

Entry	Compound	SMILES	MOLM -13	Saos -2	MRC -9	HL -60	Hel 92.1.7	Telom erase	SAOS2 /M OLM13	MRC9 /M OLM13	Metabolic stability (Mouse)	Plasma stability (60 mins)	Plasma stability (120 mins)
1		<chem>CN1C=C(C(N1)C2=CC(Cl)=CC=C2C3OC(C(C3)=C)=O)</chem>	1.749	411.6	7541				235.33	4311.61	25.1	13	2
2		<chem>O=C(C(C1)=C)OC1C2=CC=CC=C2C3=CC(NC(=O)C3)=O</chem>	98.87	7038	9006				71.18	91.09	80.77		
3		<chem>C=C(C(N1C)=O)CC1C2=CC=C(C(Cl)C=C2C3=CC(NC(=O)C3)=O</chem>	350.8	10000	100000				28.51	285.06		93.82	102.71
4		<chem>O=S(N[C@@H](C1=CC=CC=C1)C(=CC=C1)C(NC2=CC=CC(C2)=O)=O</chem>	1096	7932	10000				7.24	9.12		7.42	0.53
5		<chem>O=S(N[C@@H](C1=CC=CC=C1)C(=CC=C1)C(N2CCCC2=O)=O</chem>	1196	10000	TBD				8.36			66.59	34.76

FIG. 1

Entry	Compound	SMILES	MOLM -13	Saos -2	MRC -9	HL -60	Hel 92. 1.7	Telom erase	SAOS2 /M OLM13	MRC9 /M OLM13	Metabolic stability (Mouse)	Plasma stability (60 mins)	Plasma stability (120 mins)
6		<chem>O=C1N(C2=CC=C(C(=C2)C(=O)N1C)C(=O)O)C</chem>	132.91	76930.57	TBD								
7		<chem>OC(CC(C(=O)O)C(=O)N1C)C(=O)N1C</chem>	1001.00	5057.00	>100000								
8		<chem>O=C1N(C2=CC=C(C(=C2)C(=O)N1C)C(=O)O)C</chem>	238.3	>10000	TBD							73.03	62.09
9		<chem>C=C1C(O)CN(C2=CC=C(C(=C2)C(=O)N1C)C(=O)O)C</chem>	2073	>100000	>100000								
10		<chem>C=C1C(O)CN(C2=CC=C(C(=C2)C(=O)N1C)C(=O)O)C</chem>	6251	>100000	>100000								

FIG. 2

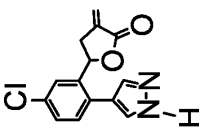
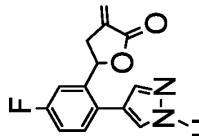
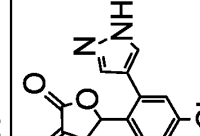
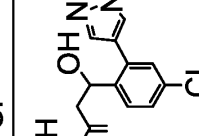
Structure	MOLM13	Saos2	MRC9	HL-60	Telomerase activity*	Fold selectivity		Mice PK	
	IC50 (nM)					Saos2/MOLM13	MRC9/MOLM13	T1/2 (h)	Exposure above MOLM-13 IC50
	11.30	478.43	1044.60	78.22	10.62	42	92	7.4	>8h
	89.32	570.15	1213.72	169.42	10.74	6	14	3.6	>8h
	7.47	151.52	589.66	9.99	10.66	20	79	1	>4h
	229	4232	>100000	TBD	10.88	18	>436	8.4	>4h

FIG. 3

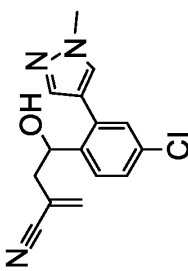
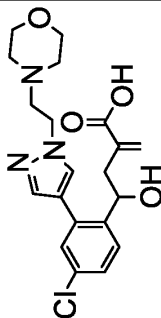
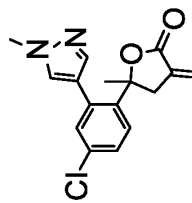
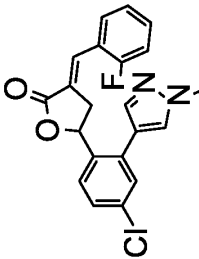
Structure	MOLM13	Saos2	MRC9	Fold selectivity		Mice PK	
	IC50 (nM)			Saos2/MOLM13	MRC9/MOLM13	T1/2 (h)	Exposure above MOLM-13 IC50
	12.7	486	753	38	59		8 h
	160.4	4426	100000	28	623	6.1 h	8 h
	54.9	643	1214.7	12	22	TBD	24 h
	221.33	>100000	>100000	>450	>450	TBD	TBD

FIG. 4

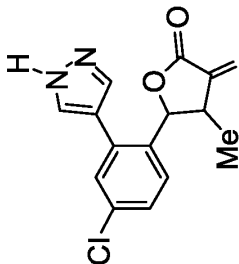
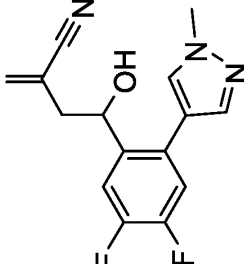
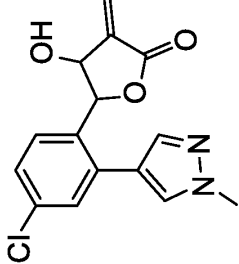
Structure	MOLM13	Saos2	MRC9	HL-60	Fold selectivity		Exposure above MOLM-13 IC50
	IC50 (nM)				Saos2/MOLM13	MRC9/MOLM13	
	2	235	268	36.8	117.5	434.0	
							
	38	1532	590		40	15	4 h
	45.38	85947	36464		1890	803	24 h

FIG. 5

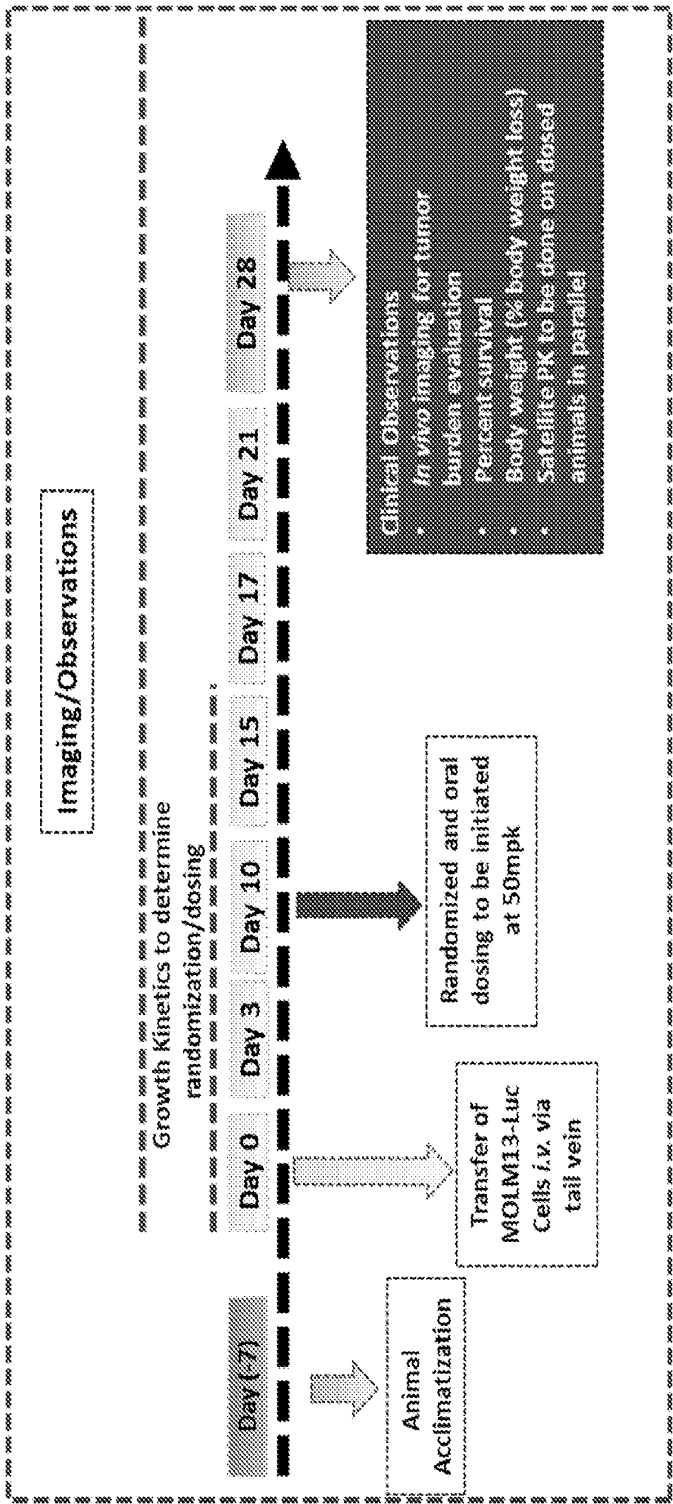


FIG. 6

Dynamic change of bioluminescent signal of Molm 13-Luc2

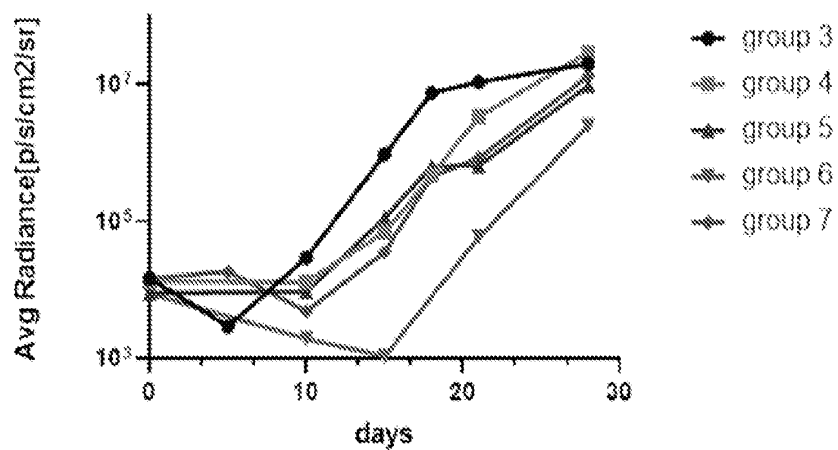


FIG. 7

Dynamic change of bioluminescent signal of Molm 13-Luc2

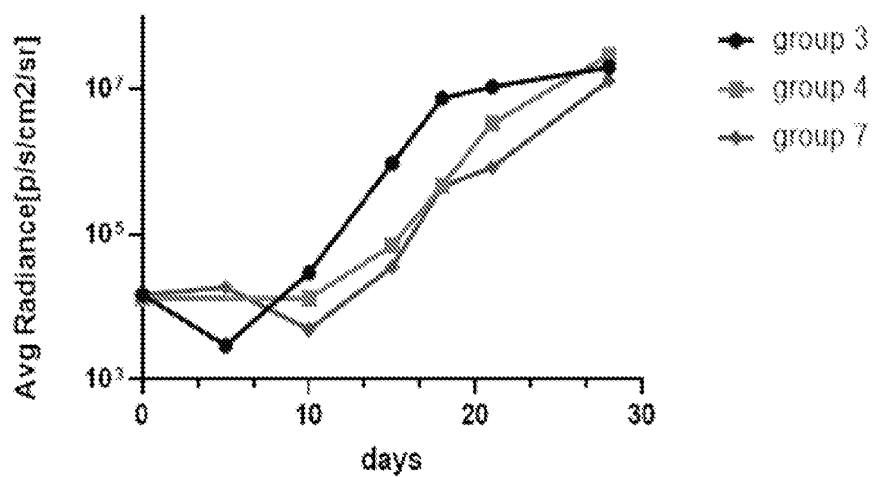


FIG. 8

Dynamic change of bioluminescent signal of Molm 13-Luc2

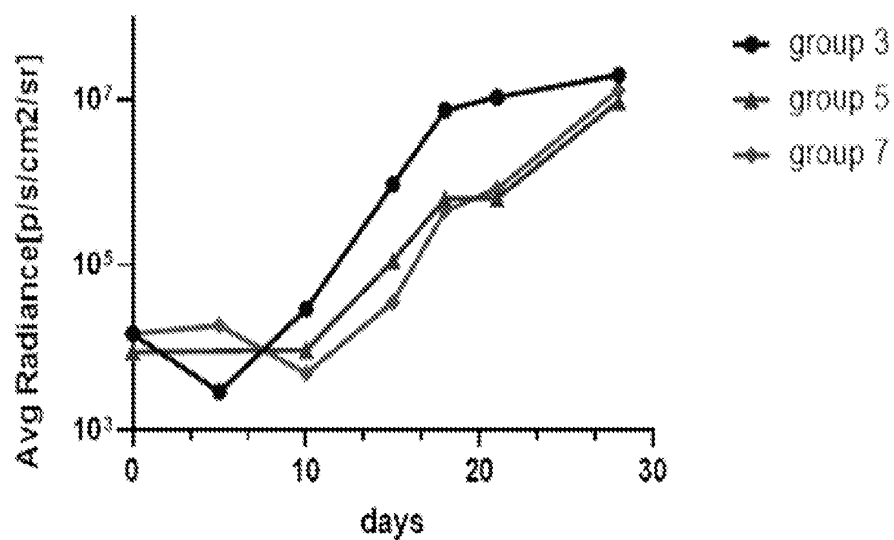


FIG. 9

Dynamic change of bioluminescent signal of Molm 13-Luc2

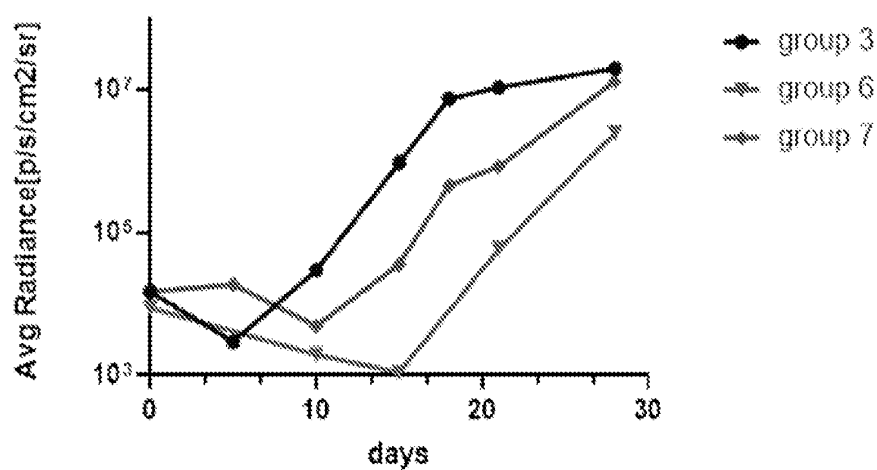
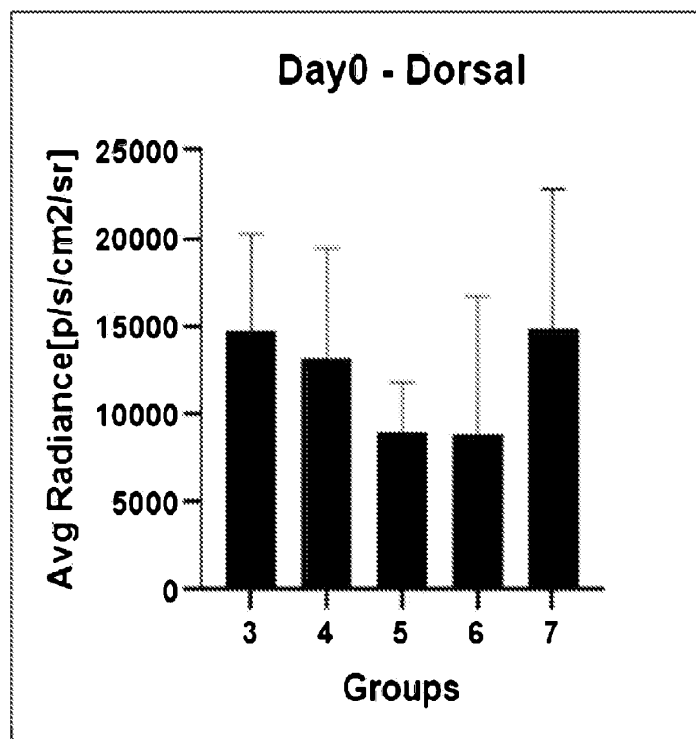


FIG. 10

FIG. 11A



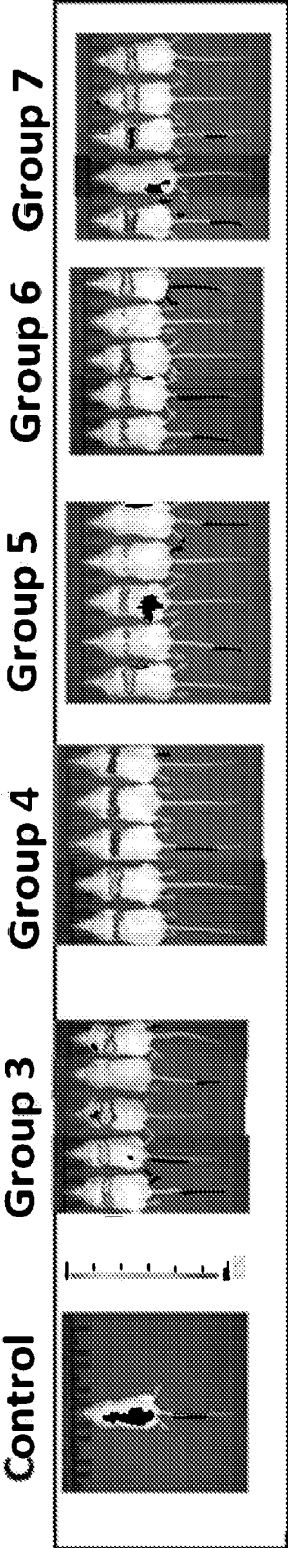
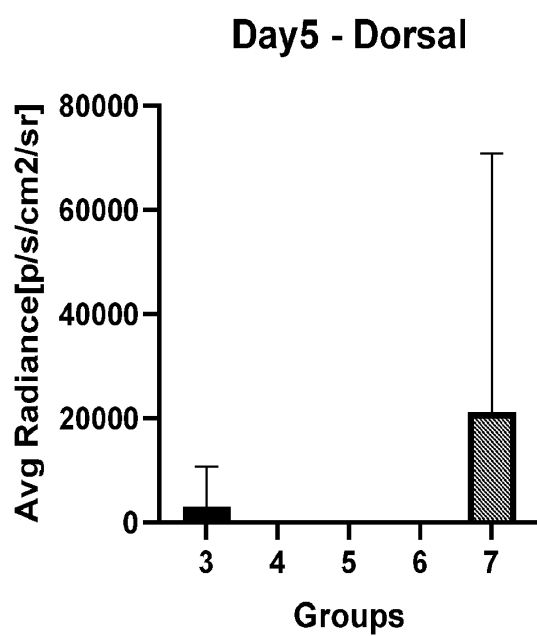


FIG. 11B

FIG. 12A



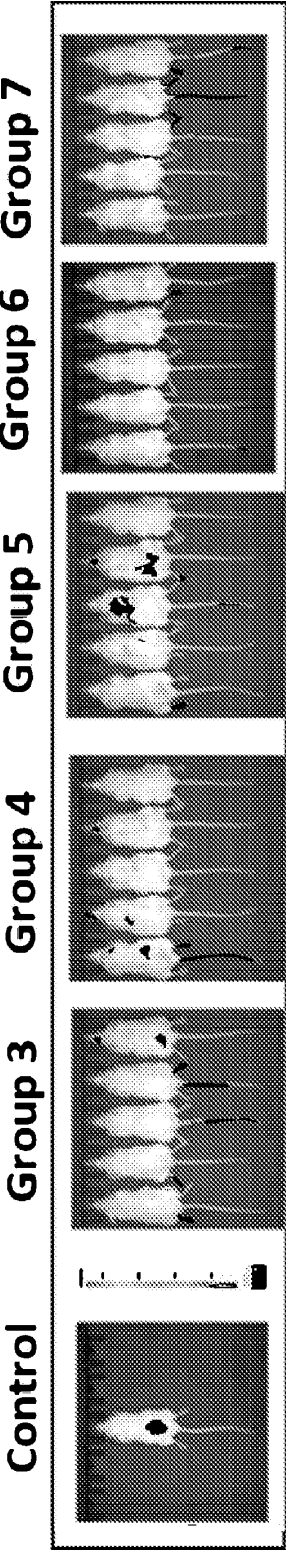
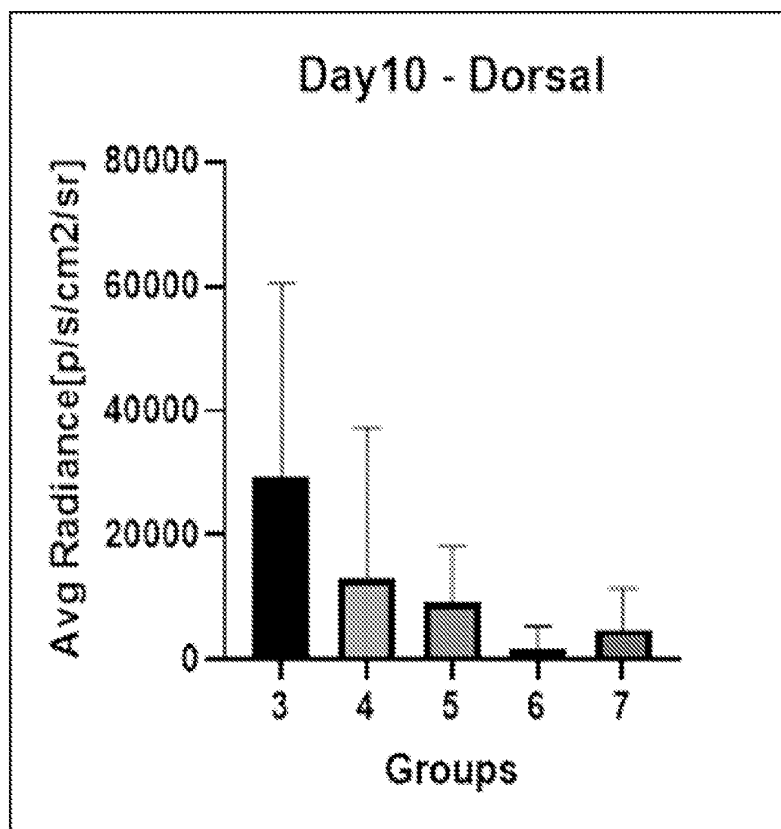


FIG. 12B

FIG. 13A



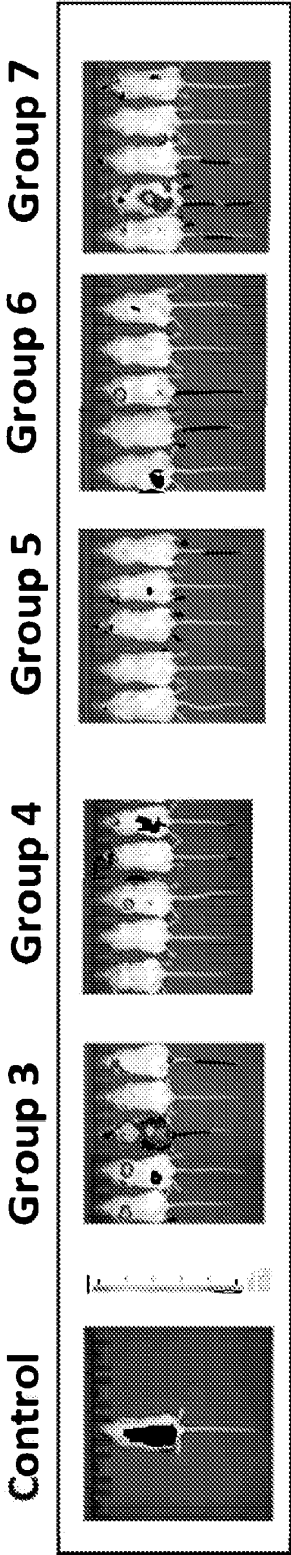
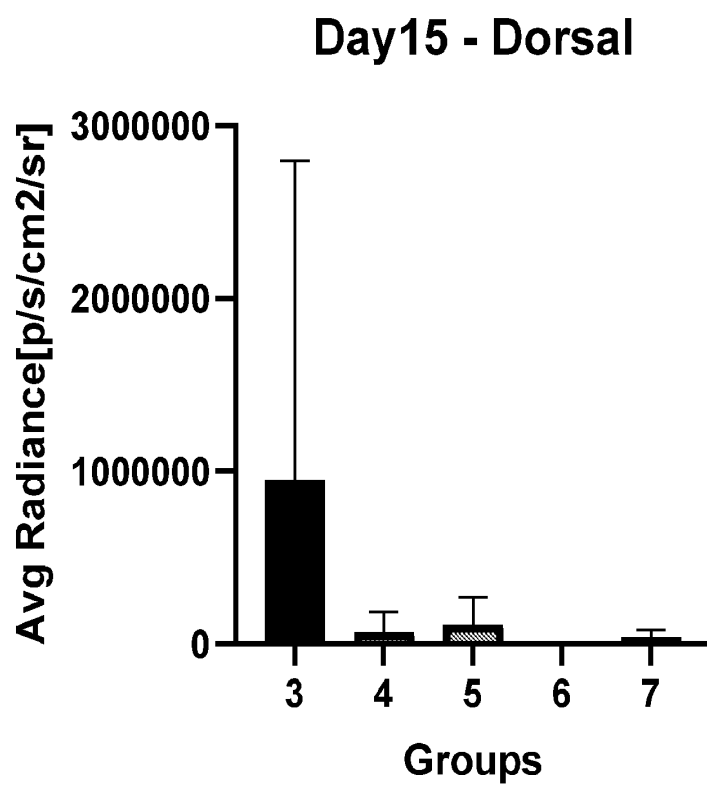


FIG. 13B

FIG. 14A



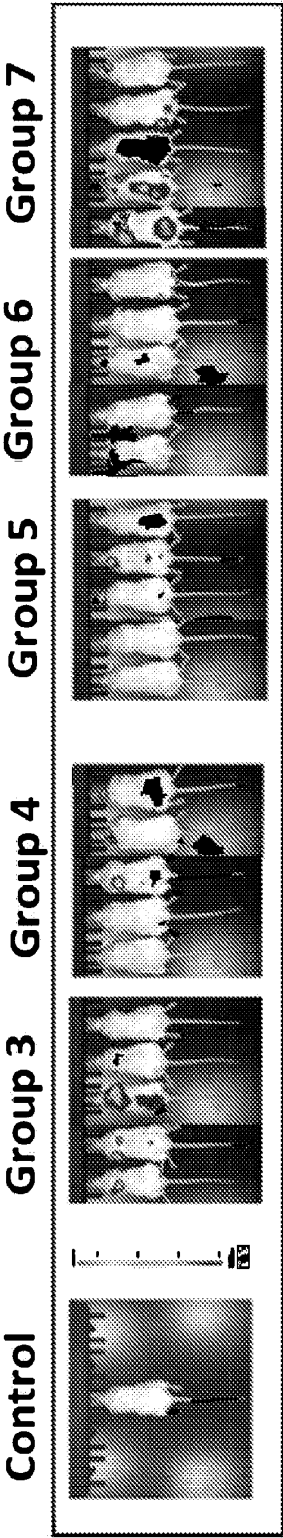
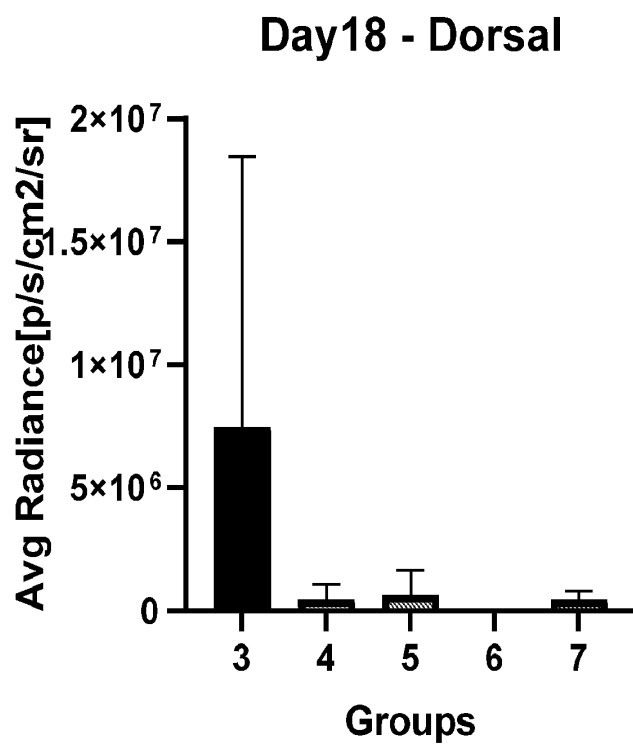


FIG. 14B

FIG. 15A



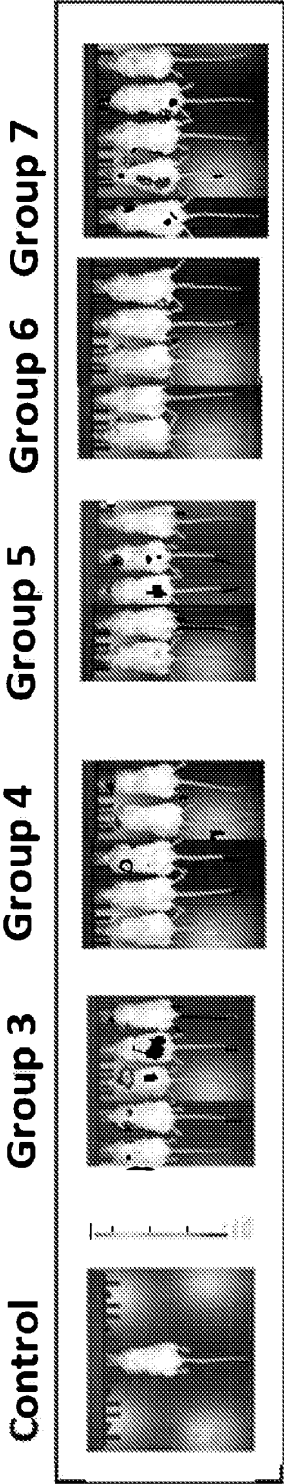
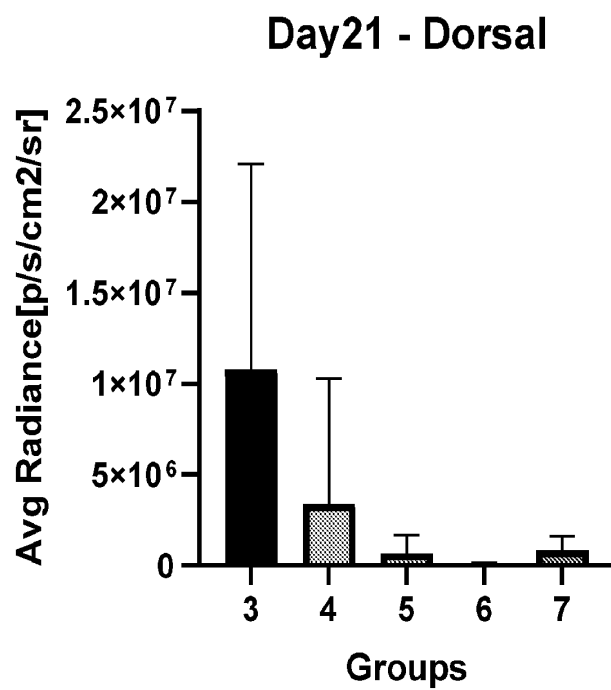


FIG. 15B

FIG. 16A



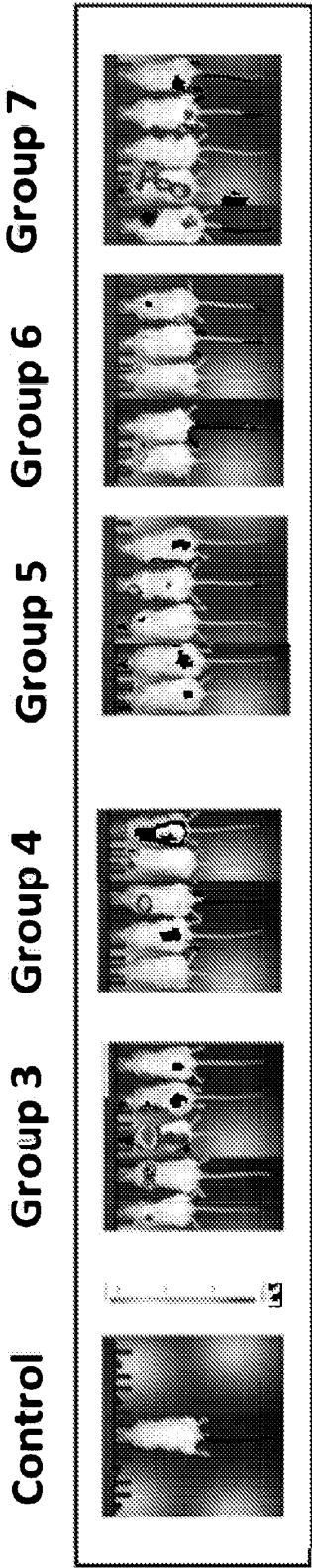
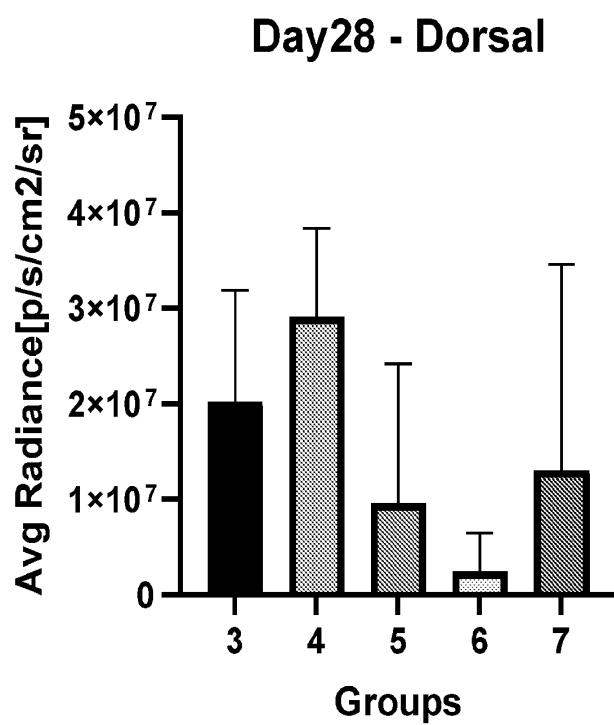
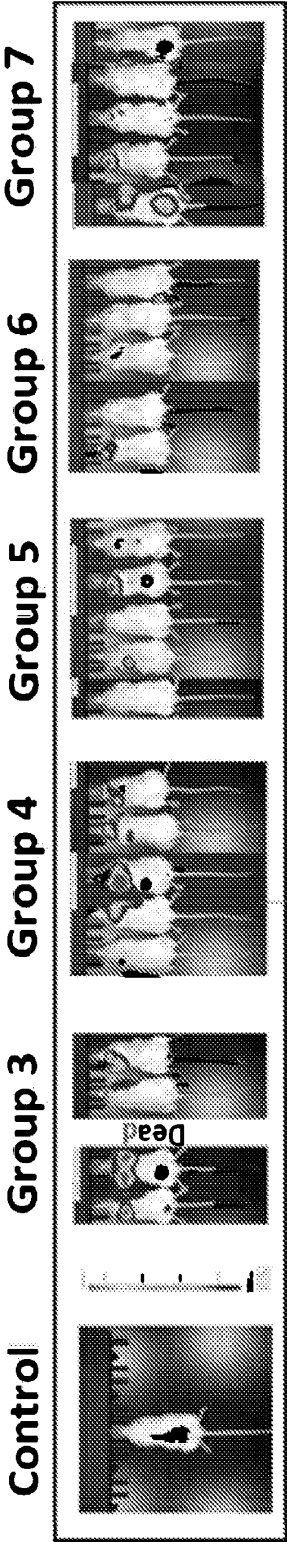


FIG. 16B

FIG. 17A





These two animals were too weak and could not be dosed last time

FIG. 17B

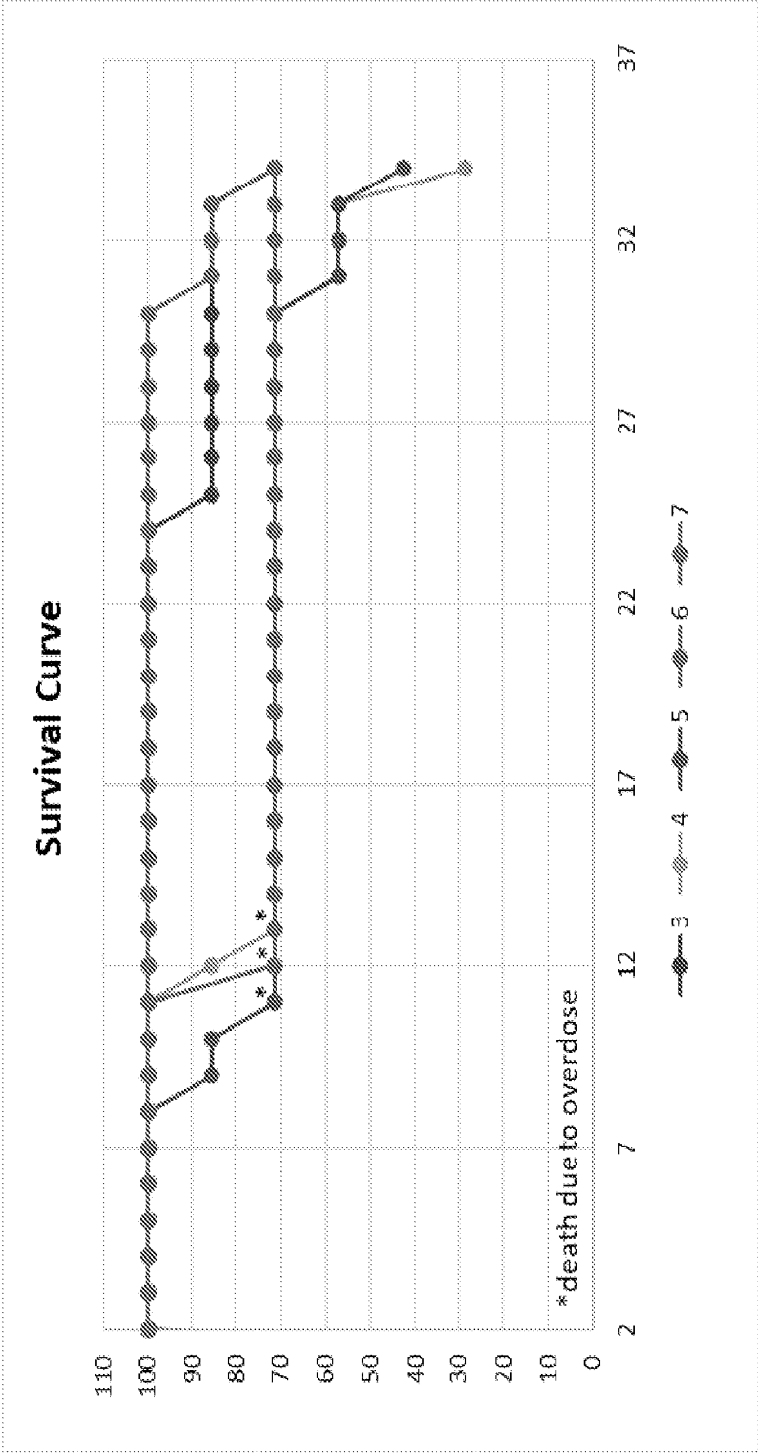


FIG. 18

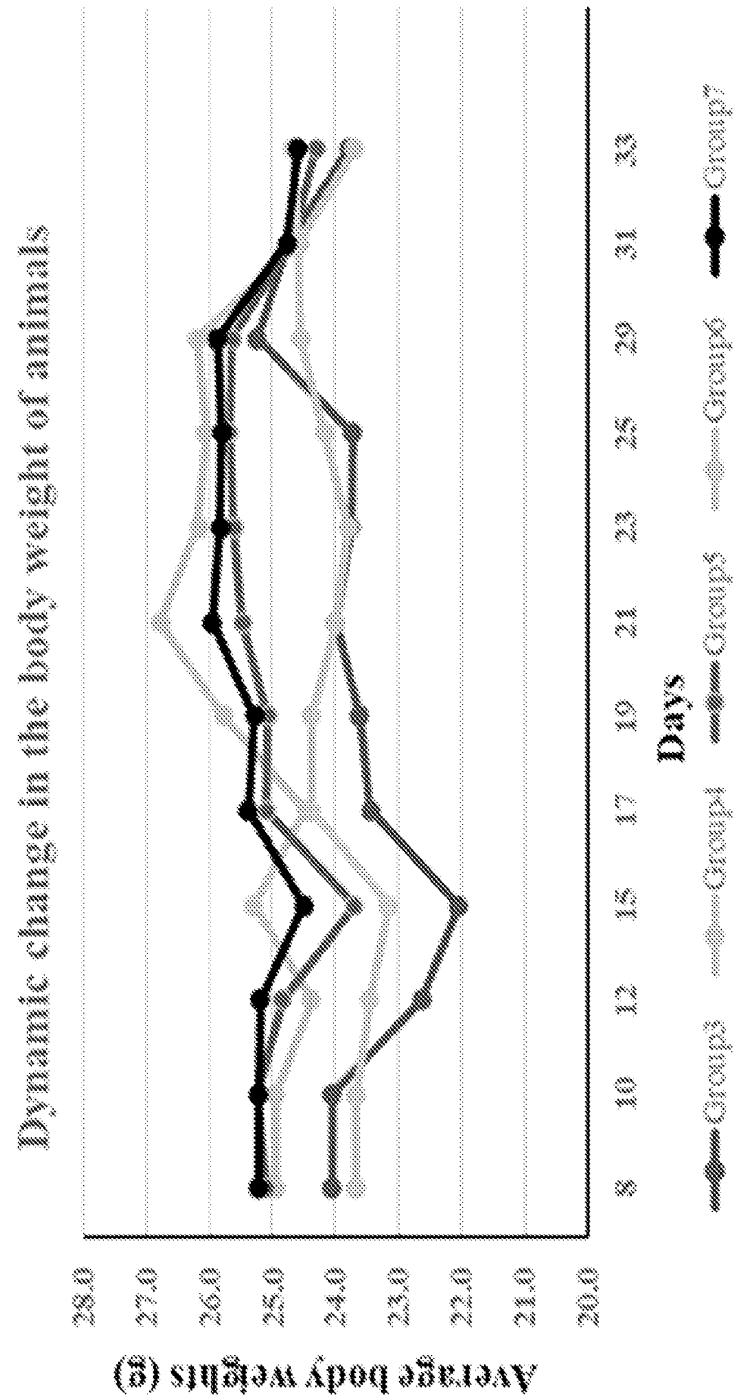


FIG. 19

FIG. 20A

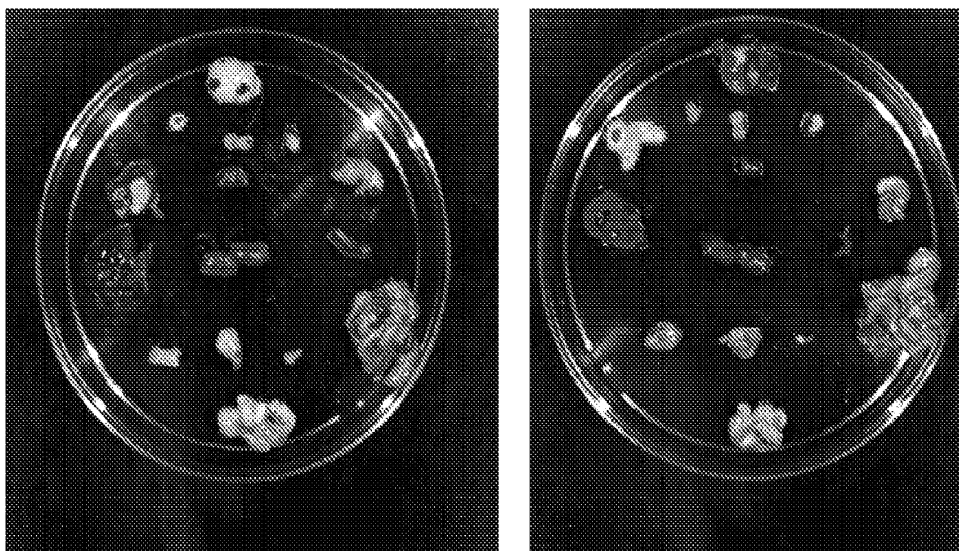


FIG. 20B

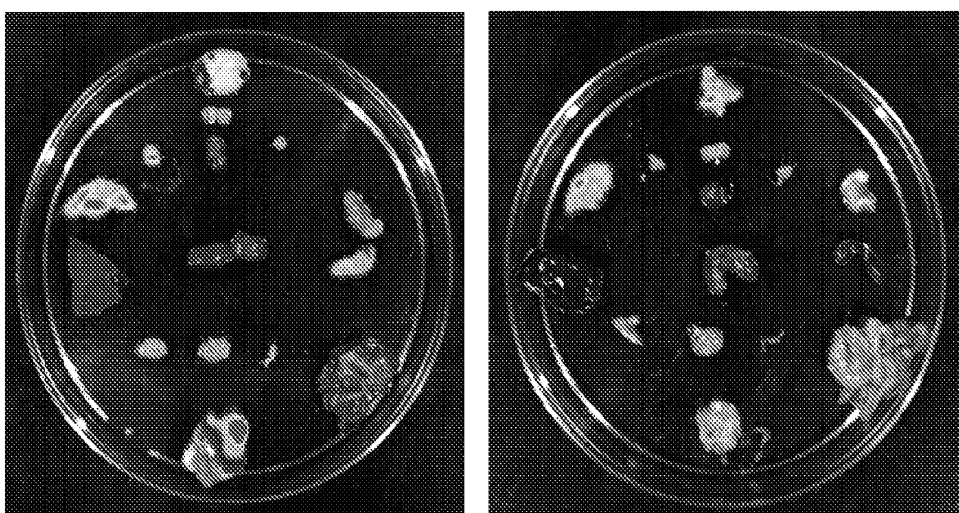


FIG. 20C

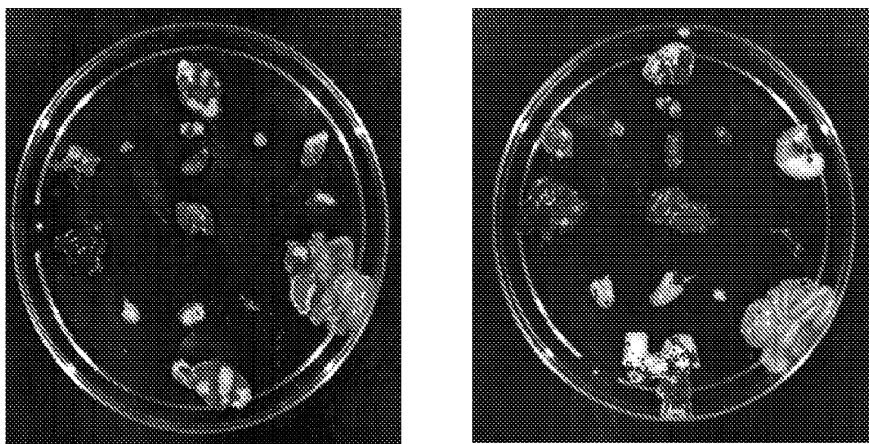


FIG. 20D

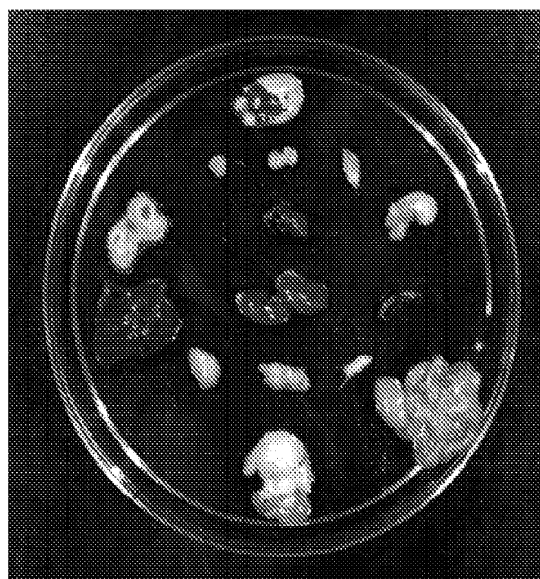


FIG. 20E

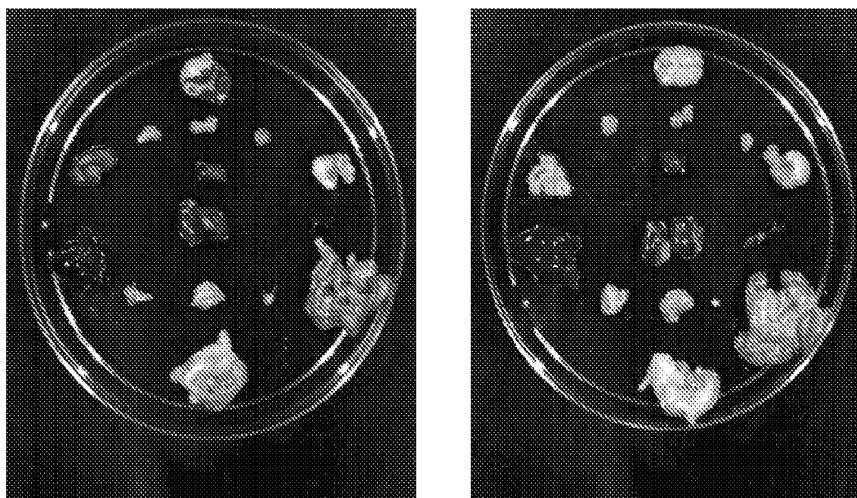


FIG. 20F

