

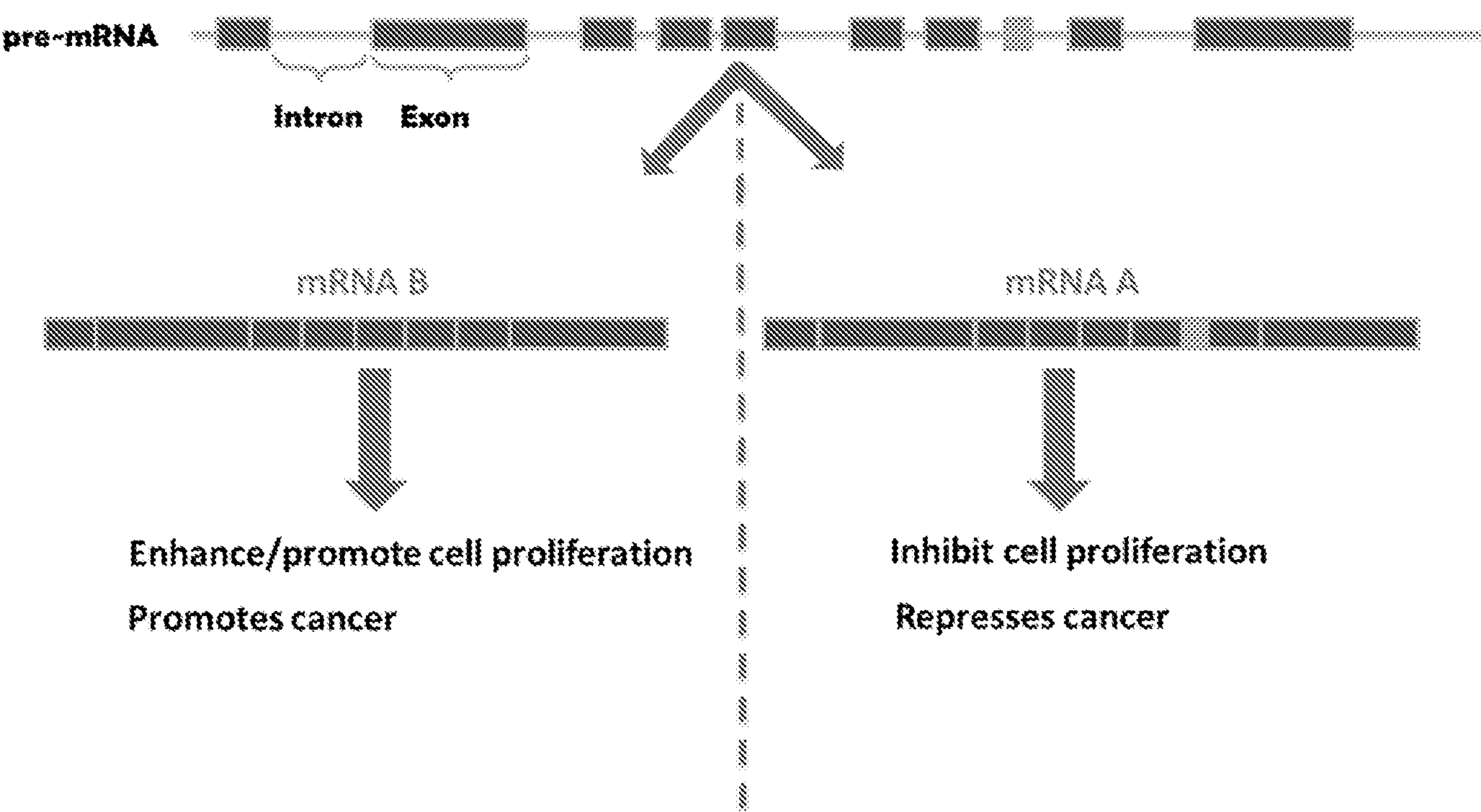


FIG. 1

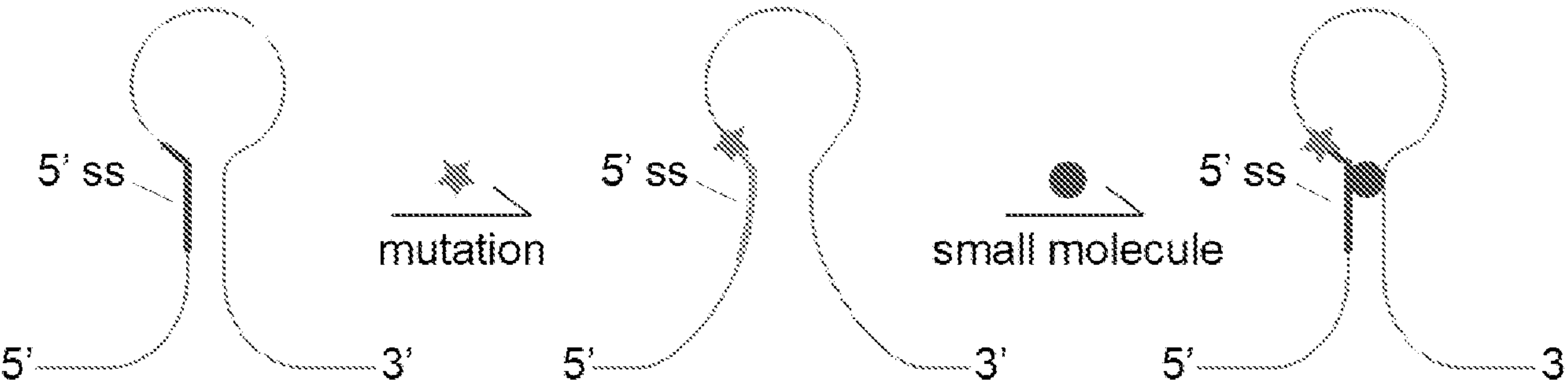
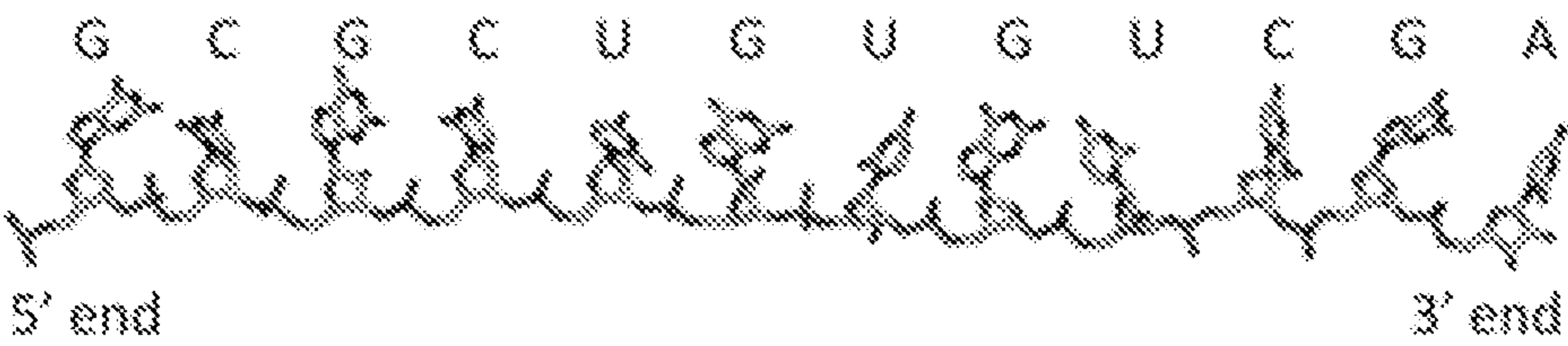


FIG. 2

Primary structure



Secondary structure



Tertiary structure

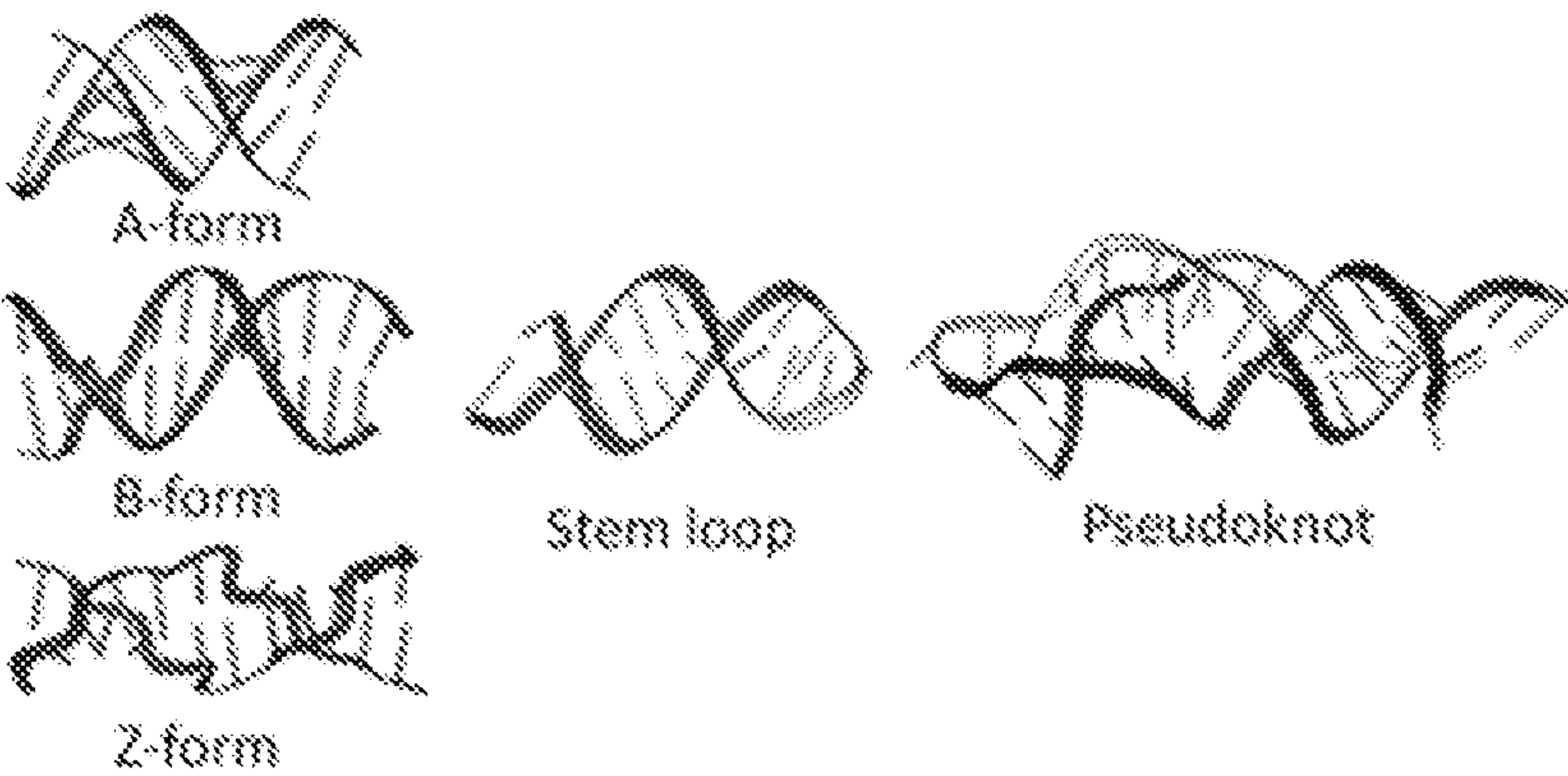


FIG. 3



FIG. 4

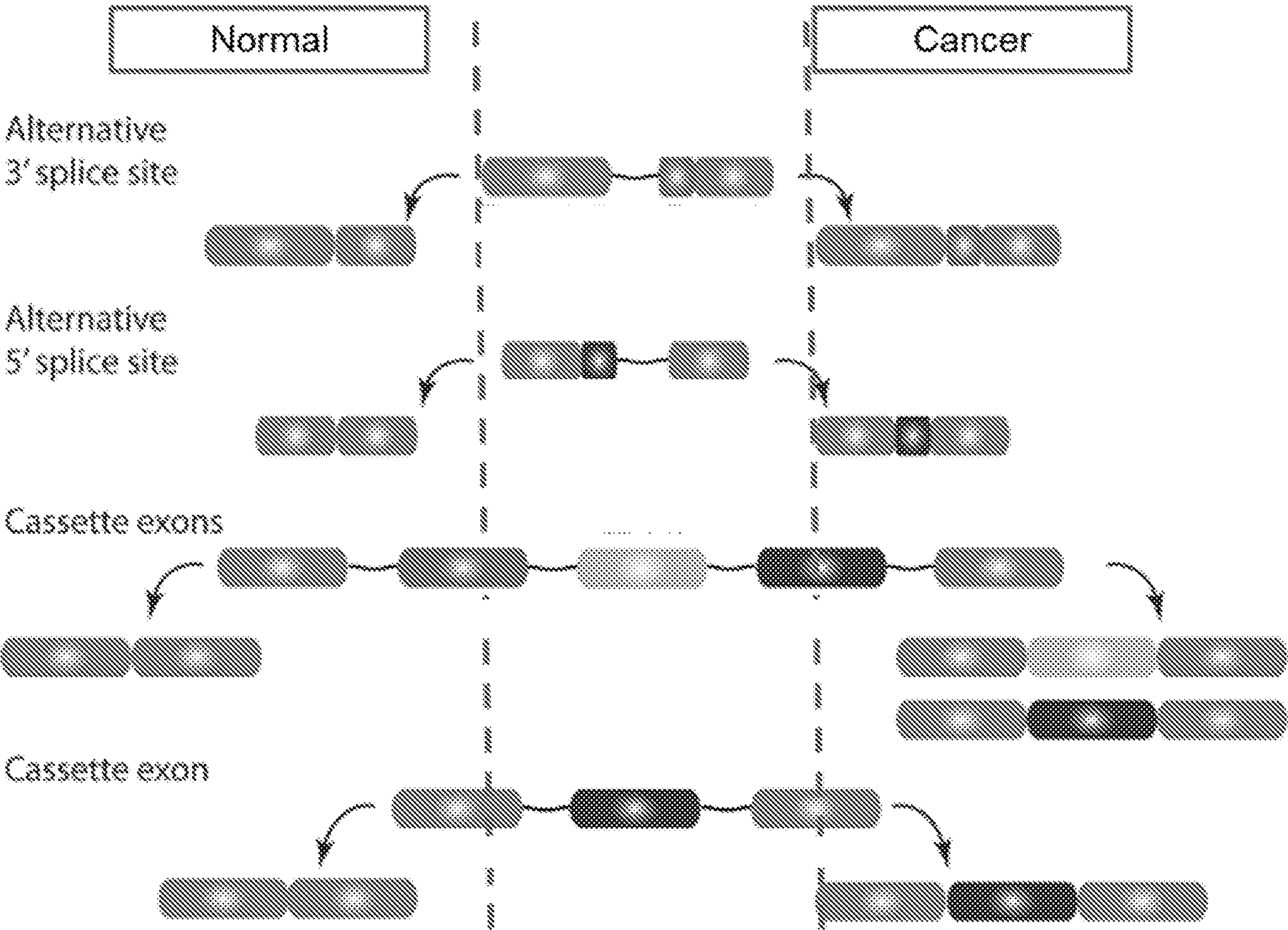


FIG. 5

5/31

SMN2

A: 3' -GUCCAΨΨCAUA-5'
 B: 5' -GGGUAAGU_ACU-3'

FIG. 6A

FOXMI

A: 3' -GUCAΨΨCAUA-5'
 B: 5' -CAGUAAGUGX_{CU}-3'

FIG. 6B

IKBKAP

A: 3' -GU^CCAΨΨCAUA-5'
 B: 5' -CA_AGUAAGUG--3'

FIG. 6C

MADD1

A: 3' -GUCCAΨΨCAA^U-5'
 B: 5' --AGGUGGGGUU-3'

FIG. 6D

KRAS

A: 3' -GUCCAΨΨCAUA-5'
 B: 5' -CAGGUAAGUA--3'

FIG. 6E

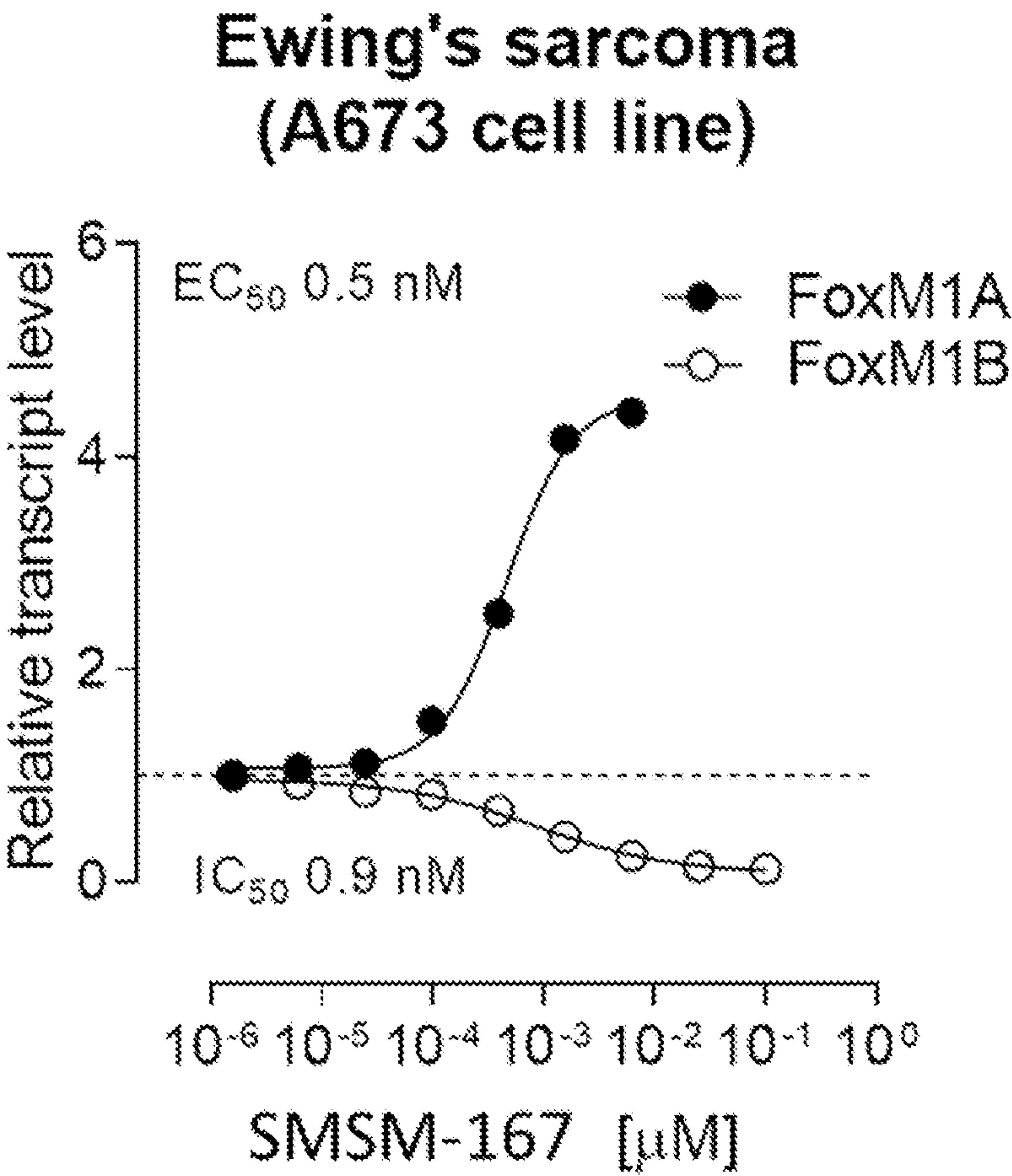


FIG. 7

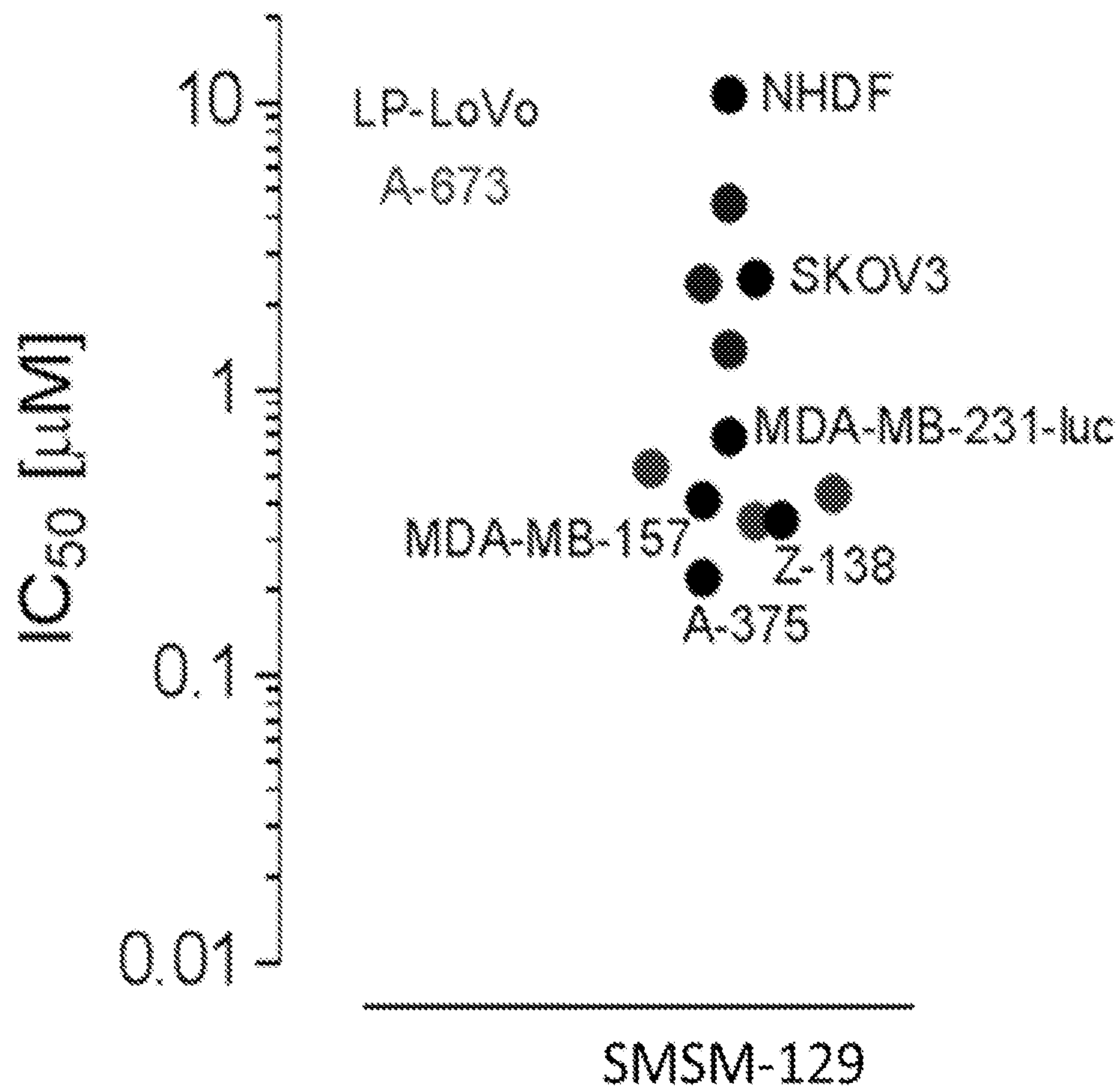


FIG. 8

8/31

A-673

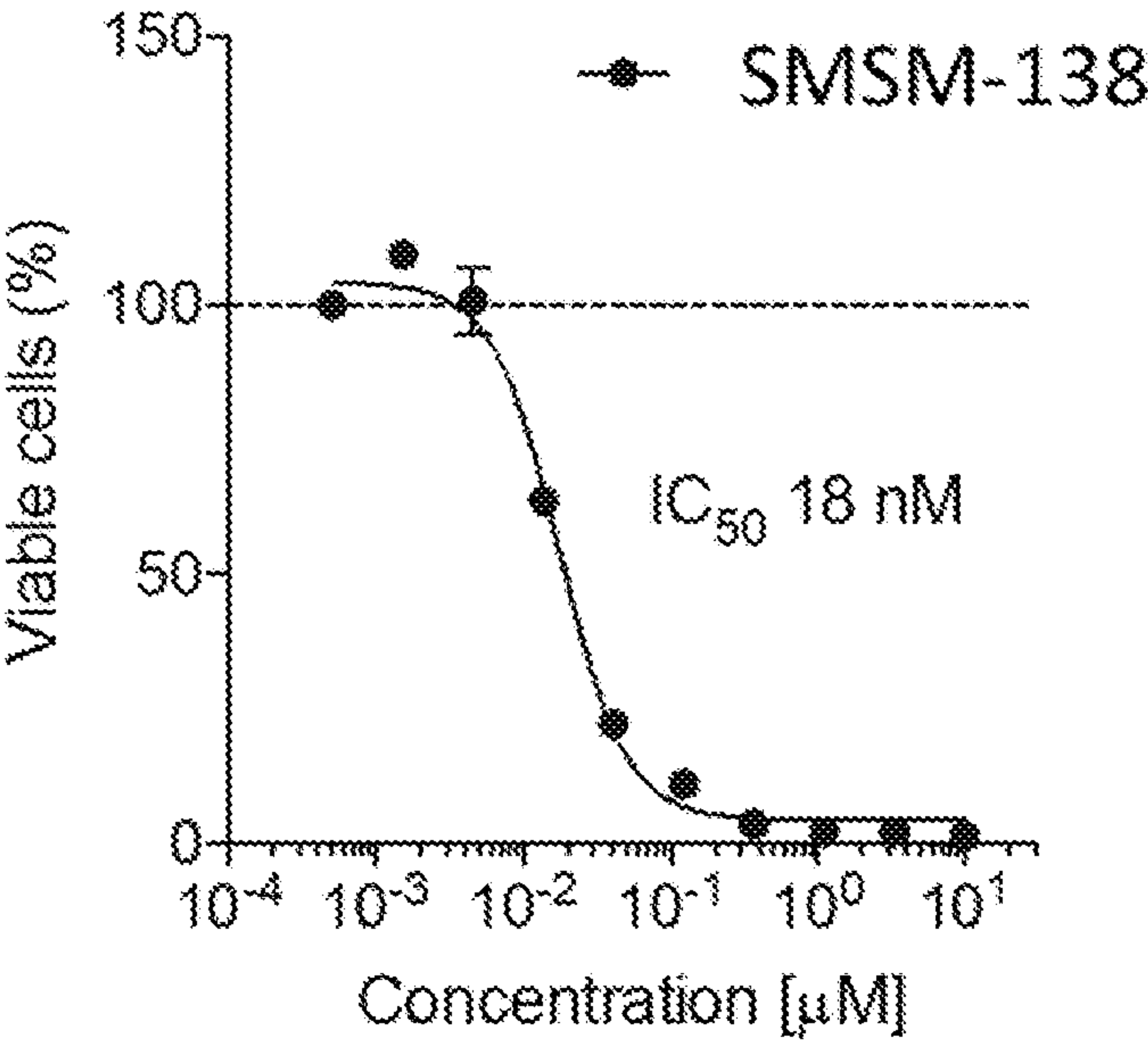


FIG. 9A

A-673

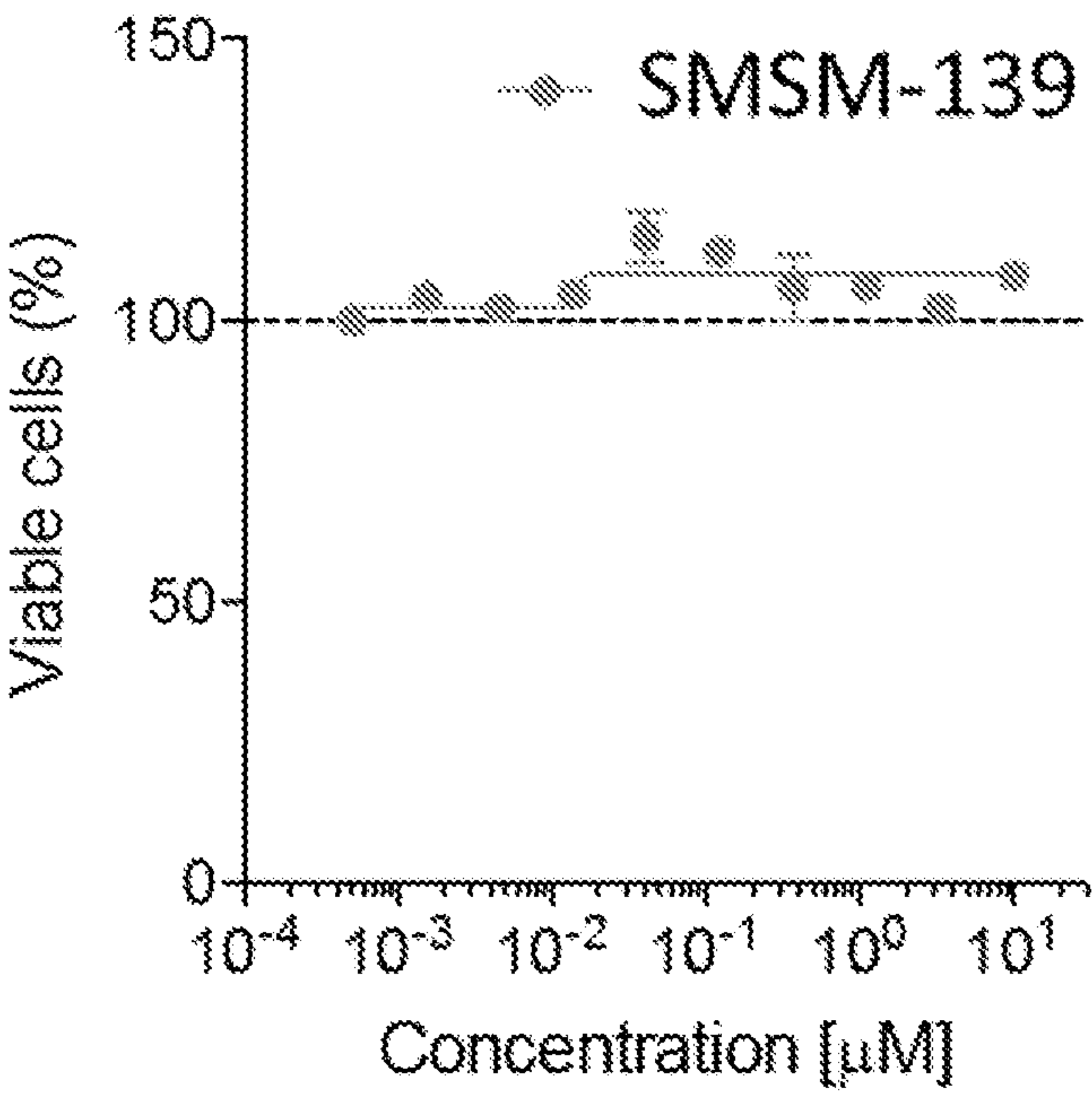


FIG. 9B

9/31

A-673

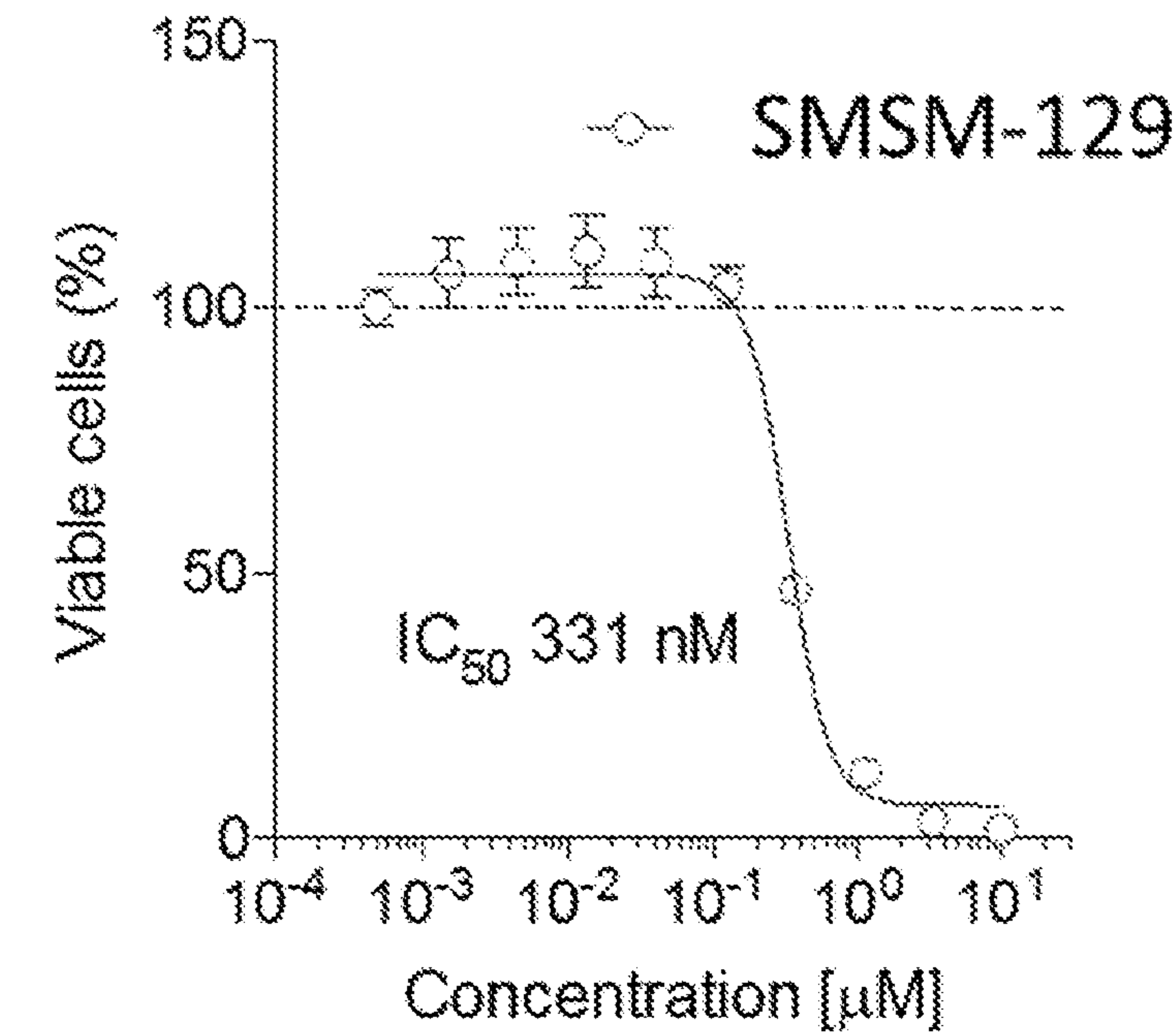


FIG. 9C

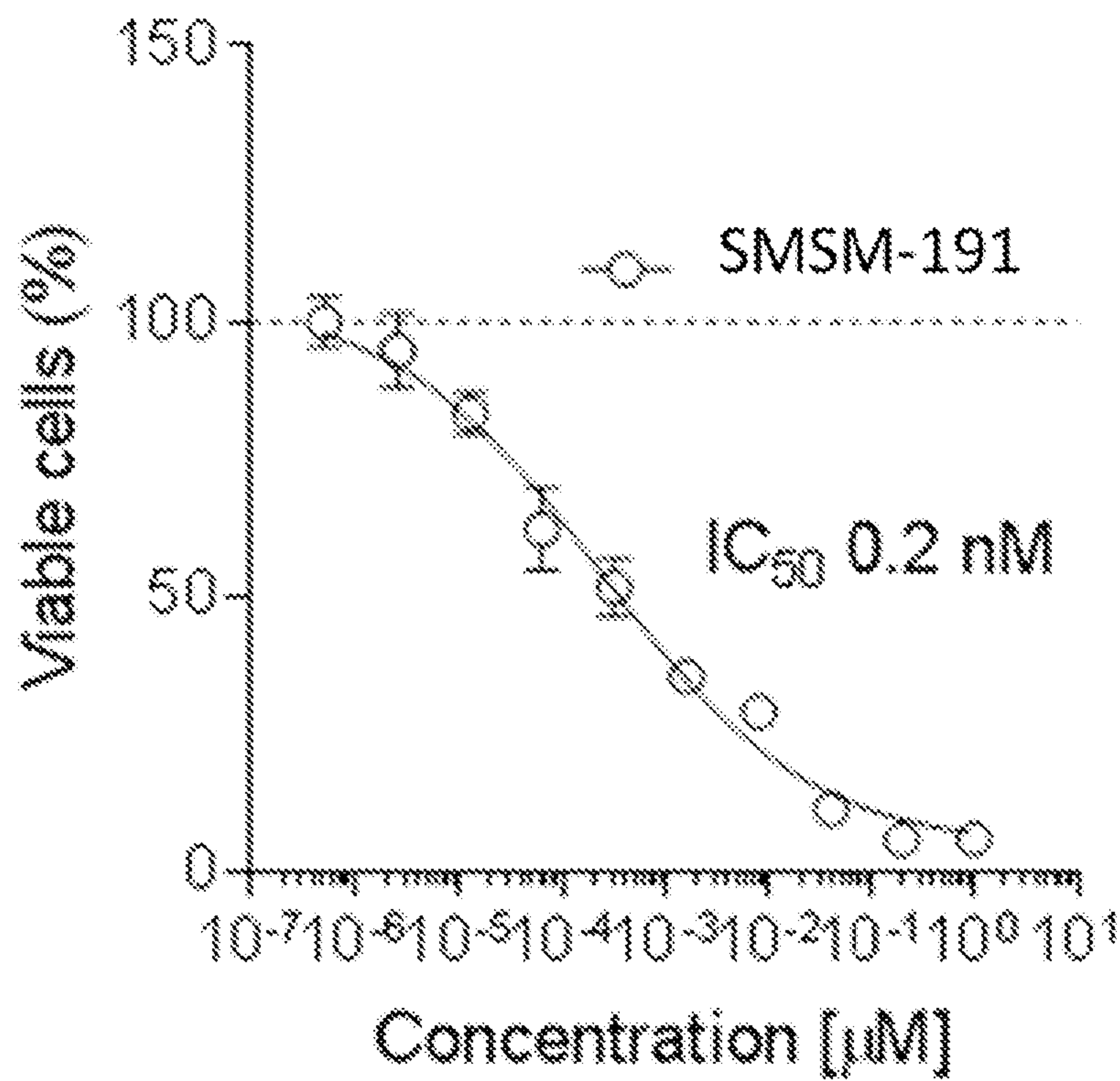


FIG. 9D

10/31

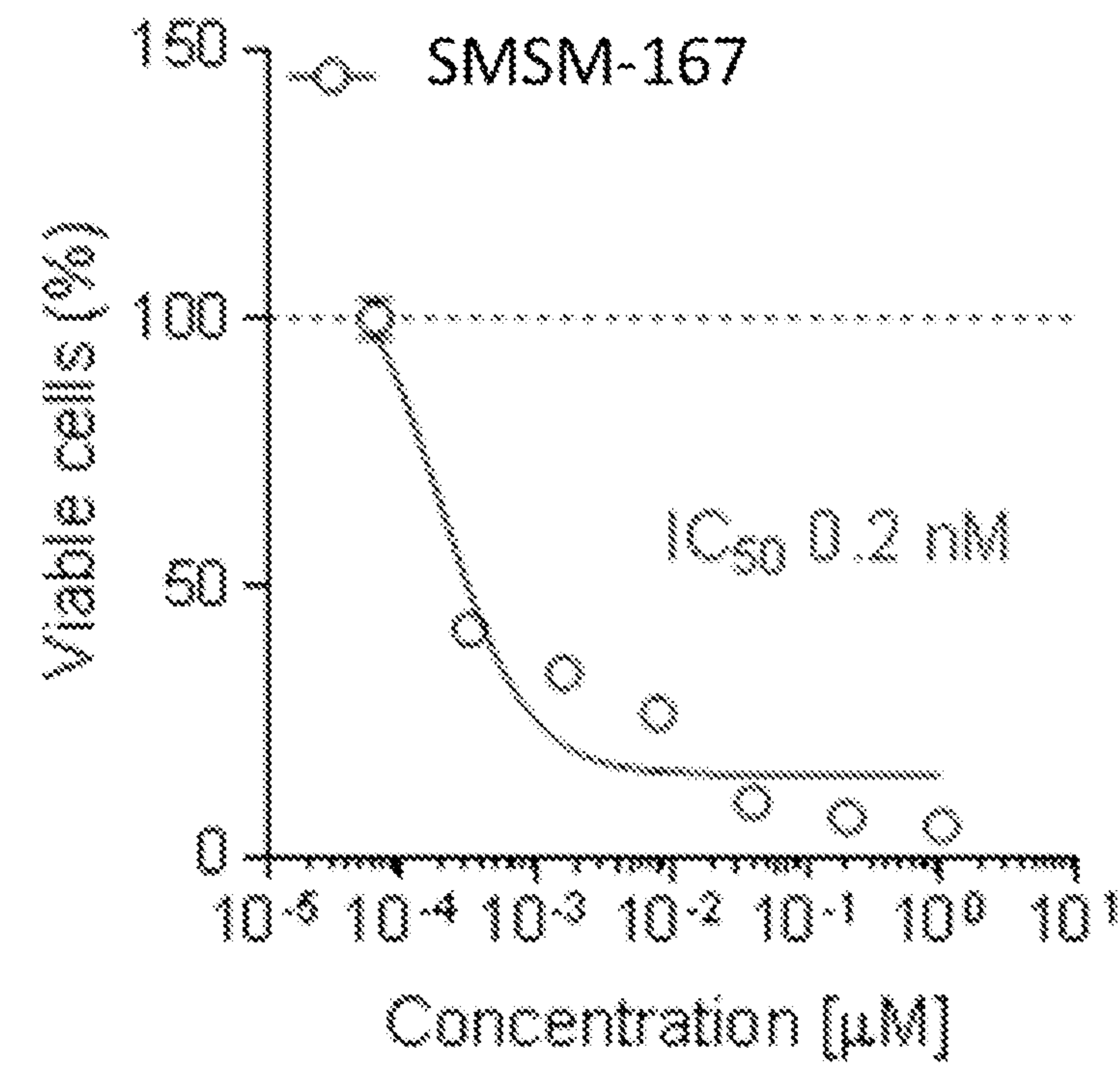


FIG. 9E

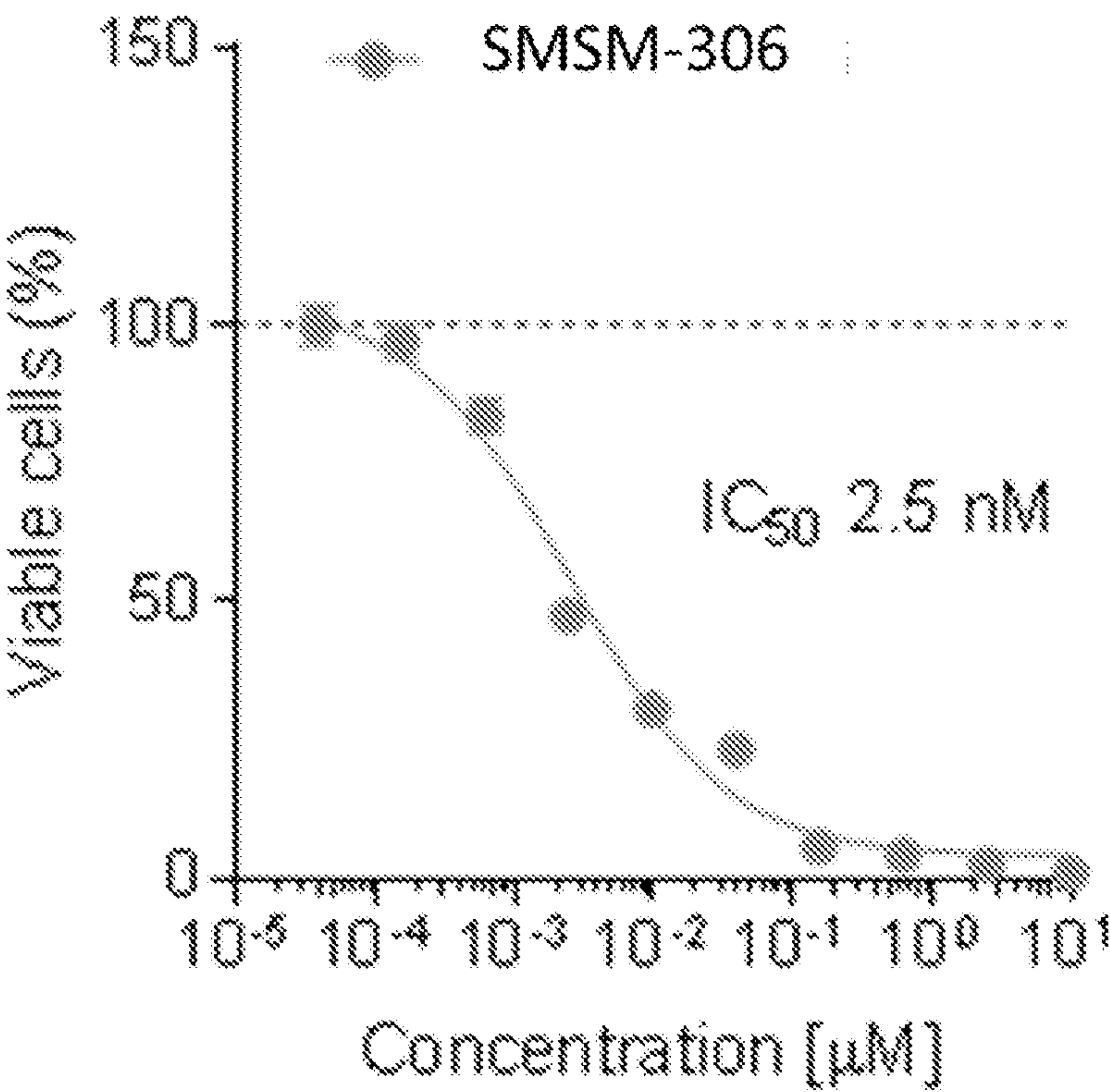


FIG. 9F

11/31

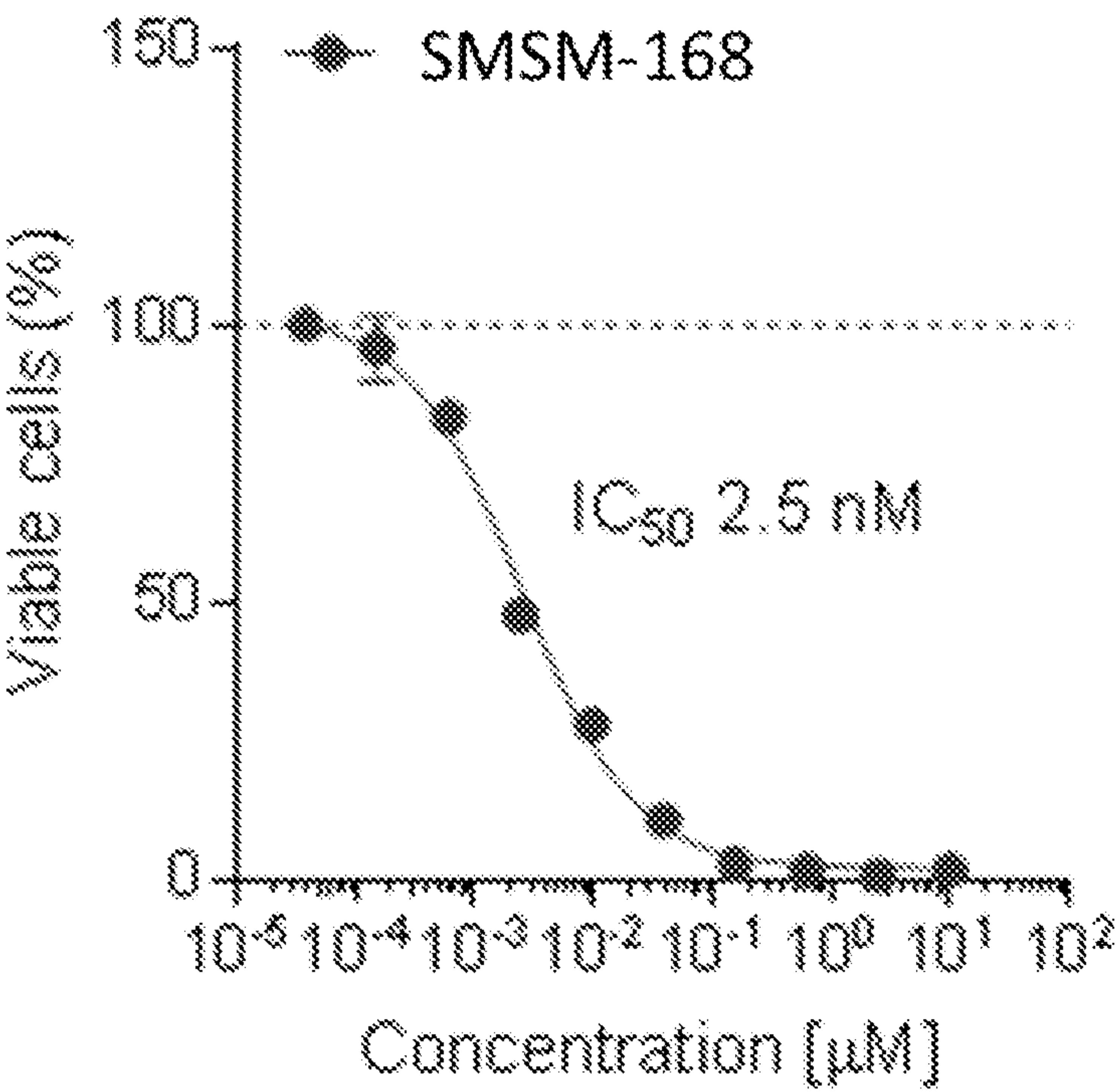


FIG. 9G

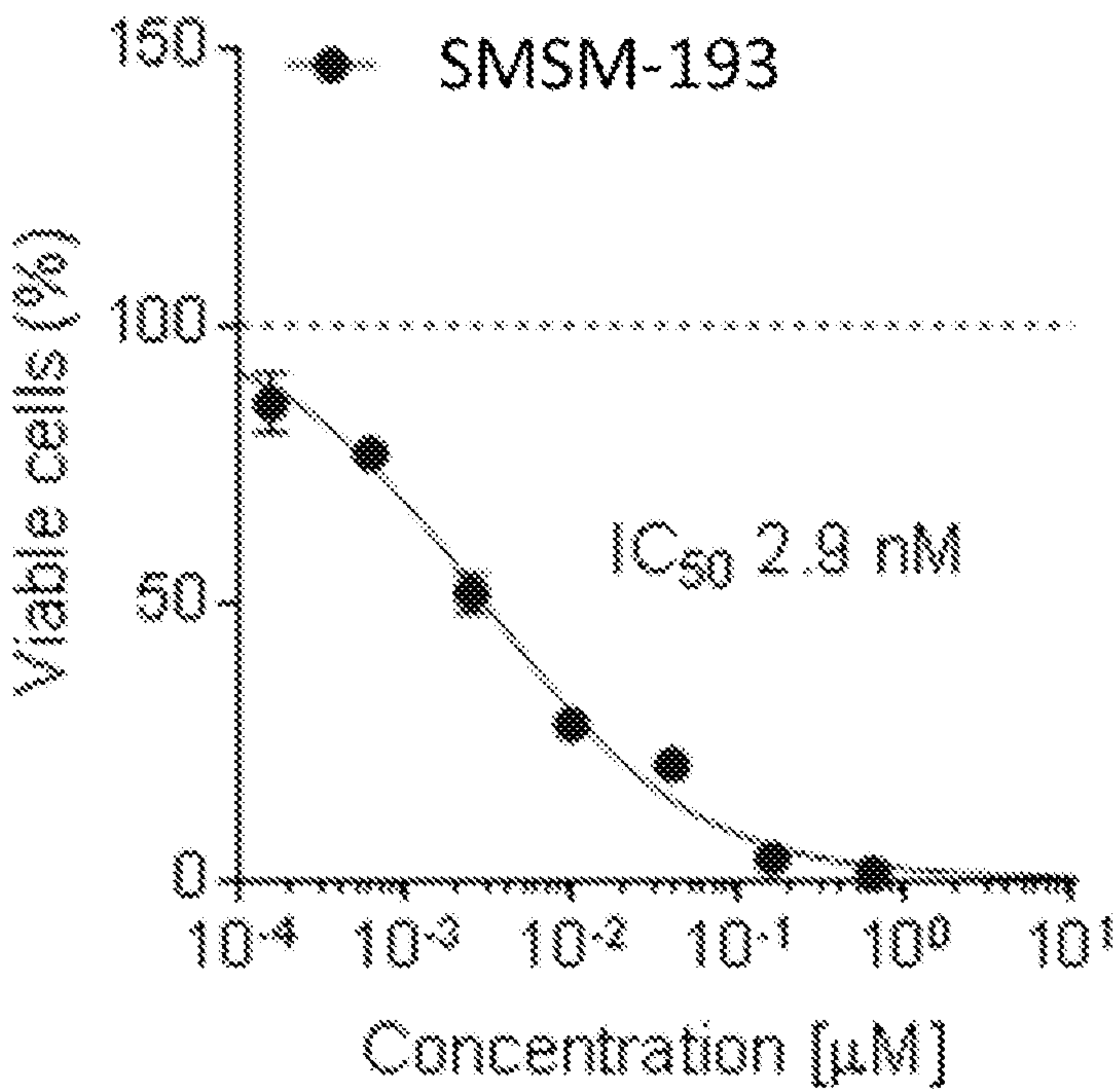


FIG. 9H

12/31

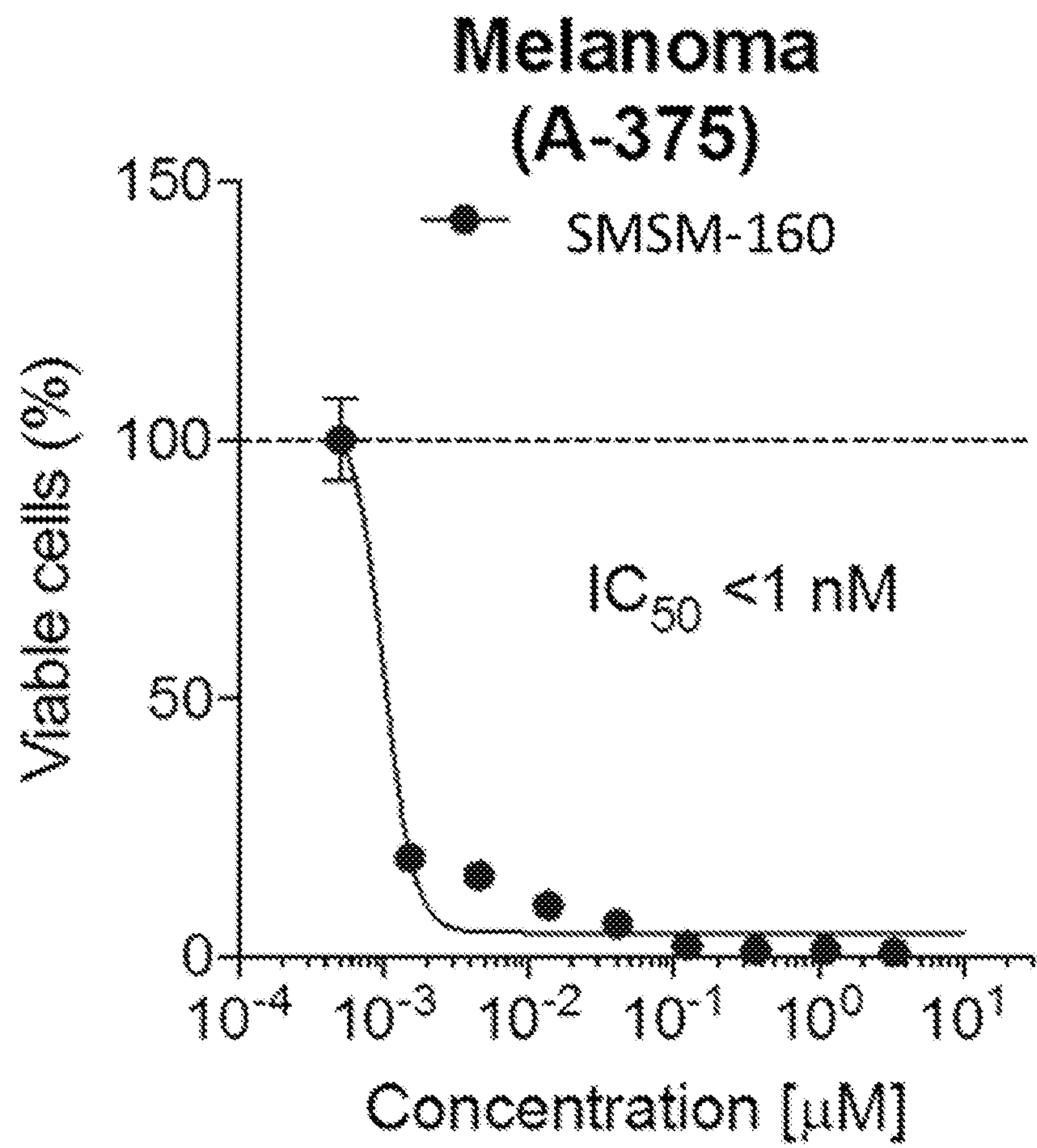


FIG. 10

NHDF

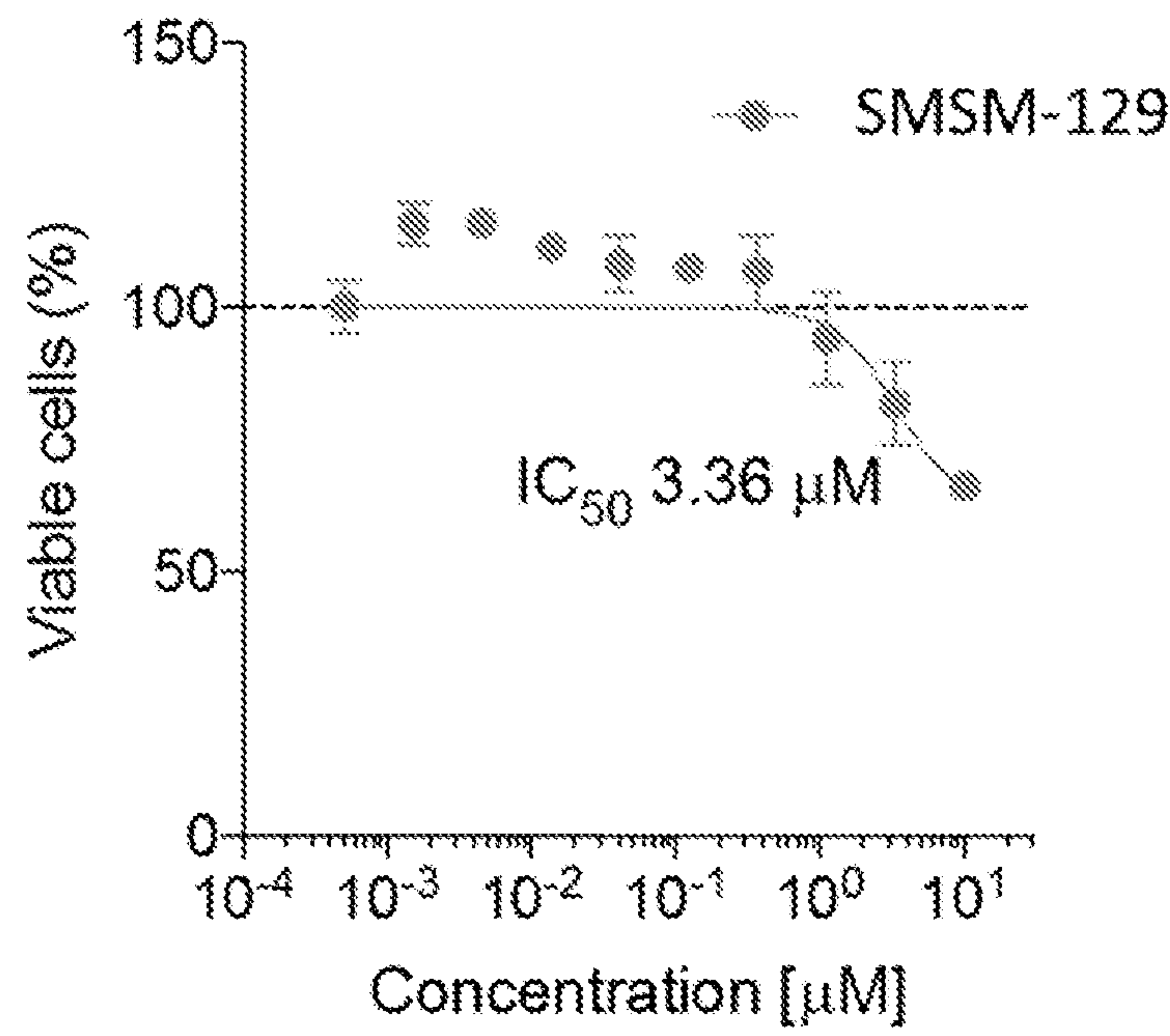


FIG. 11A

NHDF

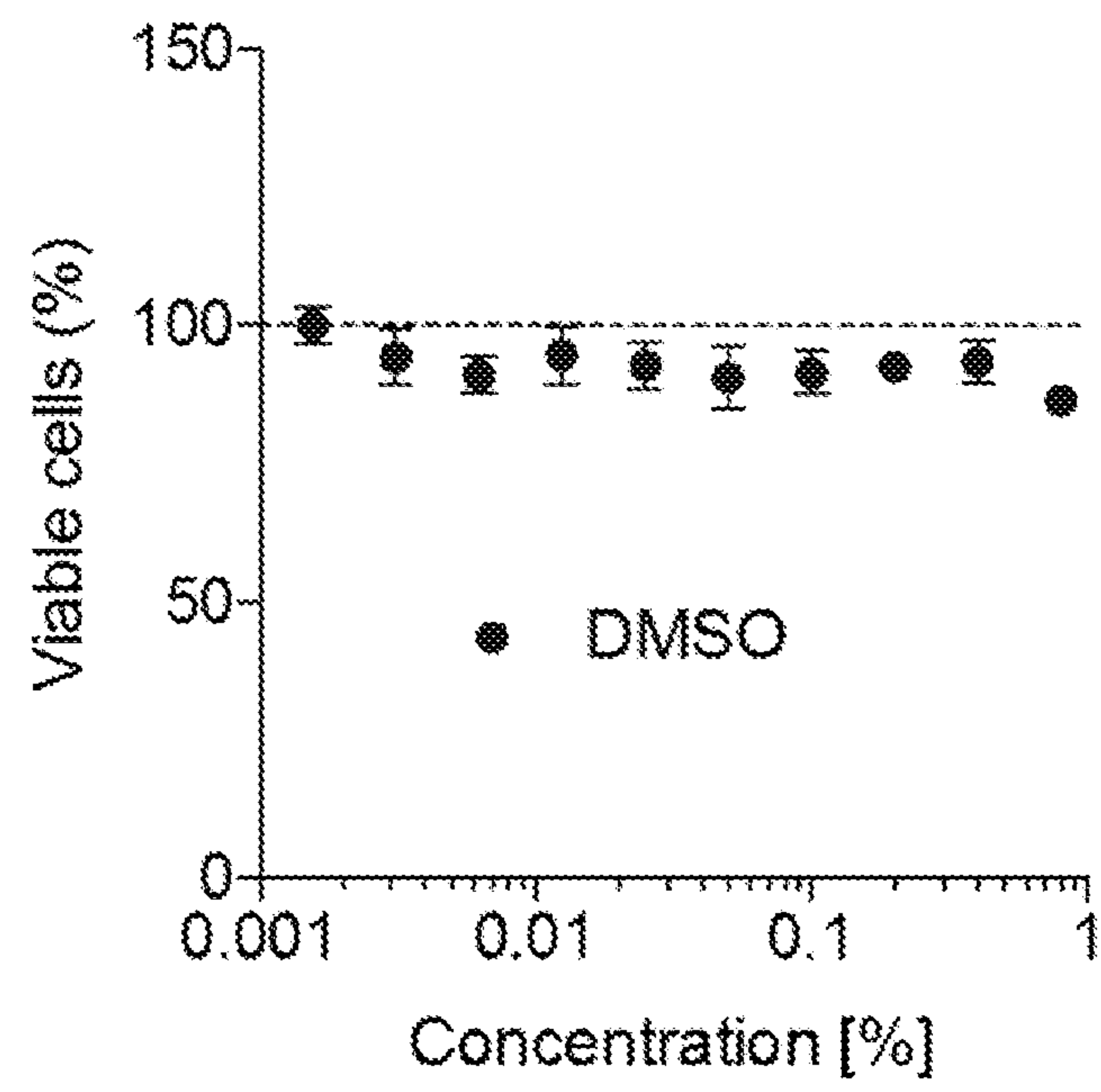


FIG. 11B

Proliferation vs. splicing

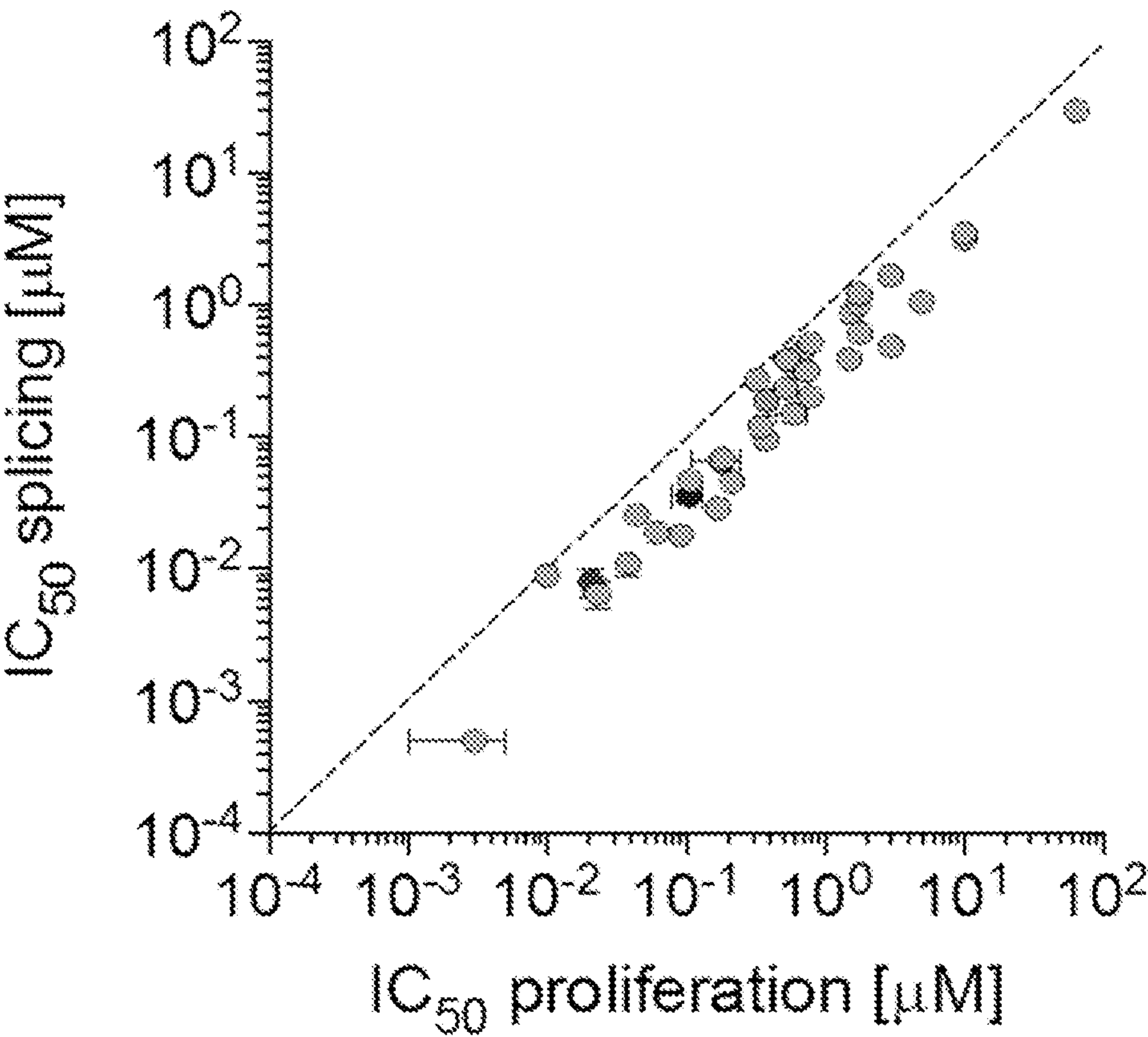
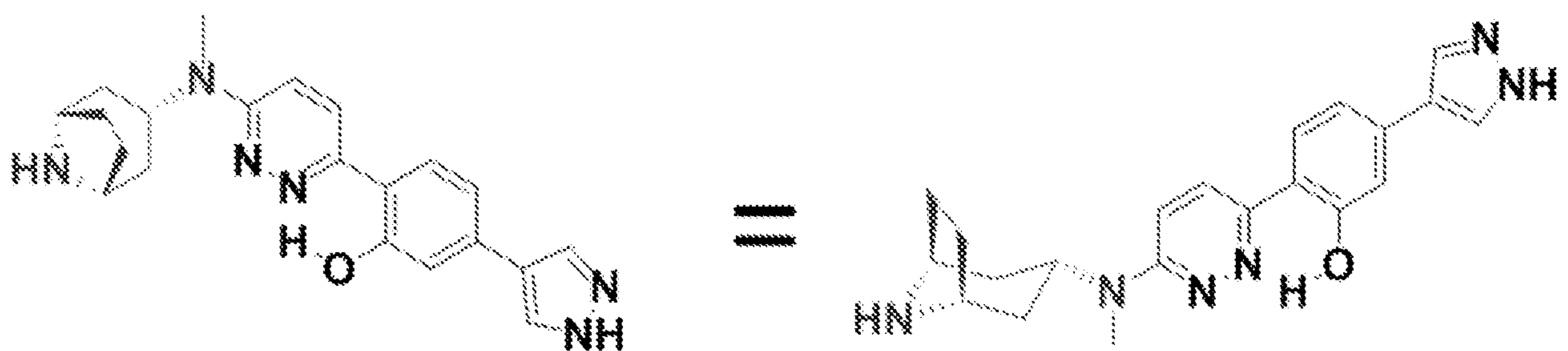


FIG. 12

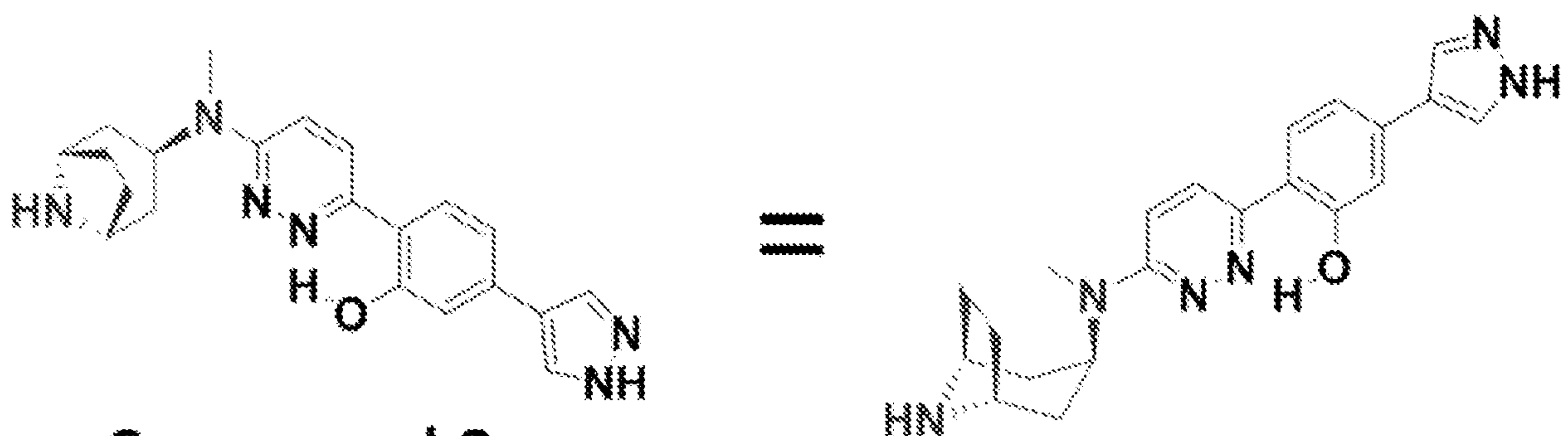
15/31

Impact of Stereochemistry on Cell Activity



Compound 1

Cell_{A-673} = 18 nM



Compound 2

Cell_{A-673} > 100 mM

FIG. 13

16/31

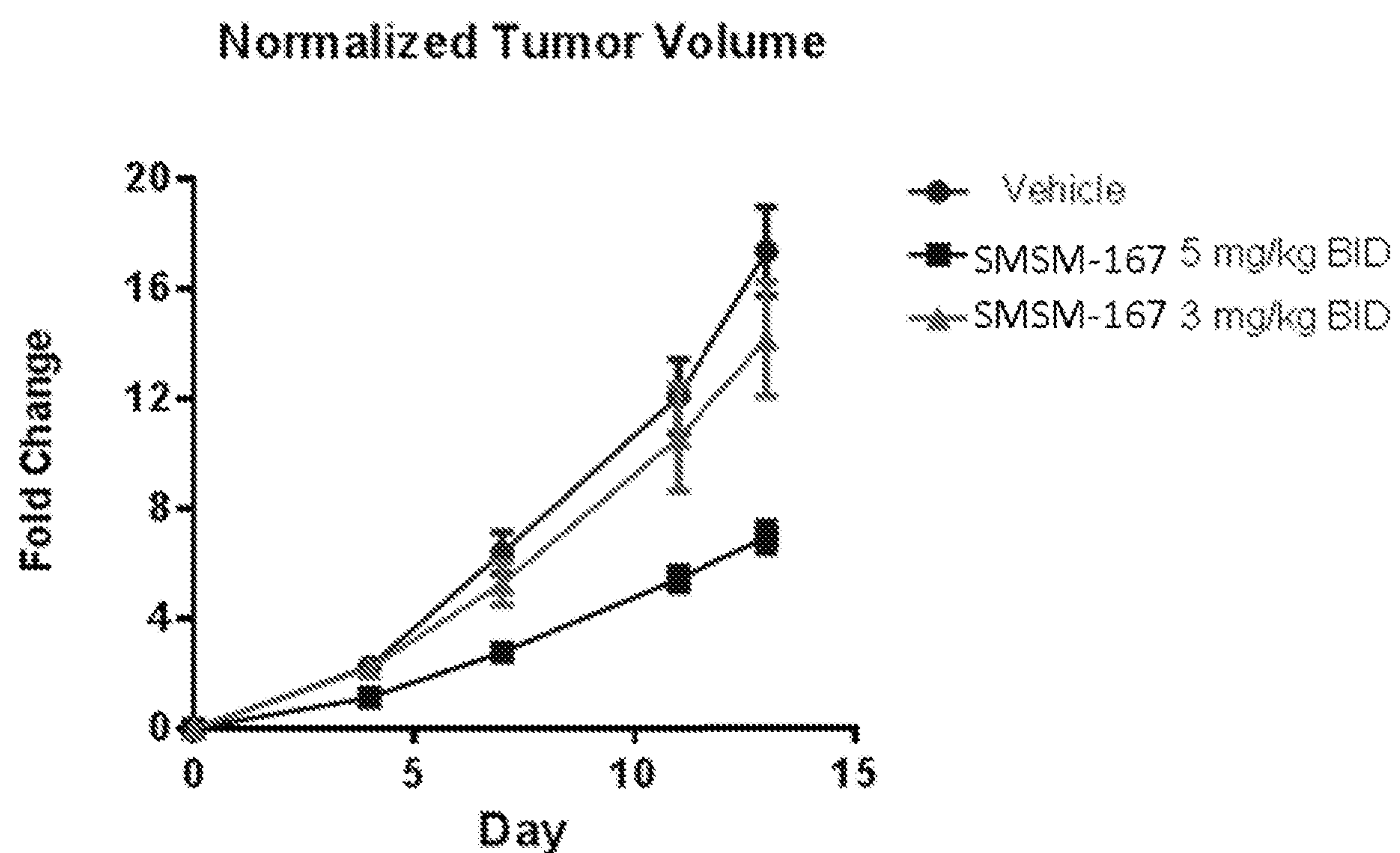


FIG. 14A

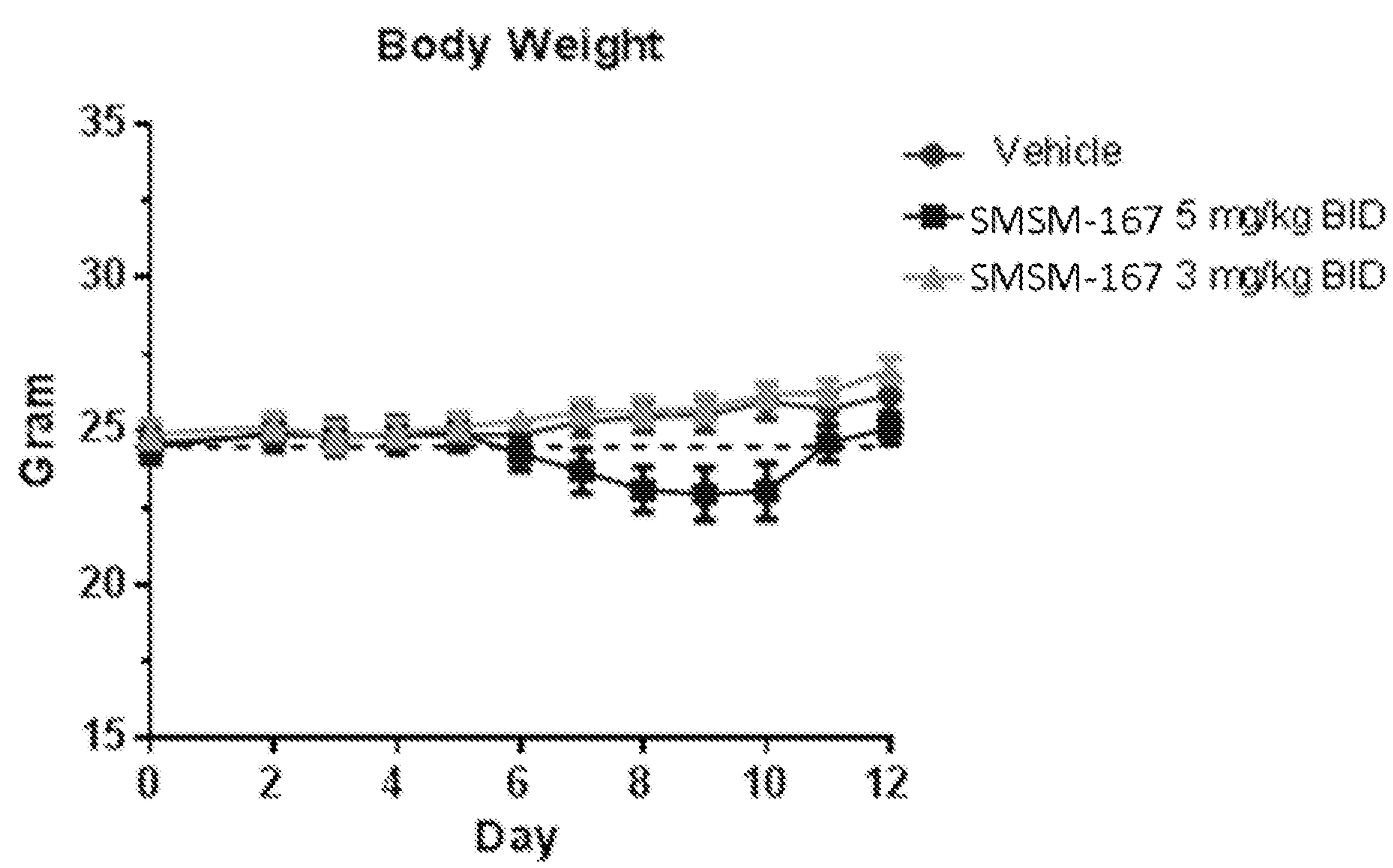


FIG. 14B

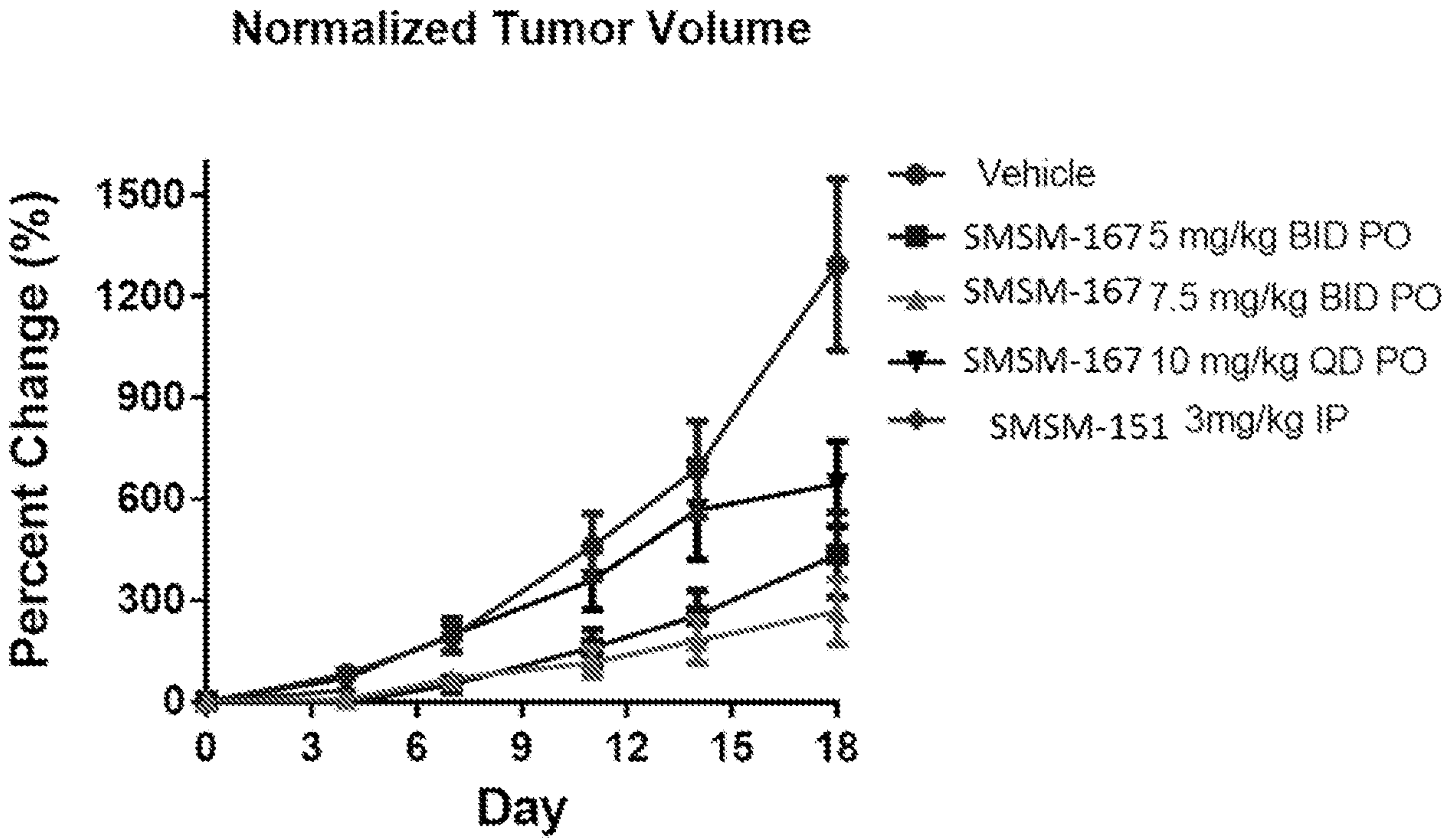


FIG. 14C

18/31

Tumor Volume

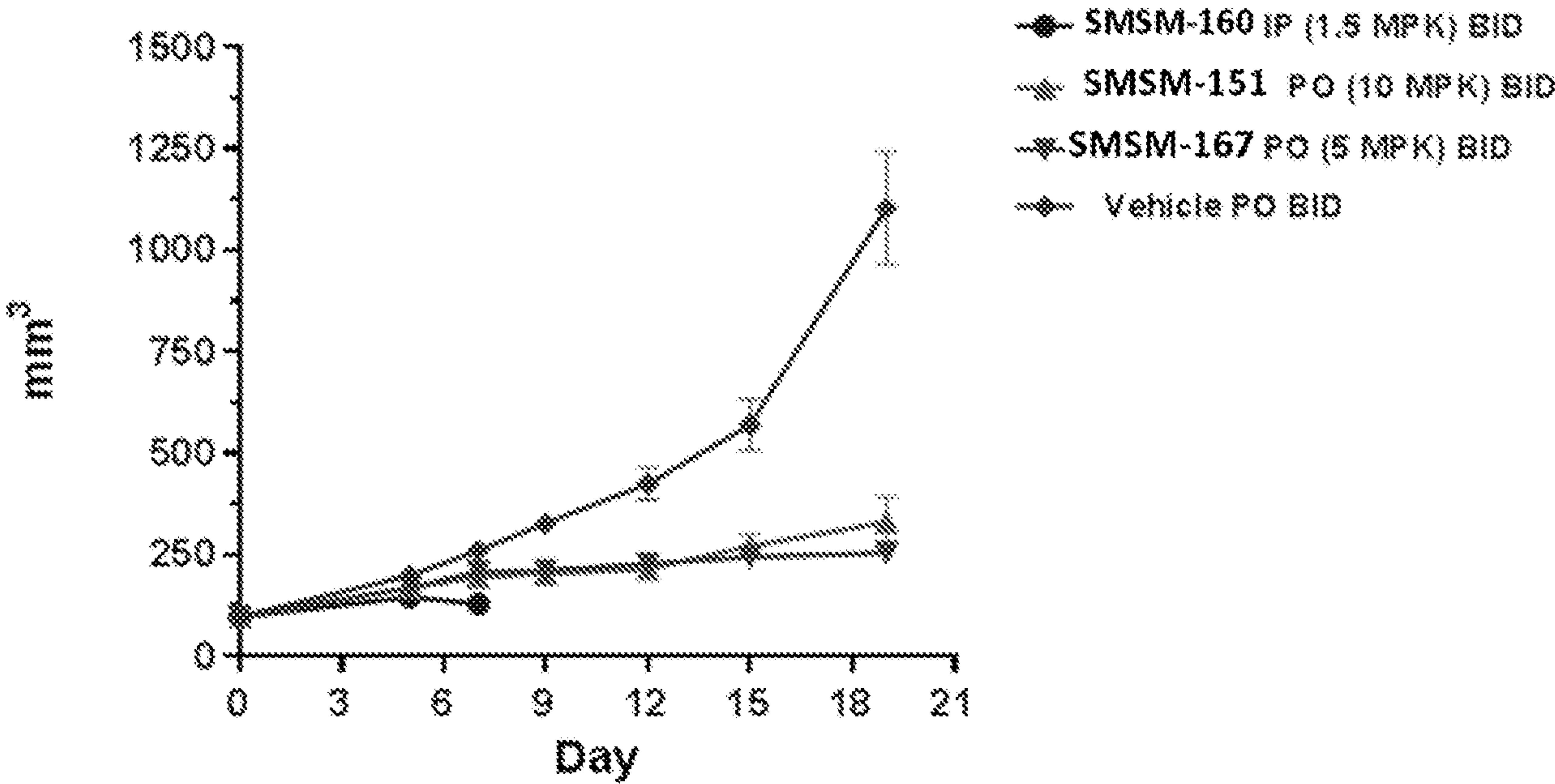


FIG. 15A

Body Weight

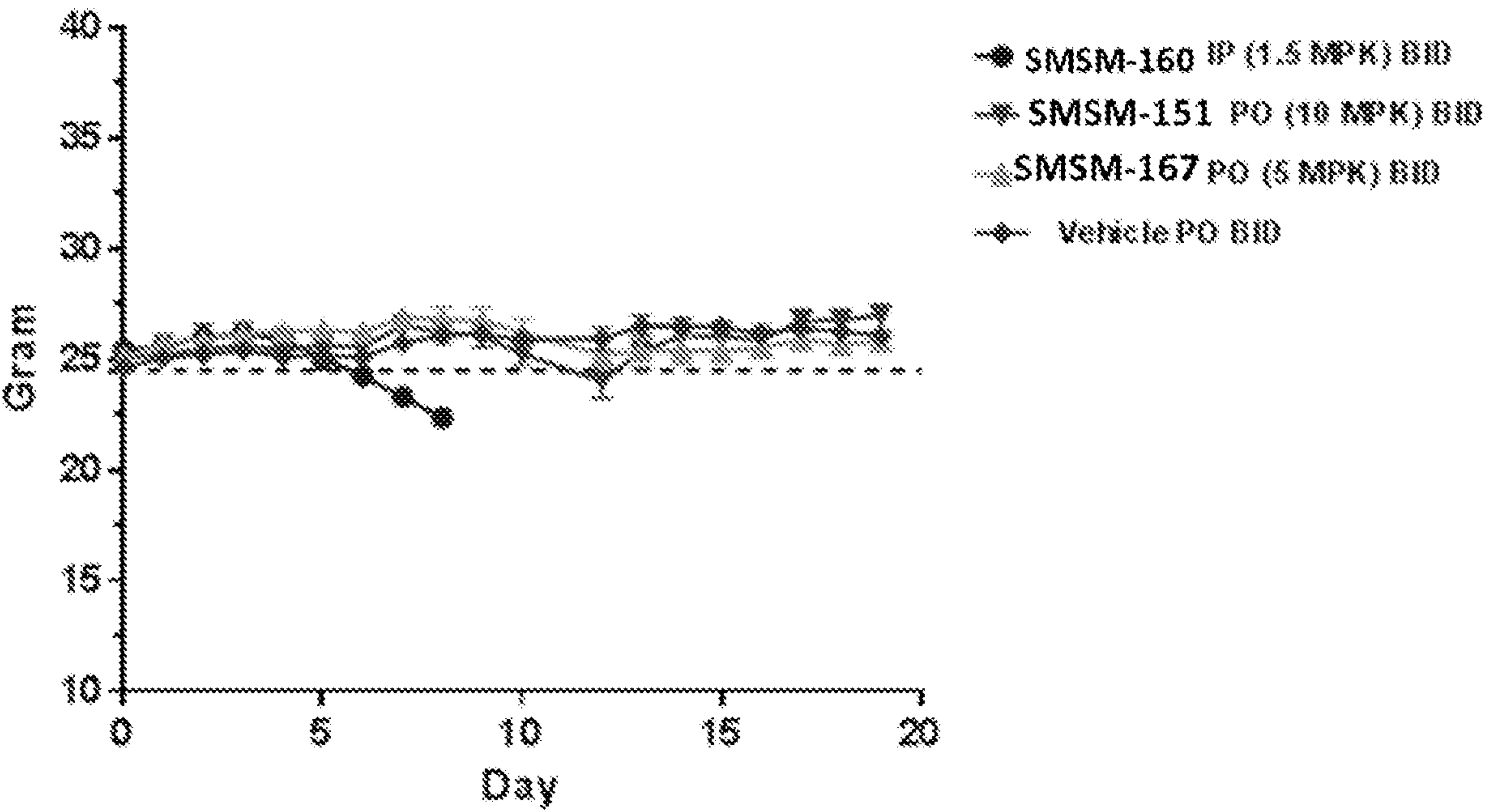


FIG. 15B

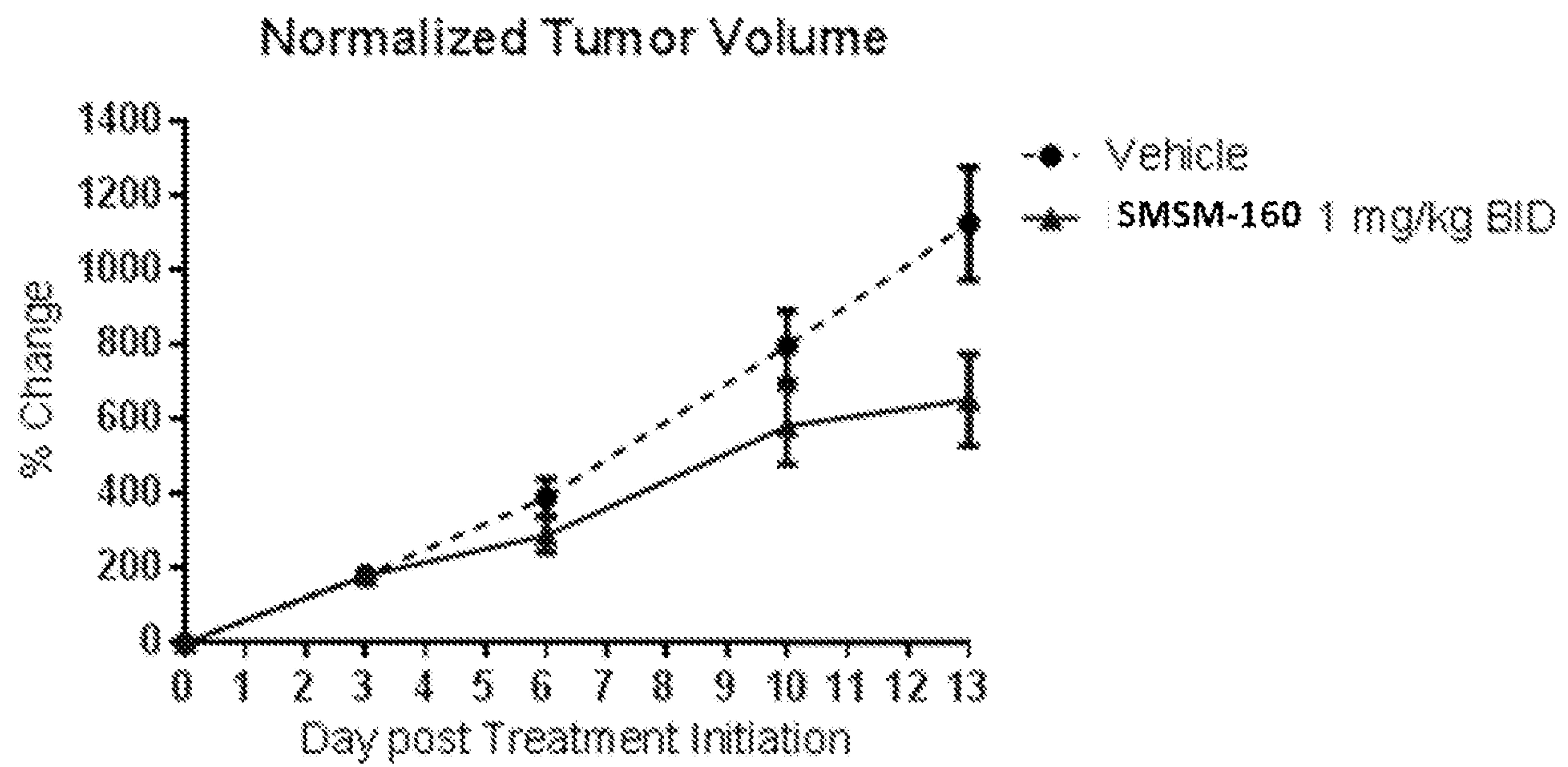


FIG. 16A

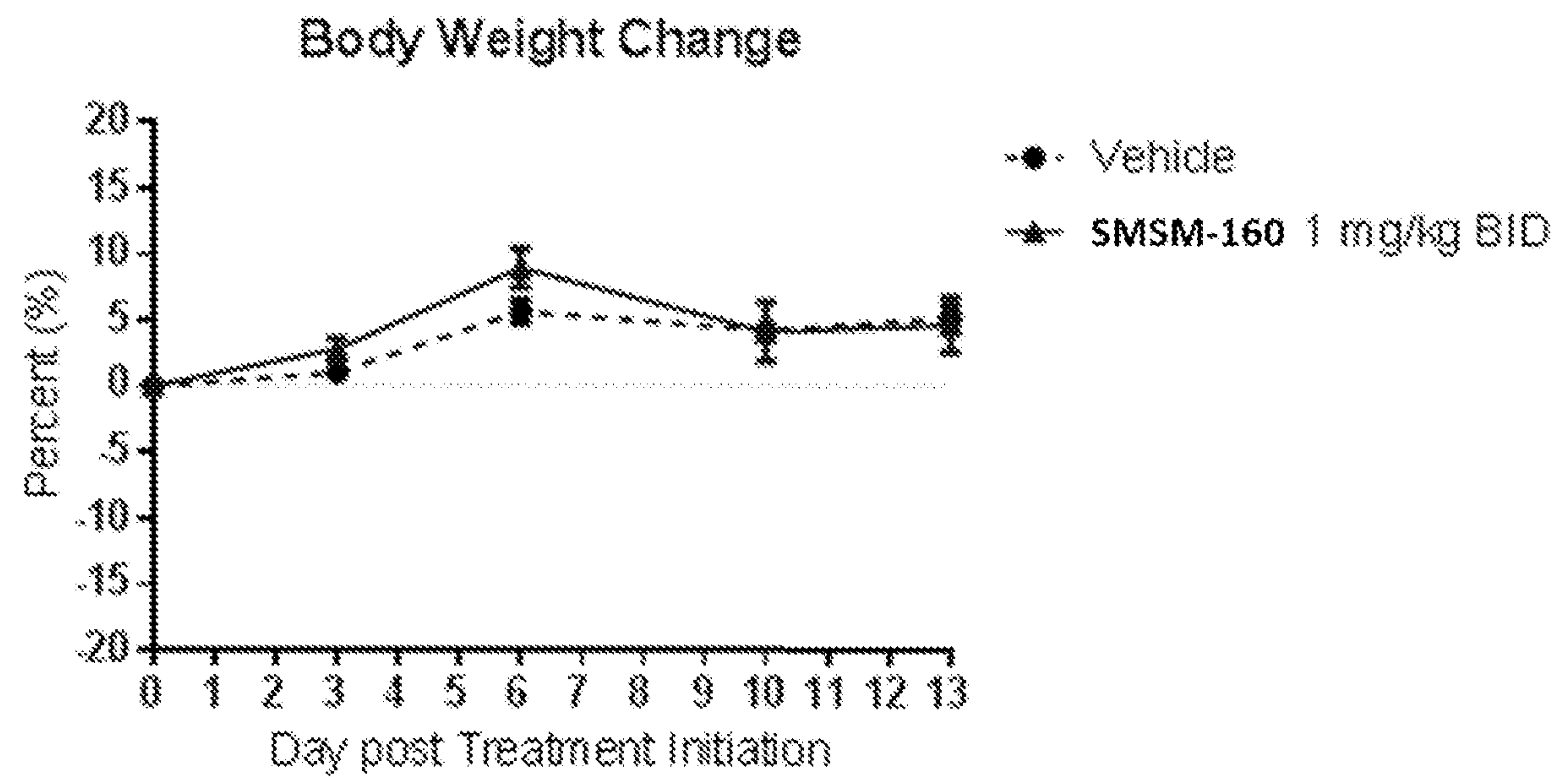


FIG. 16B

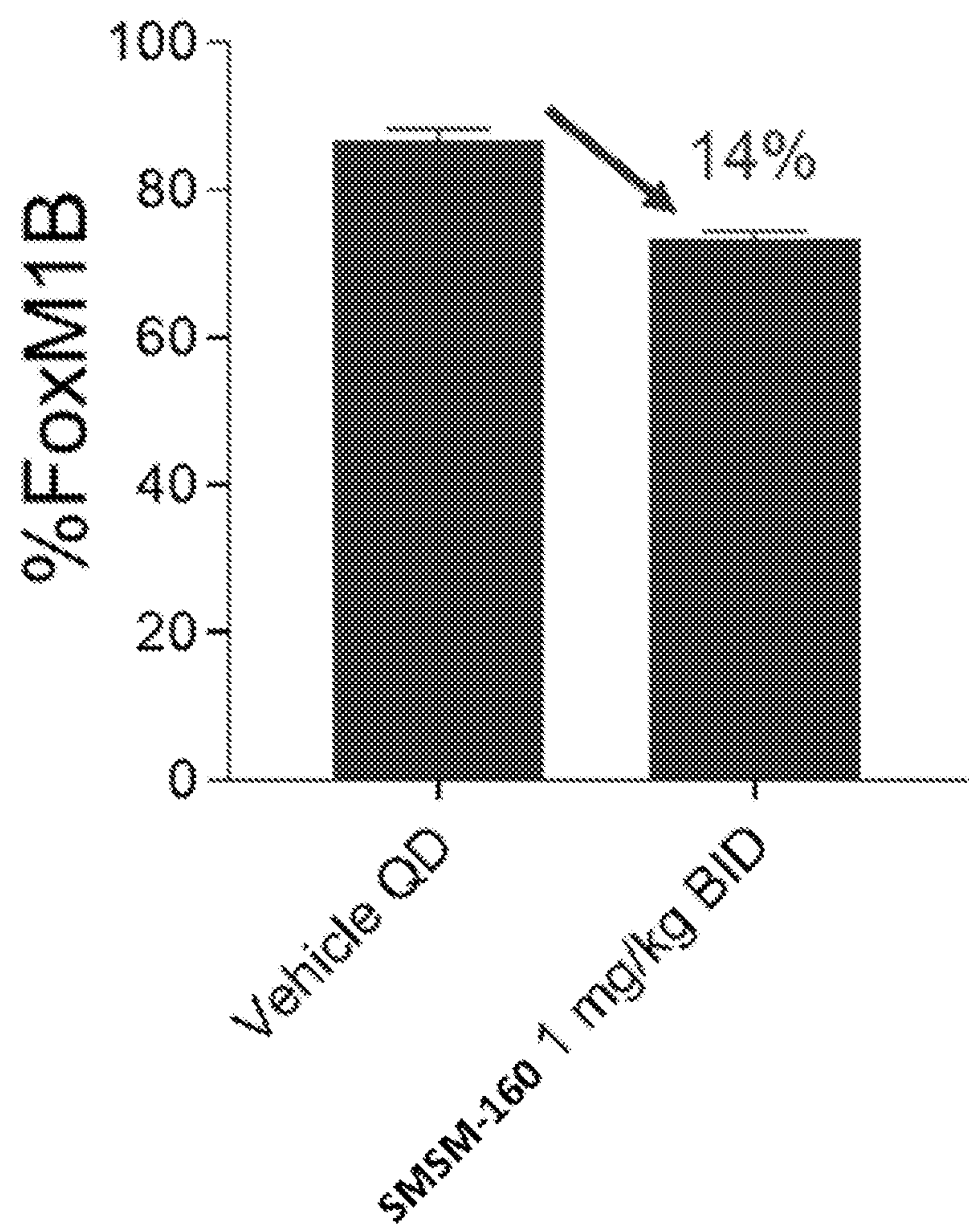


FIG. 16C

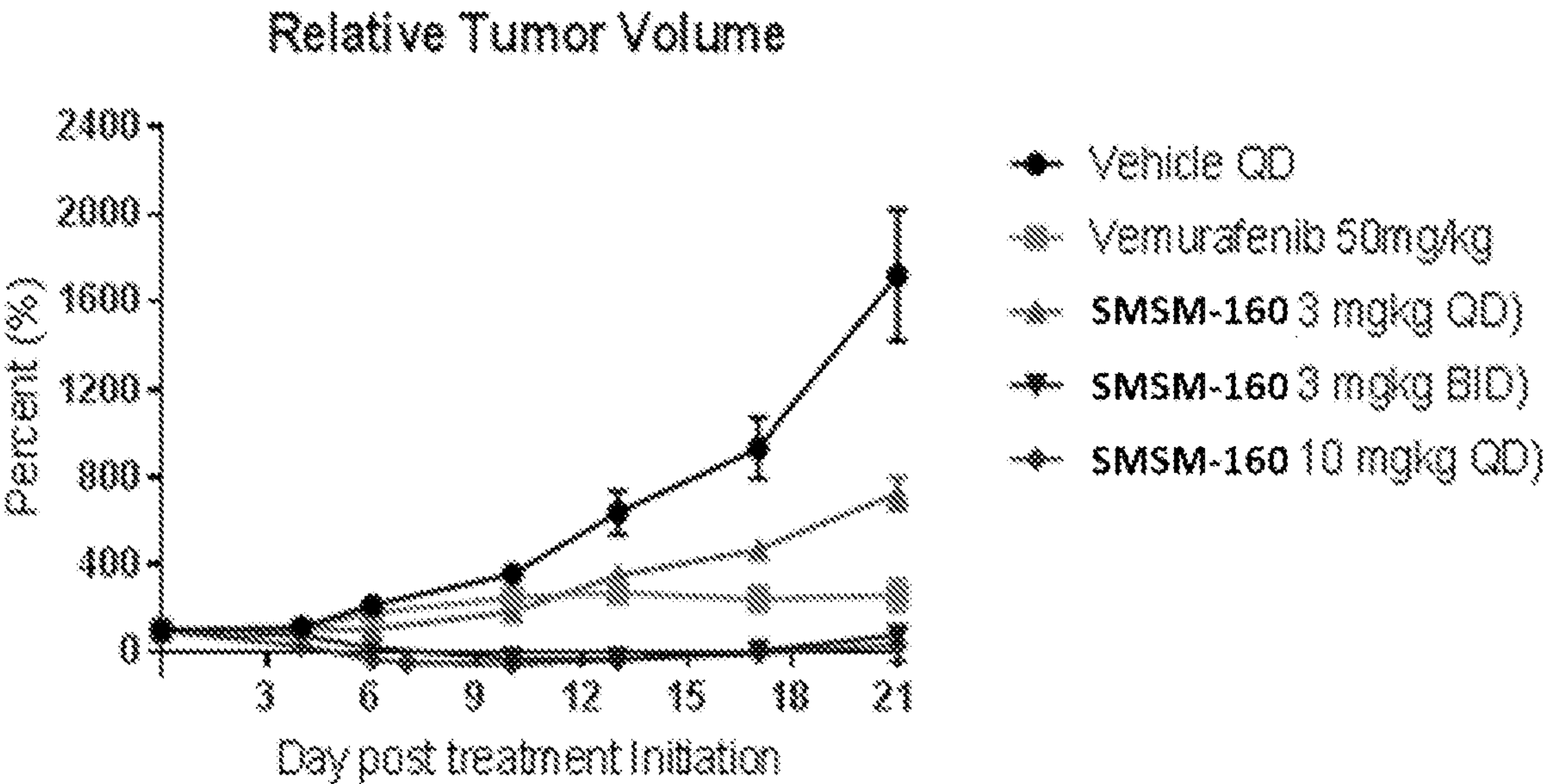


FIG. 17A

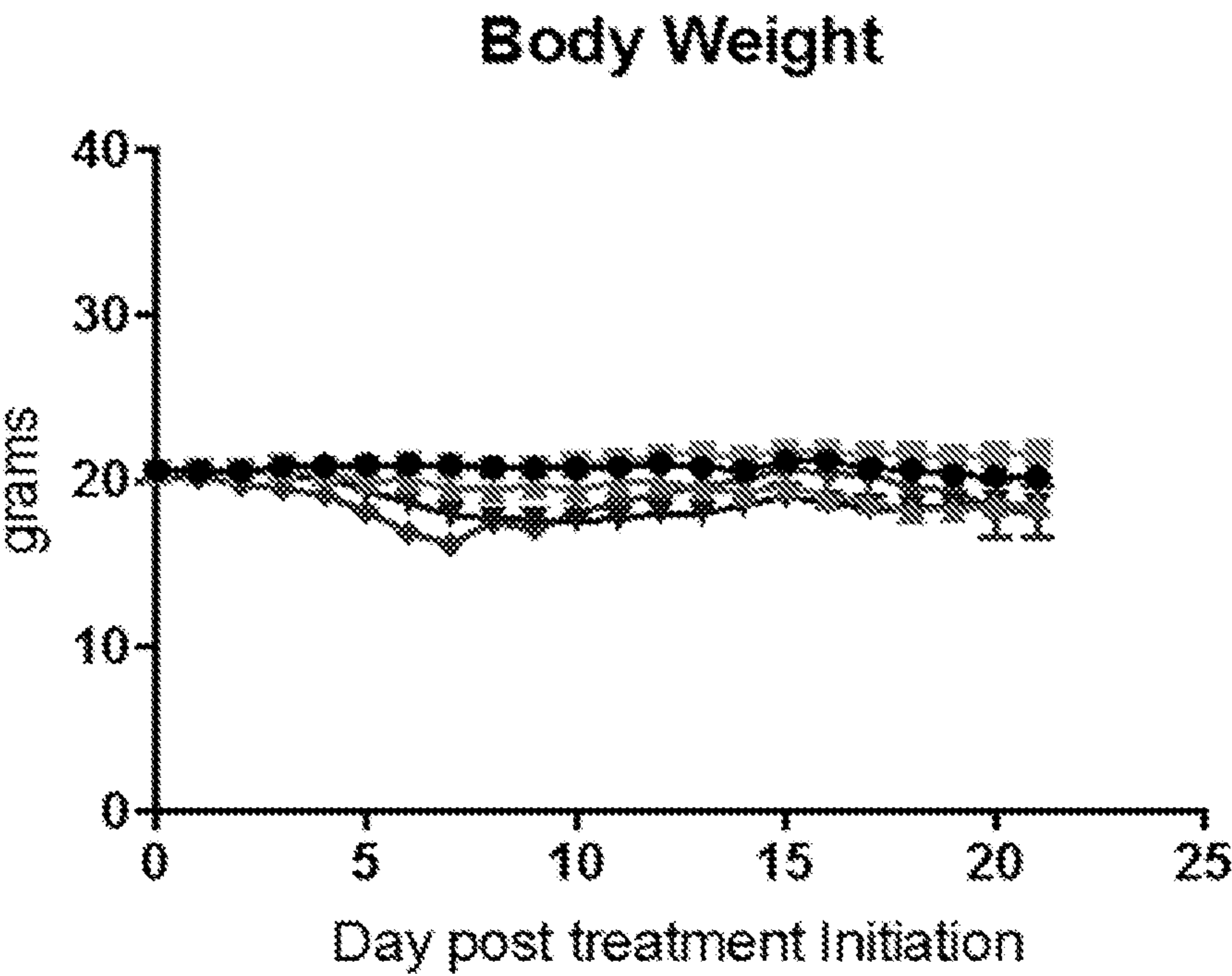


FIG. 17B

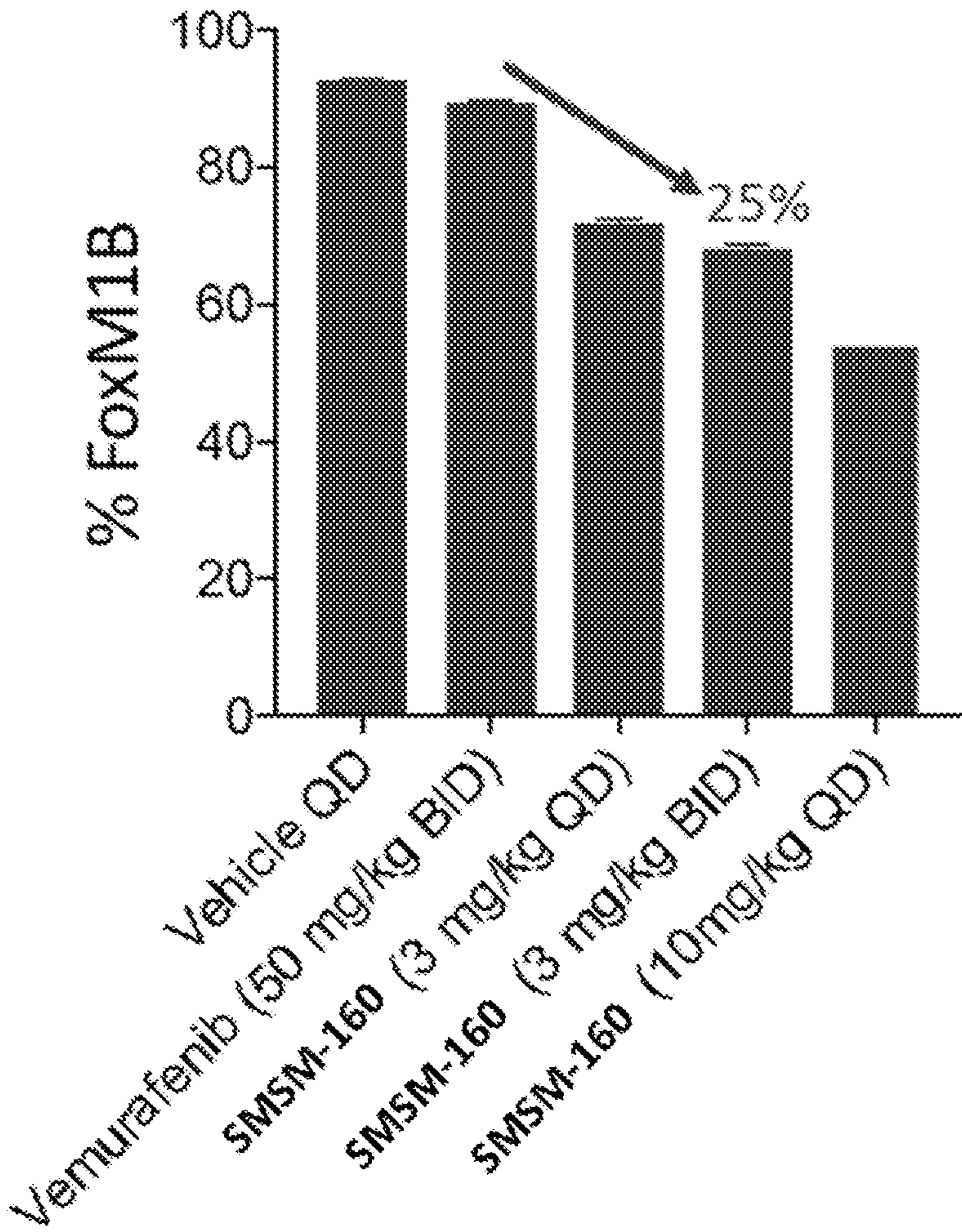


FIG. 17C

23/31

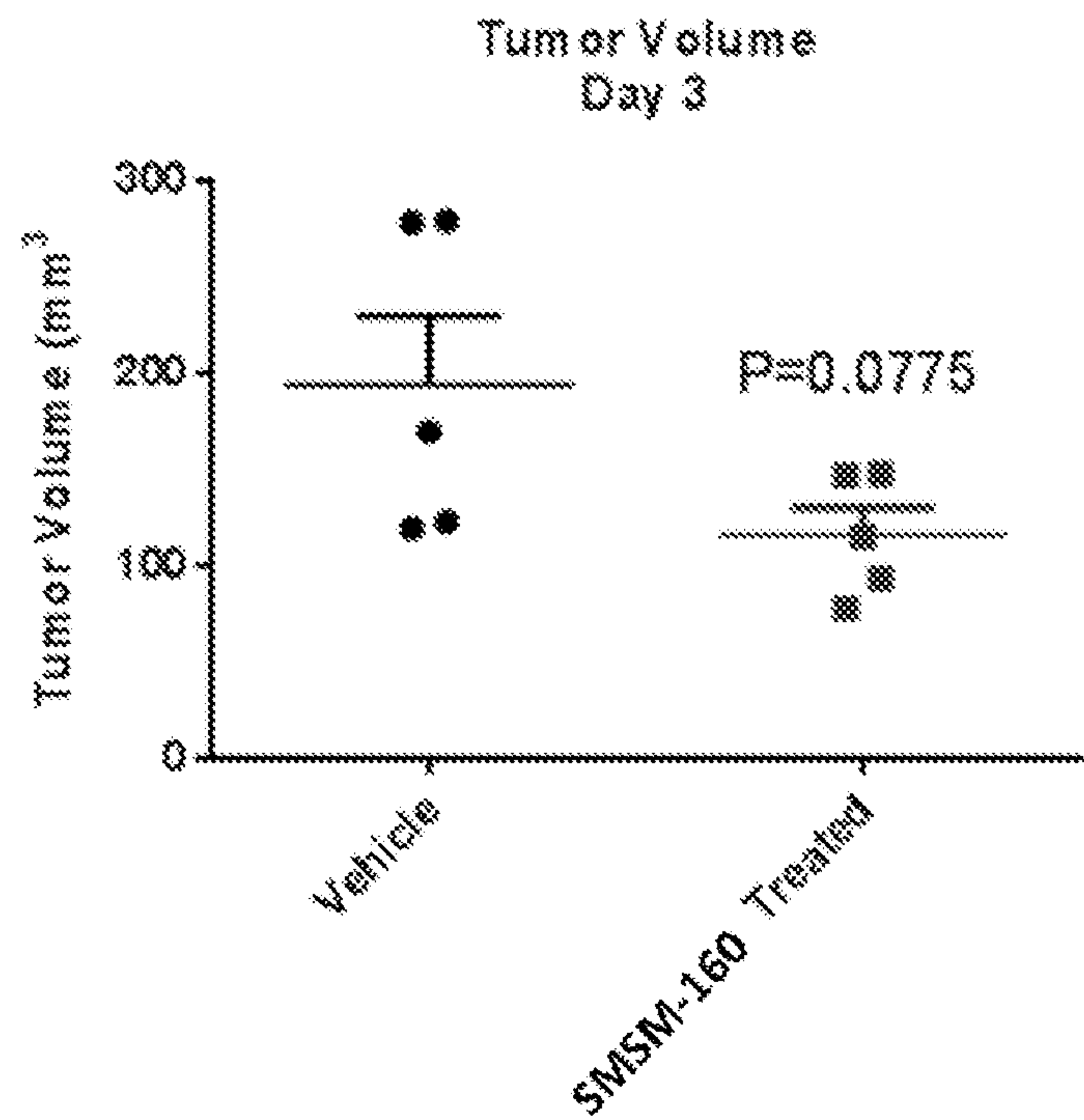


FIG. 18A

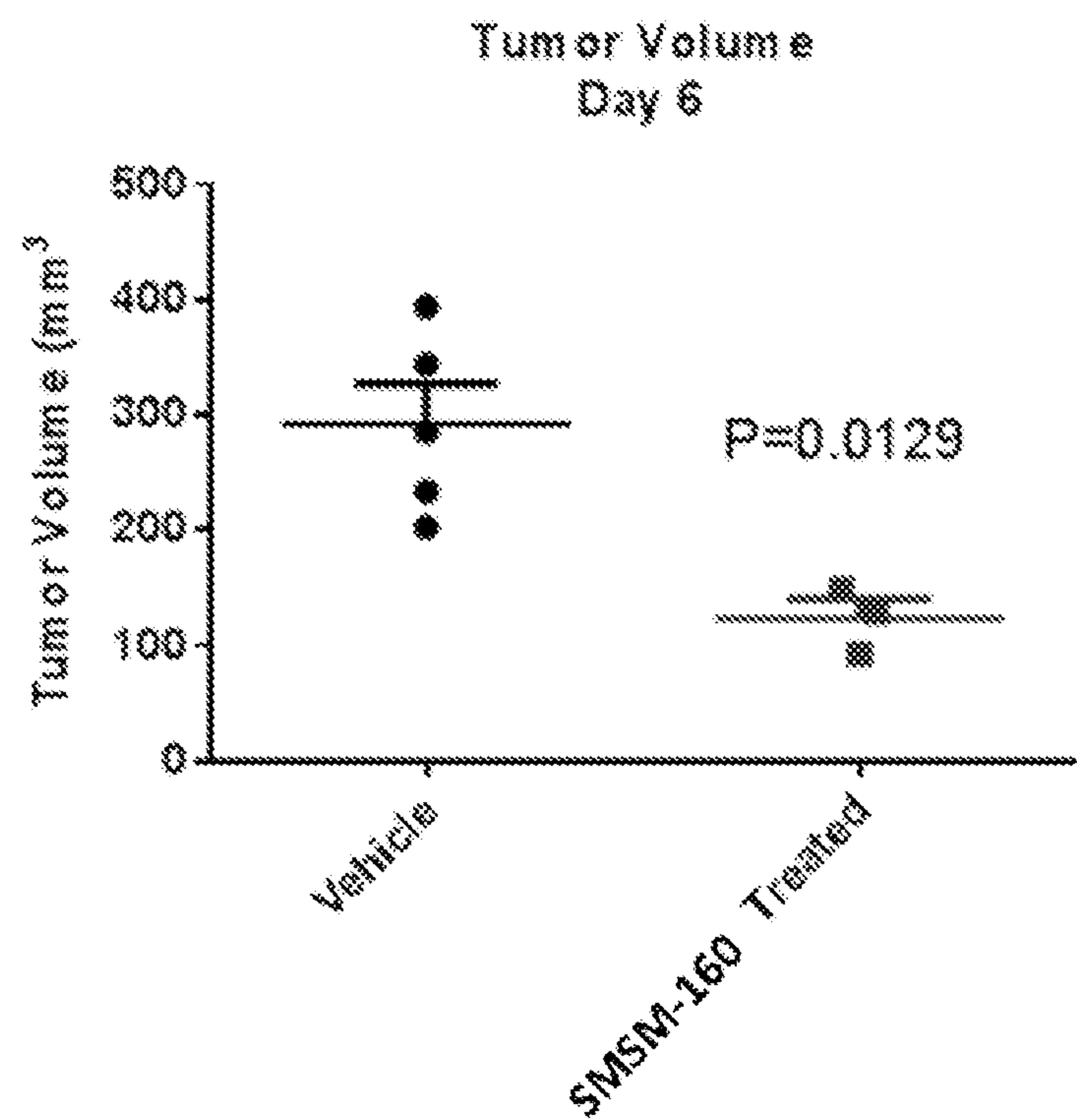


FIG. 18B

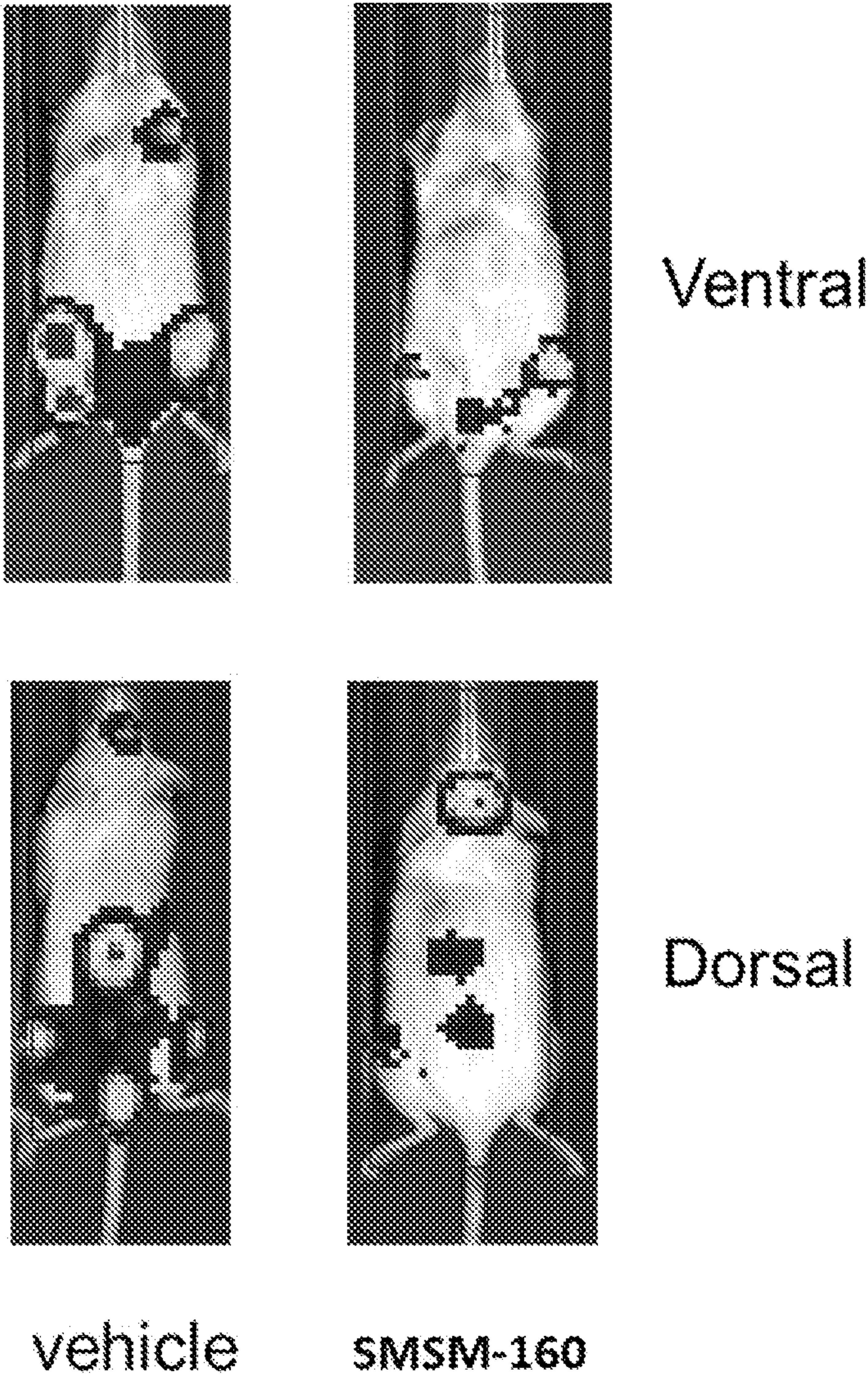


FIG. 19A

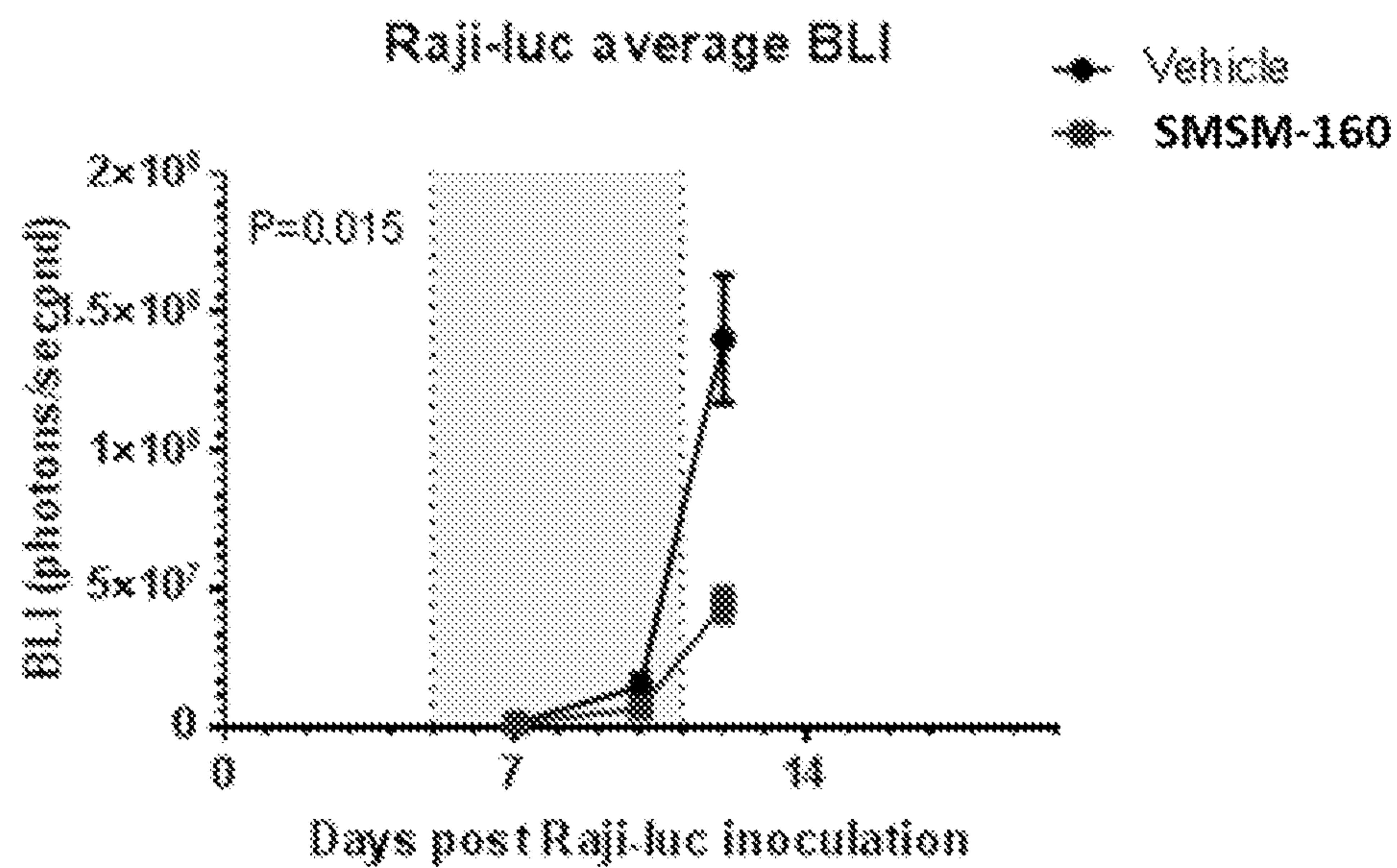


FIG. 19B

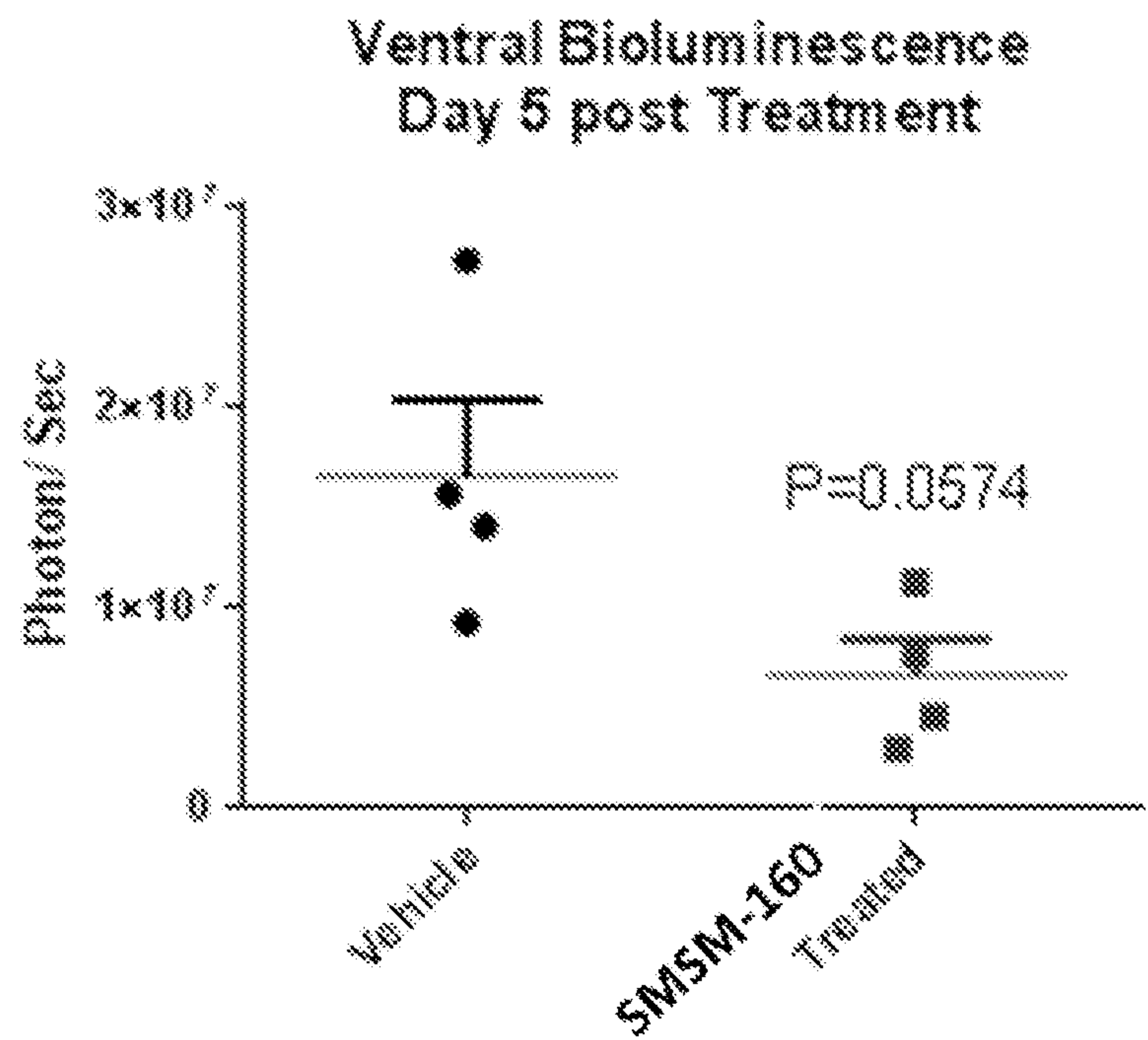


FIG. 19C

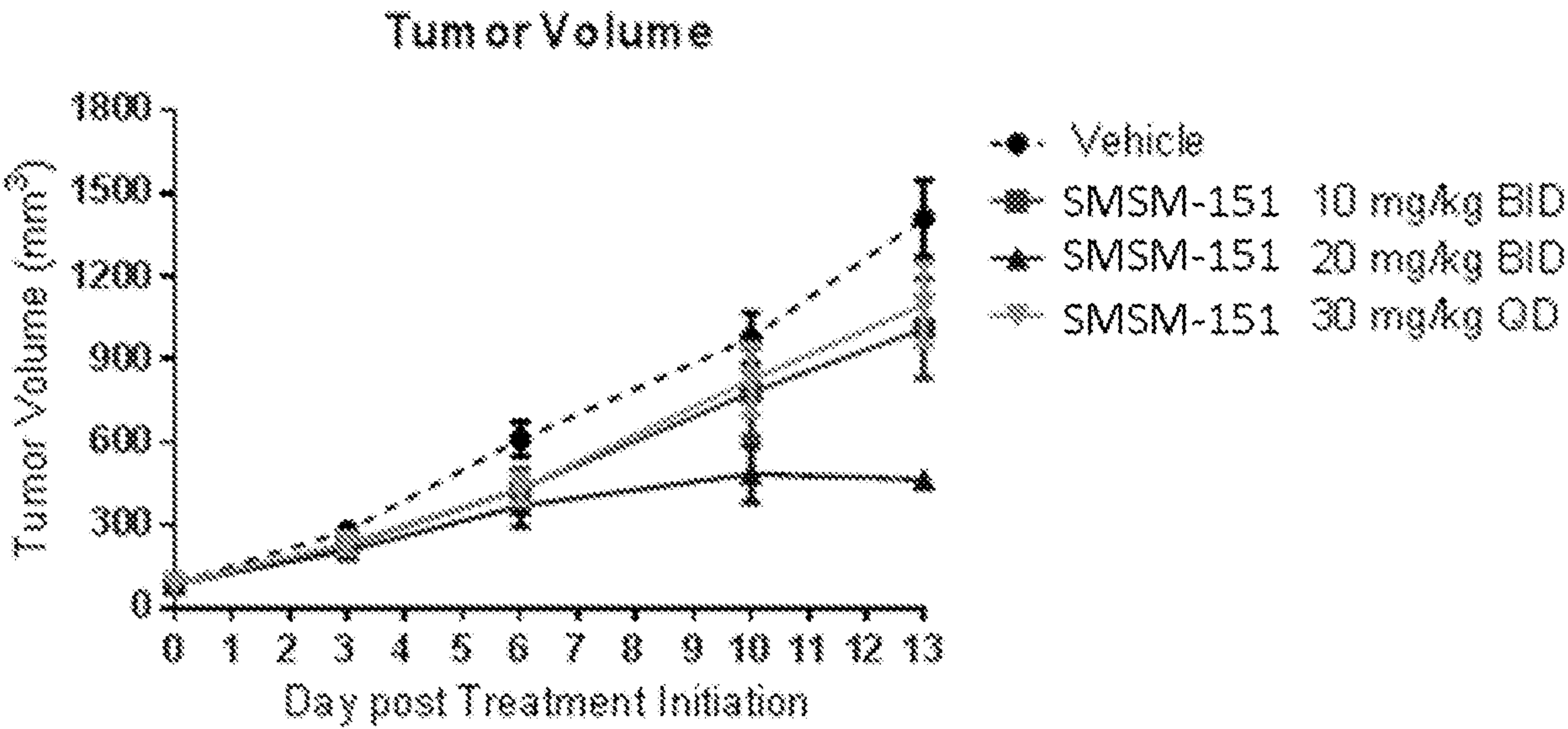


FIG. 20A

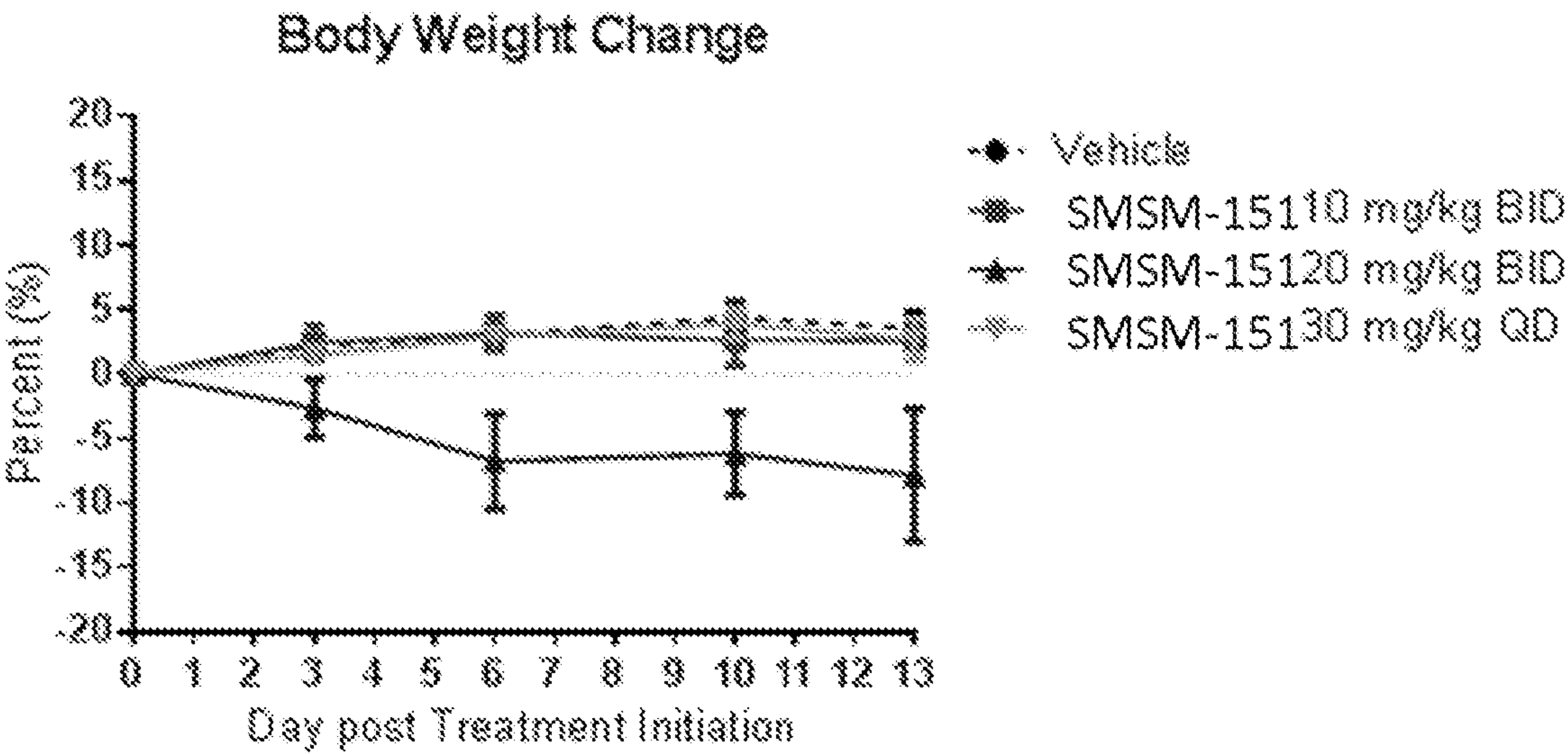


FIG. 20B

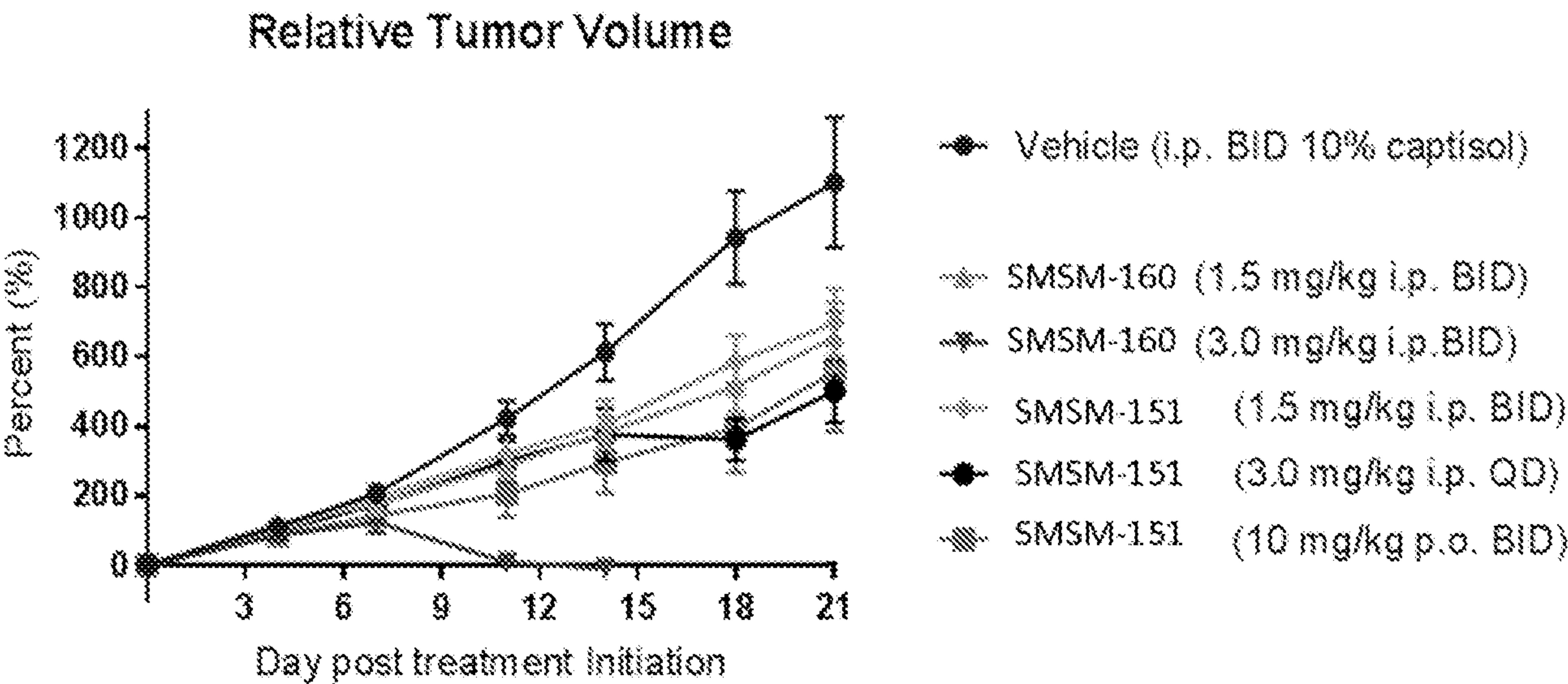


FIG. 21A

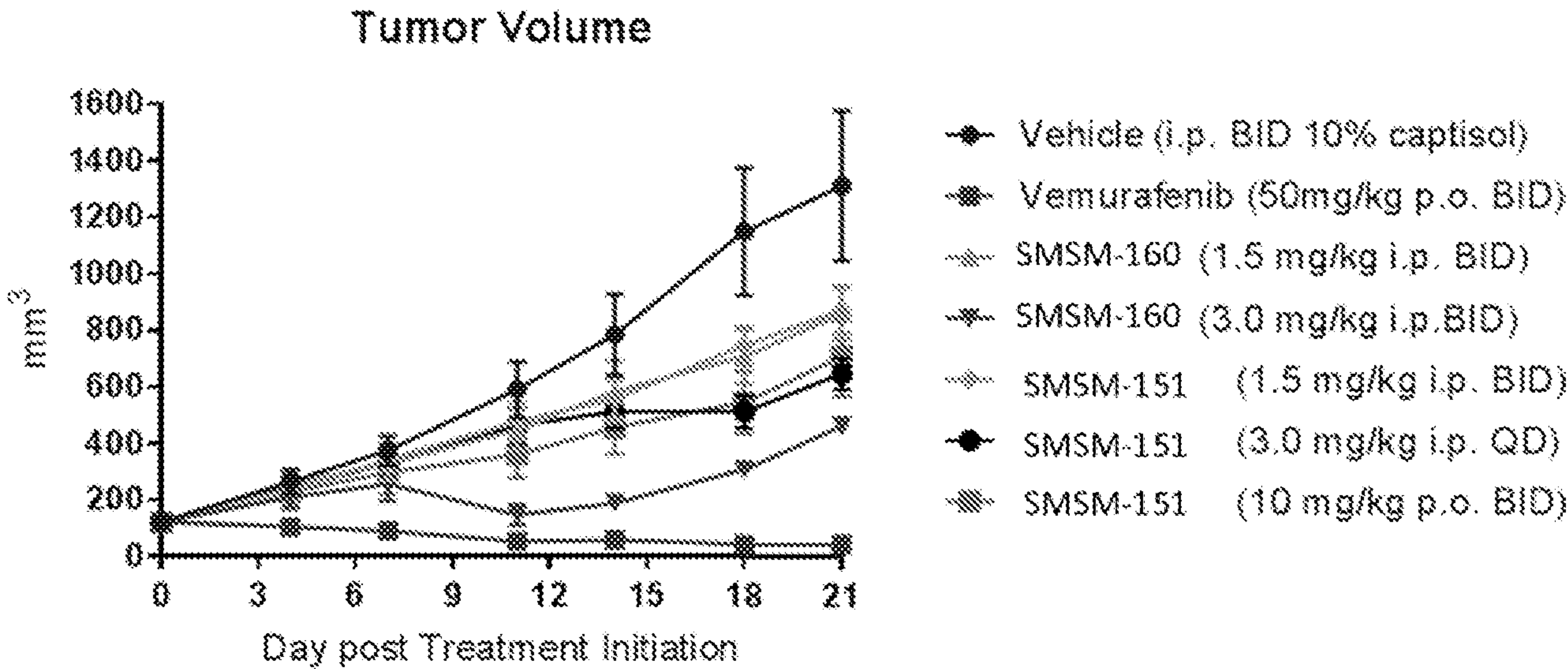


FIG. 21B

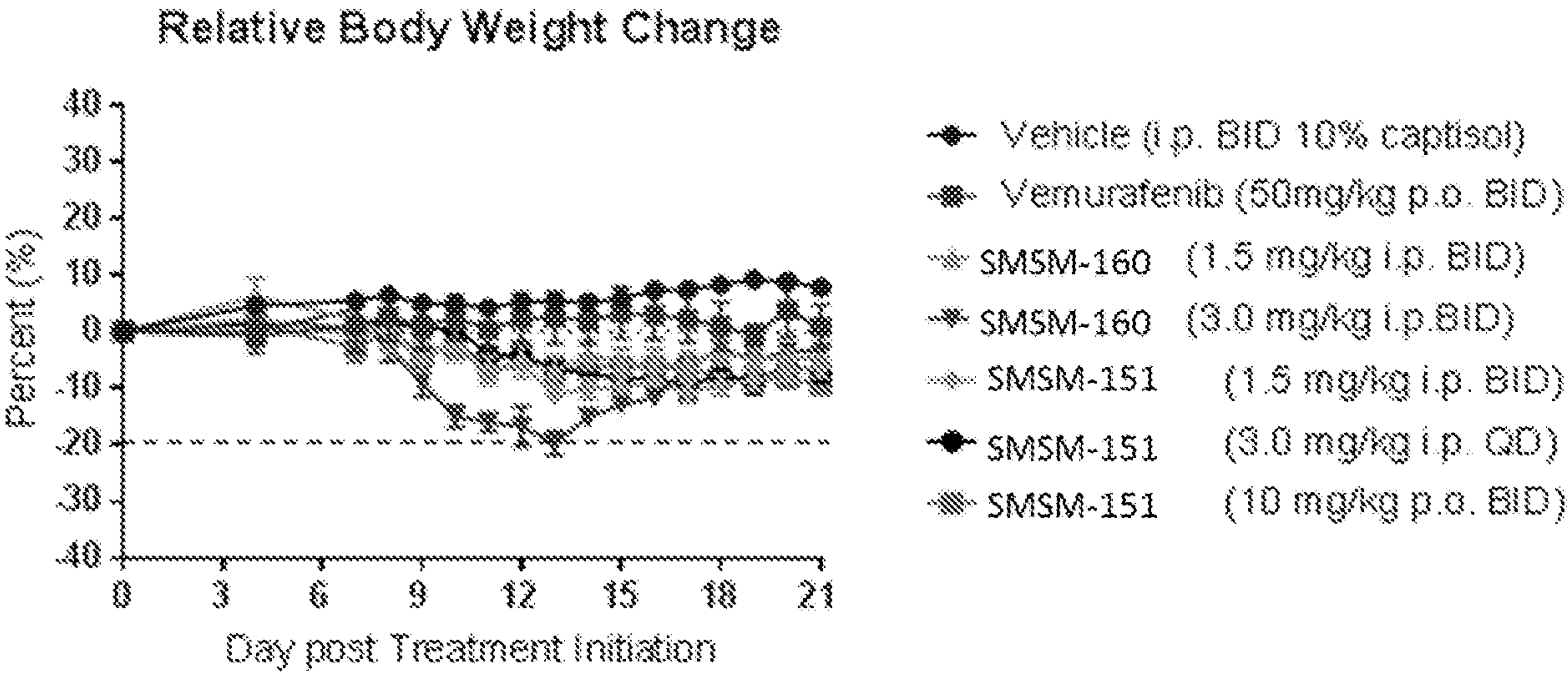
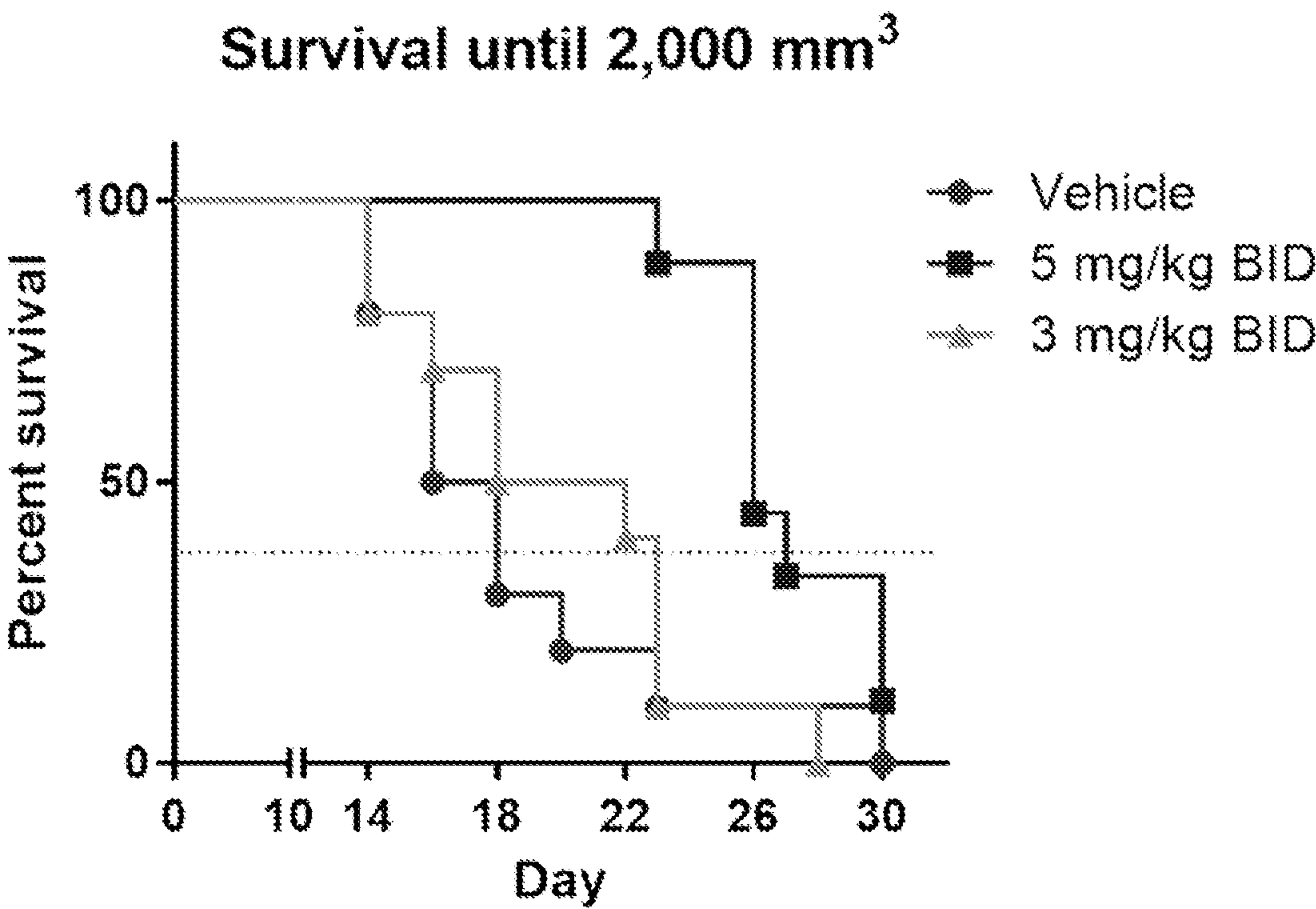


FIG. 21C



Median Survival to 2,000 mm³ tumors:

Vehicle	16 Days
SMSM-167 3 mg/kg	18 Days
SMSM-167 5 mg/kg	26 Days

FIG. 22

30/31

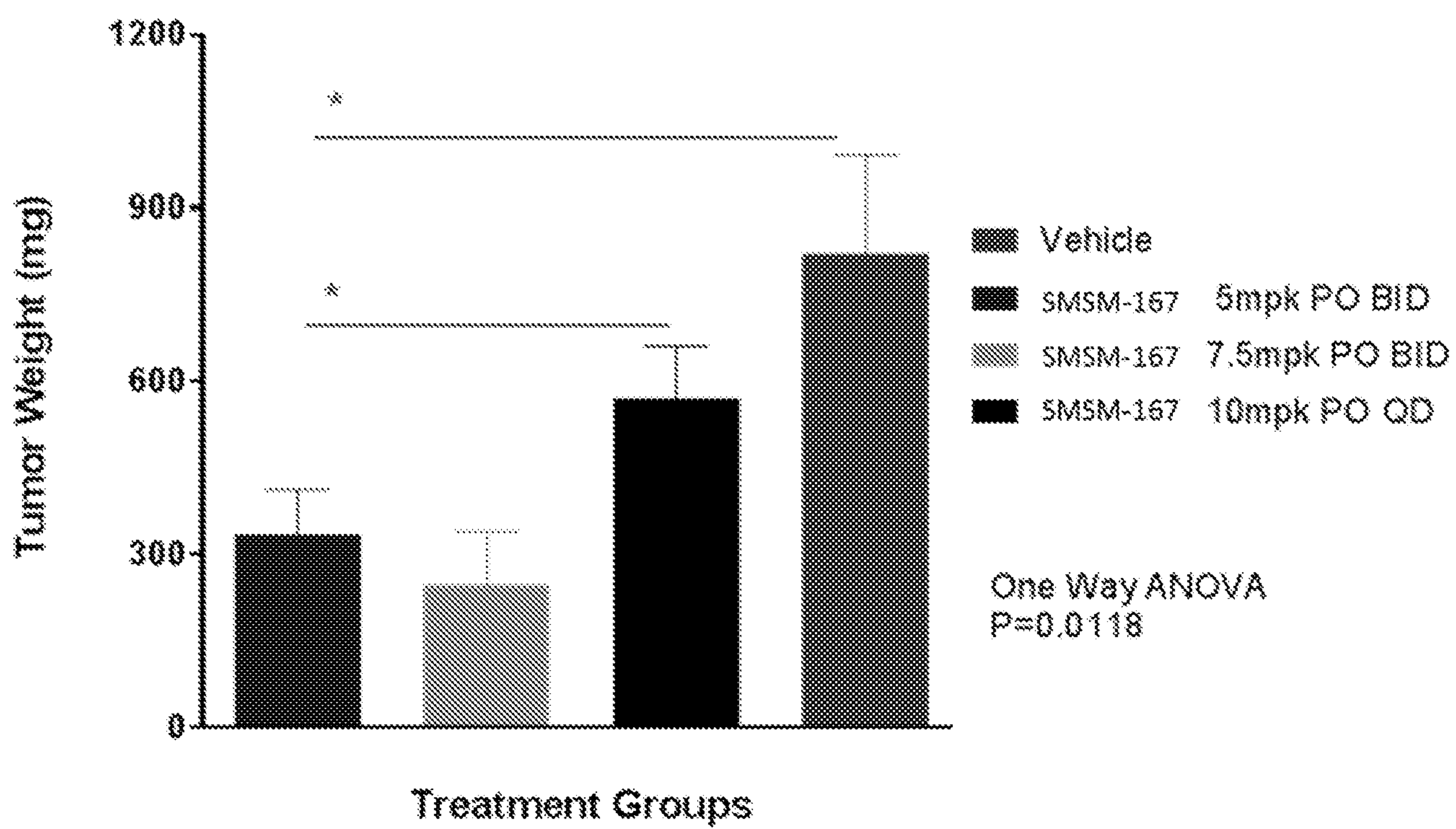


FIG. 23

31/31

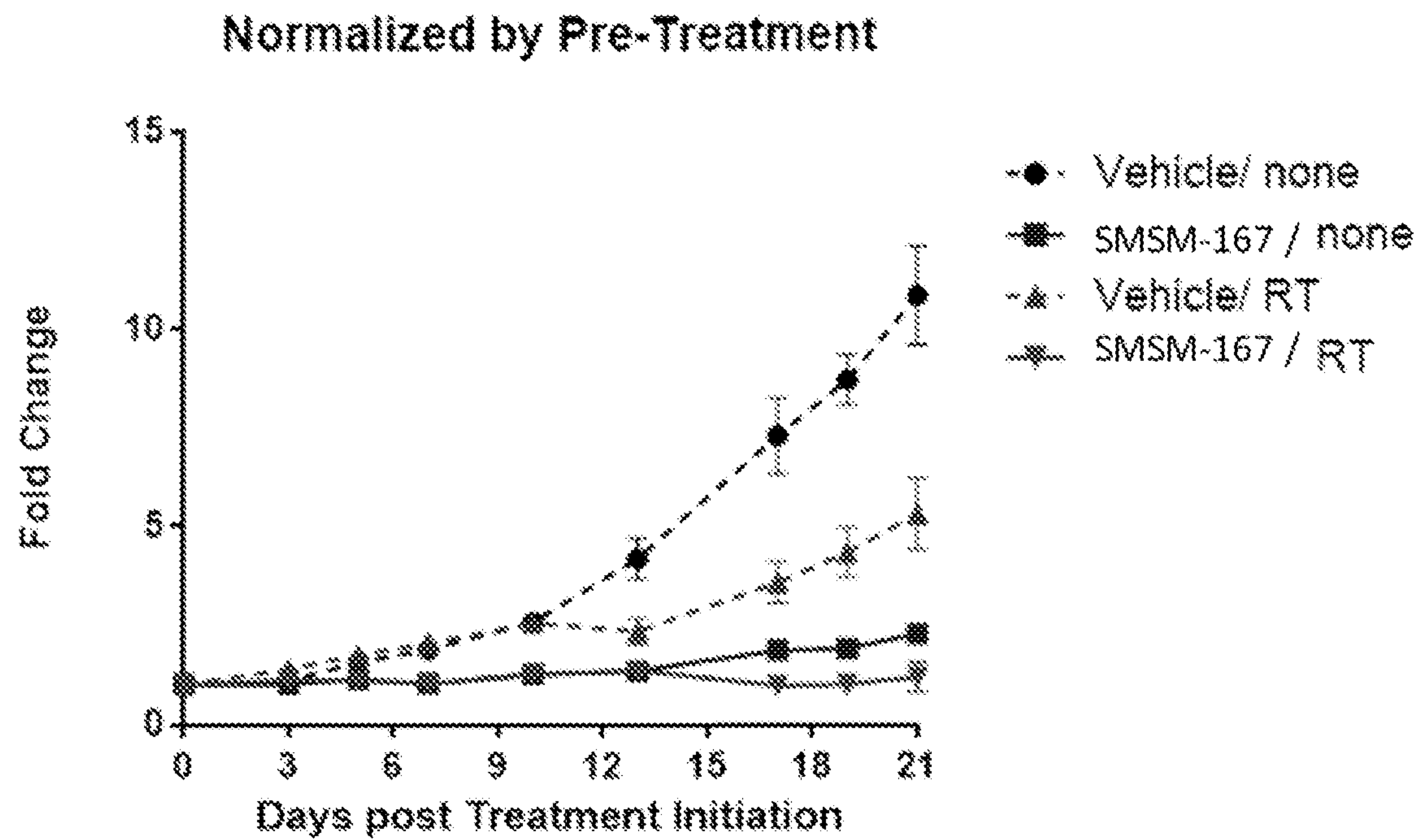


FIG. 24A

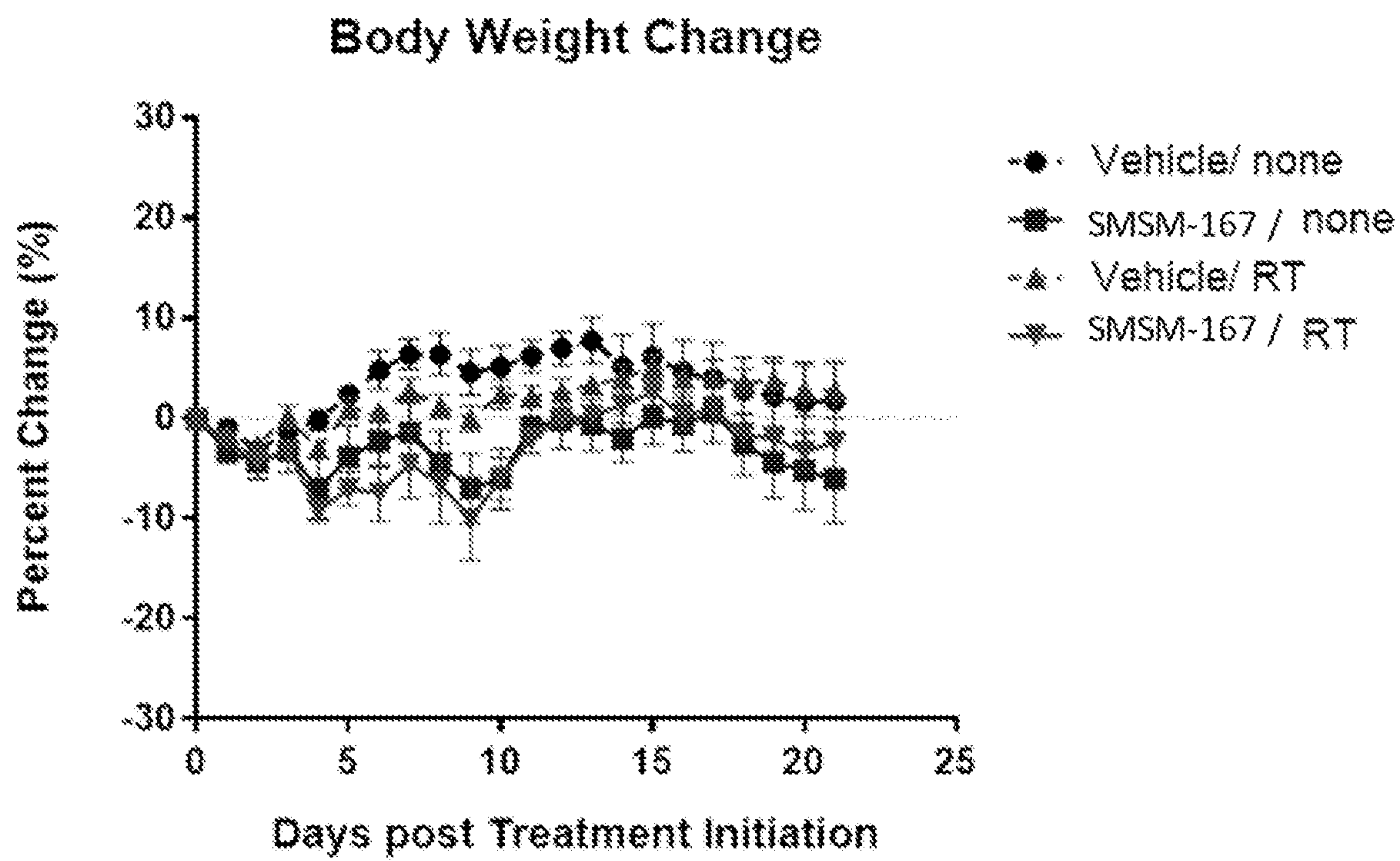


FIG. 24B