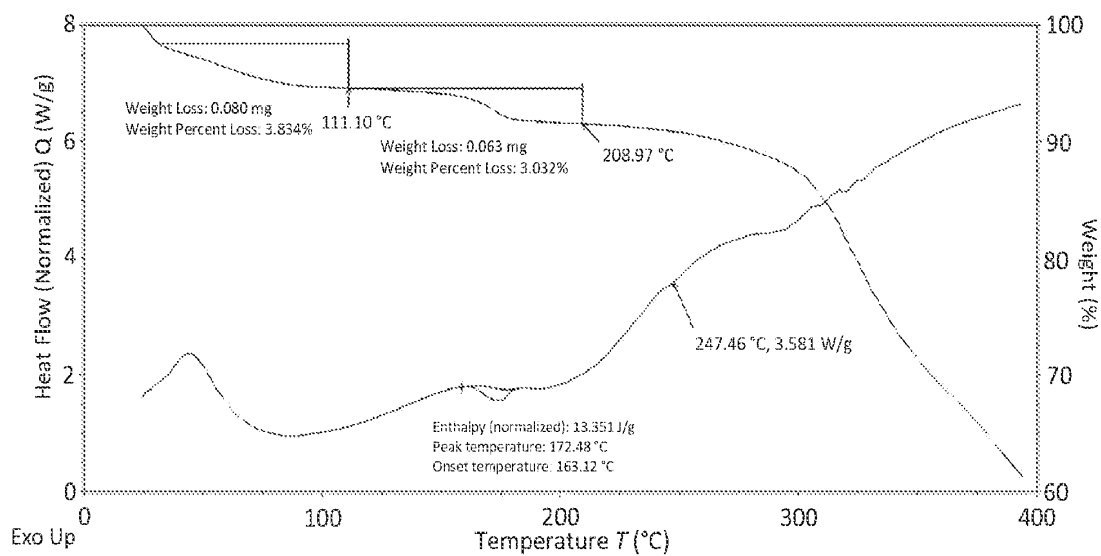


FIG. 109

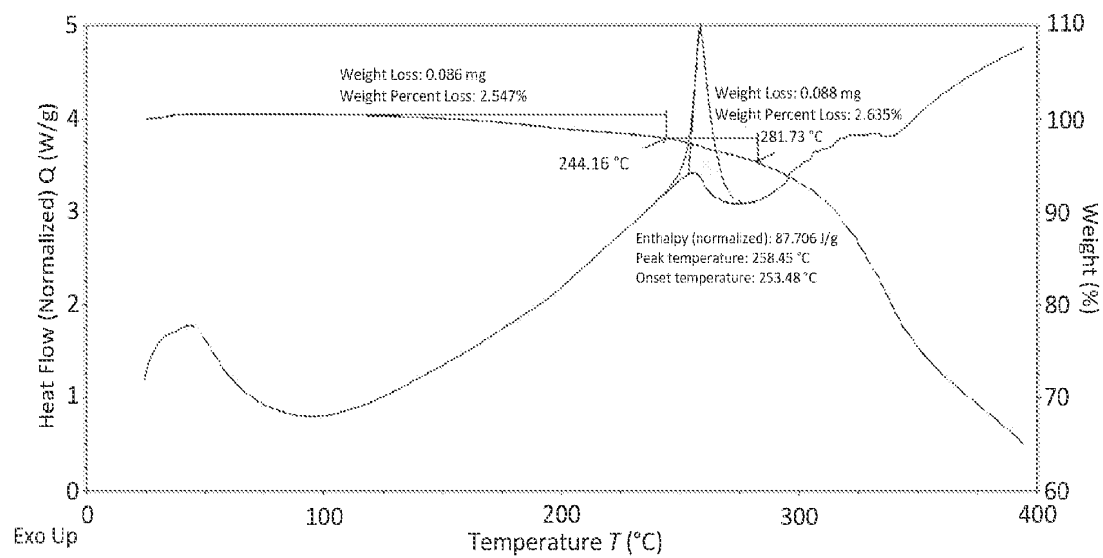


FIG. 110

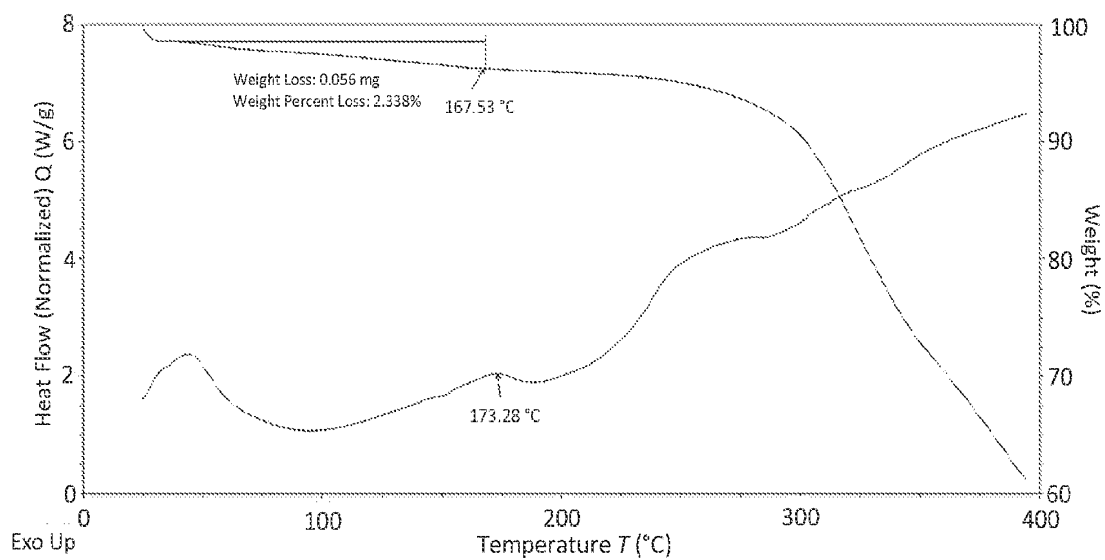


FIG. 111

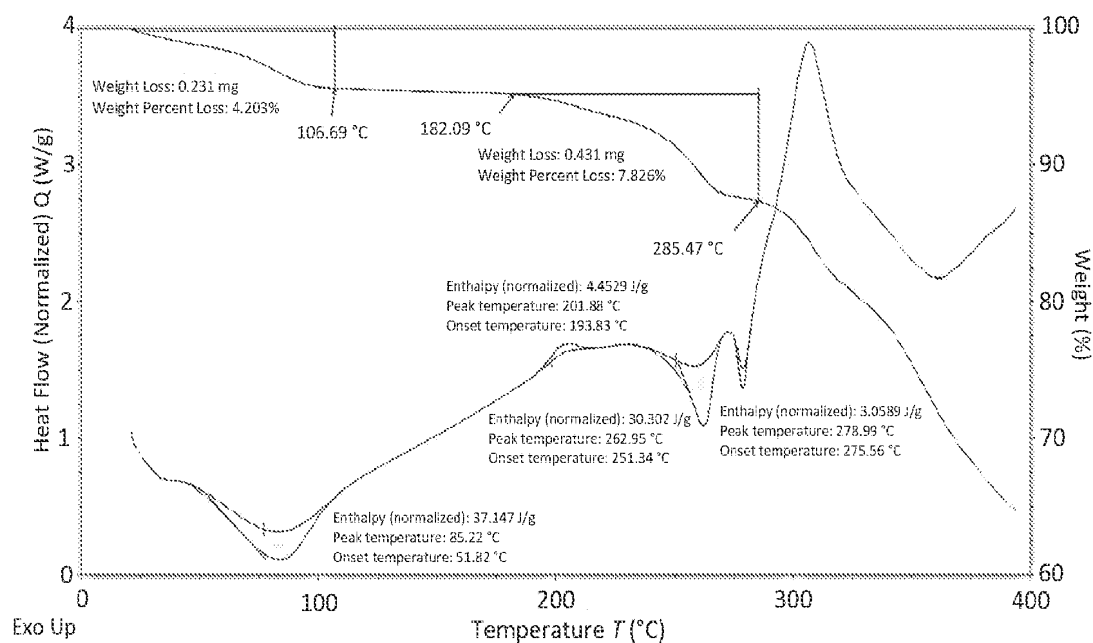


FIG. 112

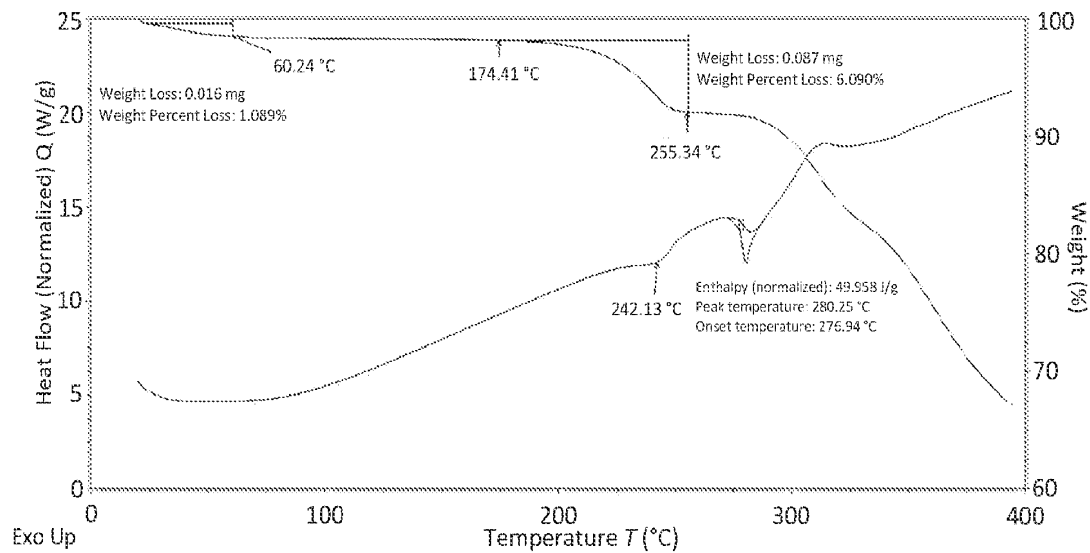


FIG. 113

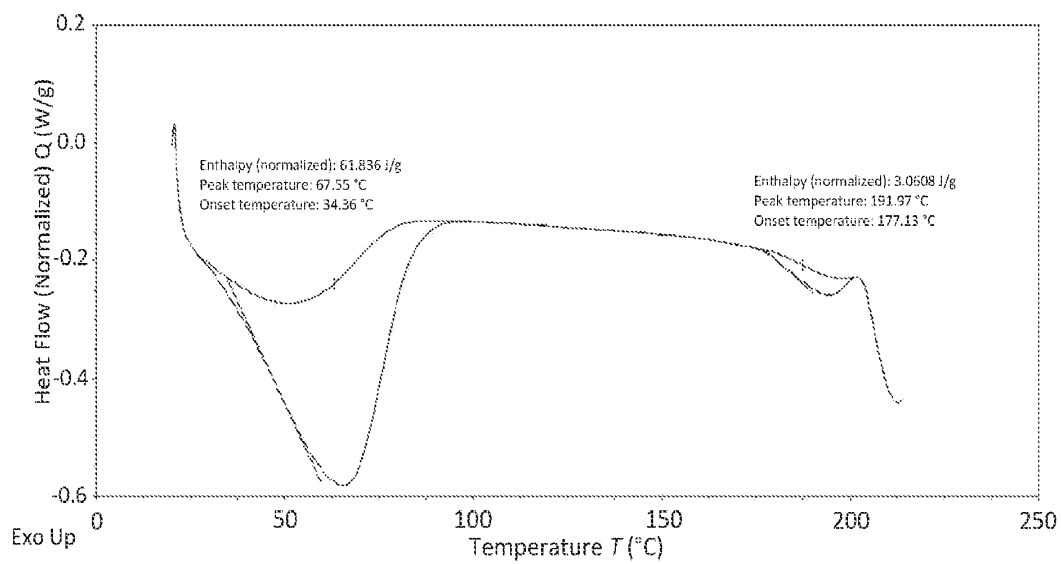


FIG. 114

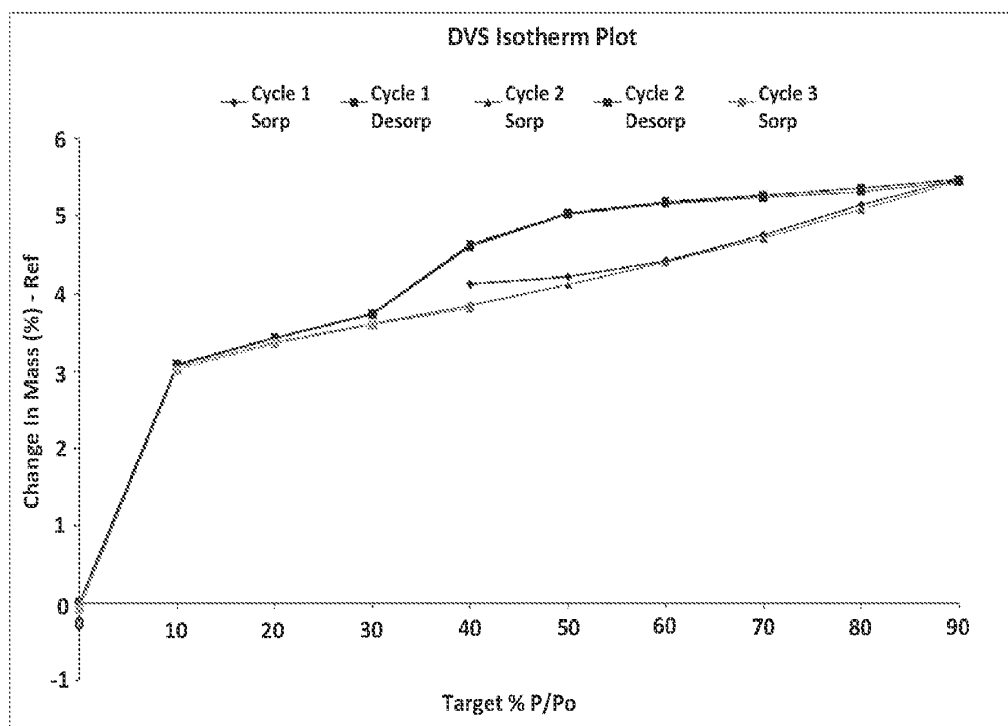


FIG. 115

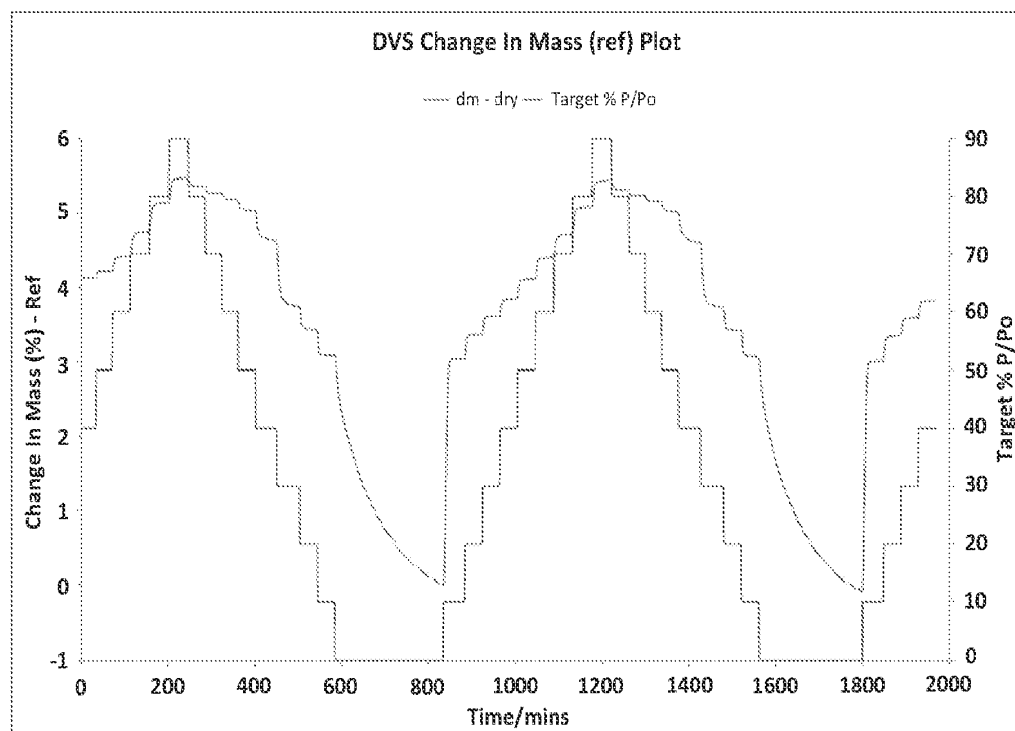


FIG. 116

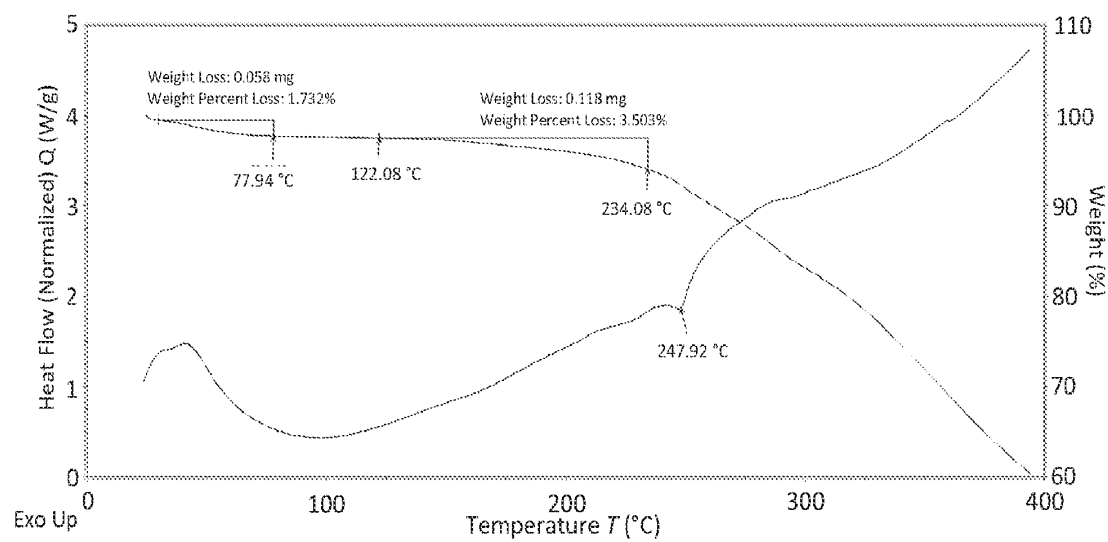


FIG. 117

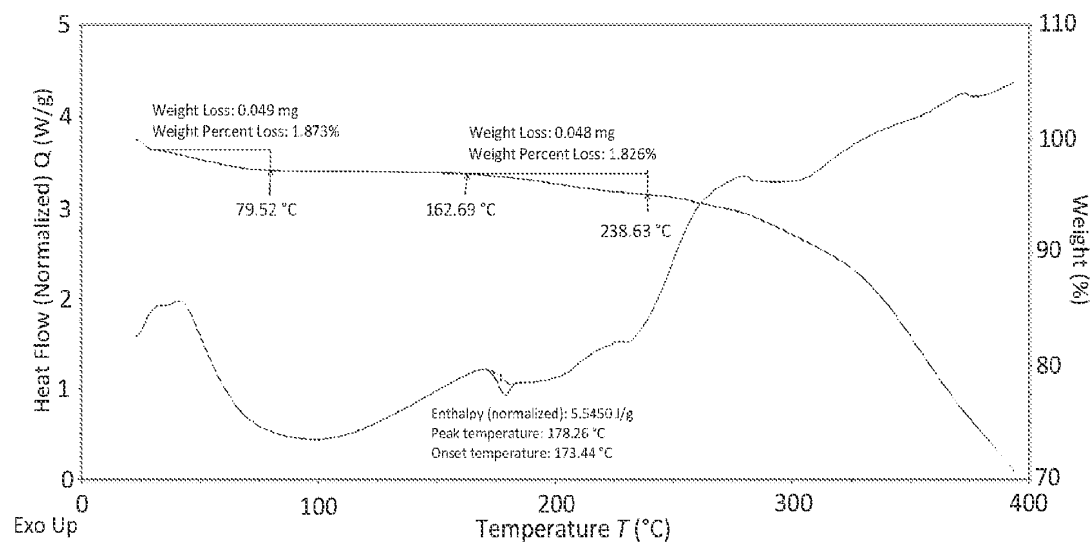


FIG. 118

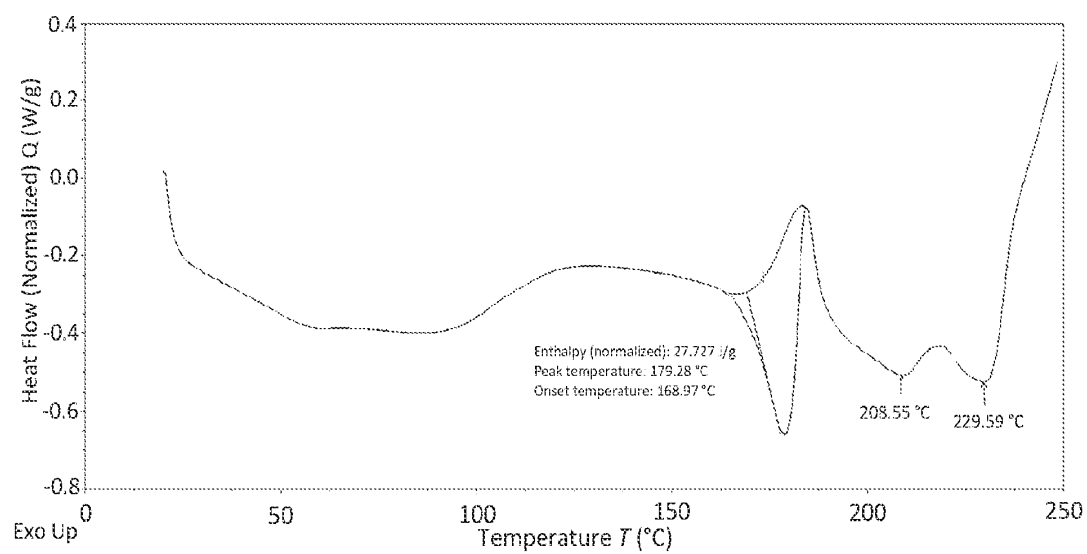


FIG. 119

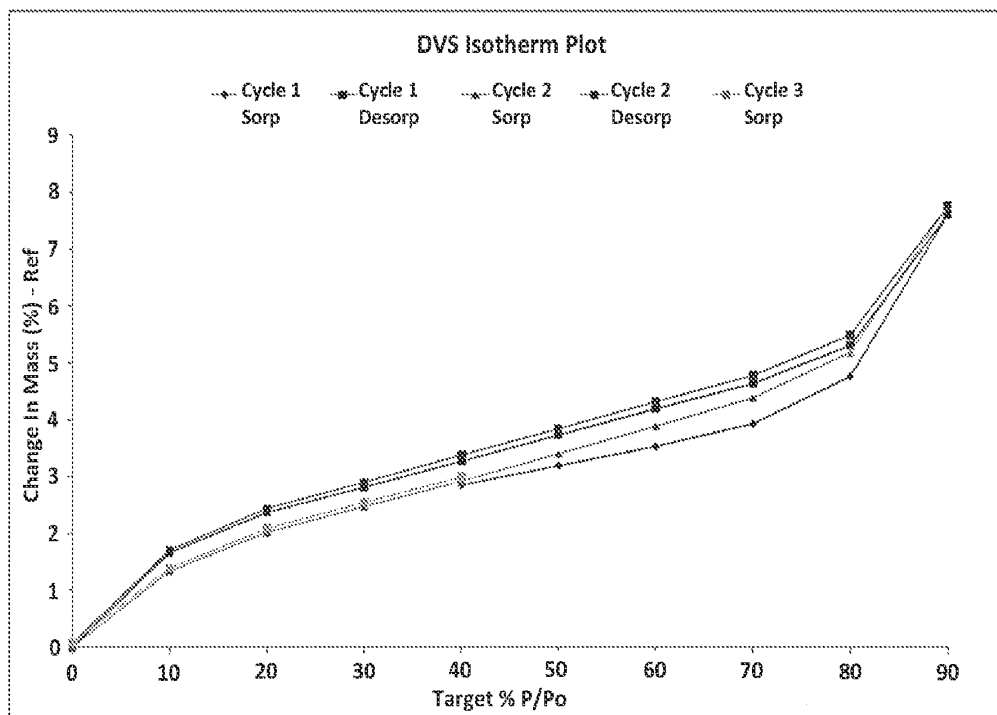


FIG. 120

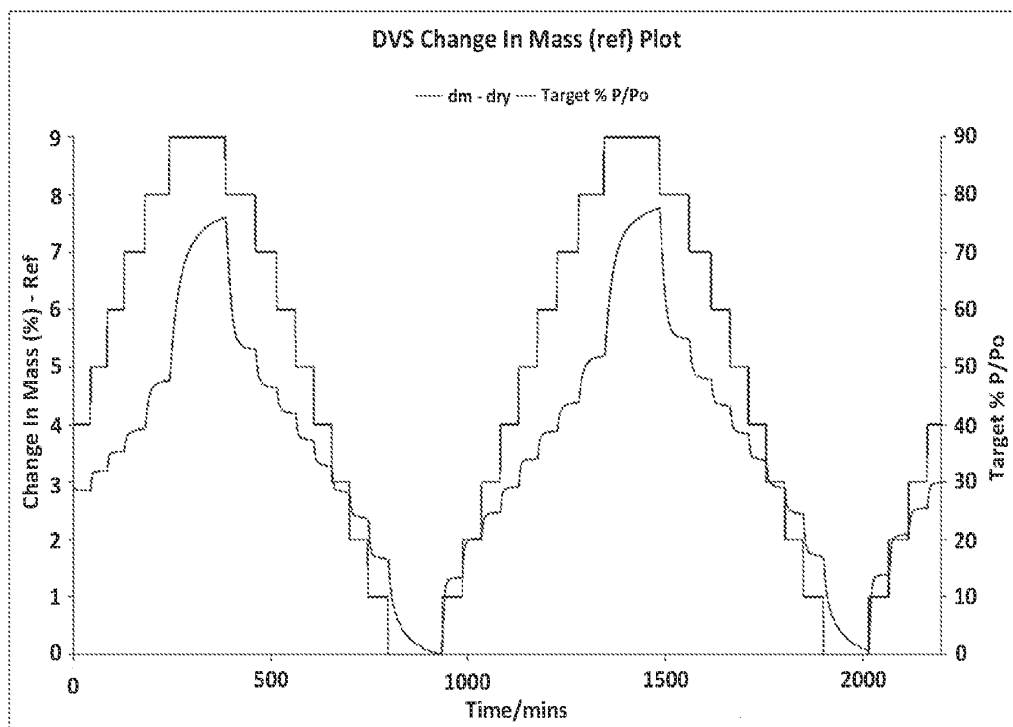


FIG. 121

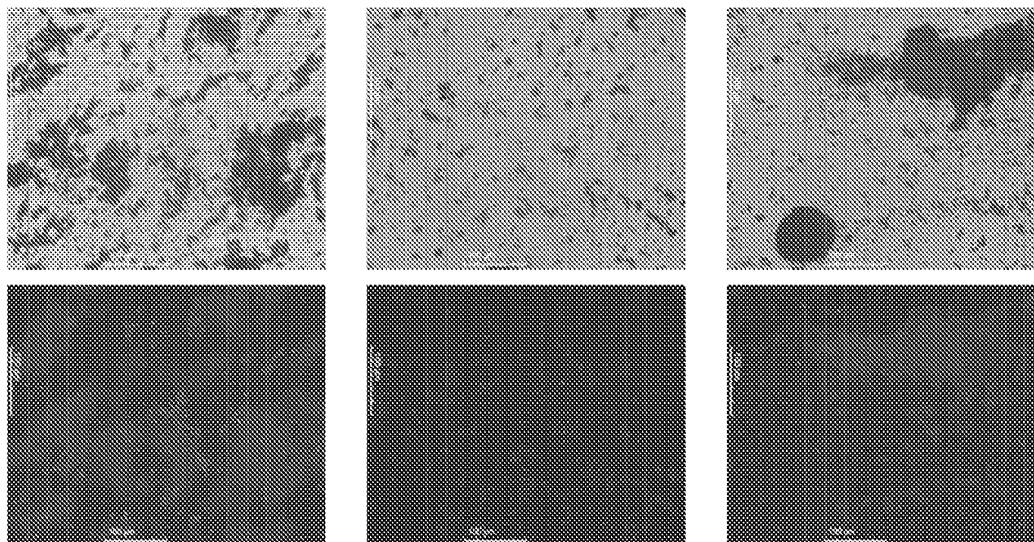


FIG. 122

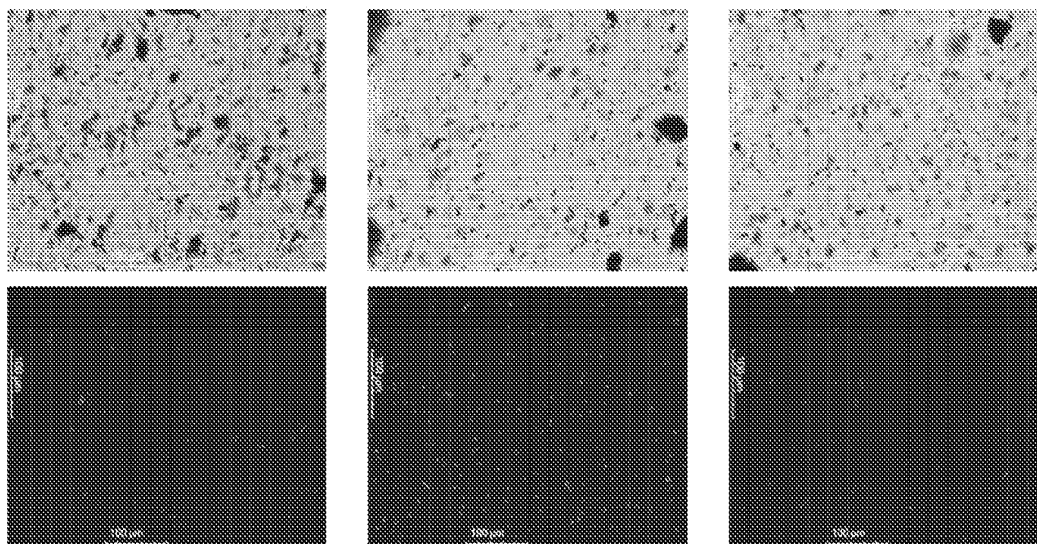


FIG. 123

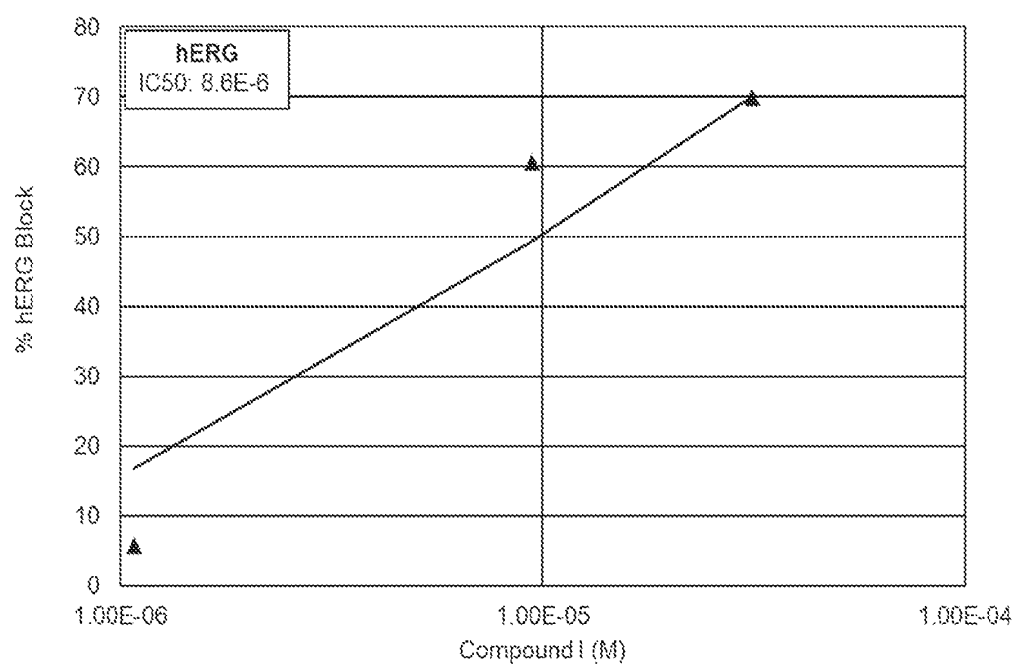


FIG. 124

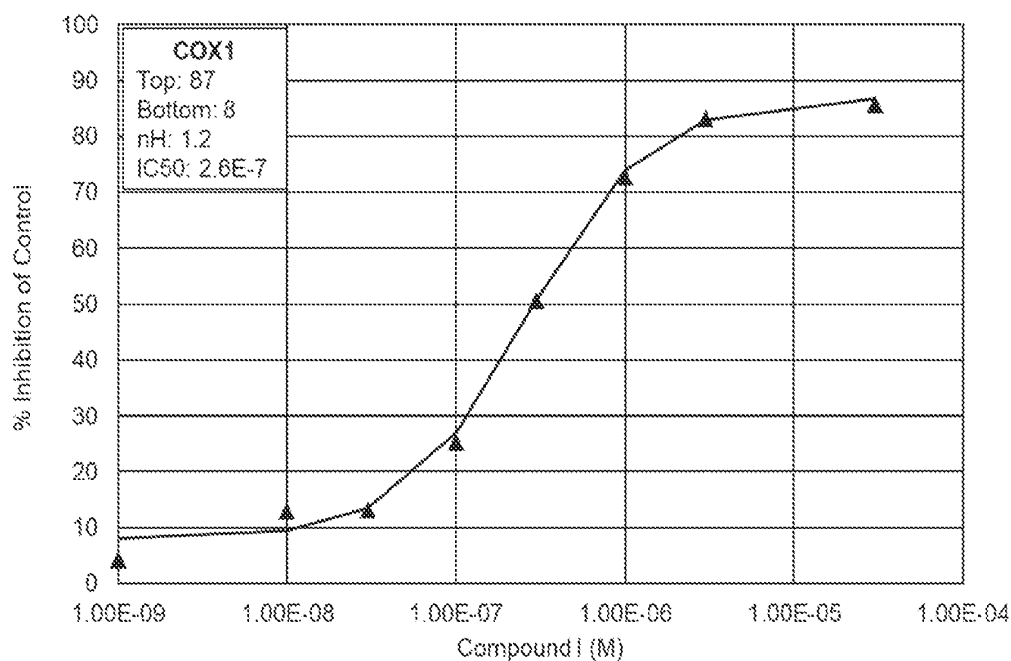


FIG. 125

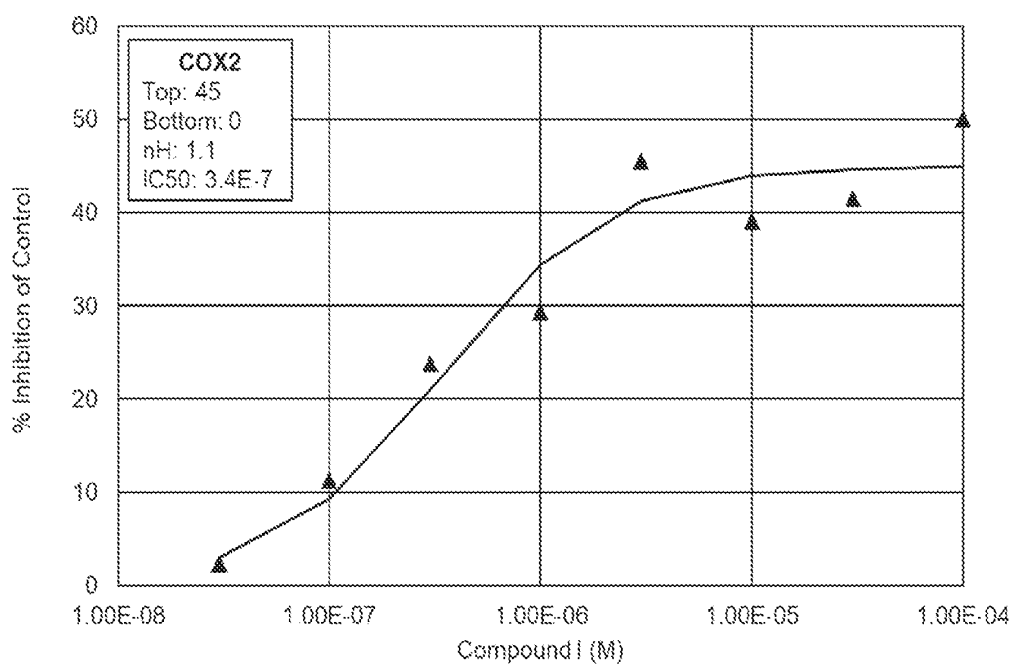


FIG. 126

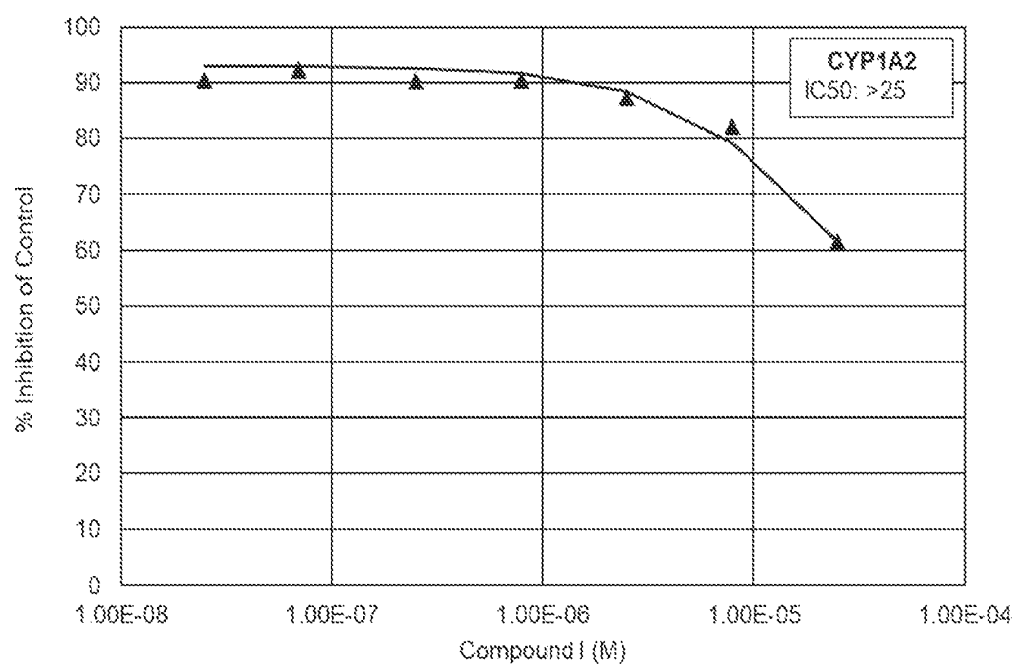


FIG. 127

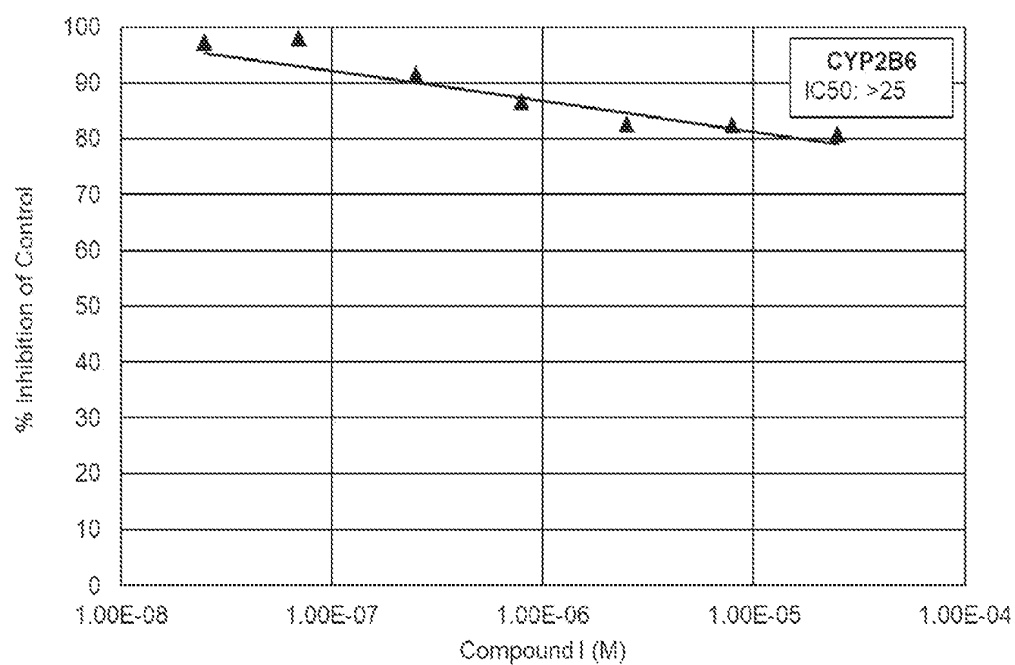


FIG. 128

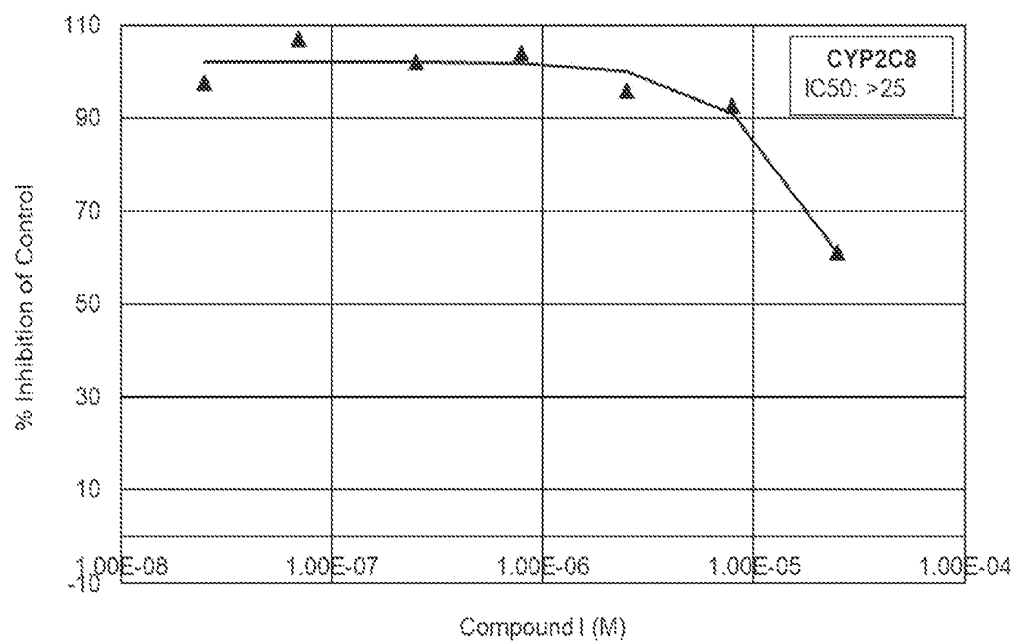


FIG. 129

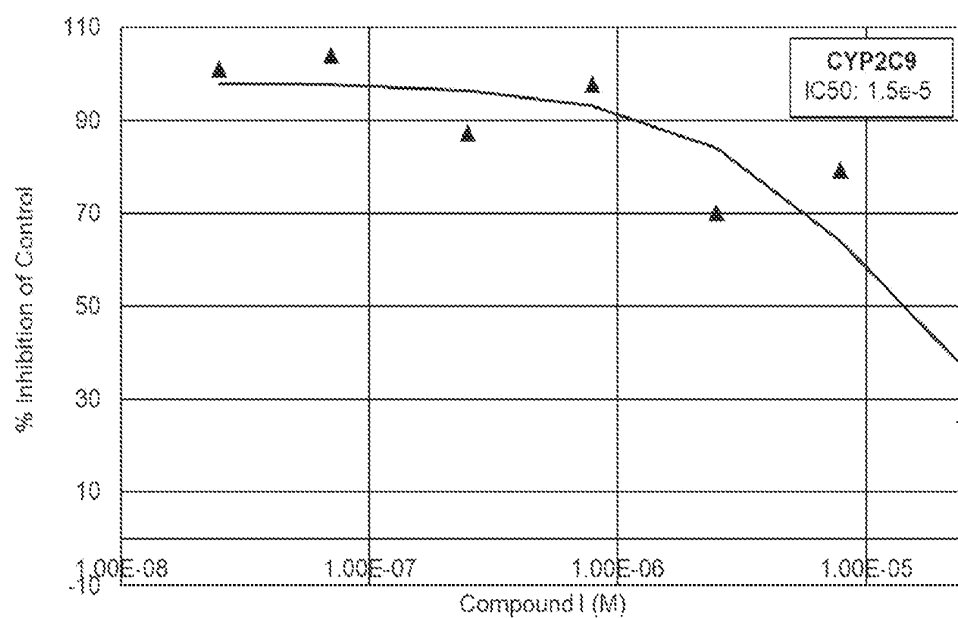


FIG. 130

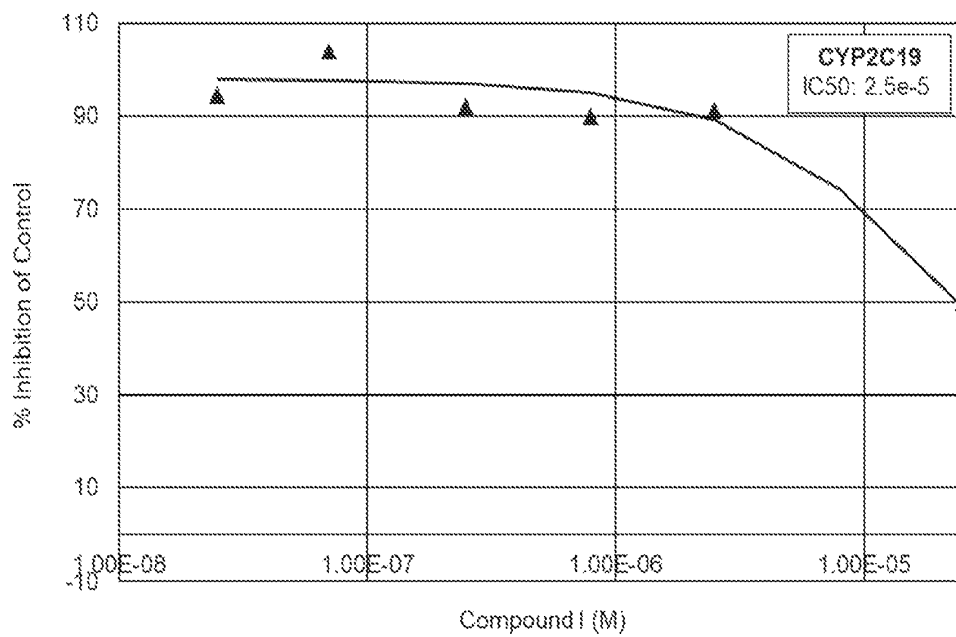


FIG. 131

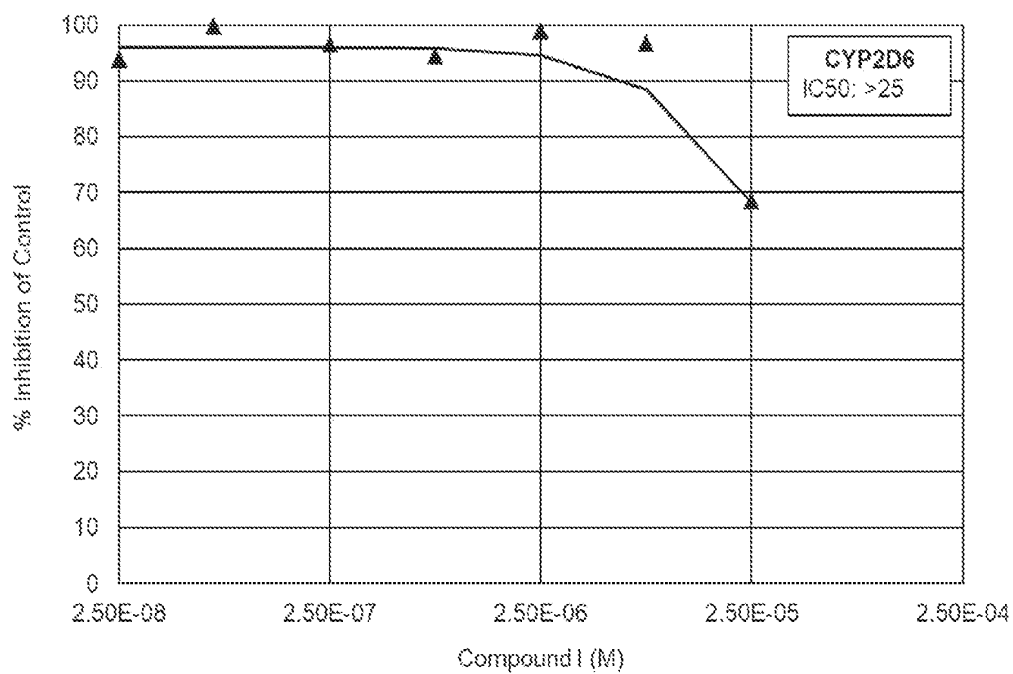


FIG. 132

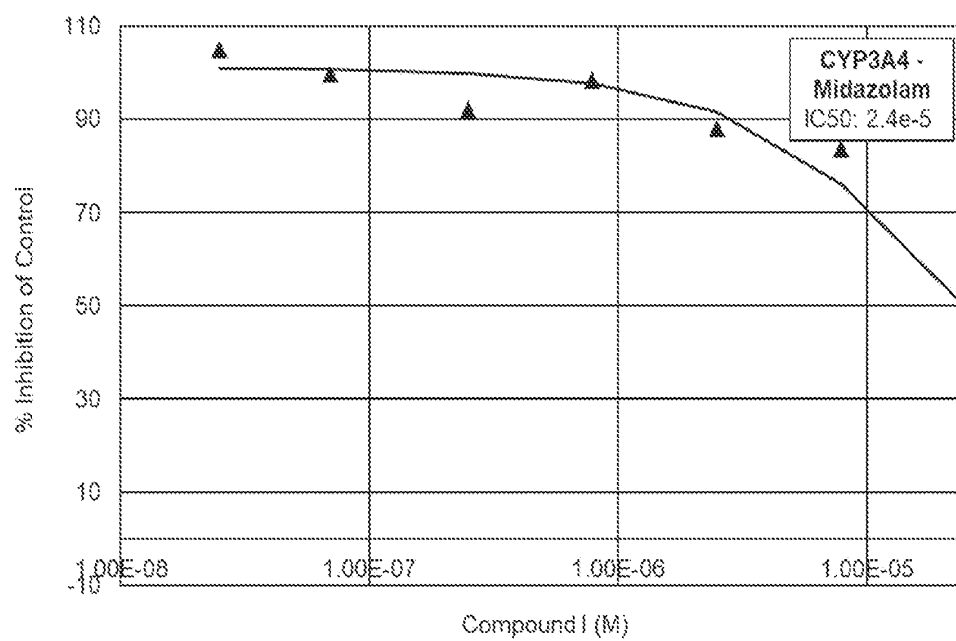


FIG. 133

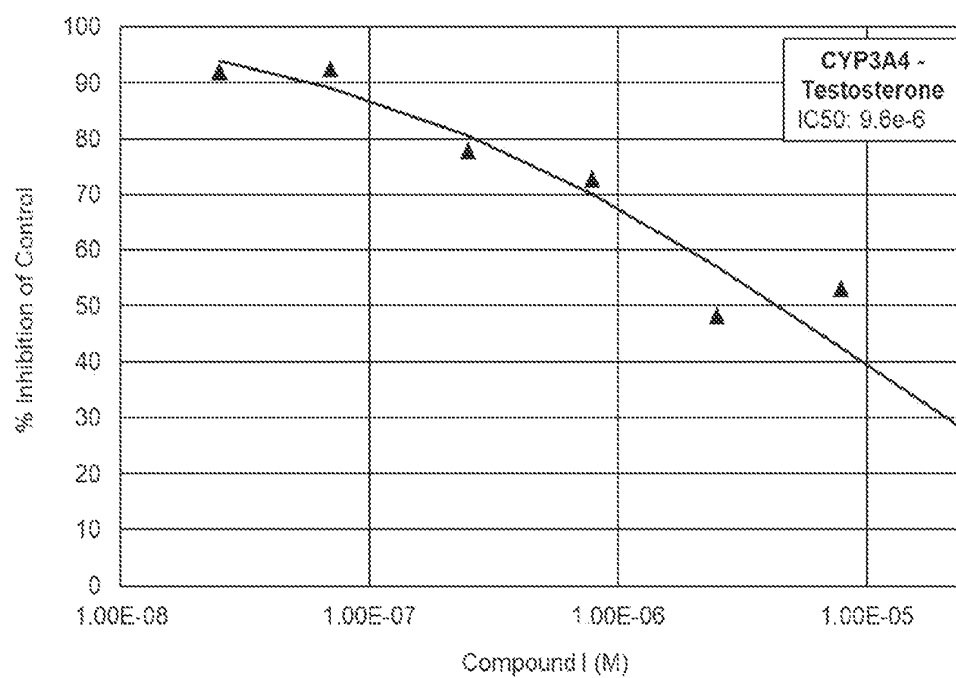


FIG. 134

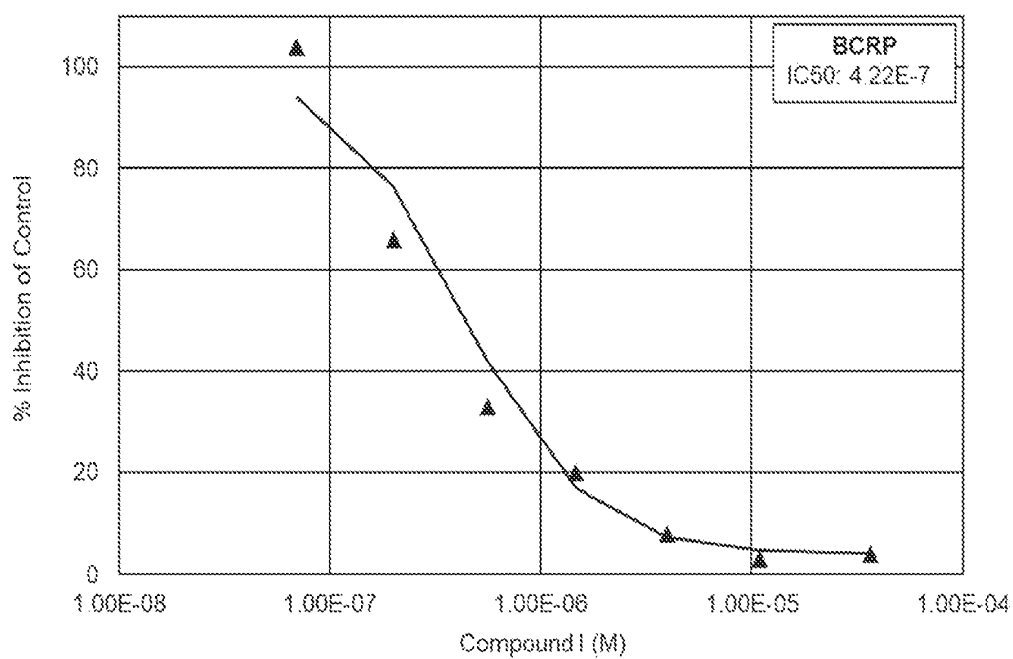


FIG. 135

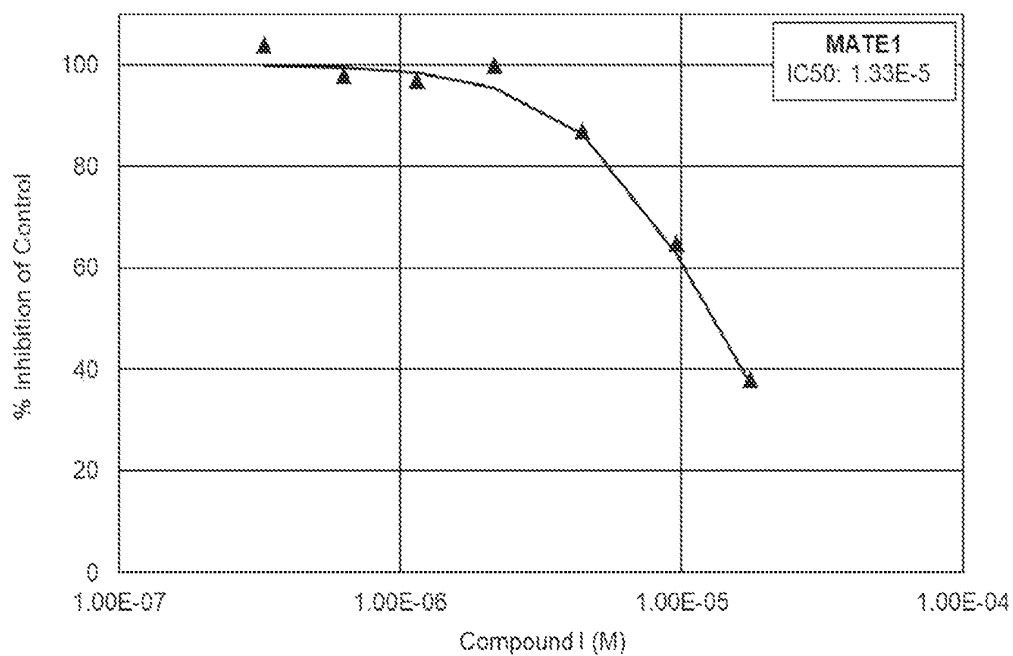


FIG. 136

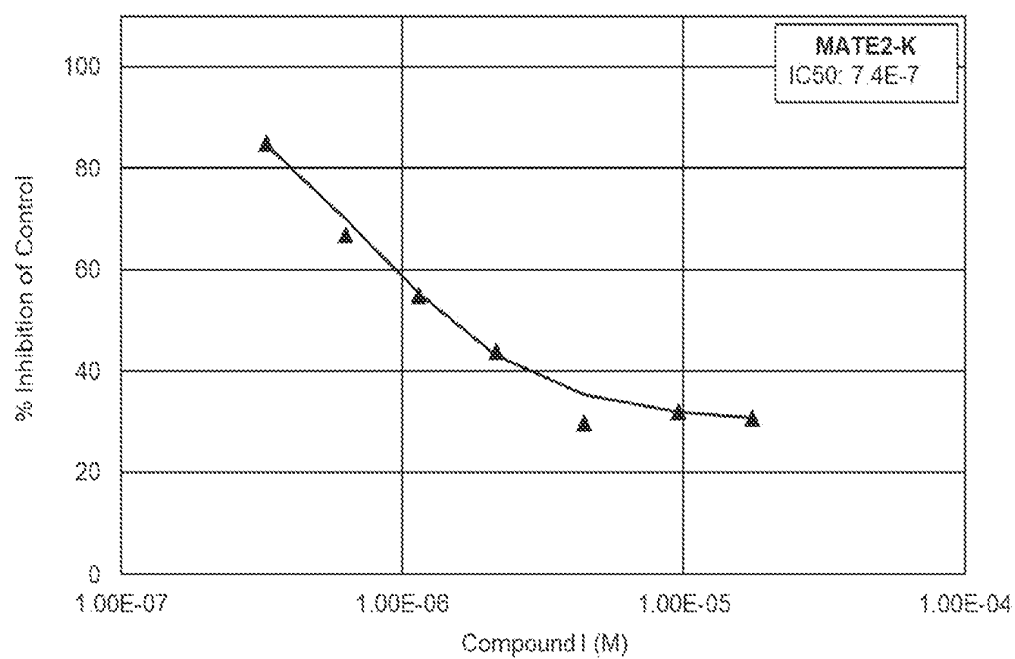


FIG. 137

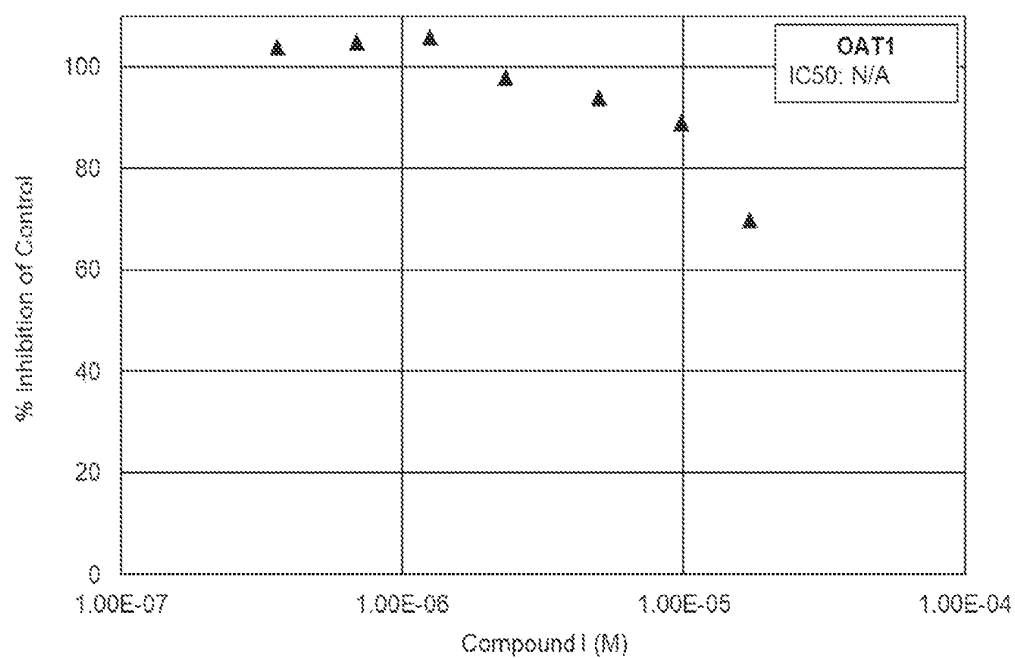


FIG. 138

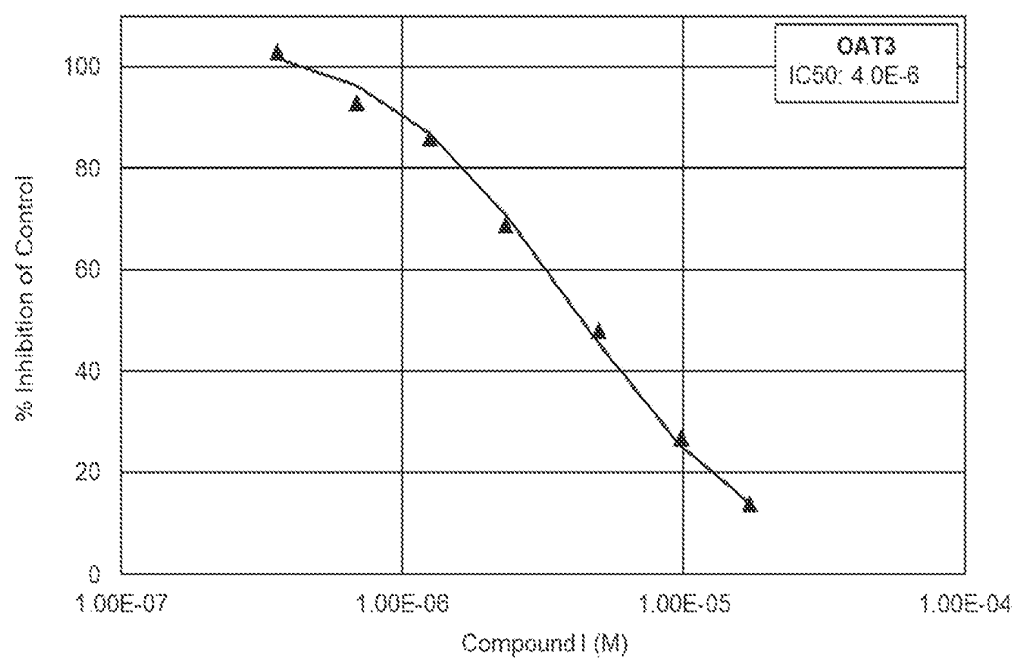


FIG. 139

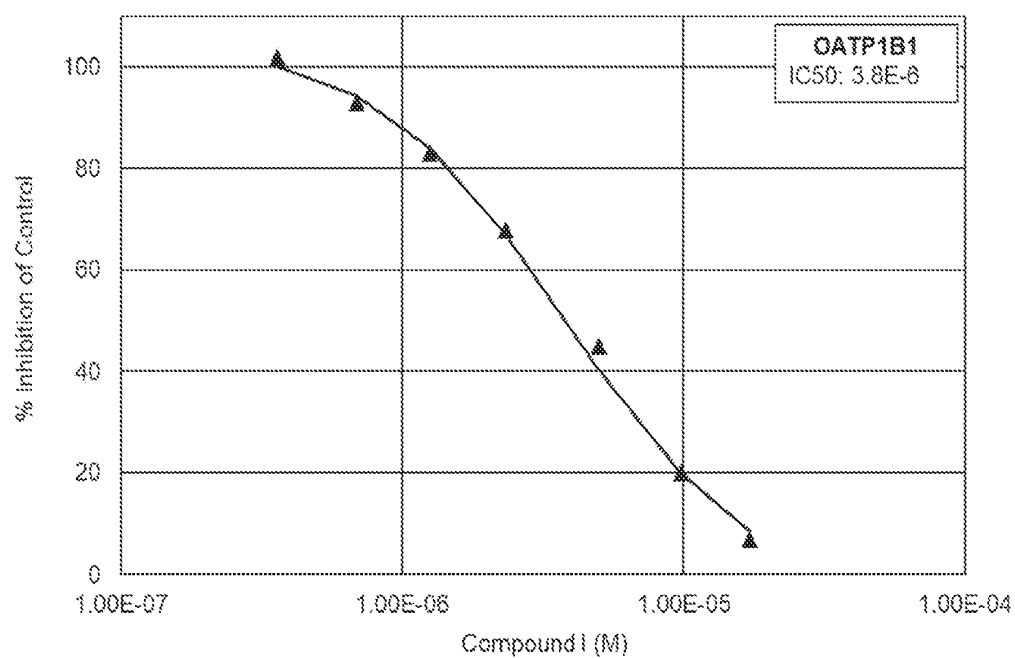


FIG. 140

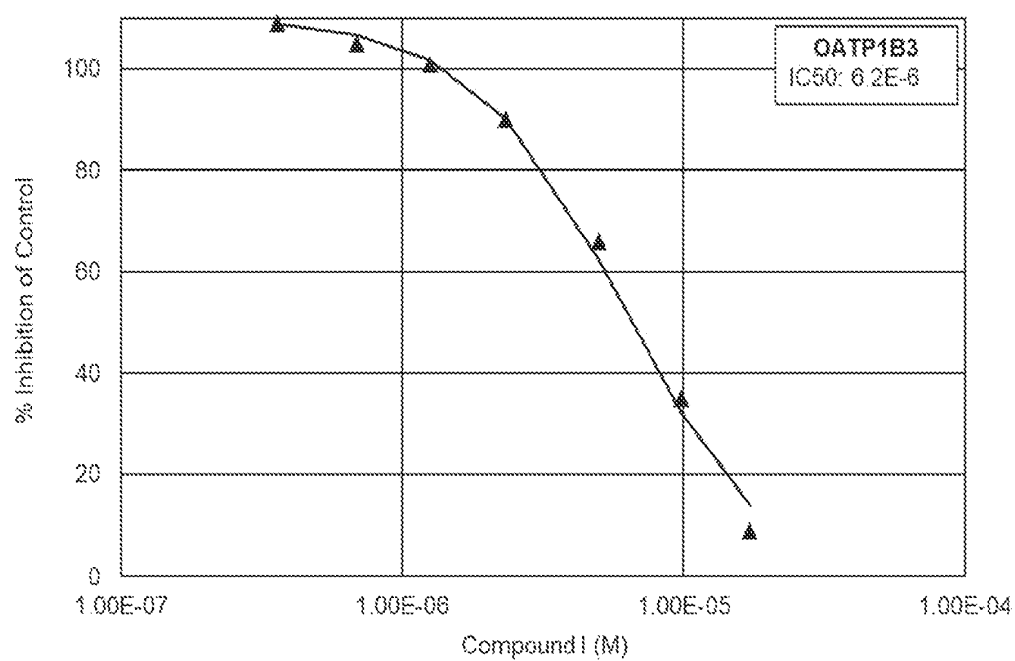


FIG. 141

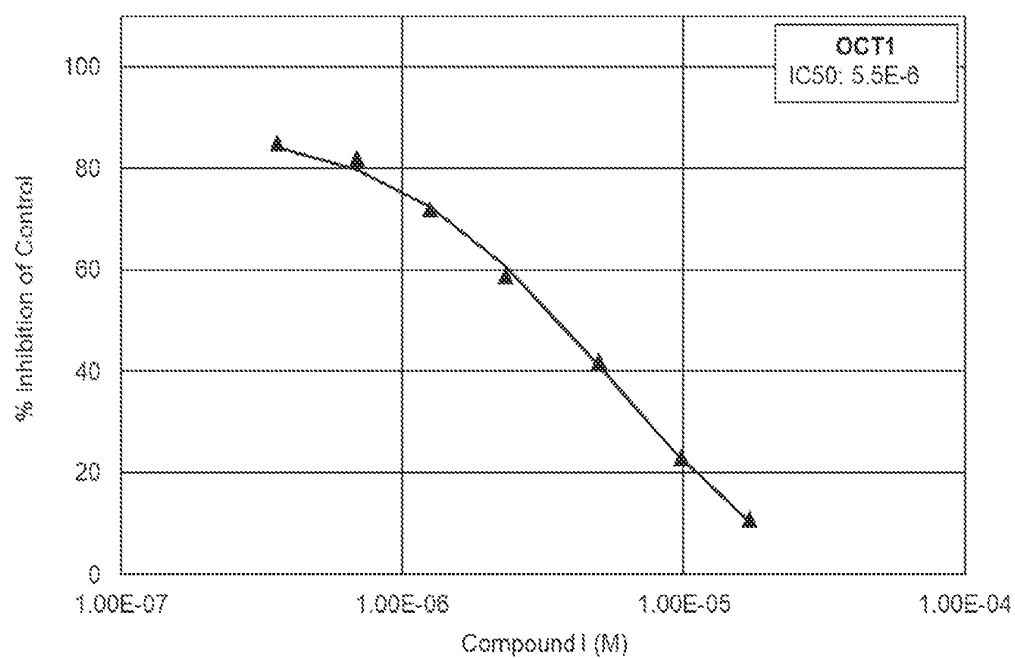


FIG. 142

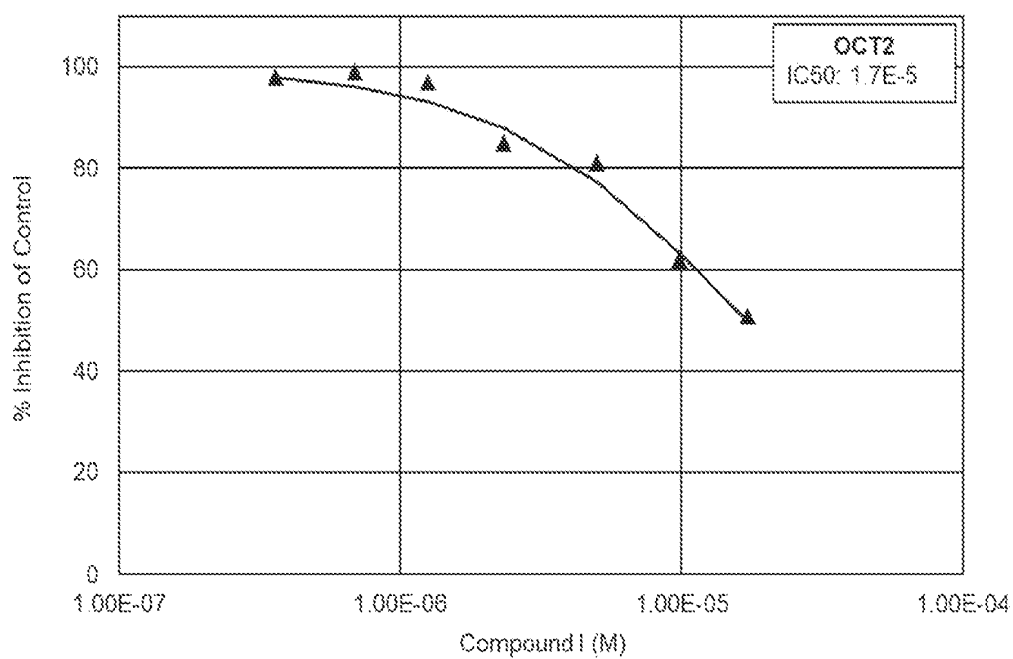


FIG. 143

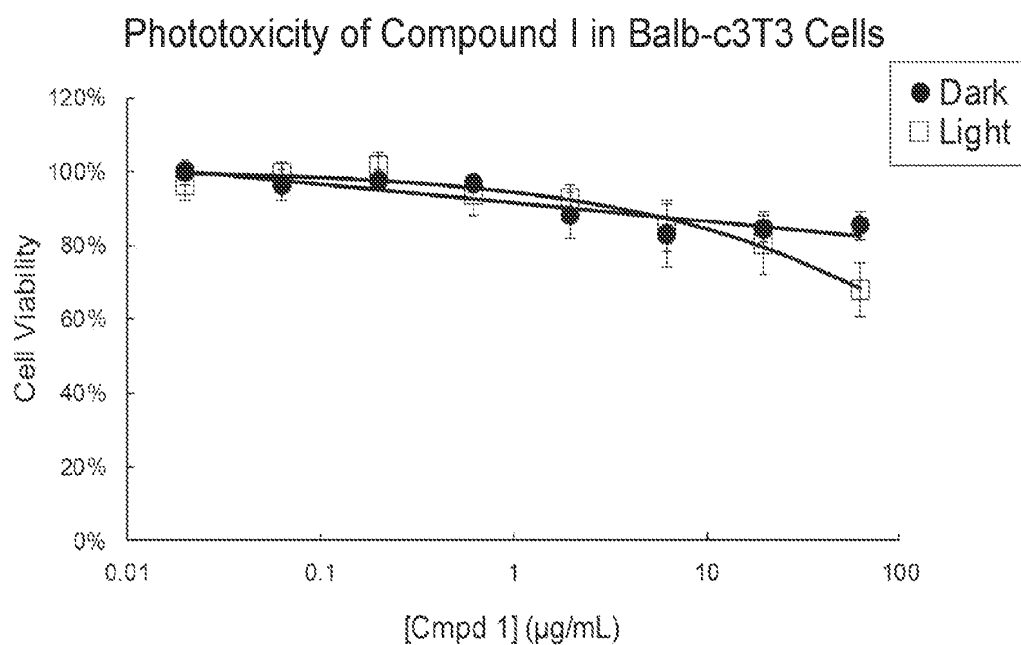


FIG. 144

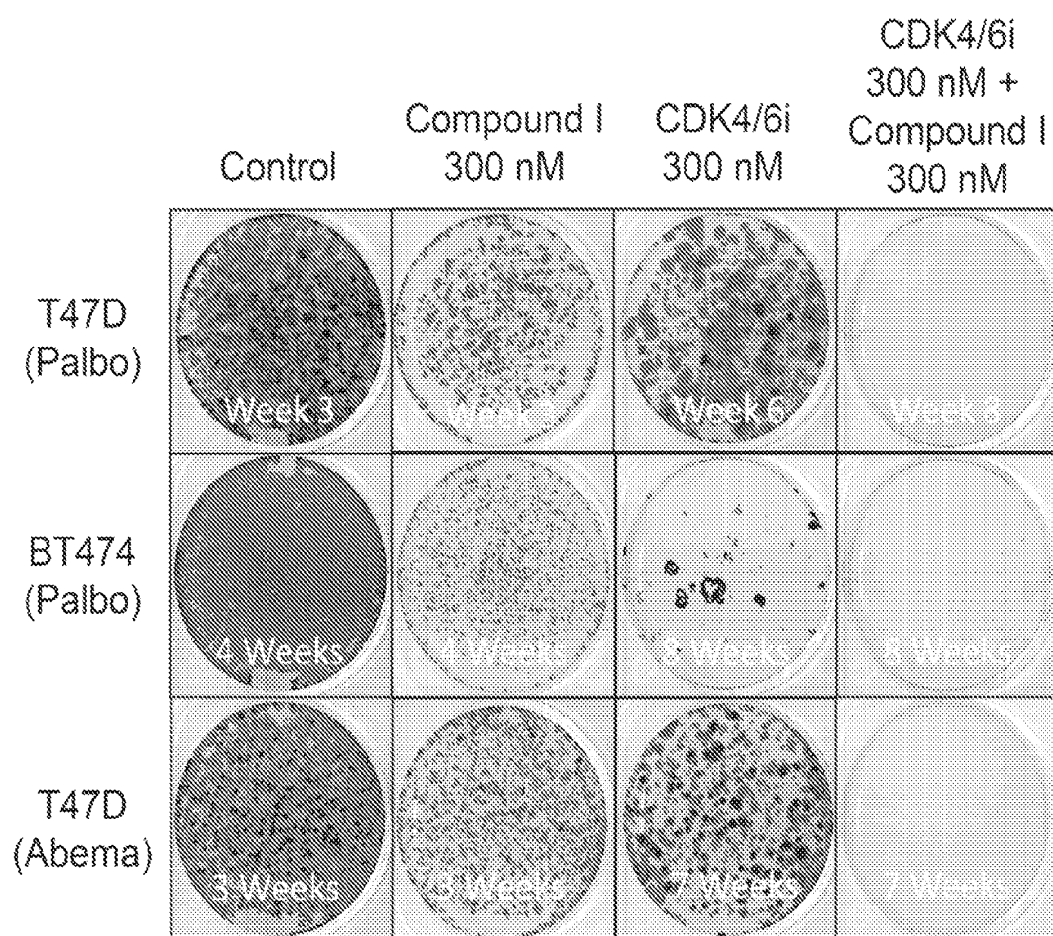


FIG. 145

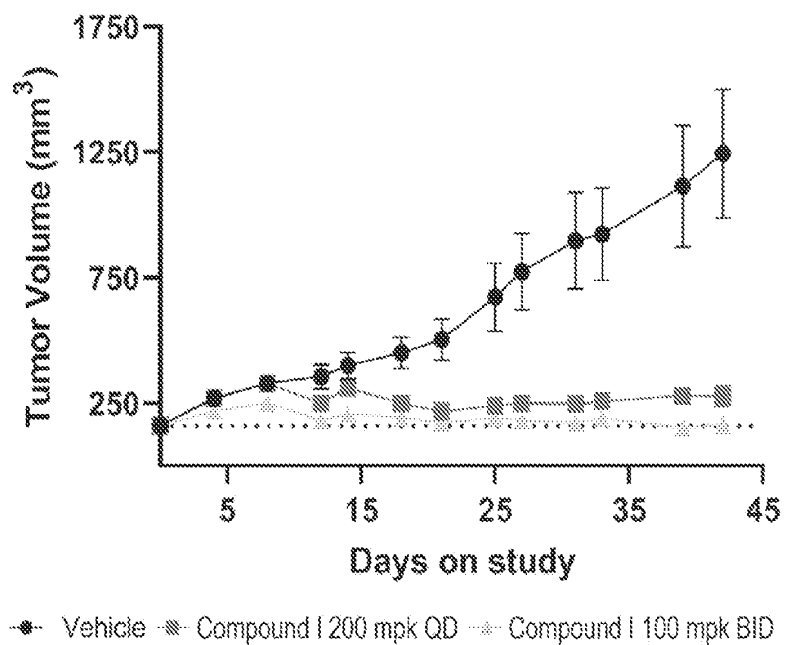
OVCAR3 CDX

FIG. 146A

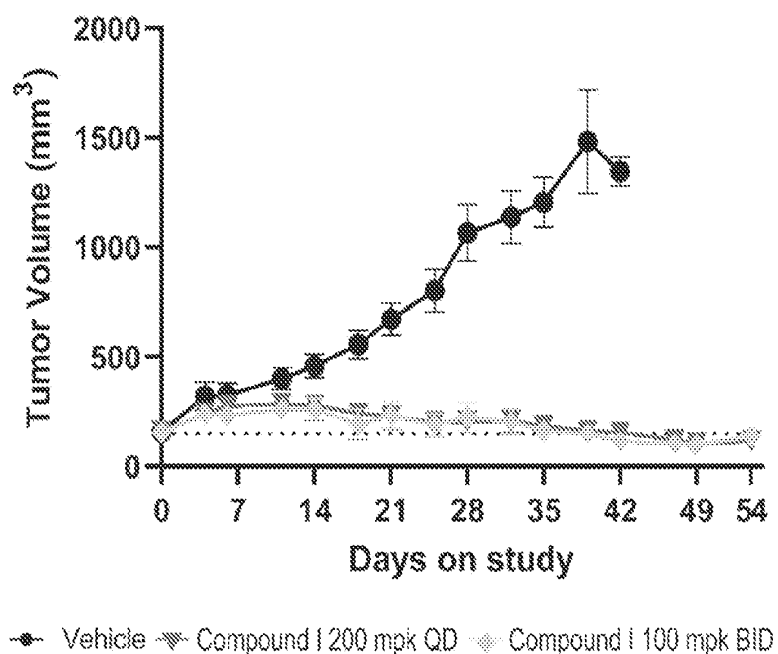
OV5398 PDX

FIG. 146B

GA0103 PDX

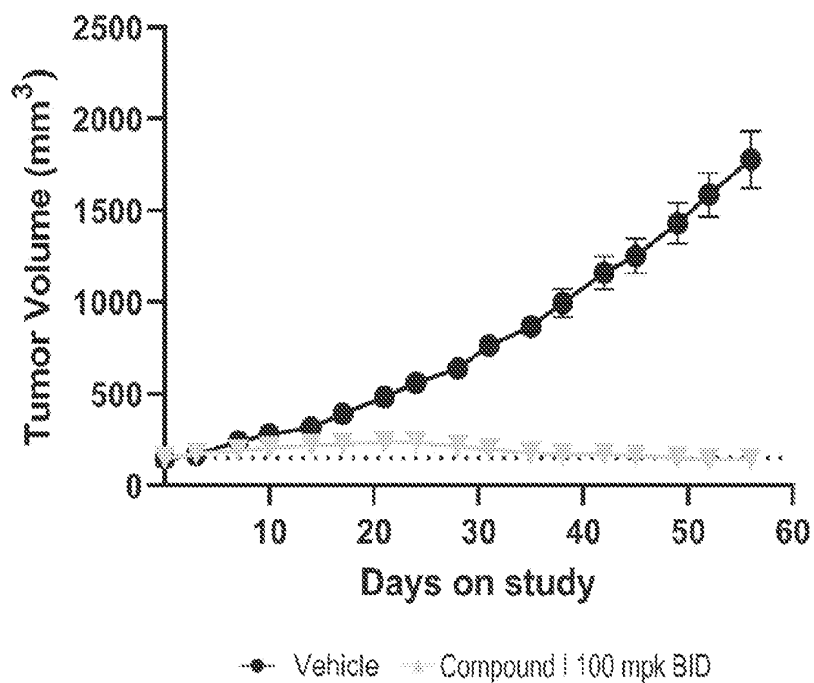


FIG. 146C

GA0114 PDX

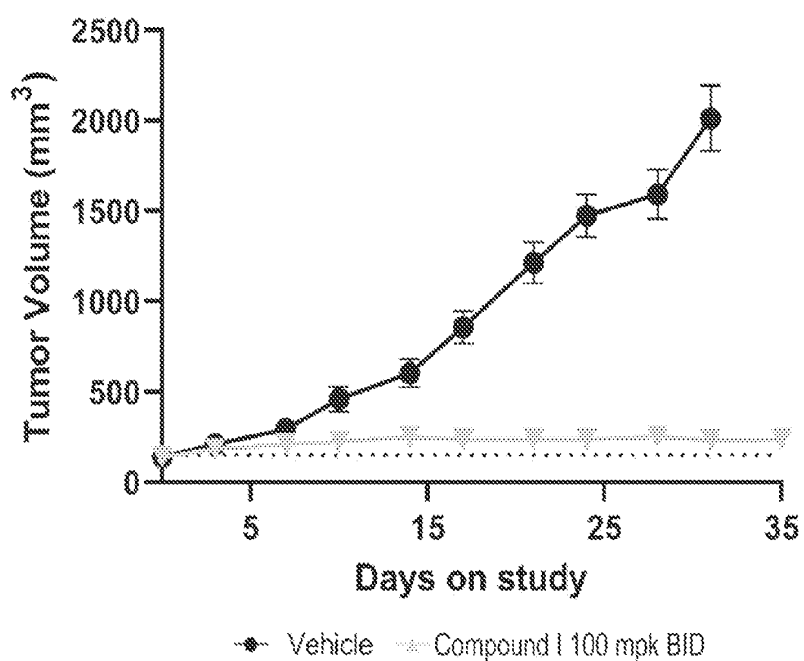


FIG. 146D

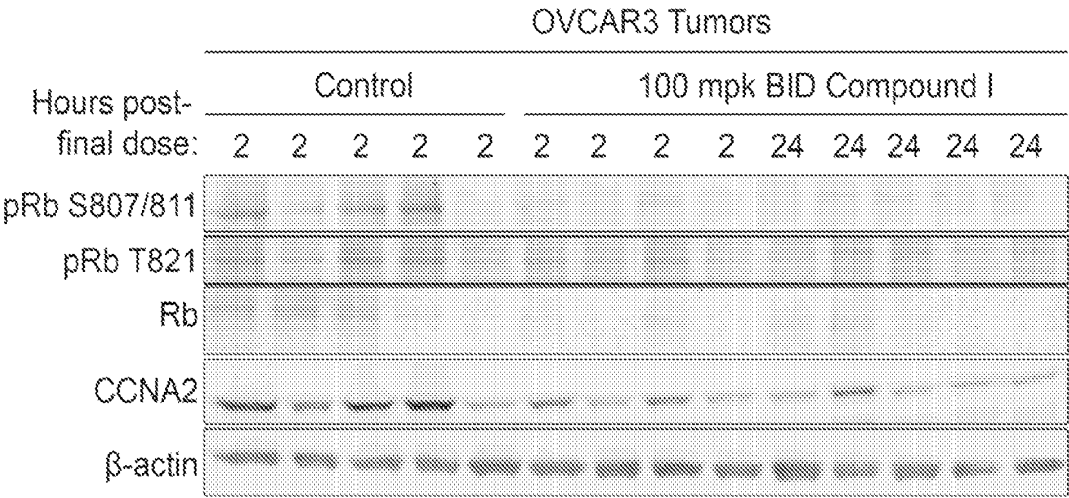


FIG. 146E

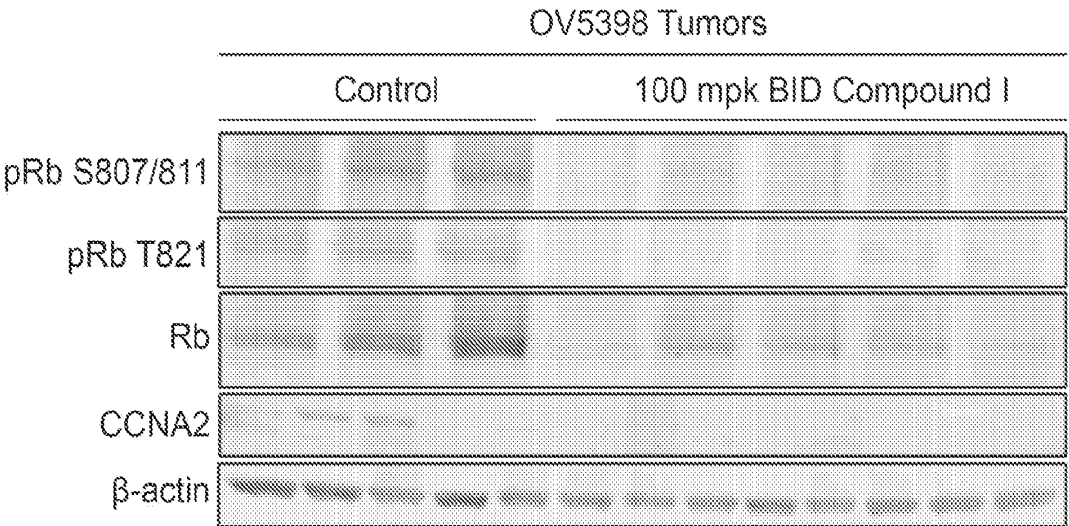


FIG. 146F

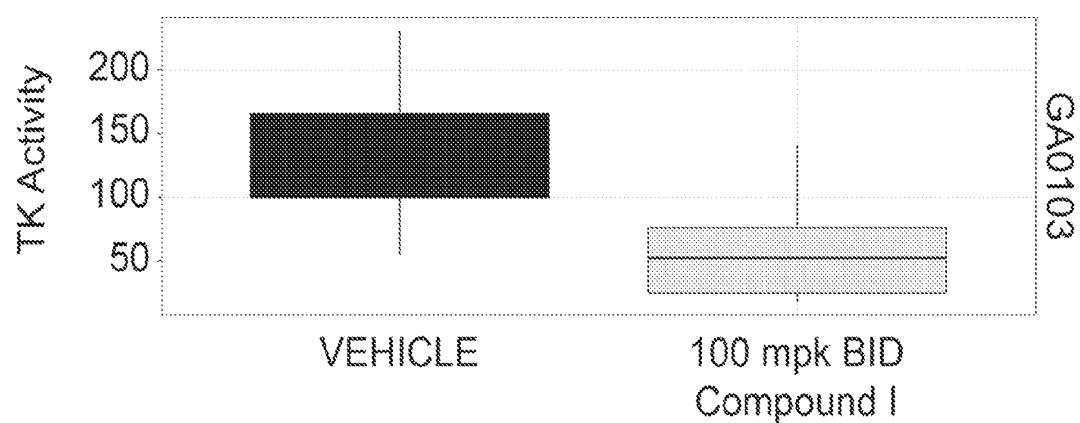


FIG. 146G

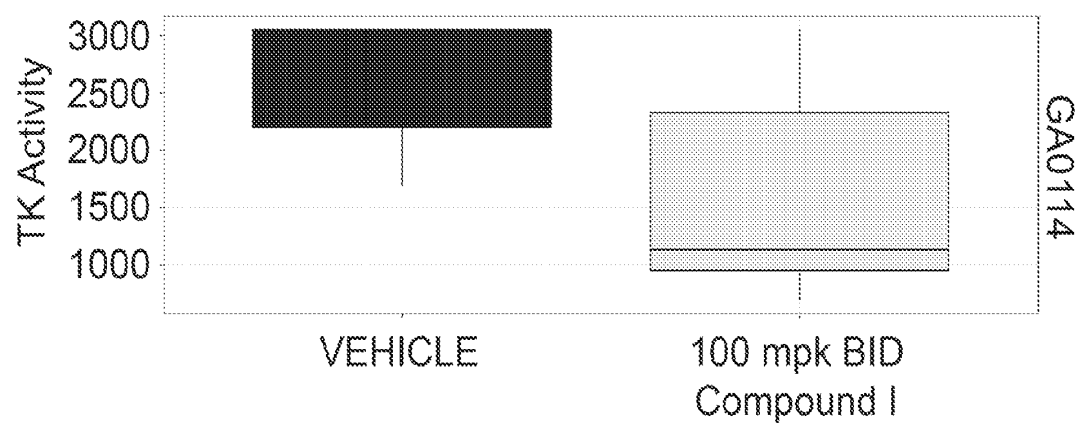


FIG. 146H

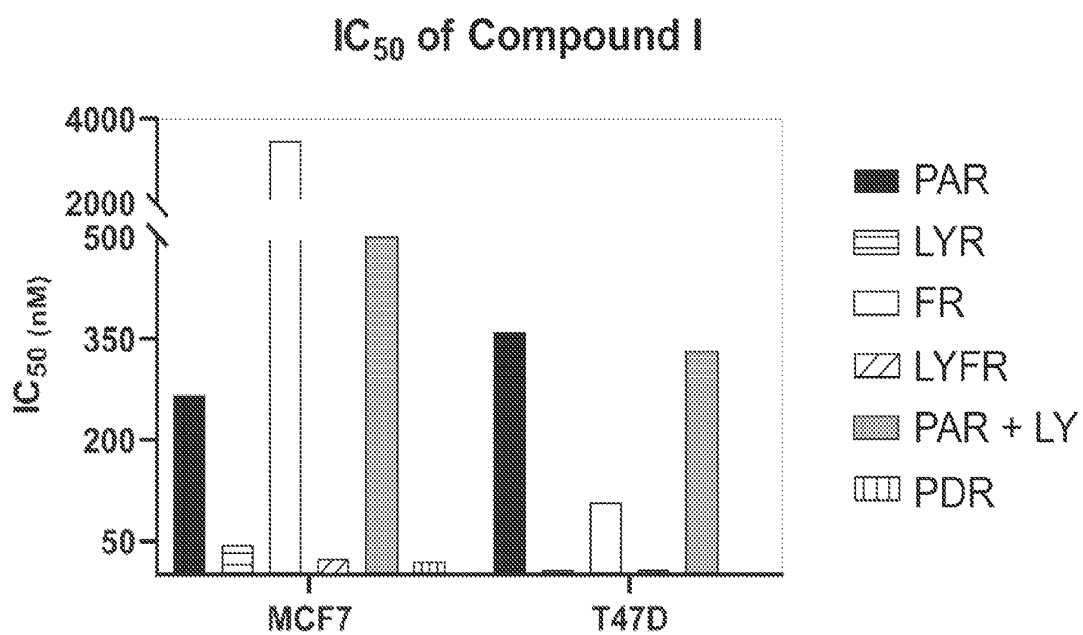


FIG. 147A

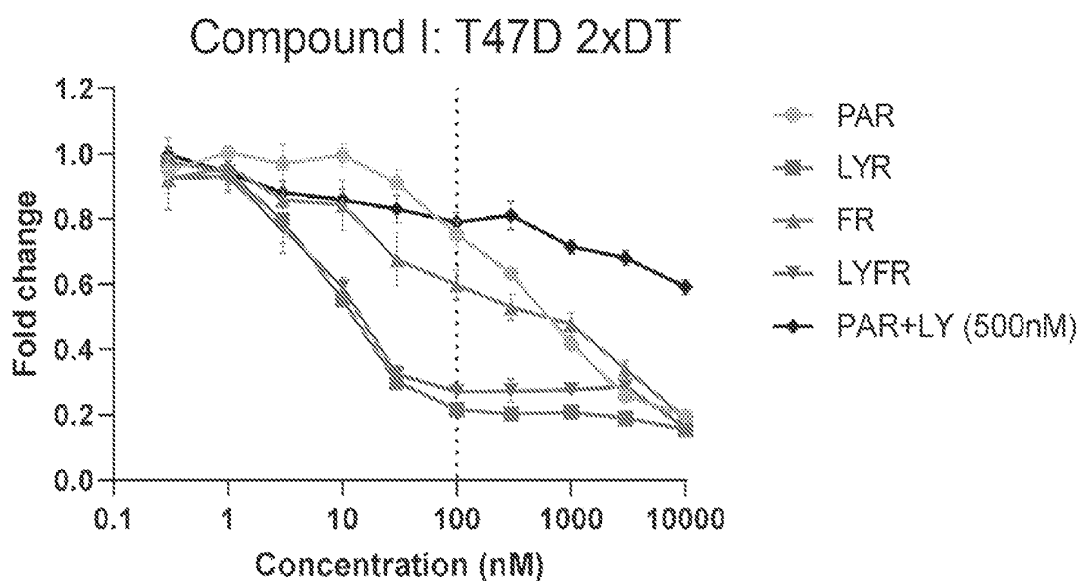


FIG. 147B

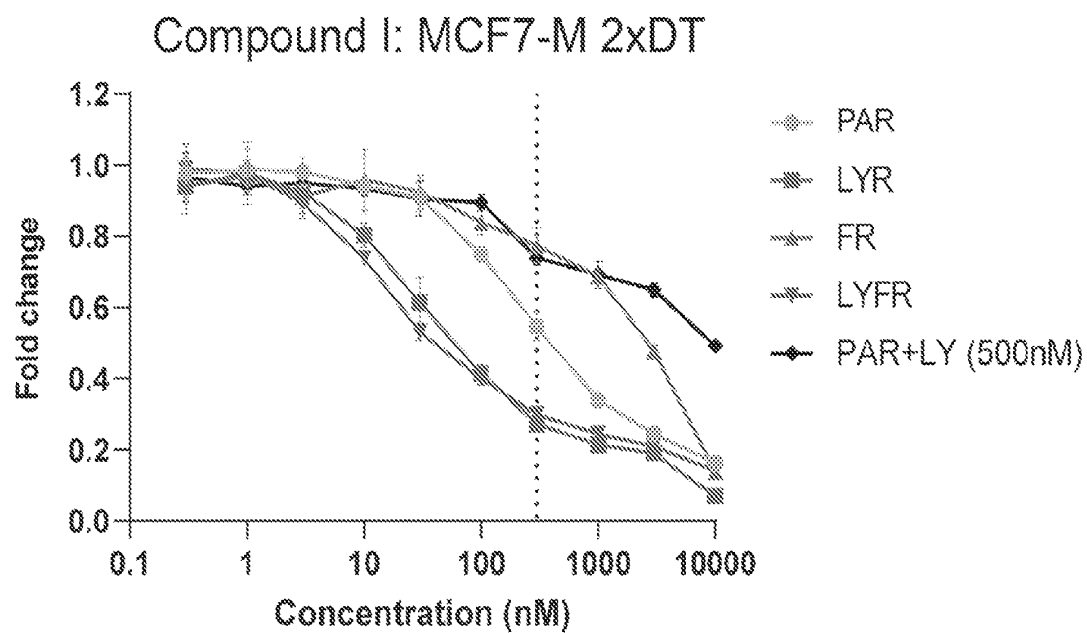


FIG. 147C

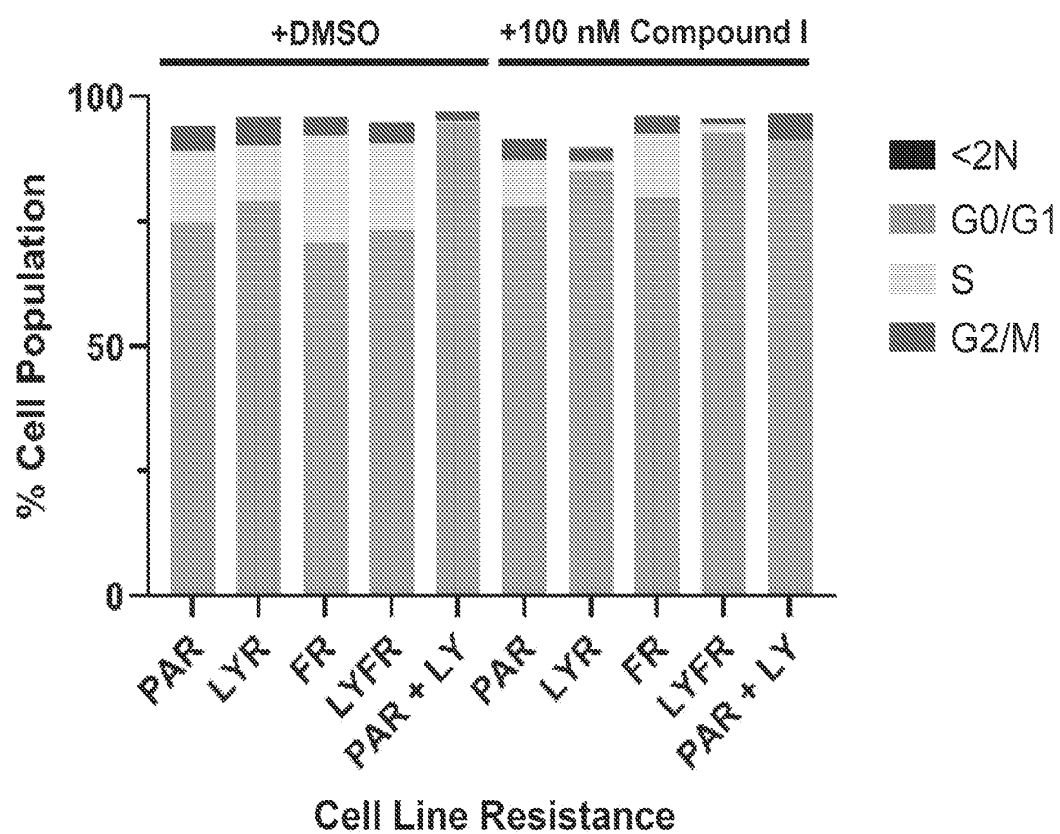


FIG. 147D

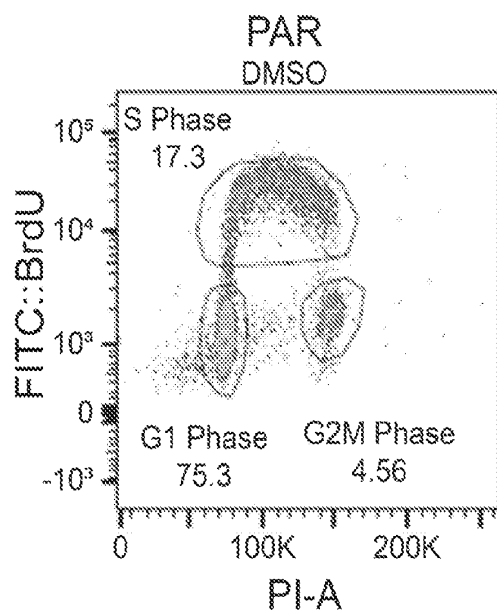


FIG. 147E

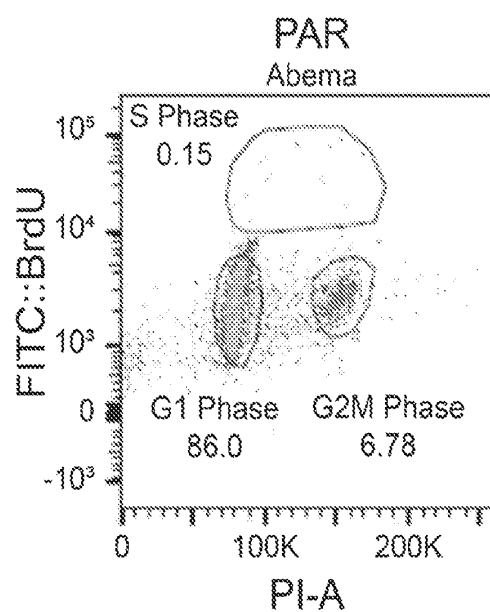


FIG. 147F

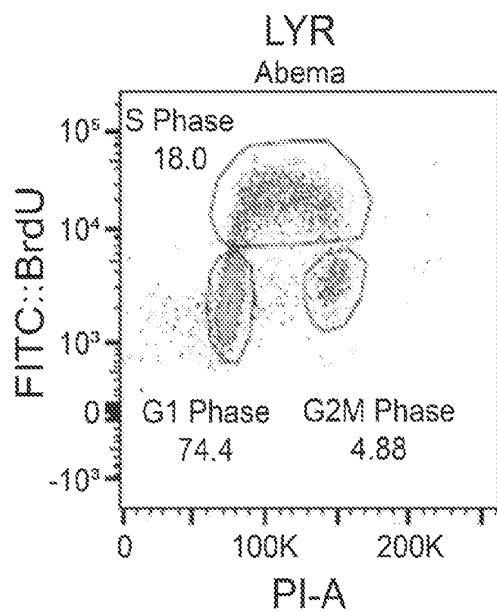


FIG. 147G

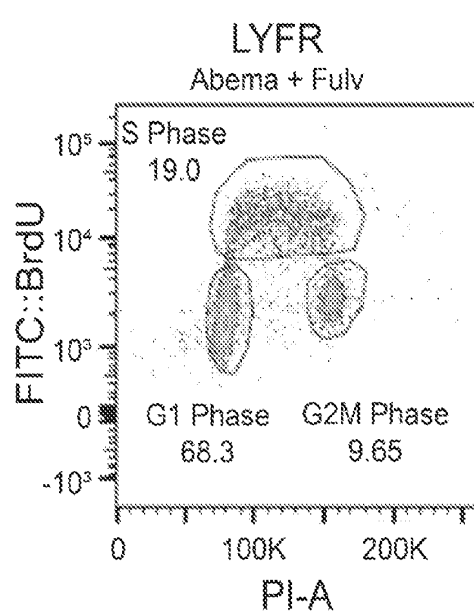


FIG. 147H

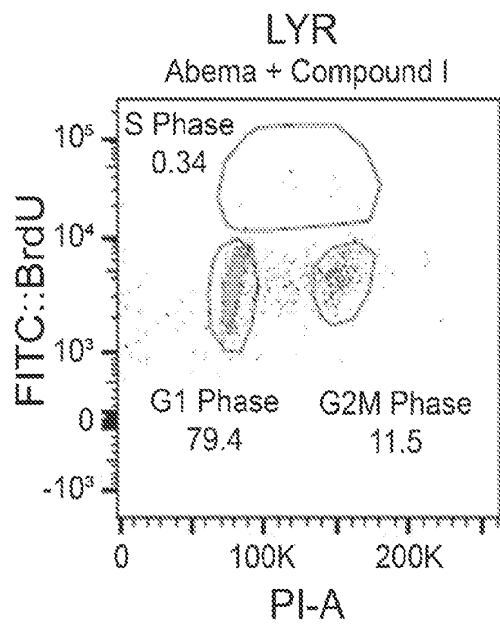


FIG. 147I

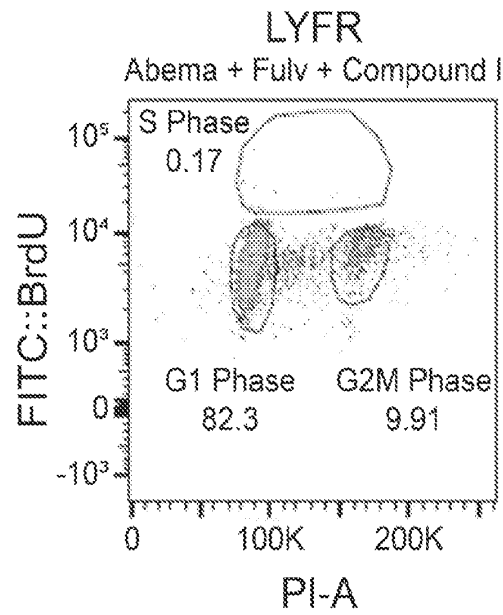


FIG. 147J

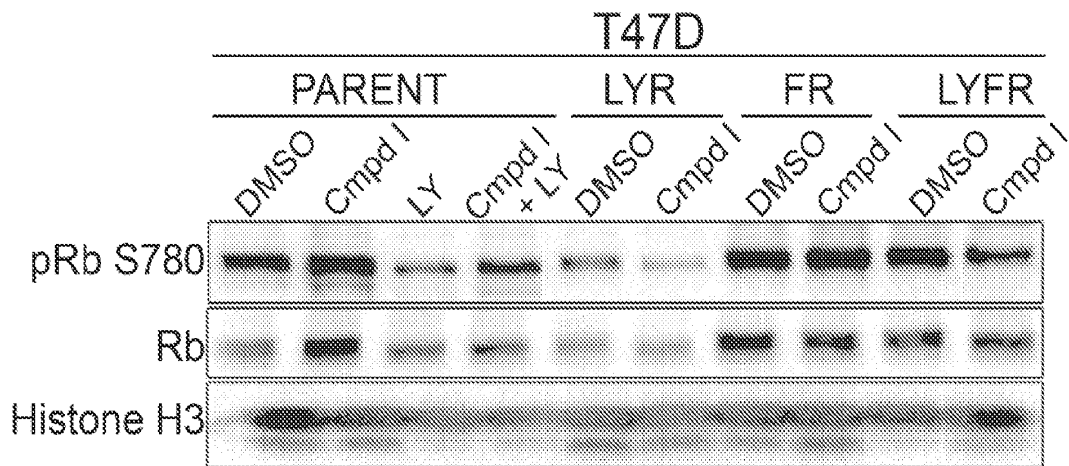


FIG. 147K

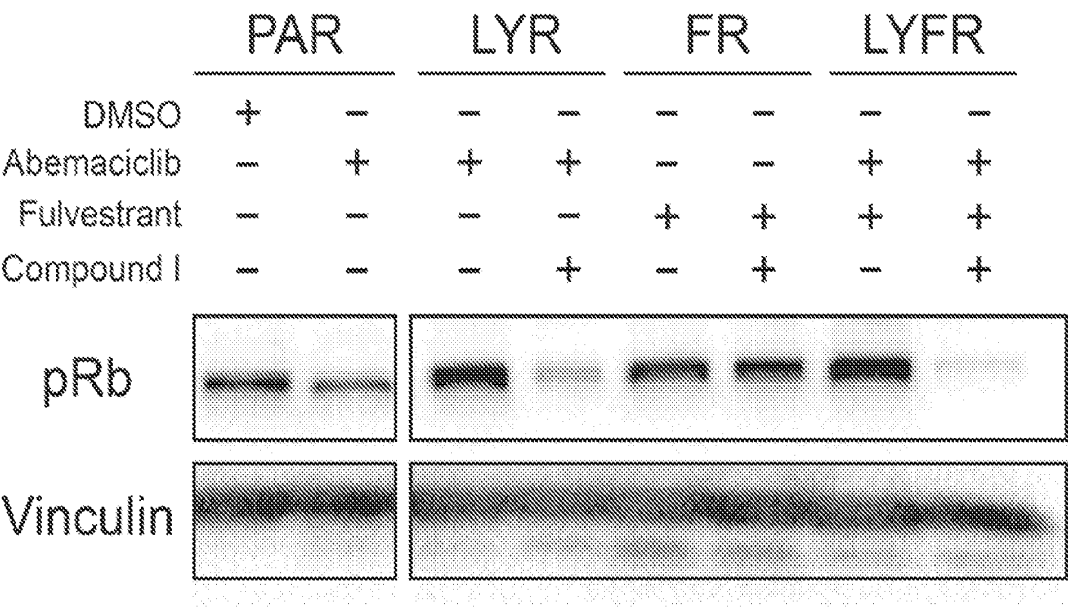


FIG. 147L

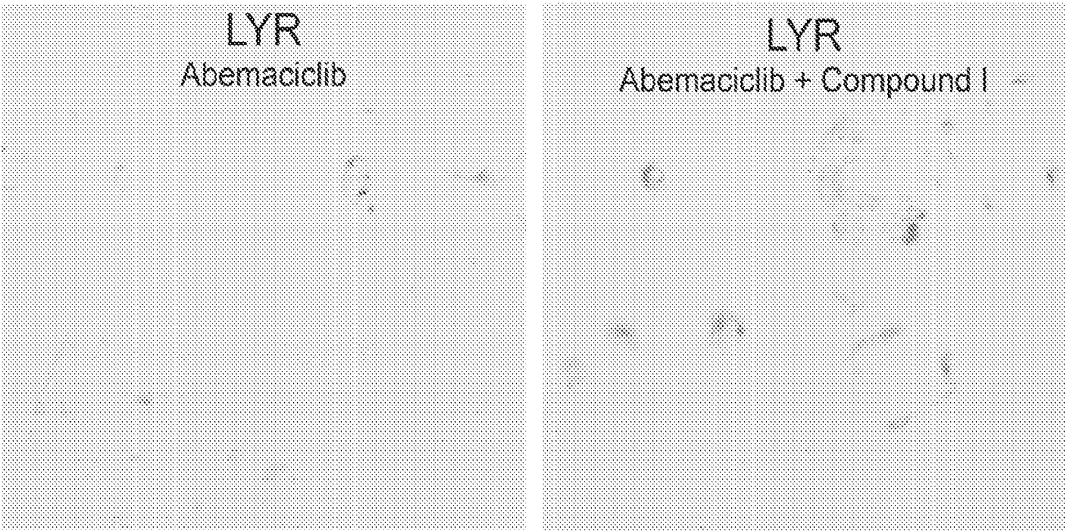


FIG. 148A

FIG. 148B

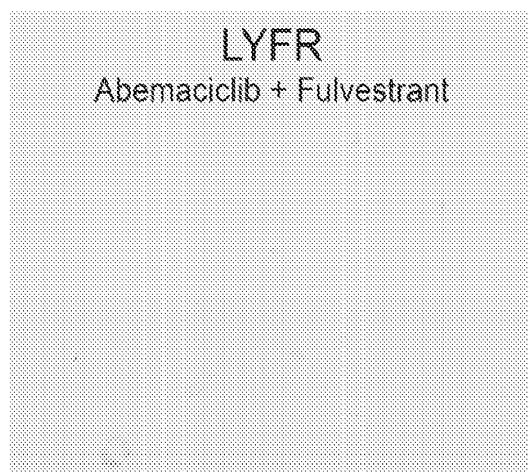


FIG. 148C

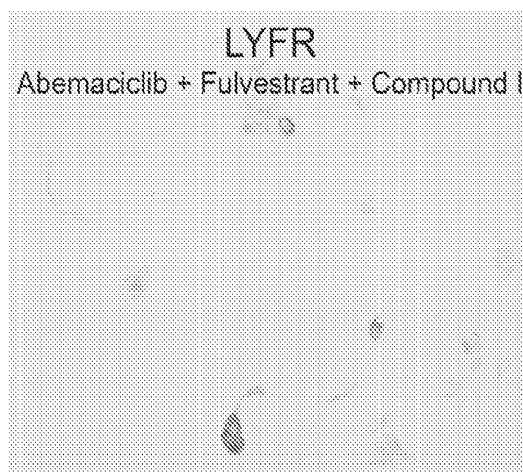


FIG. 148D

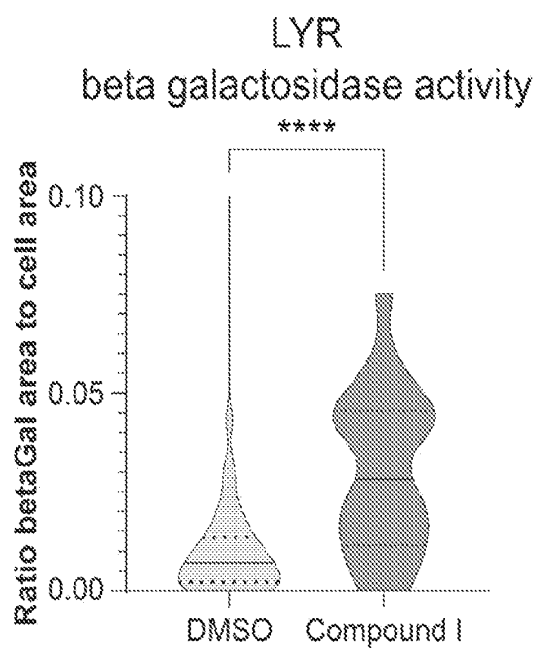


FIG. 148E

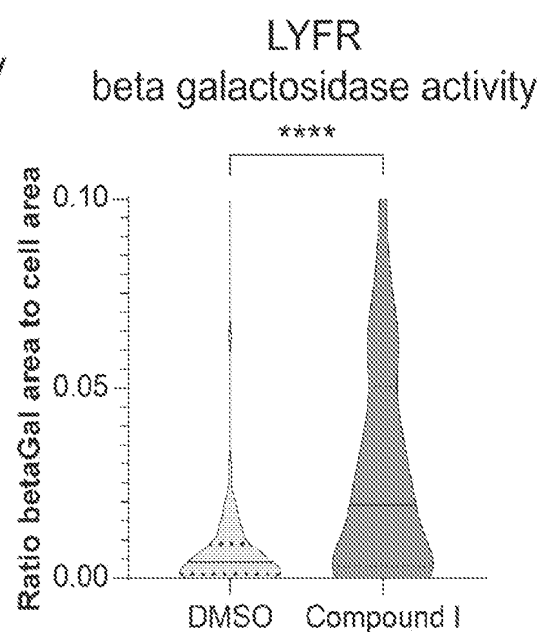


FIG. 148F

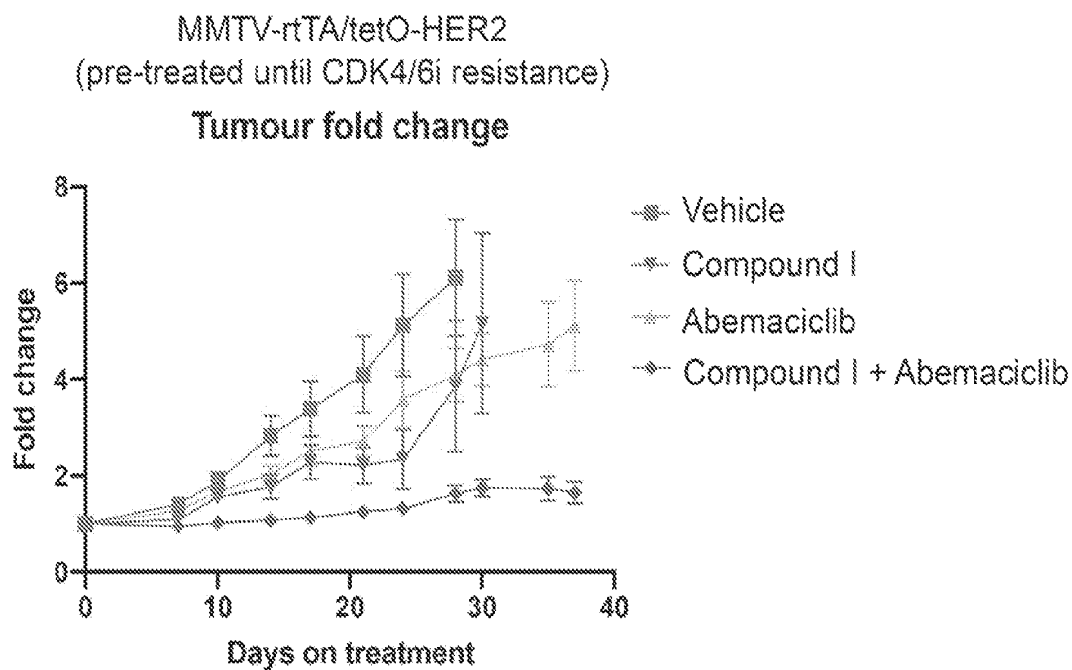


FIG. 149

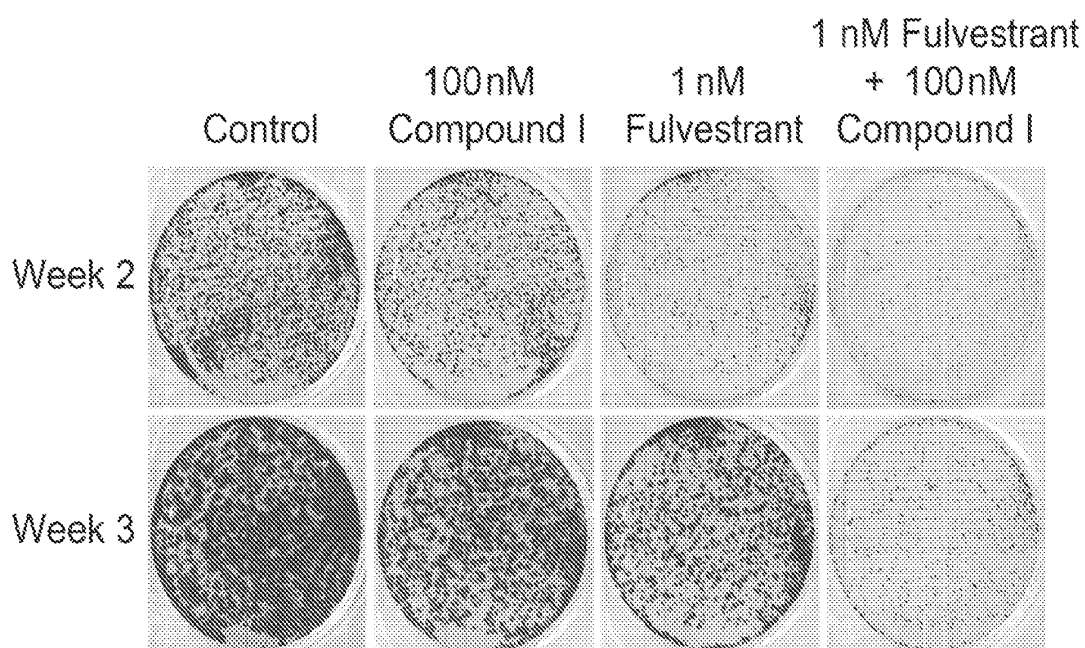


FIG. 150A



FIG. 150B

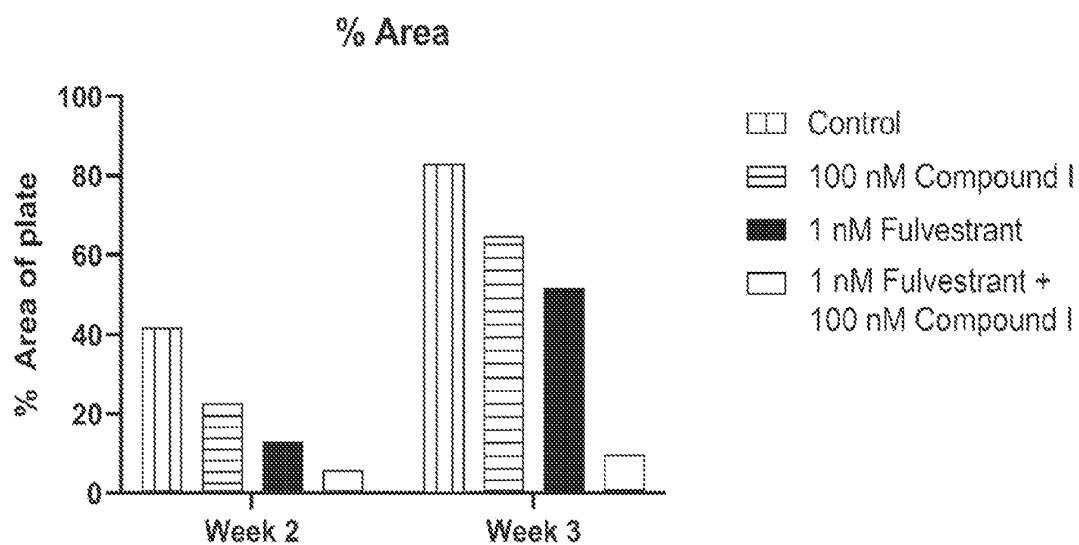


FIG. 150C

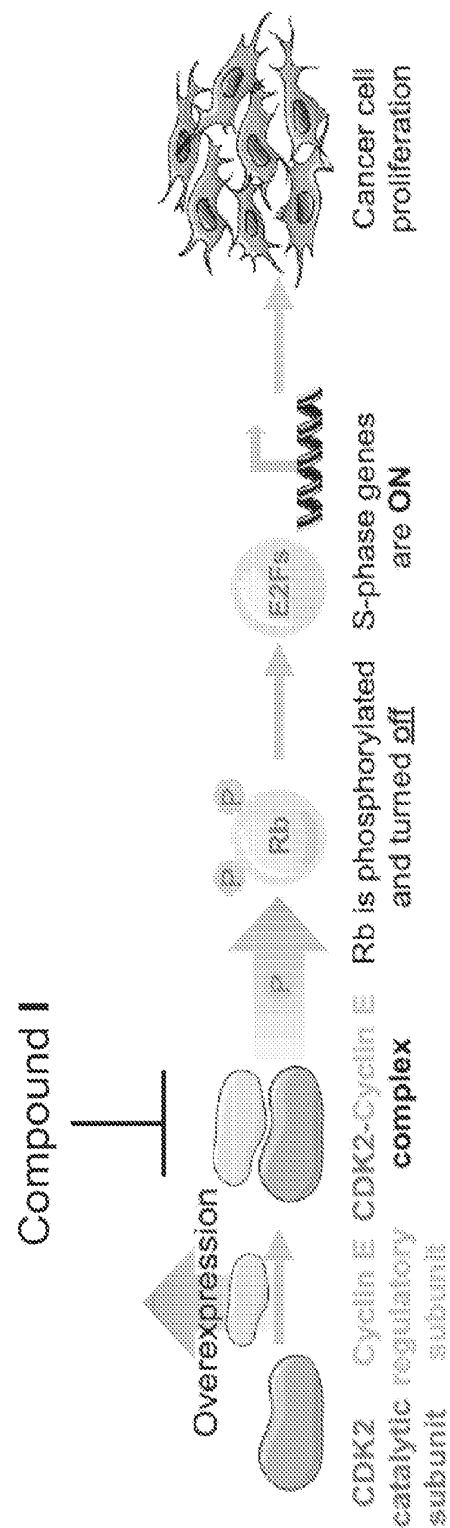


FIG. 151