

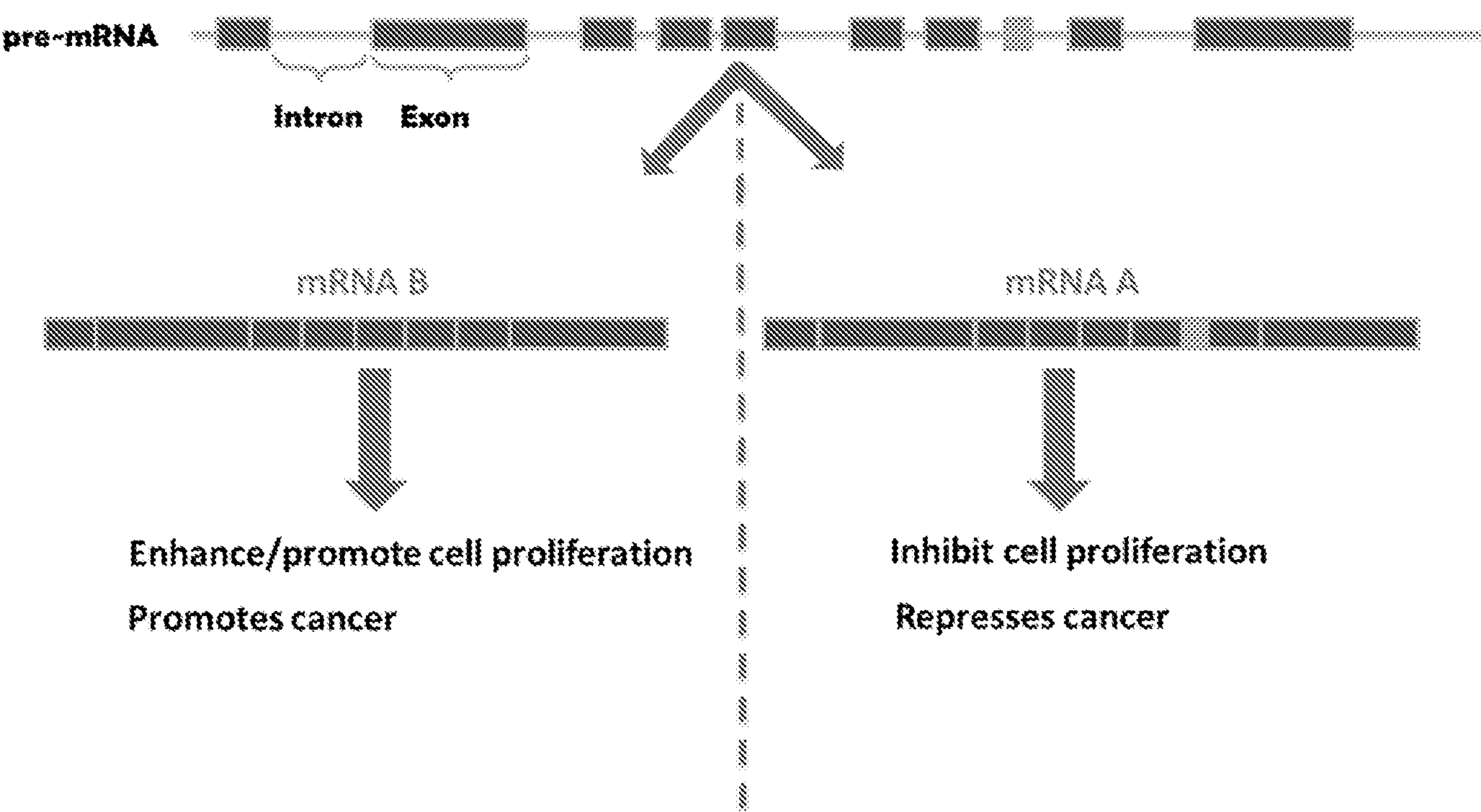


FIG. 1

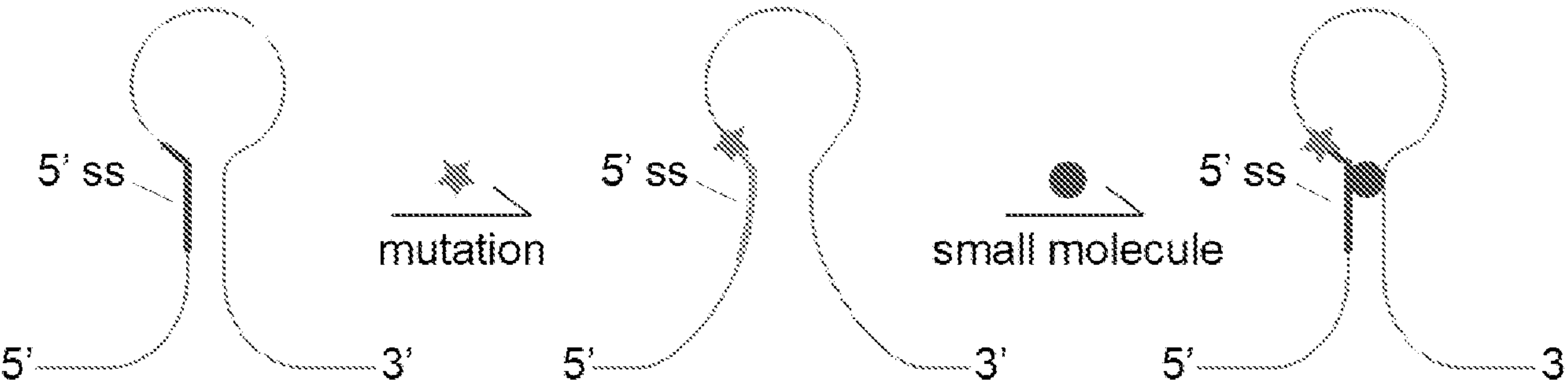
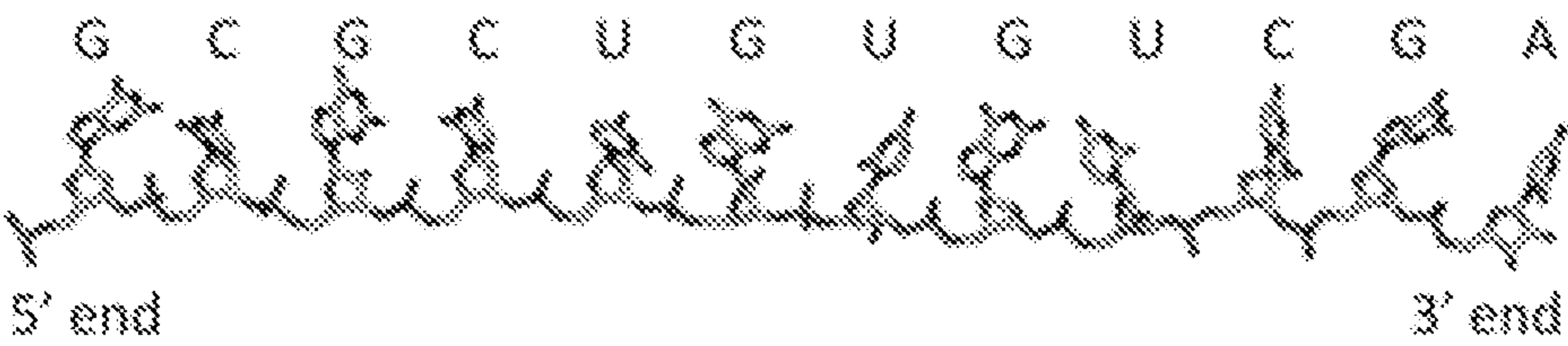


FIG. 2

Primary structure



Secondary structure



Tertiary structure

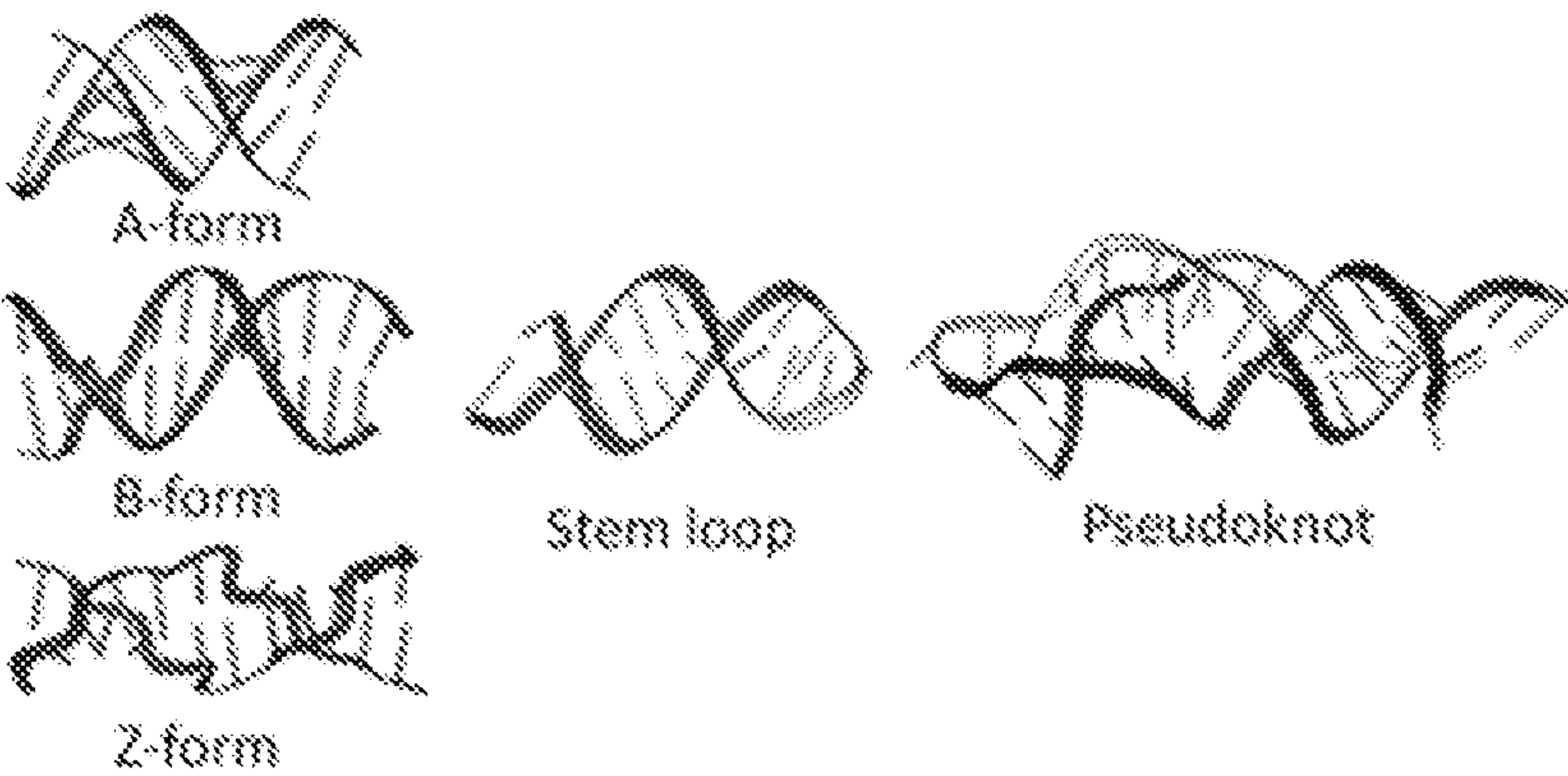


FIG. 3



FIG. 4

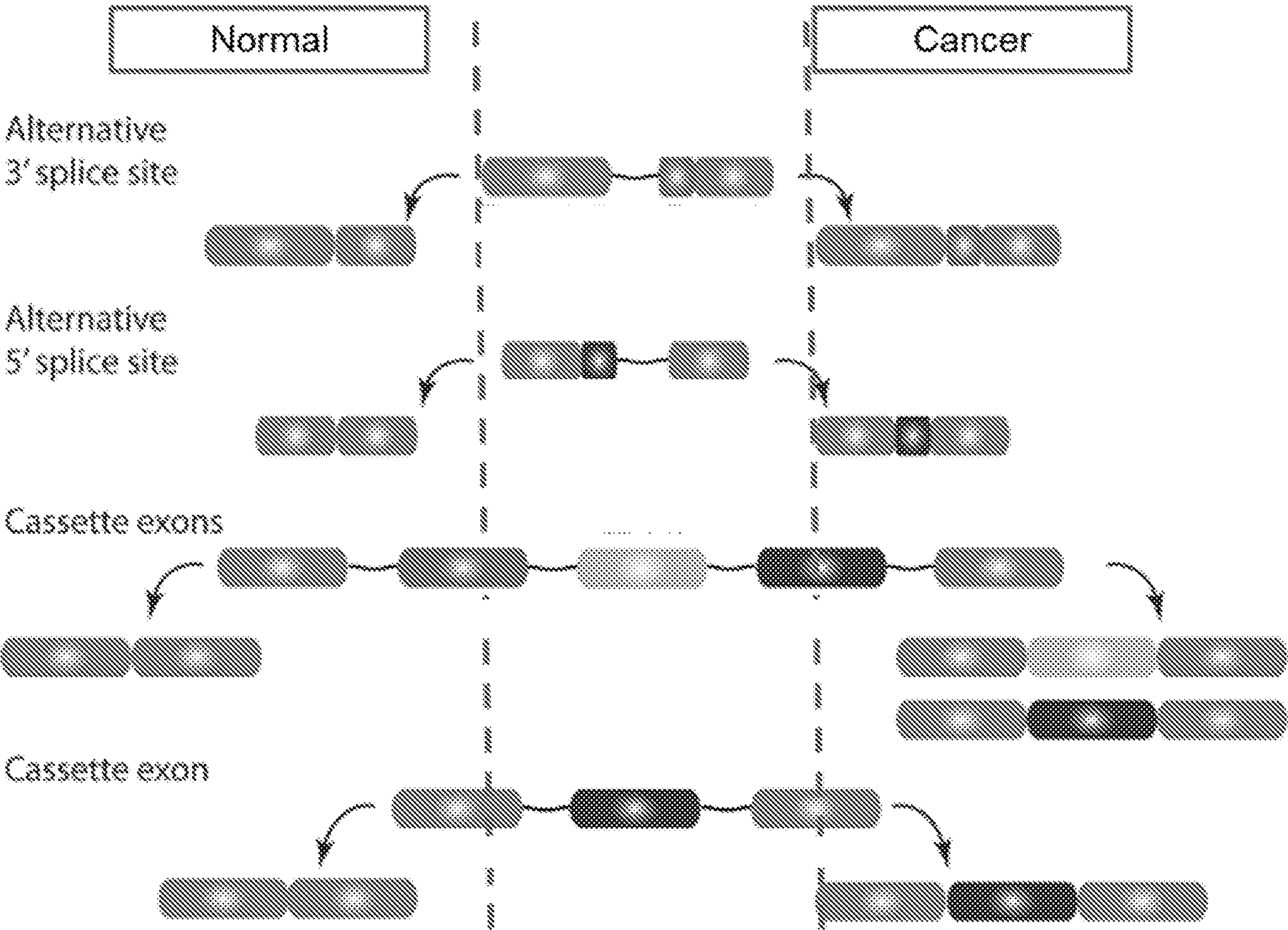


FIG. 5

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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

Ewing's sarcoma
(A673 cell line)

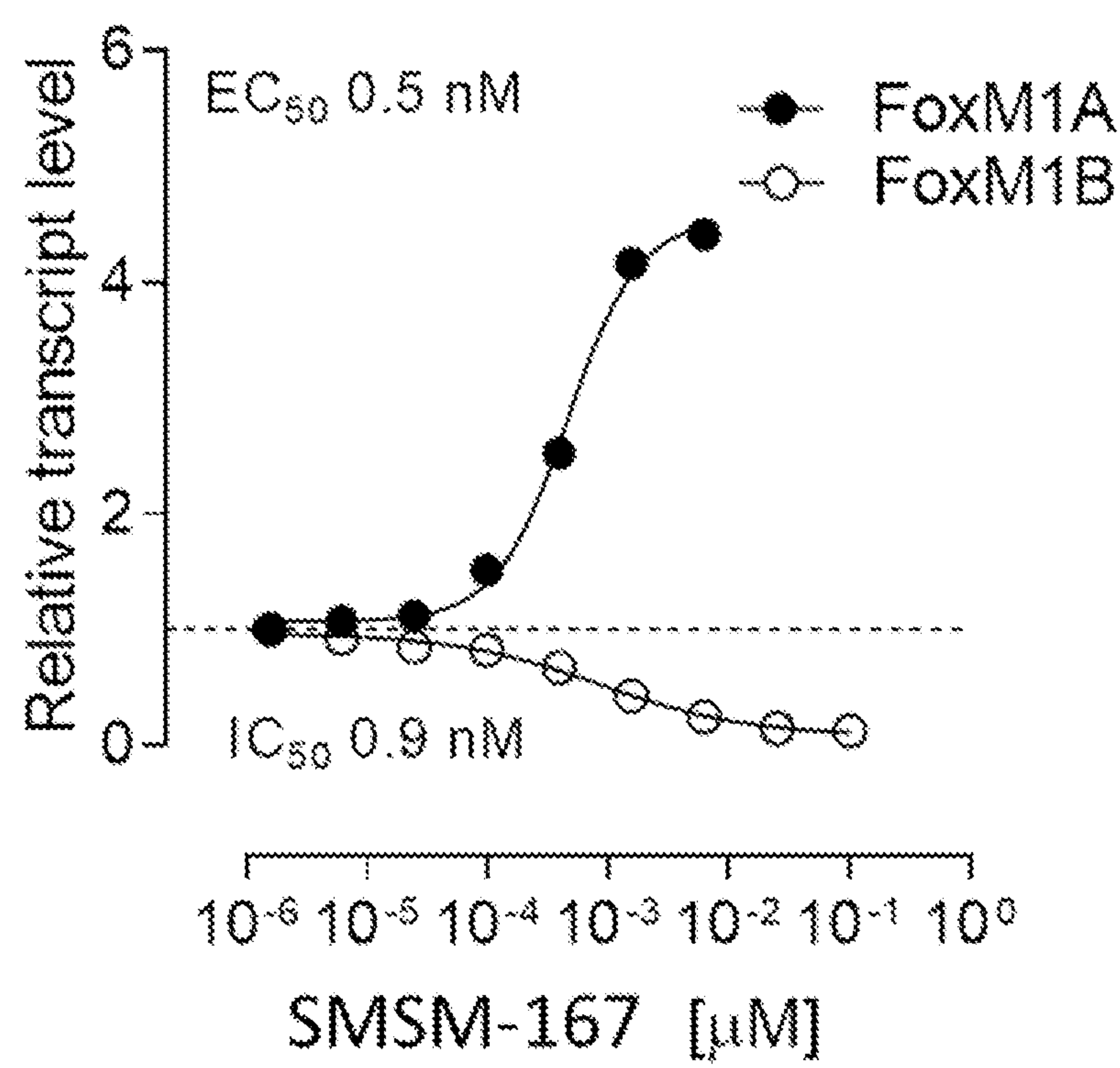


FIG. 7

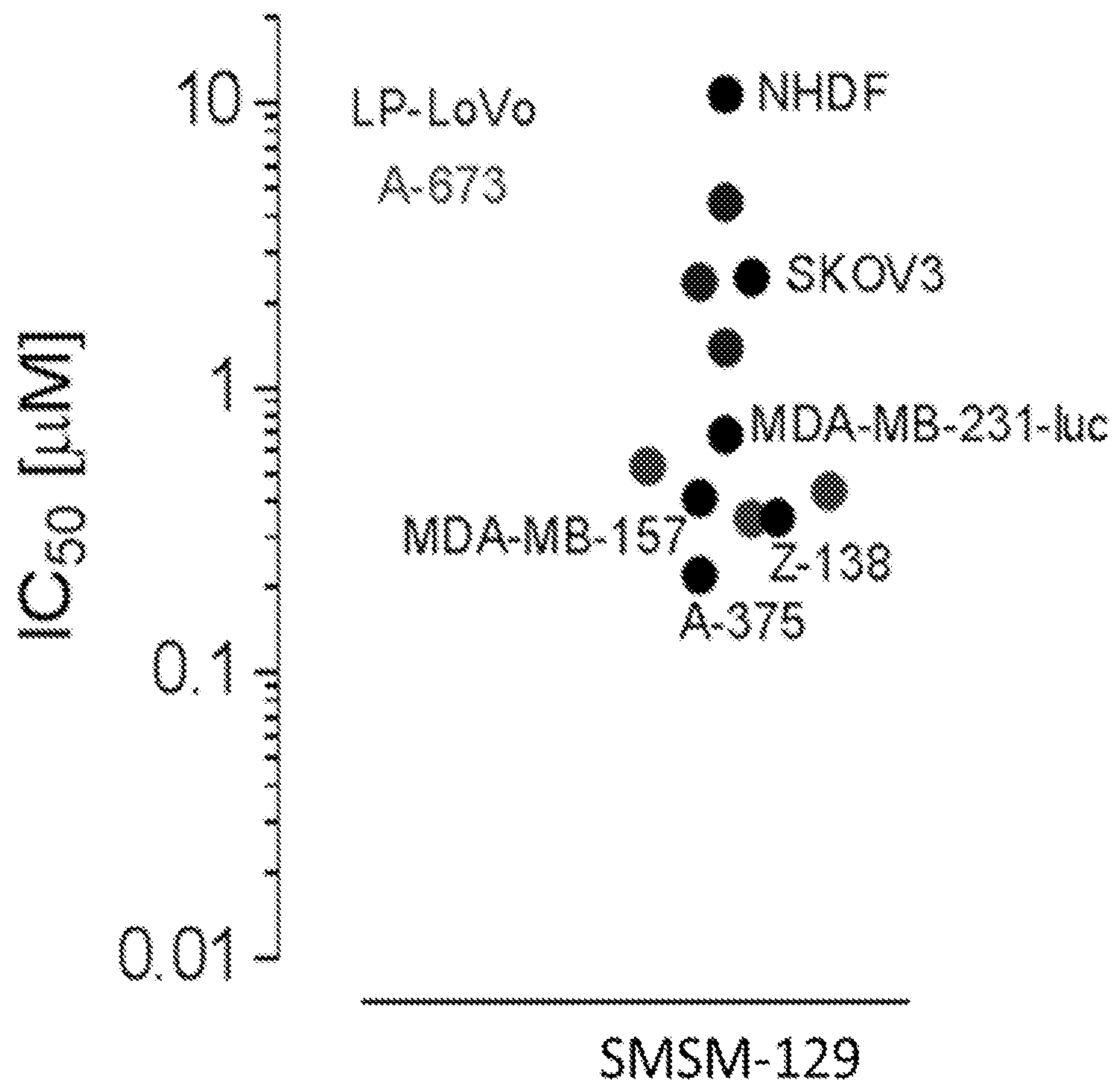


FIG. 8

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A-673

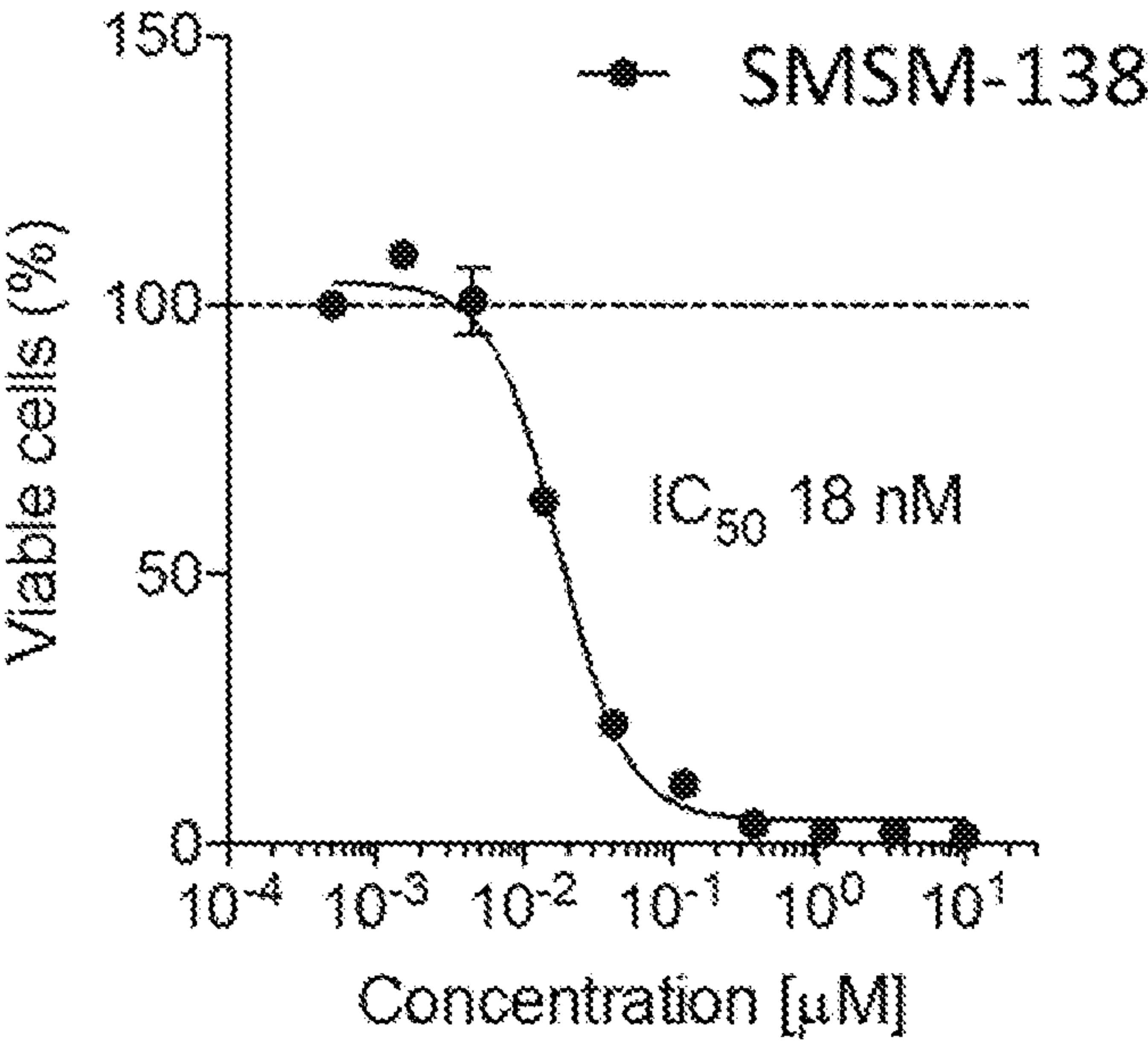


FIG. 9A

A-673

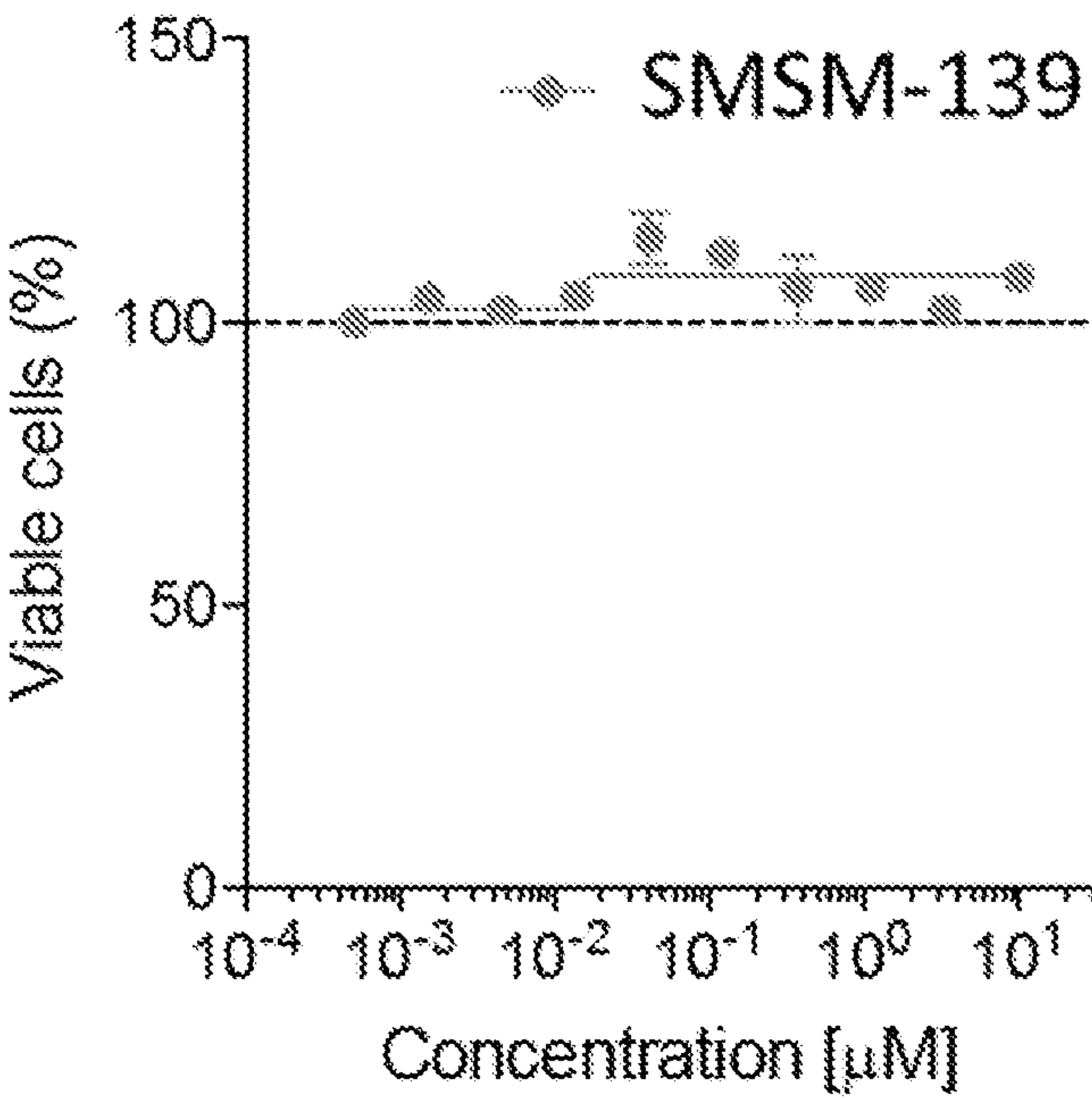


FIG. 9B

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A-673

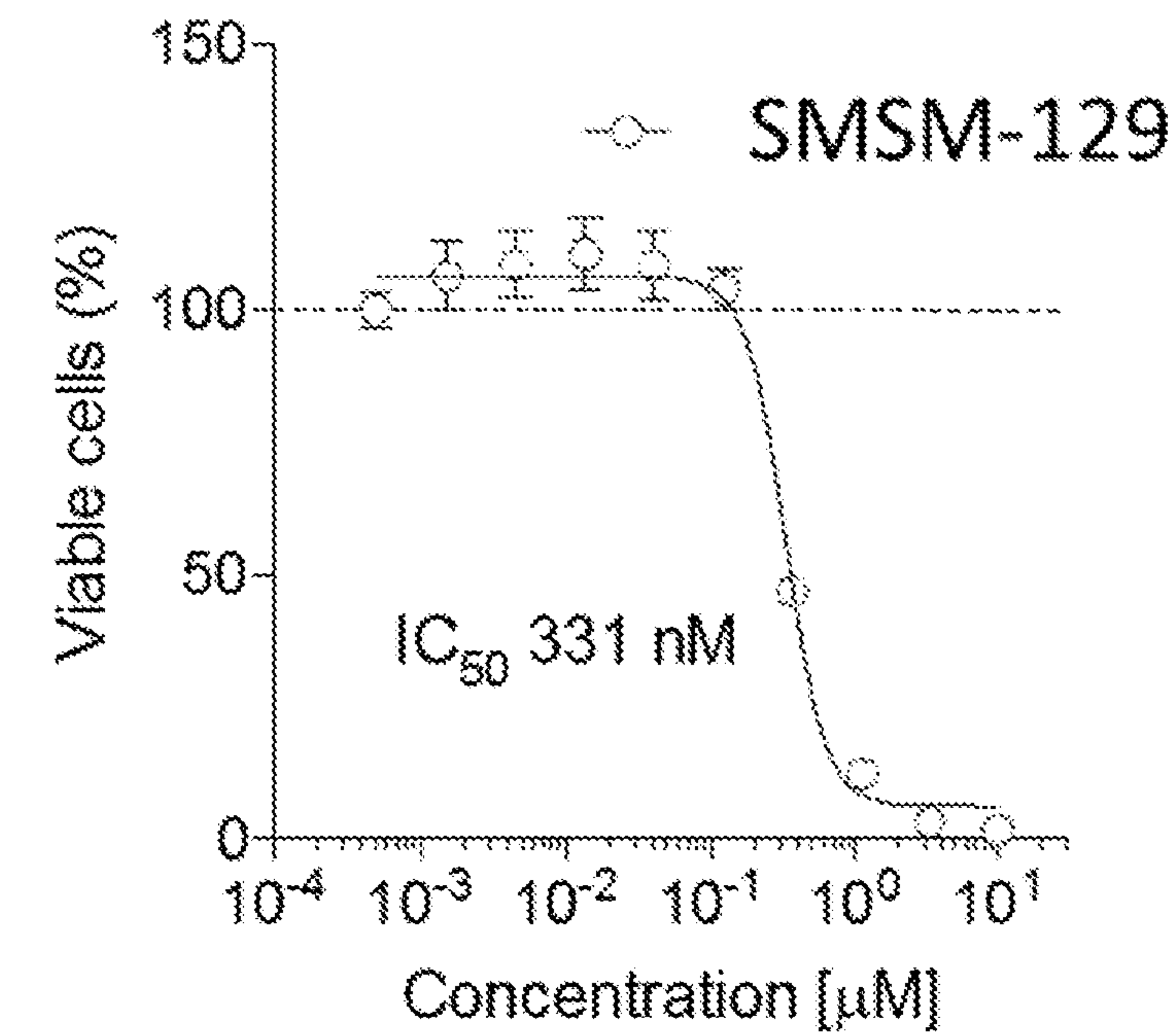


FIG. 9C

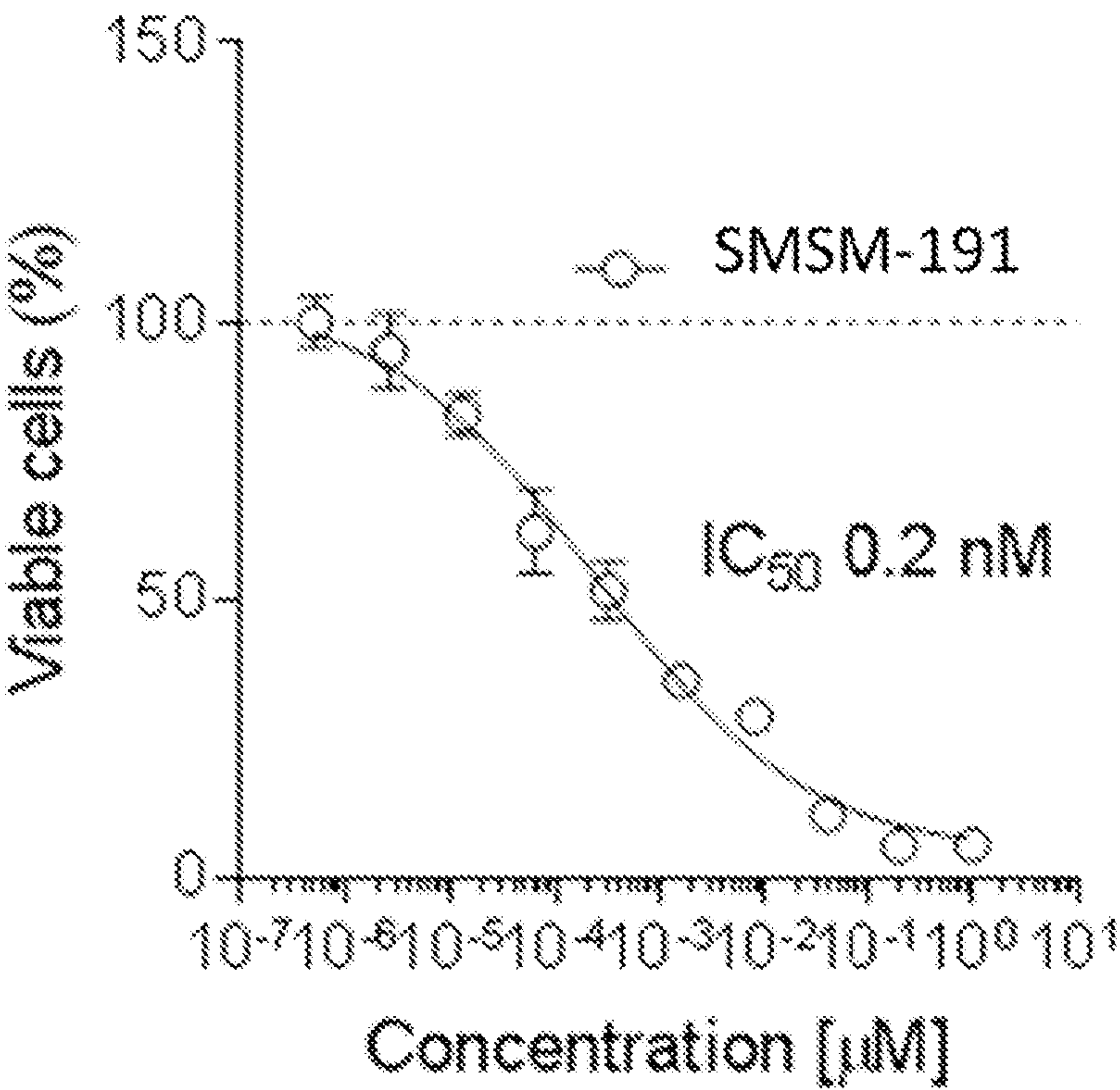


FIG. 9D

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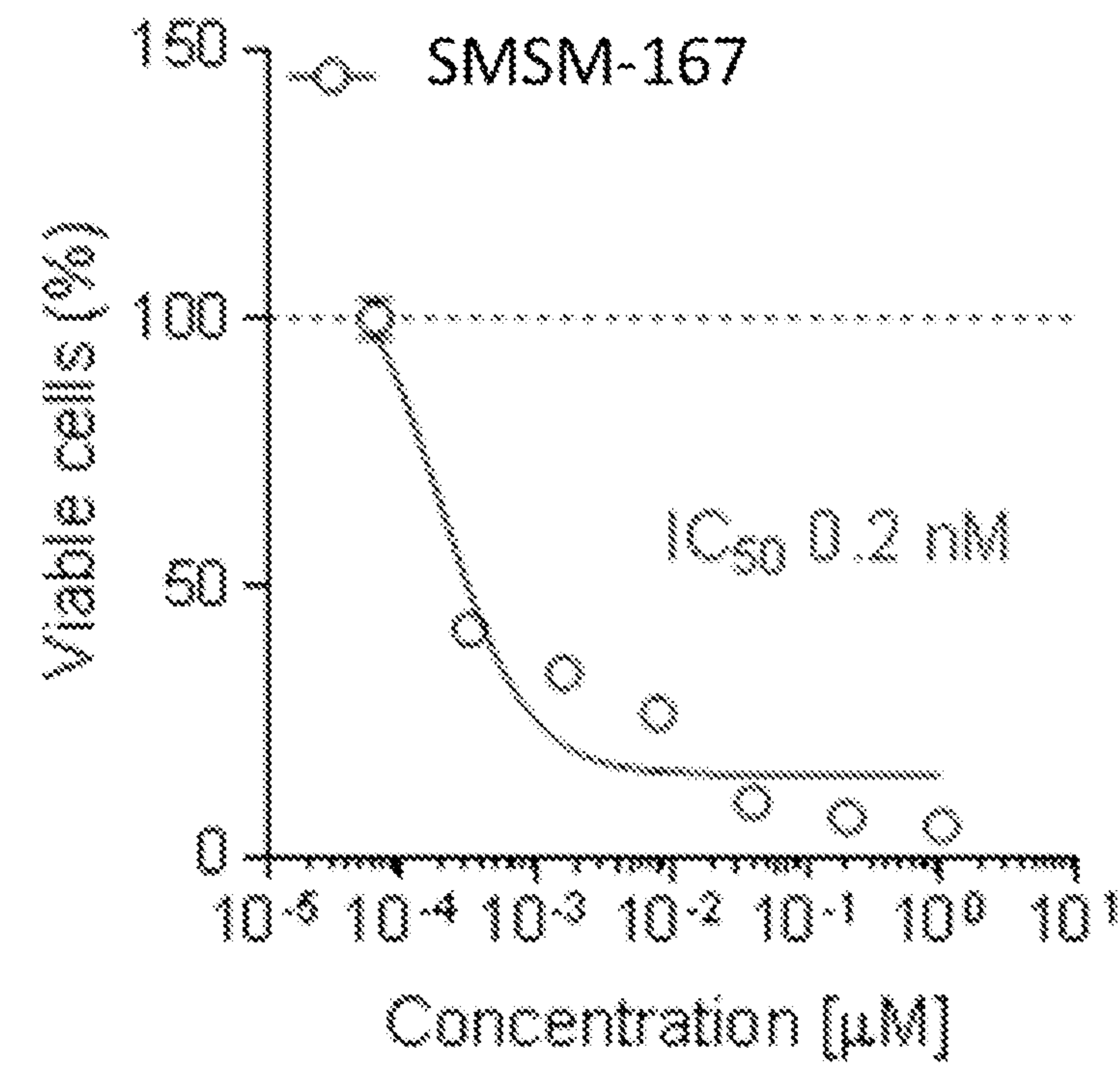


FIG. 9E

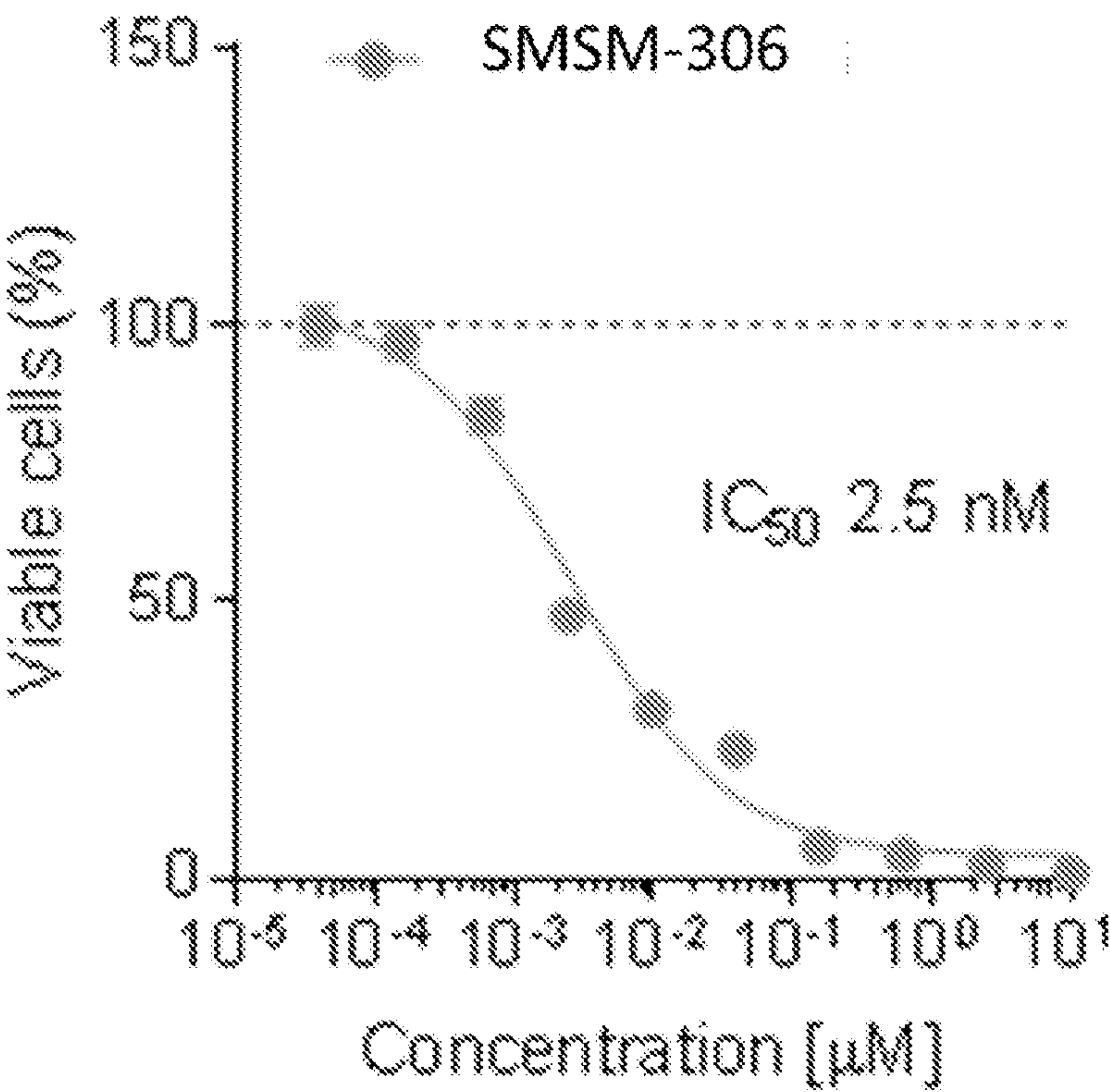


FIG. 9F

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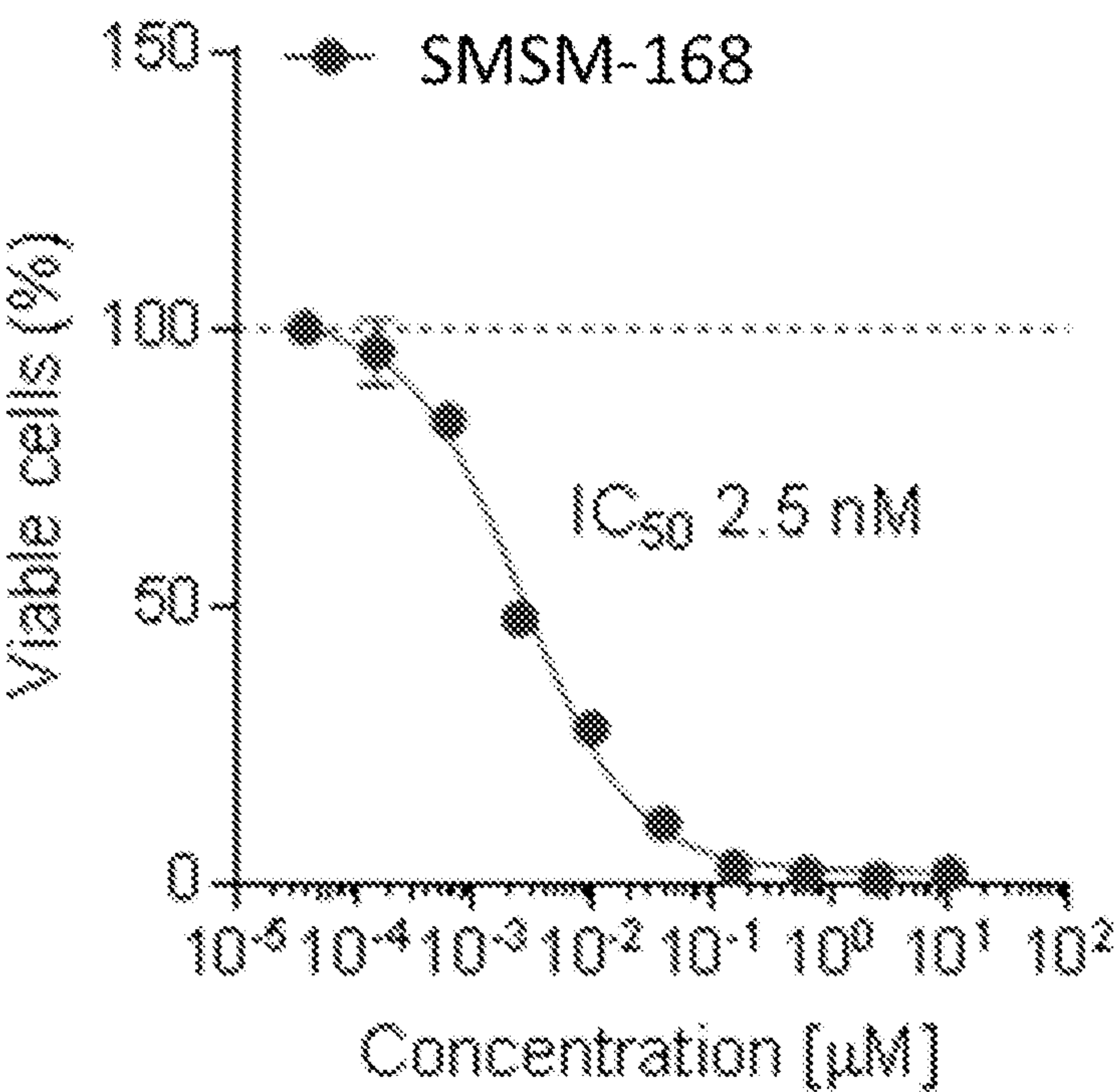


FIG. 9G

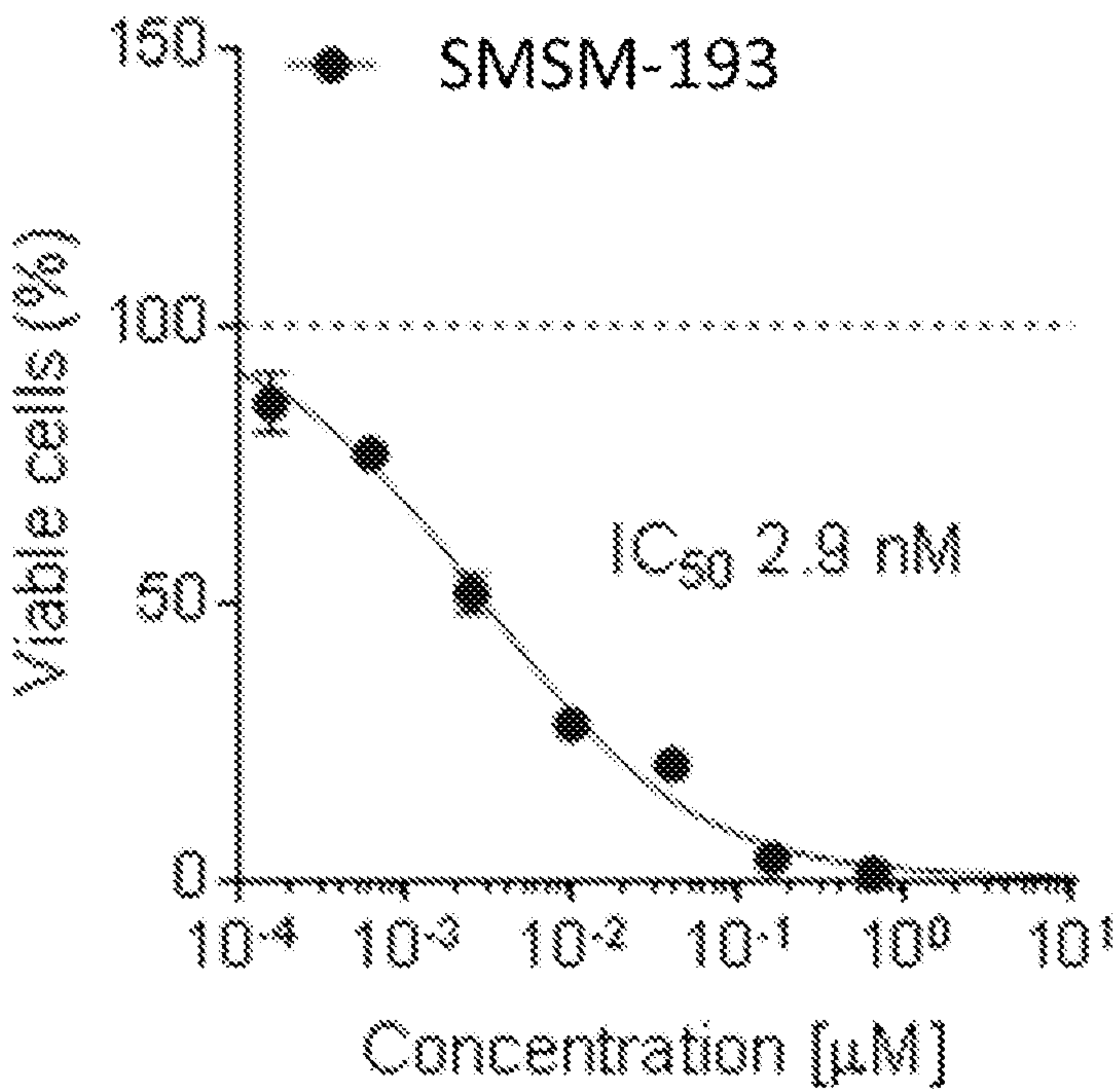


FIG. 9H

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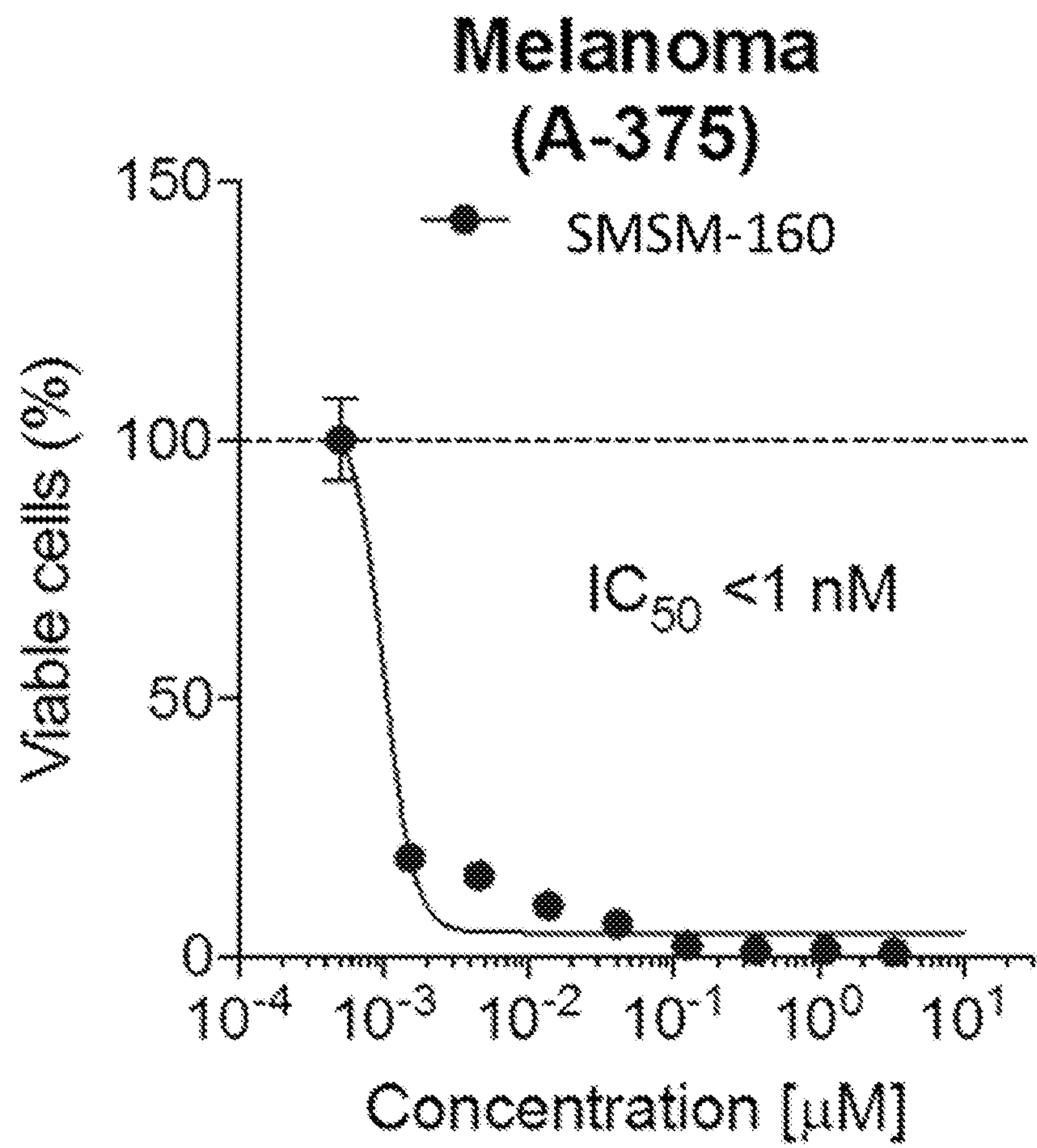


FIG. 10

NHDF

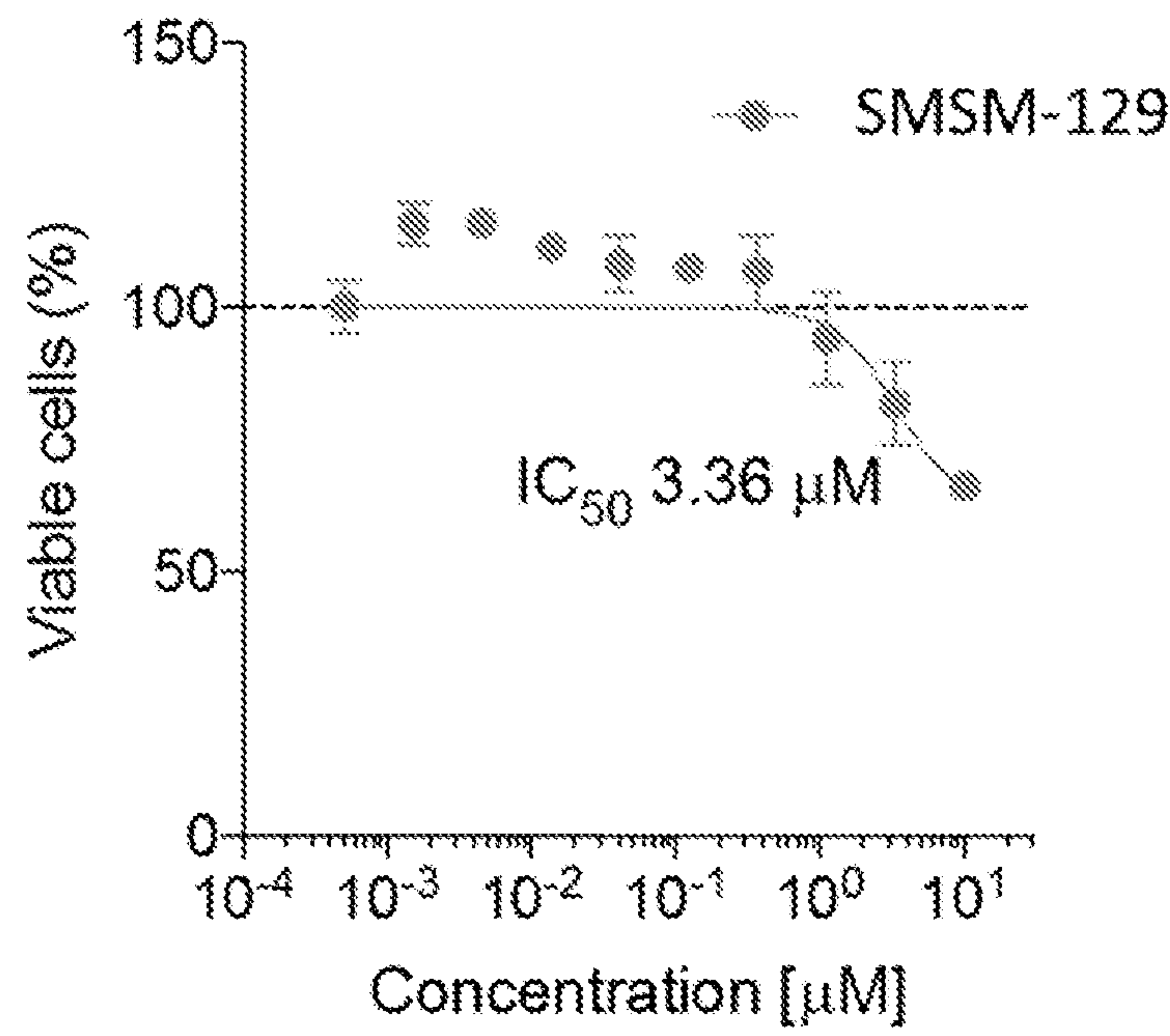


FIG. 11A

NHDF

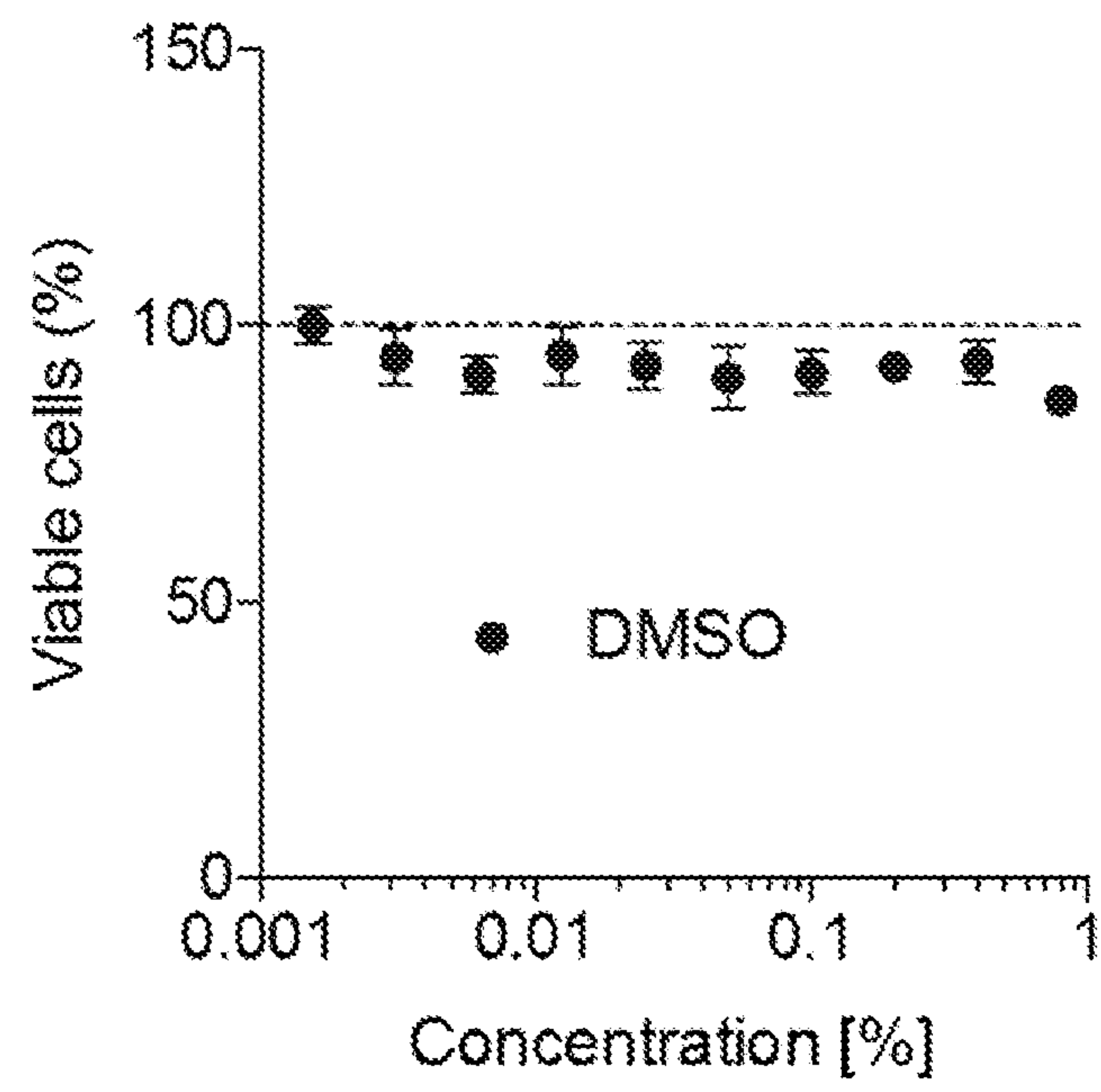


FIG. 11B

Proliferation vs. splicing

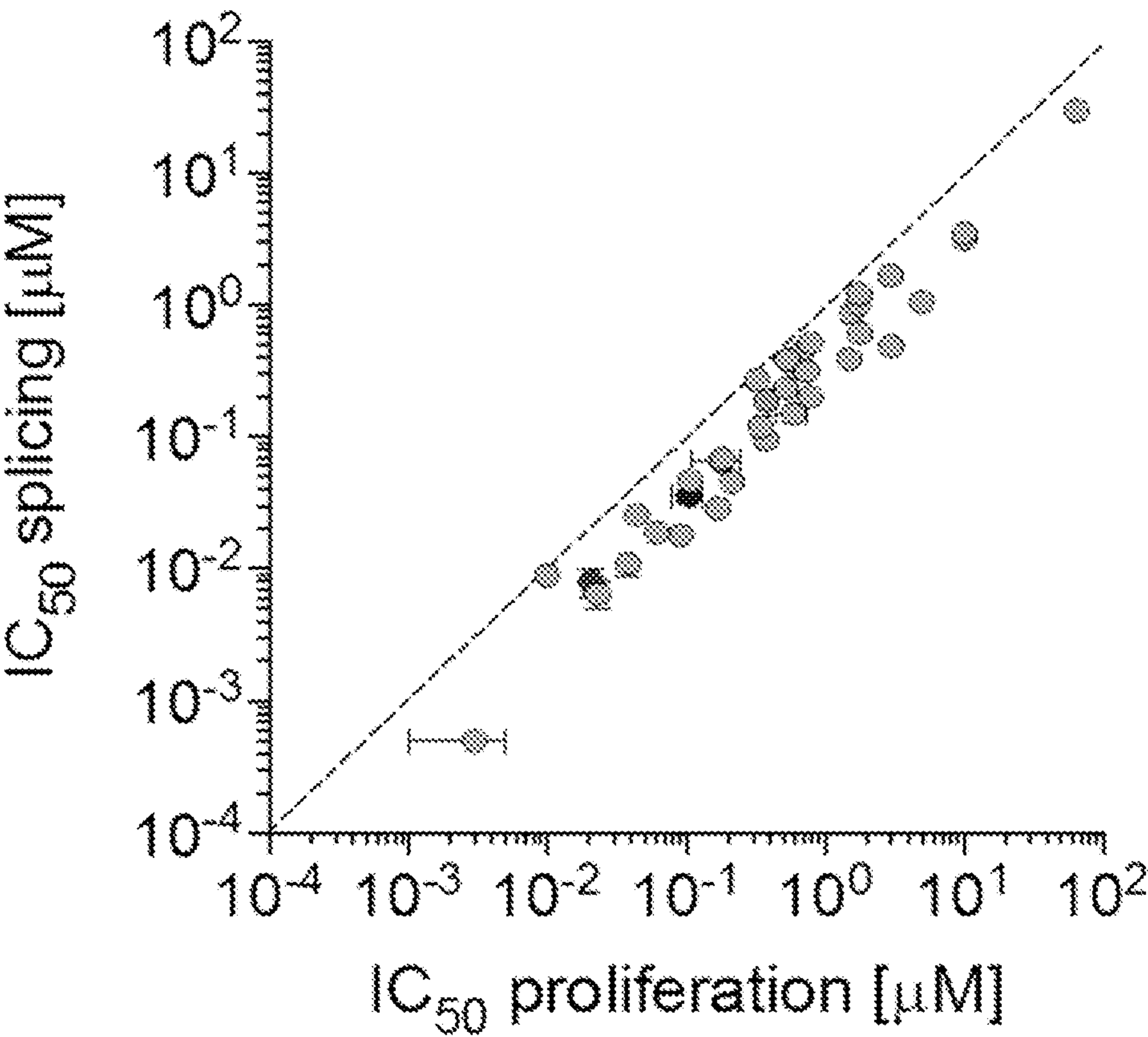
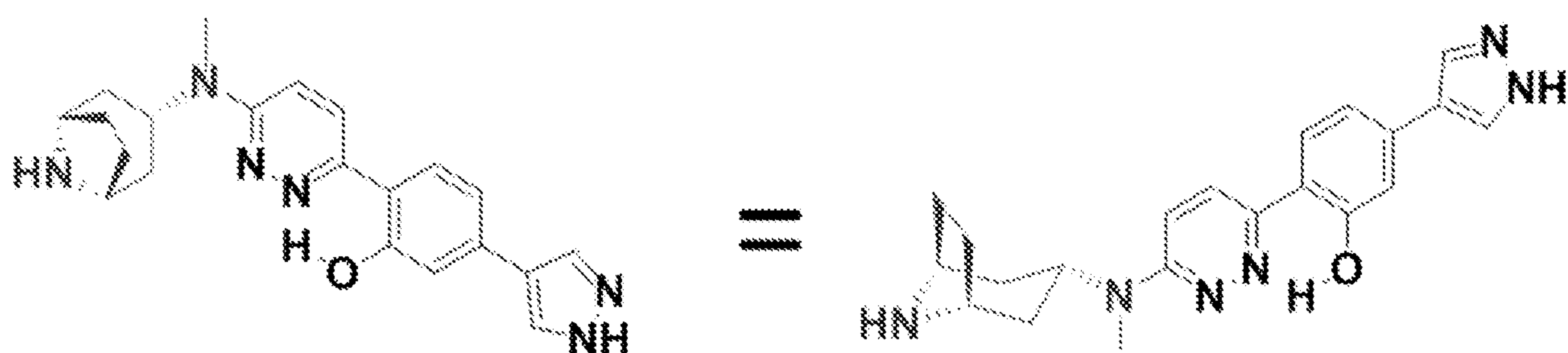


FIG. 12

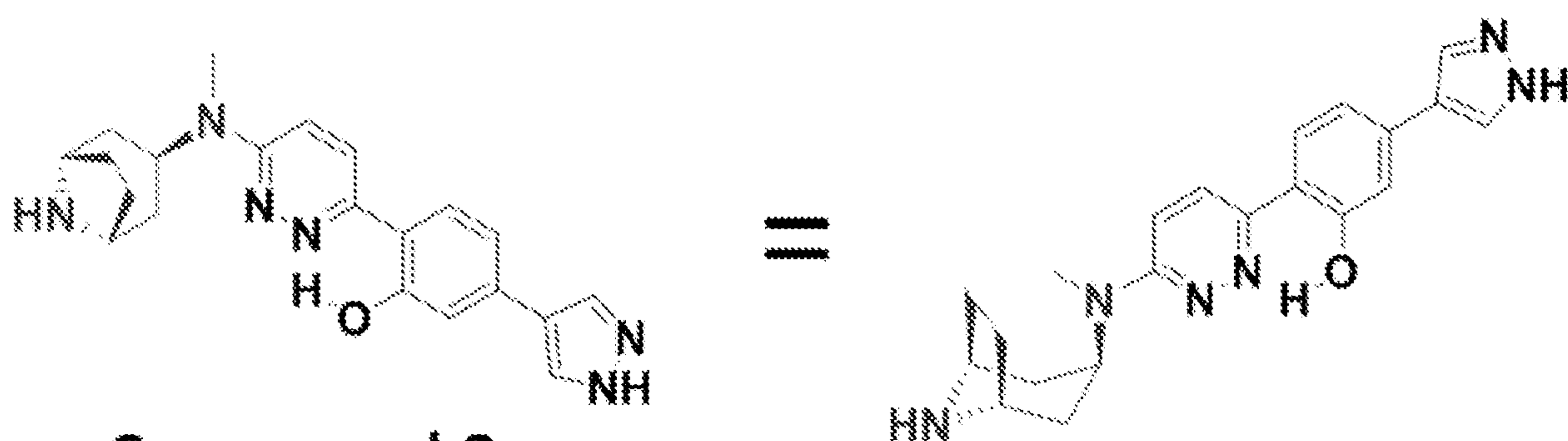
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Impact of Stereochemistry on Cell Activity



Compound 1

Cell_{A-673} = 18 nM



Compound 2

Cell_{A-673} > 100 mM

FIG. 13

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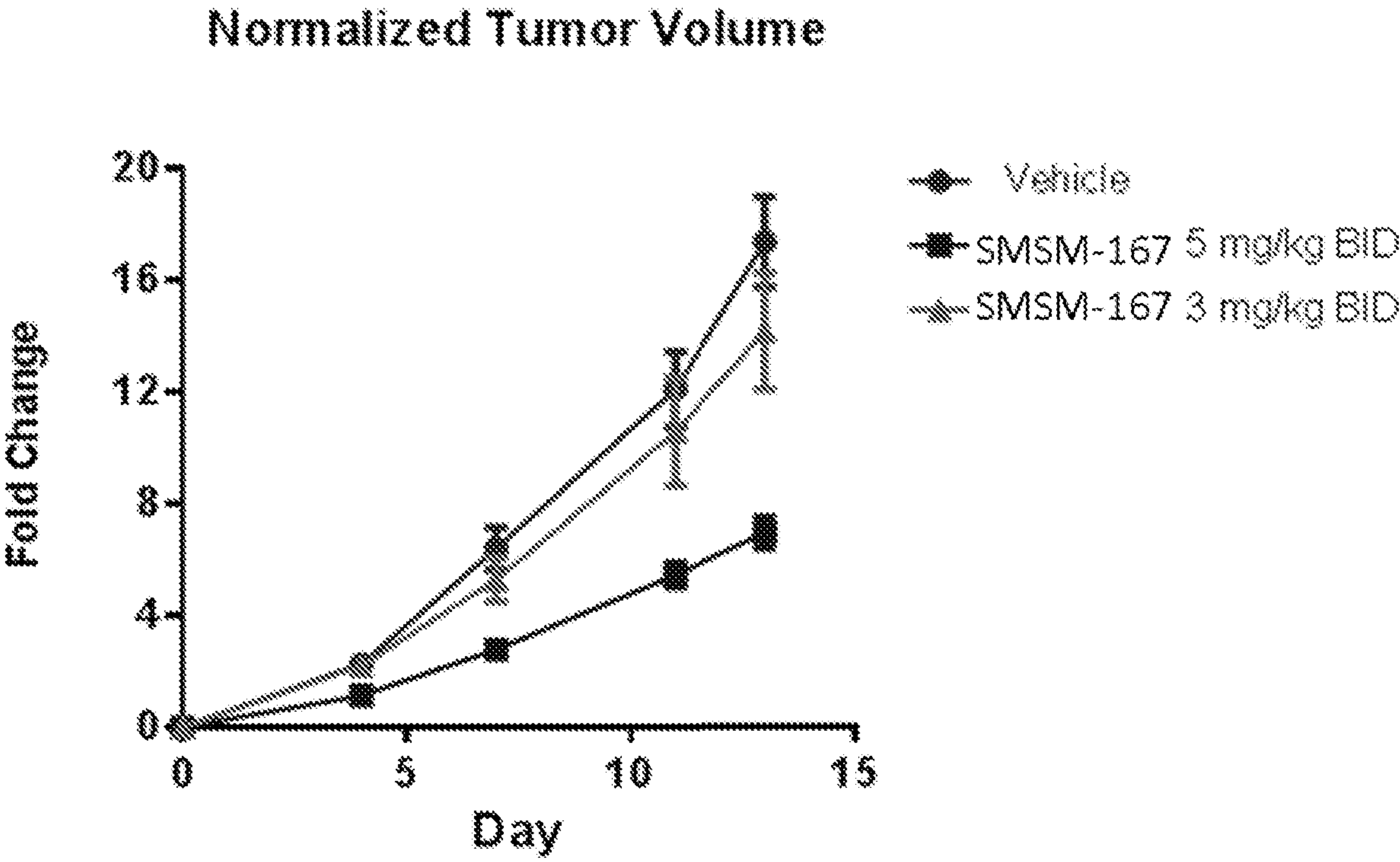


FIG. 14A

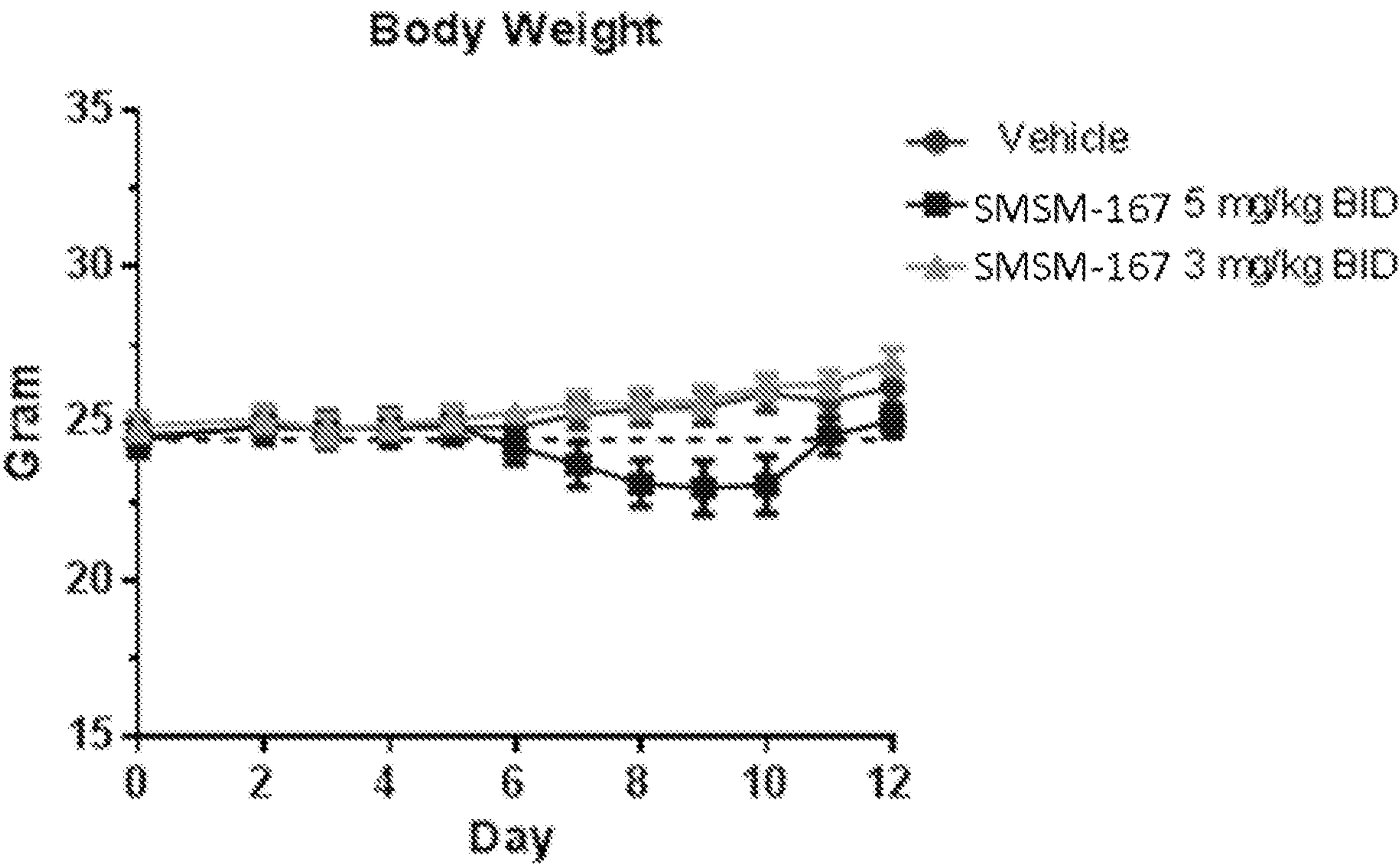


FIG. 14B

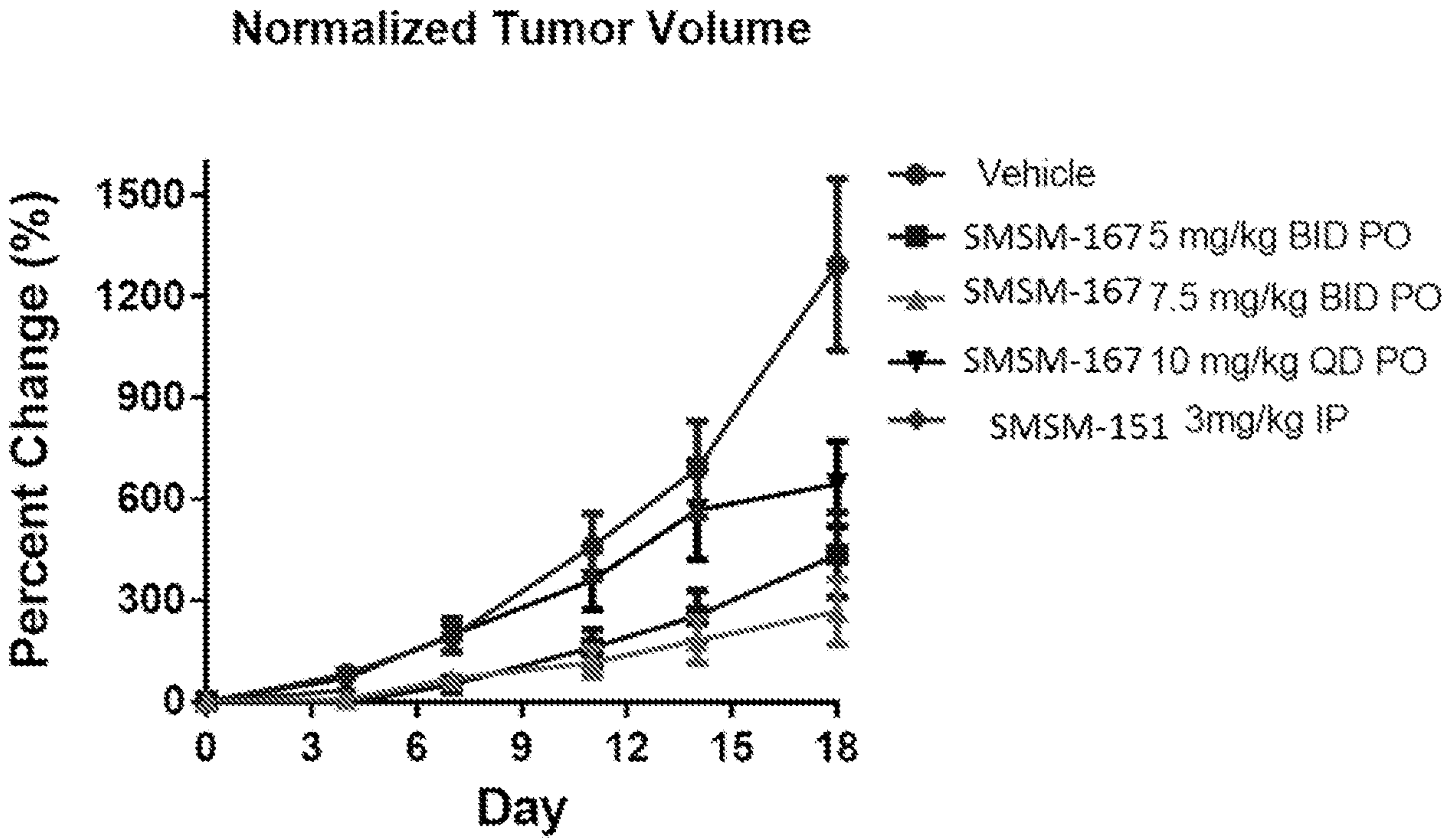


FIG. 14C

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Tumor Volume

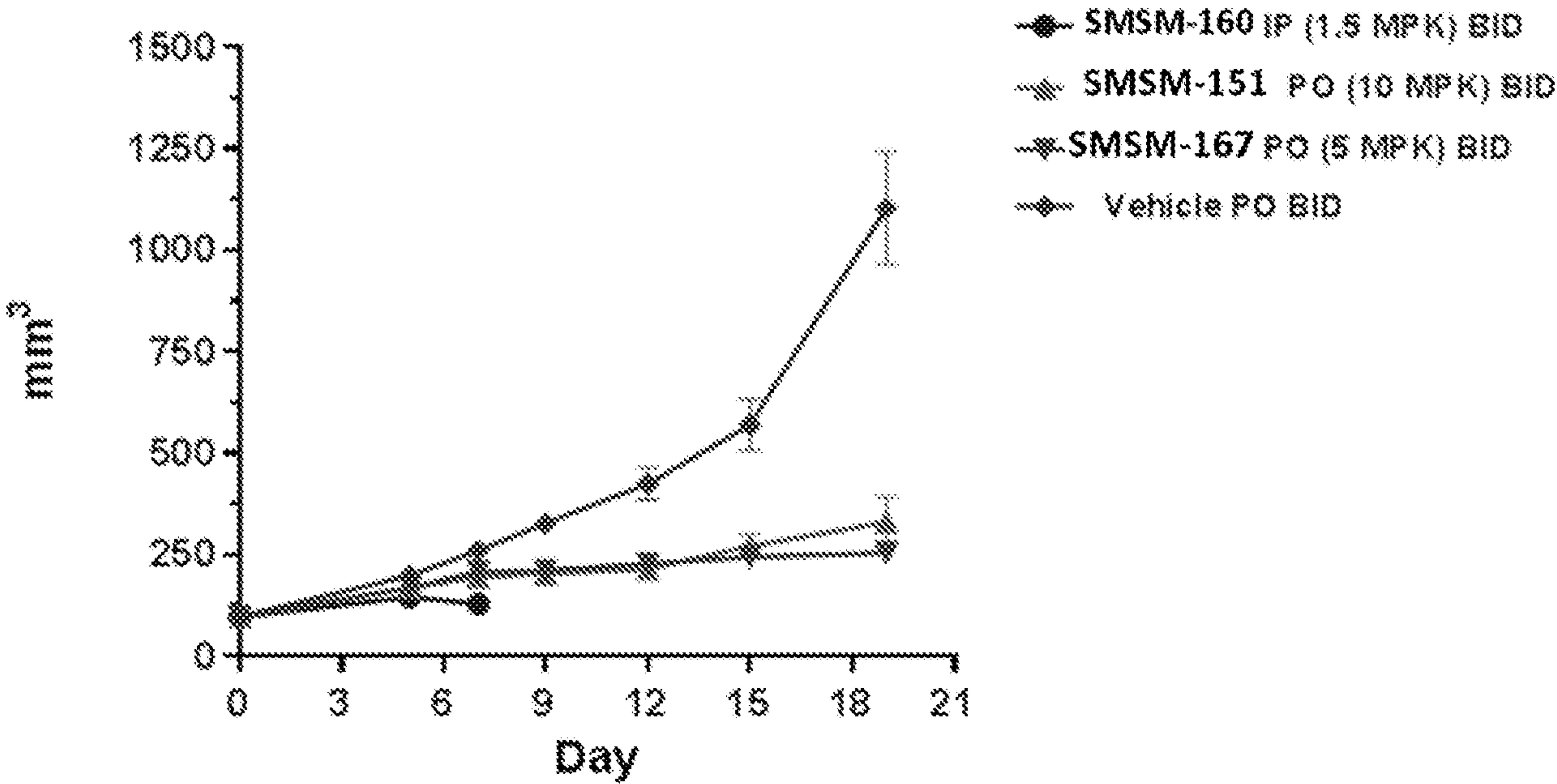


FIG. 15A

Body Weight

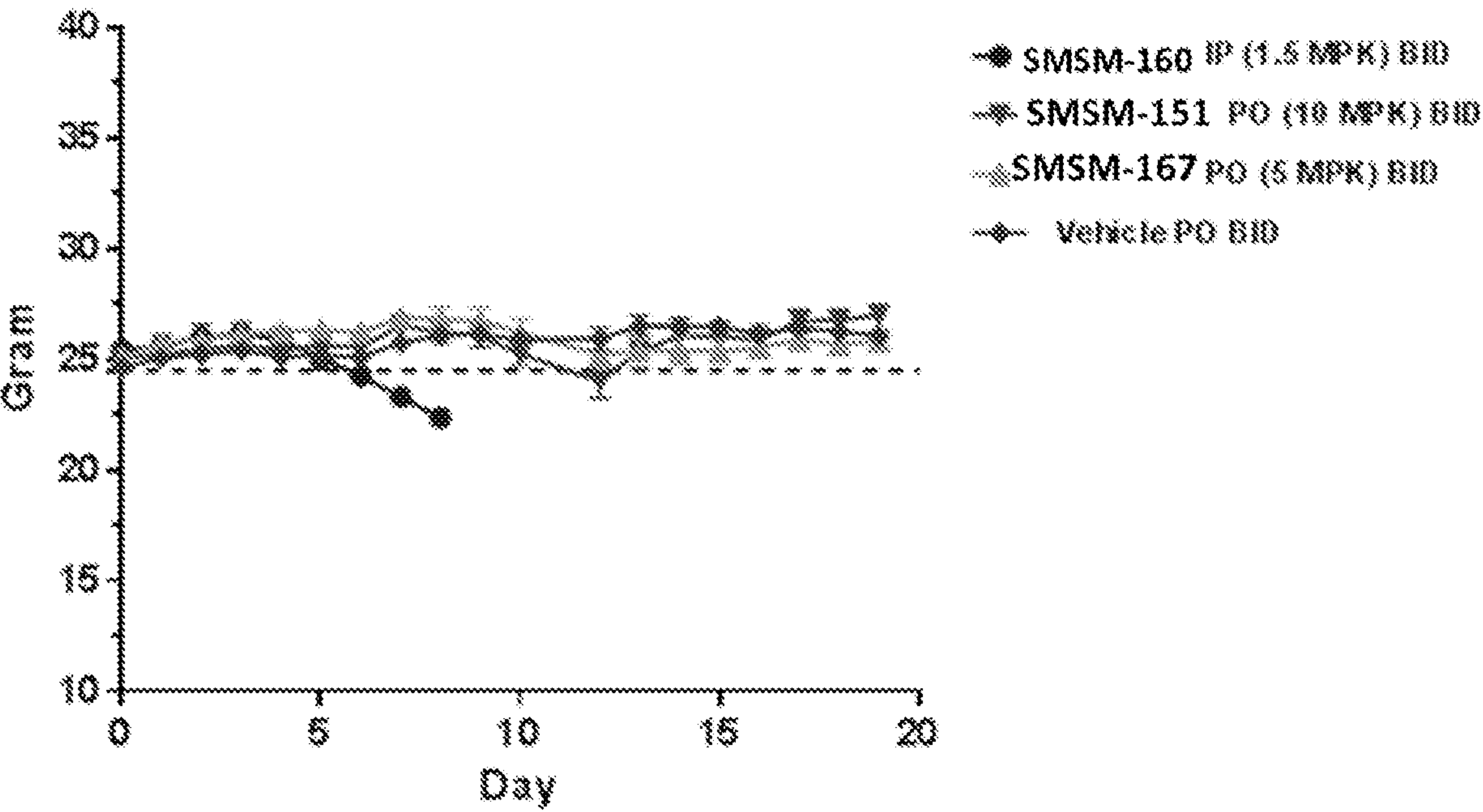


FIG. 15B

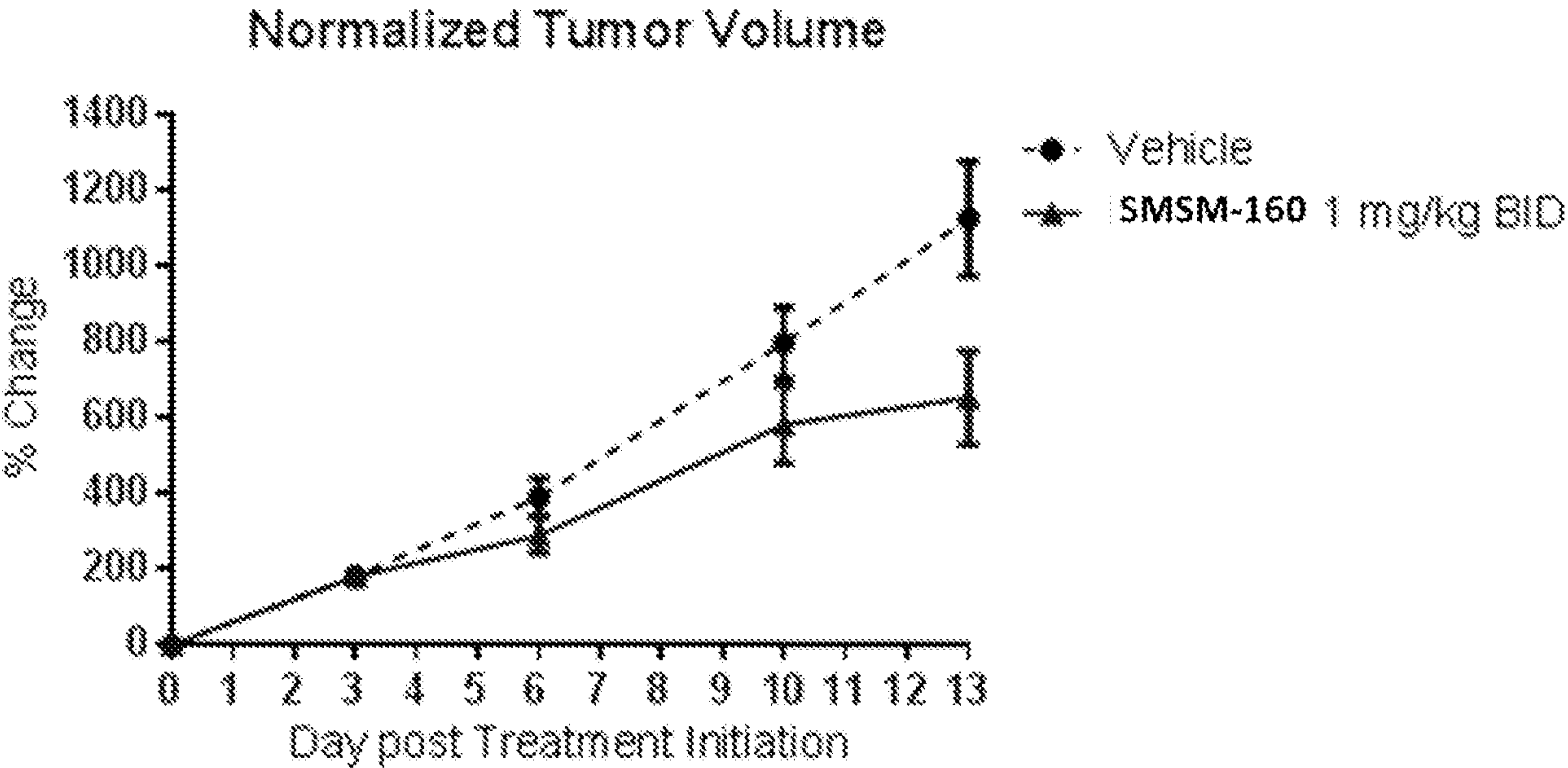


FIG. 16A

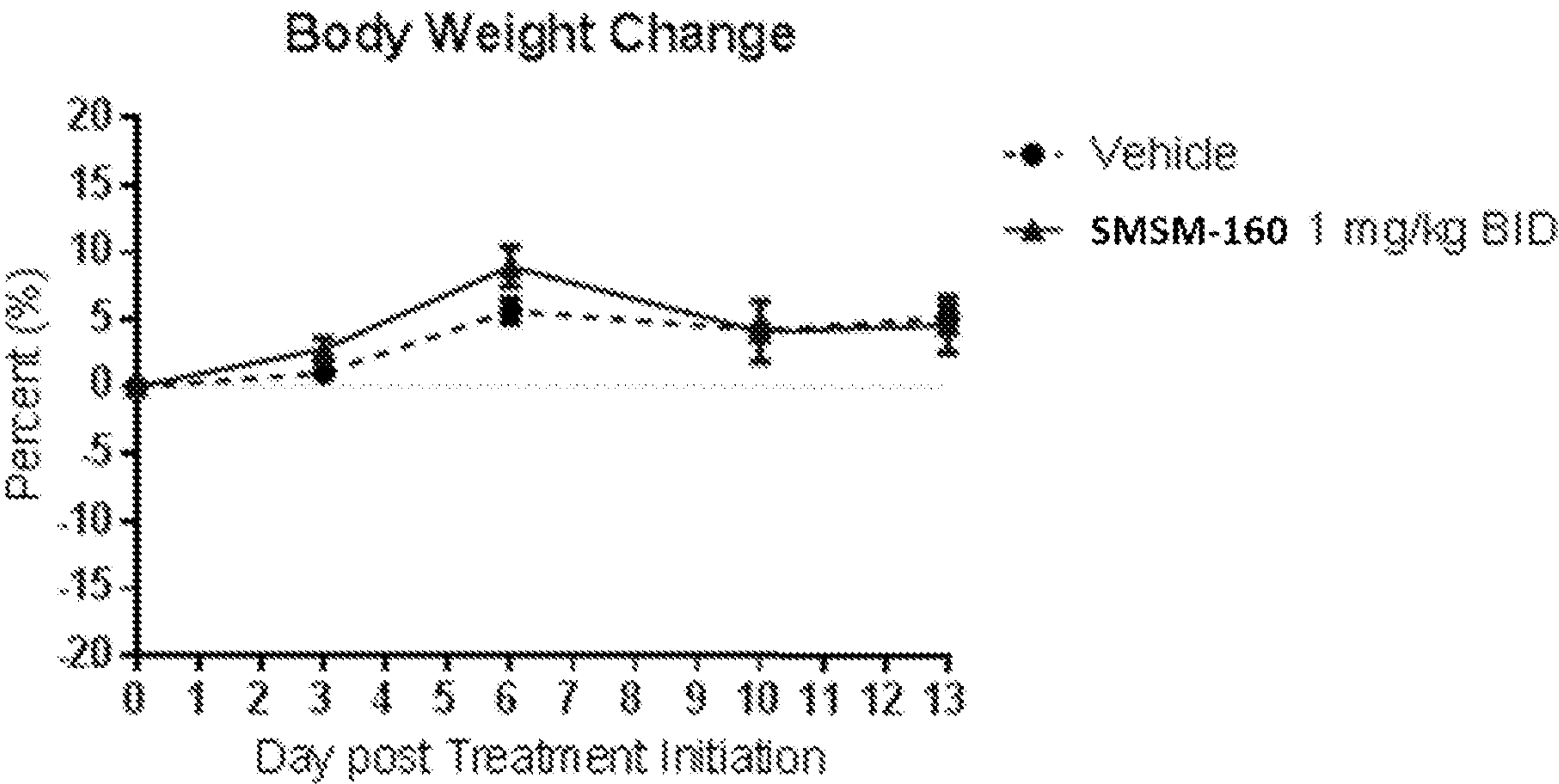


FIG. 16B

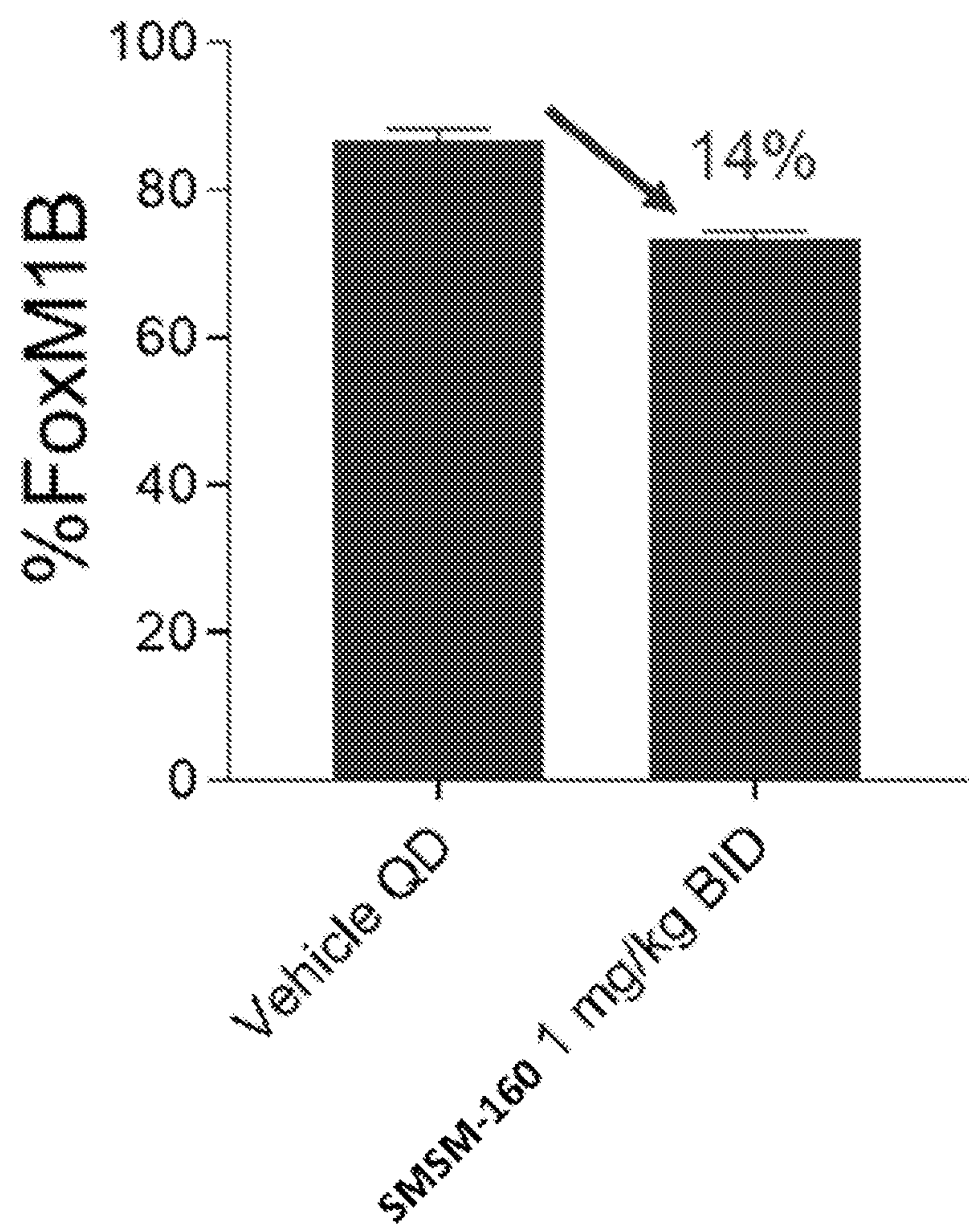


FIG. 16C

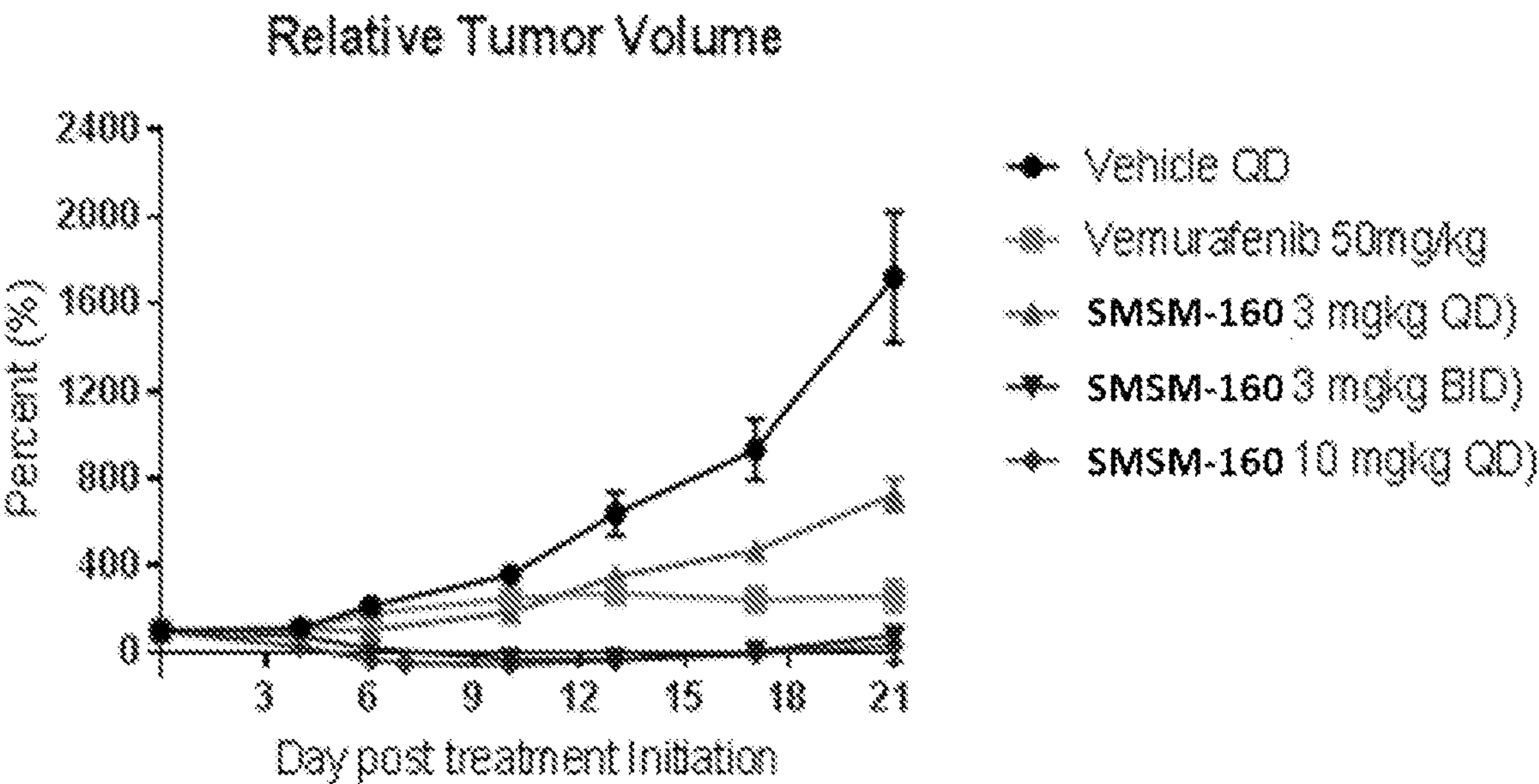


FIG. 17A

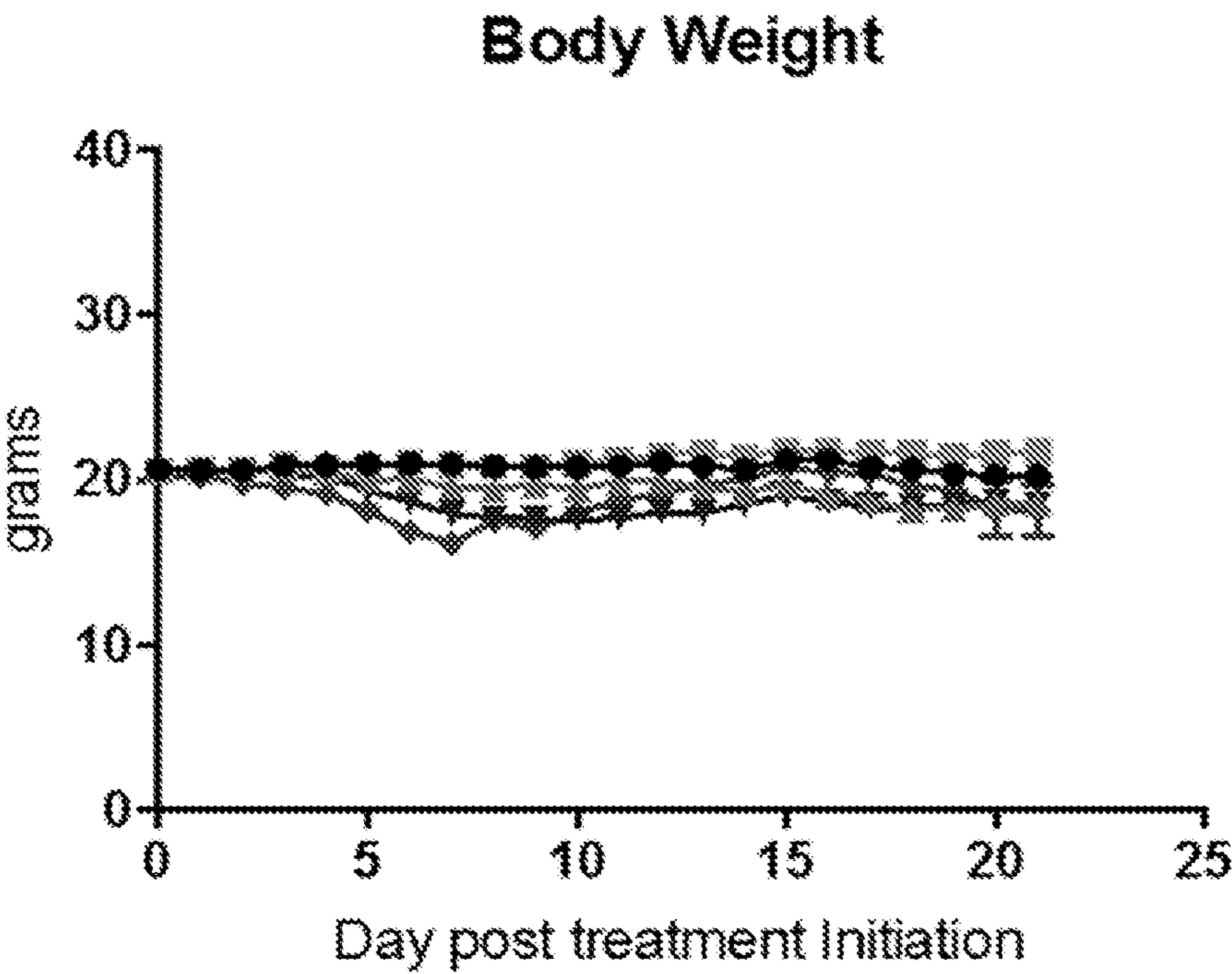


FIG. 17B

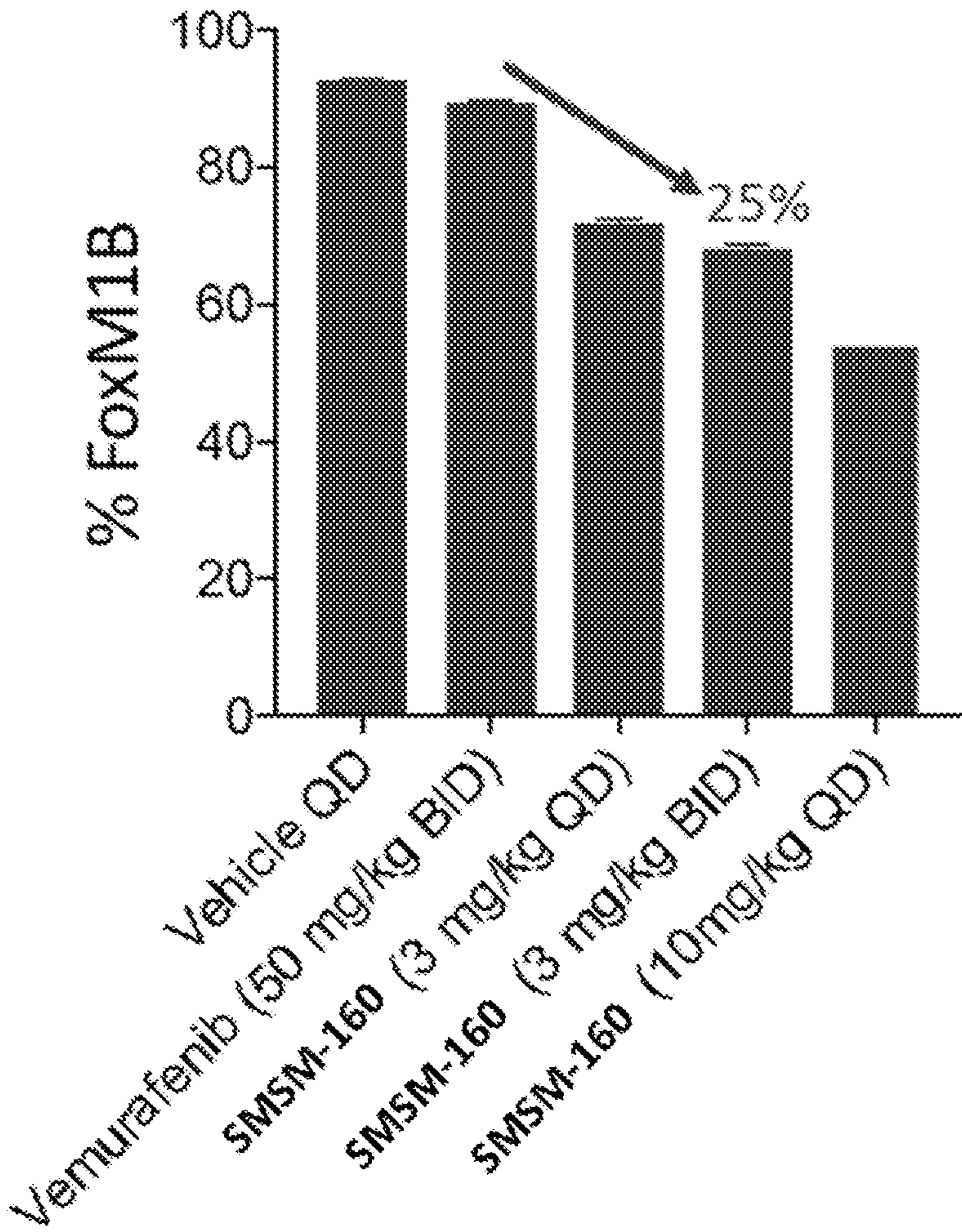


FIG. 17C

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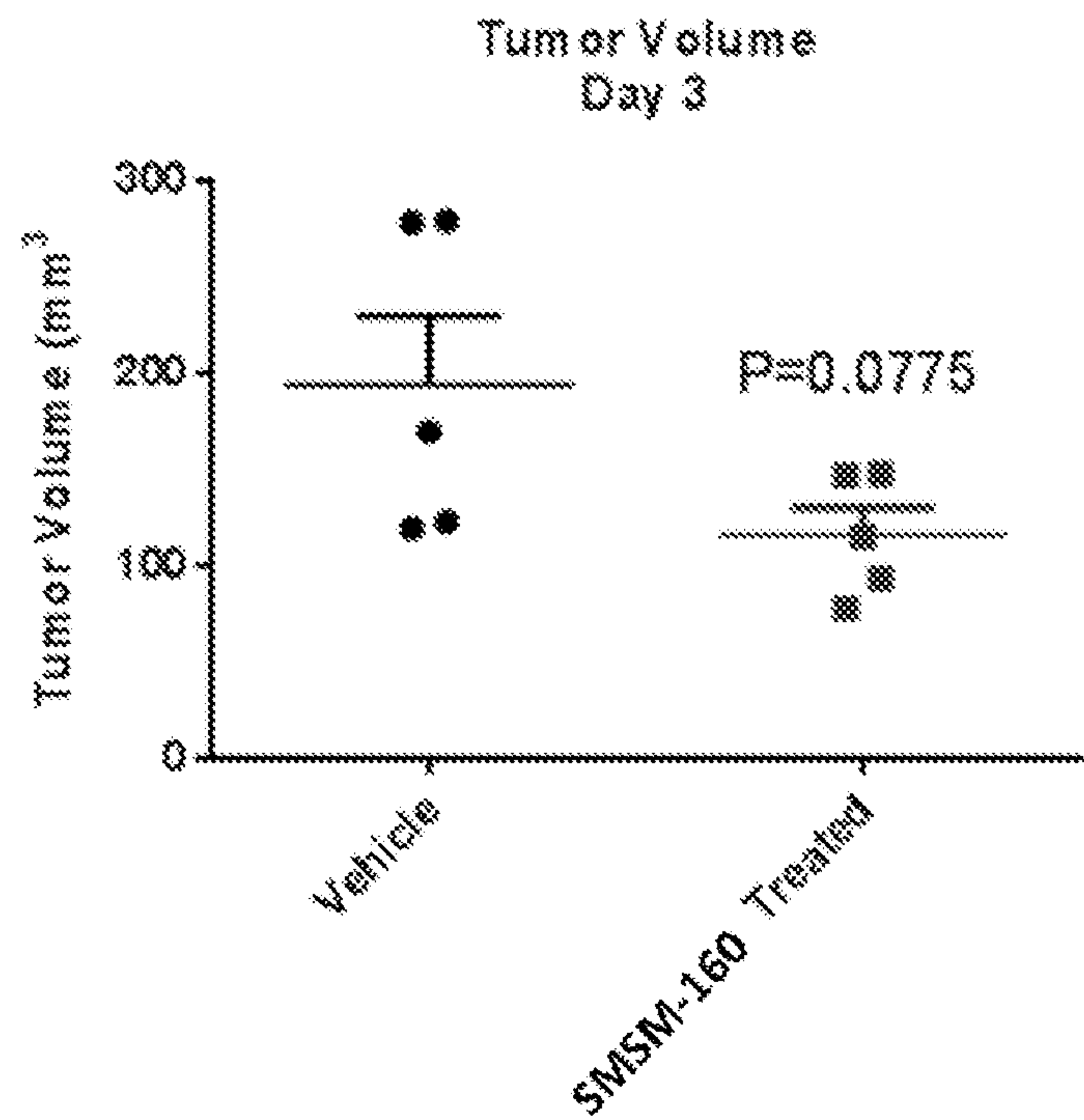


FIG. 18A

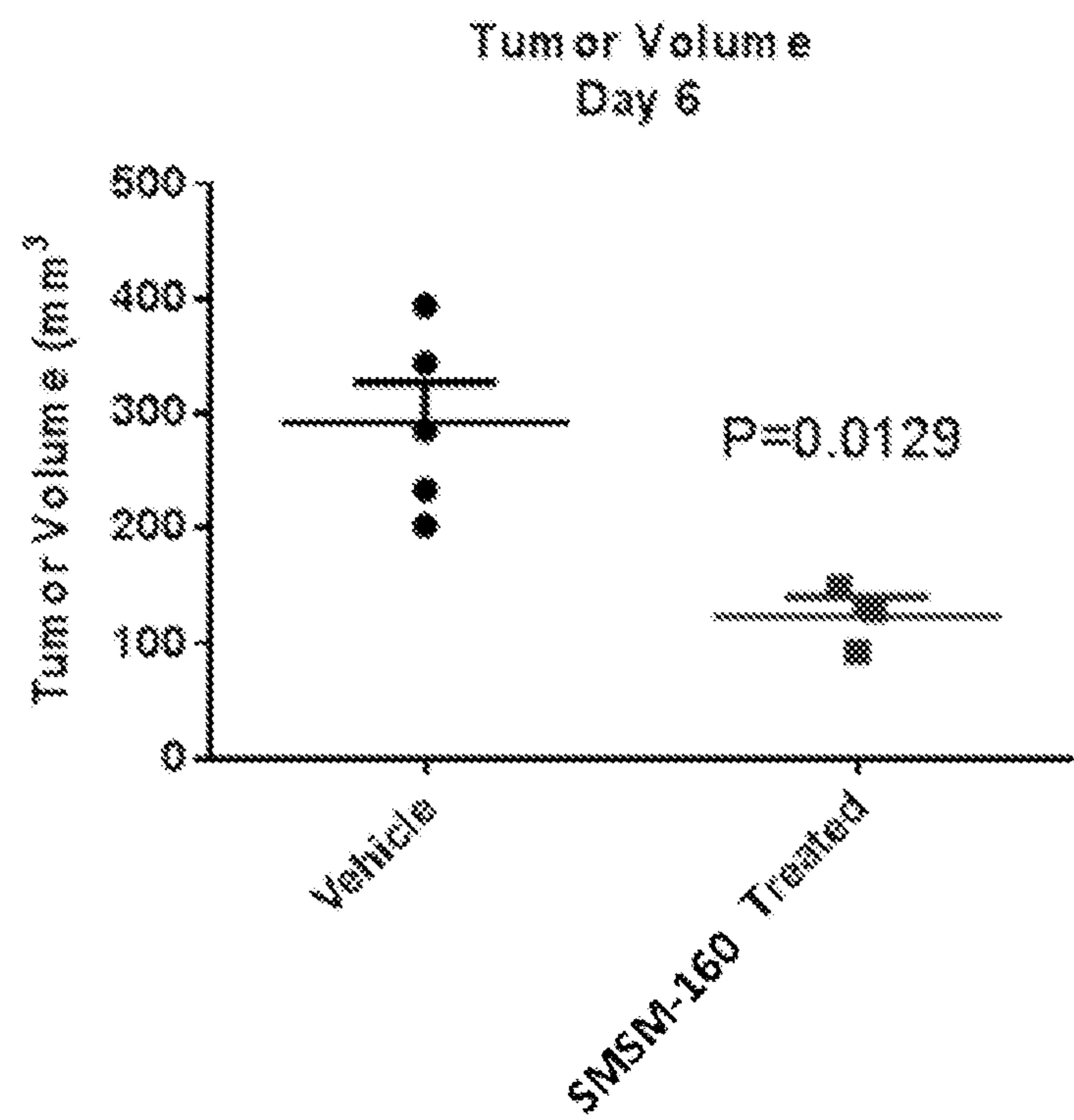


FIG. 18B

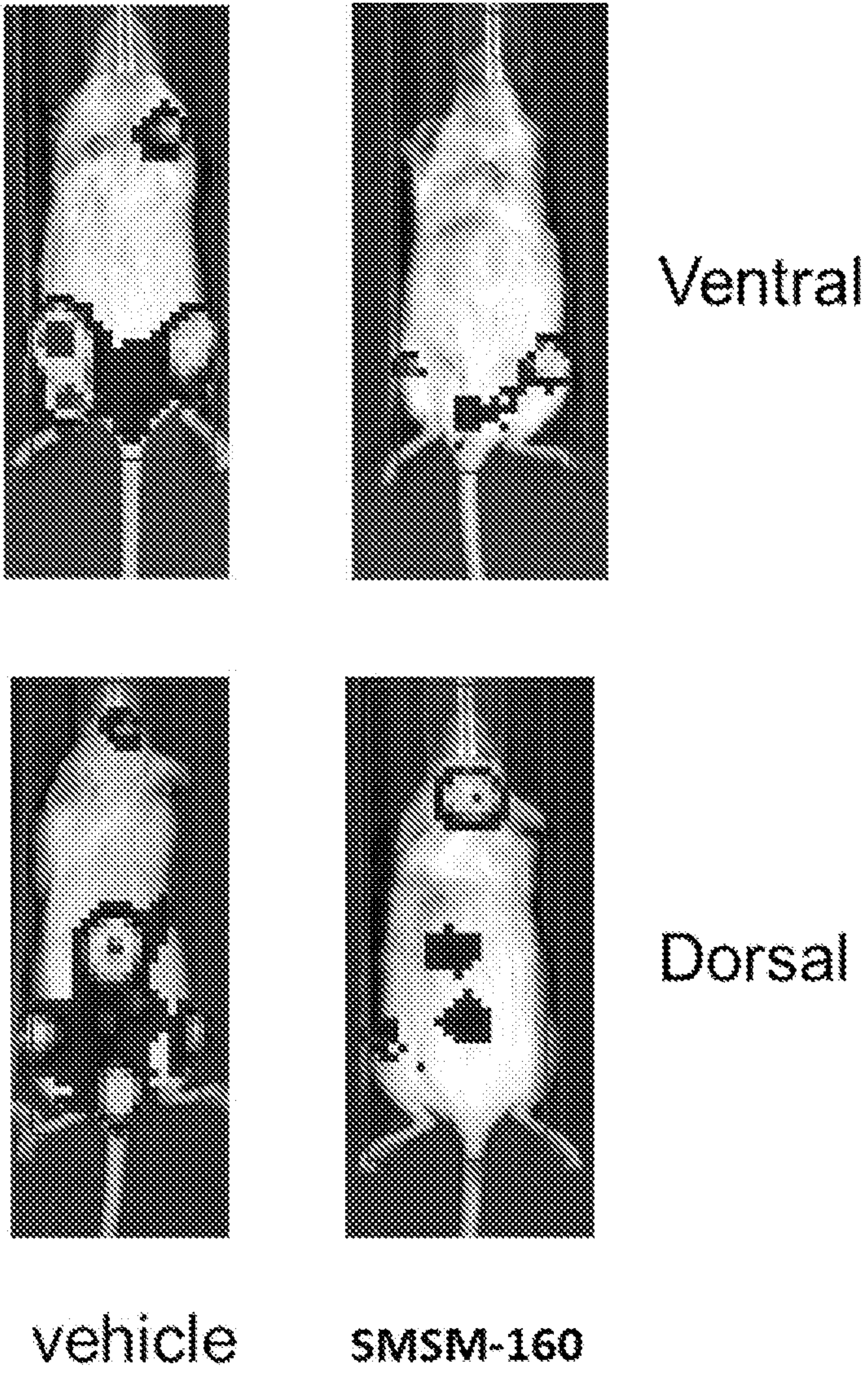


FIG. 19A

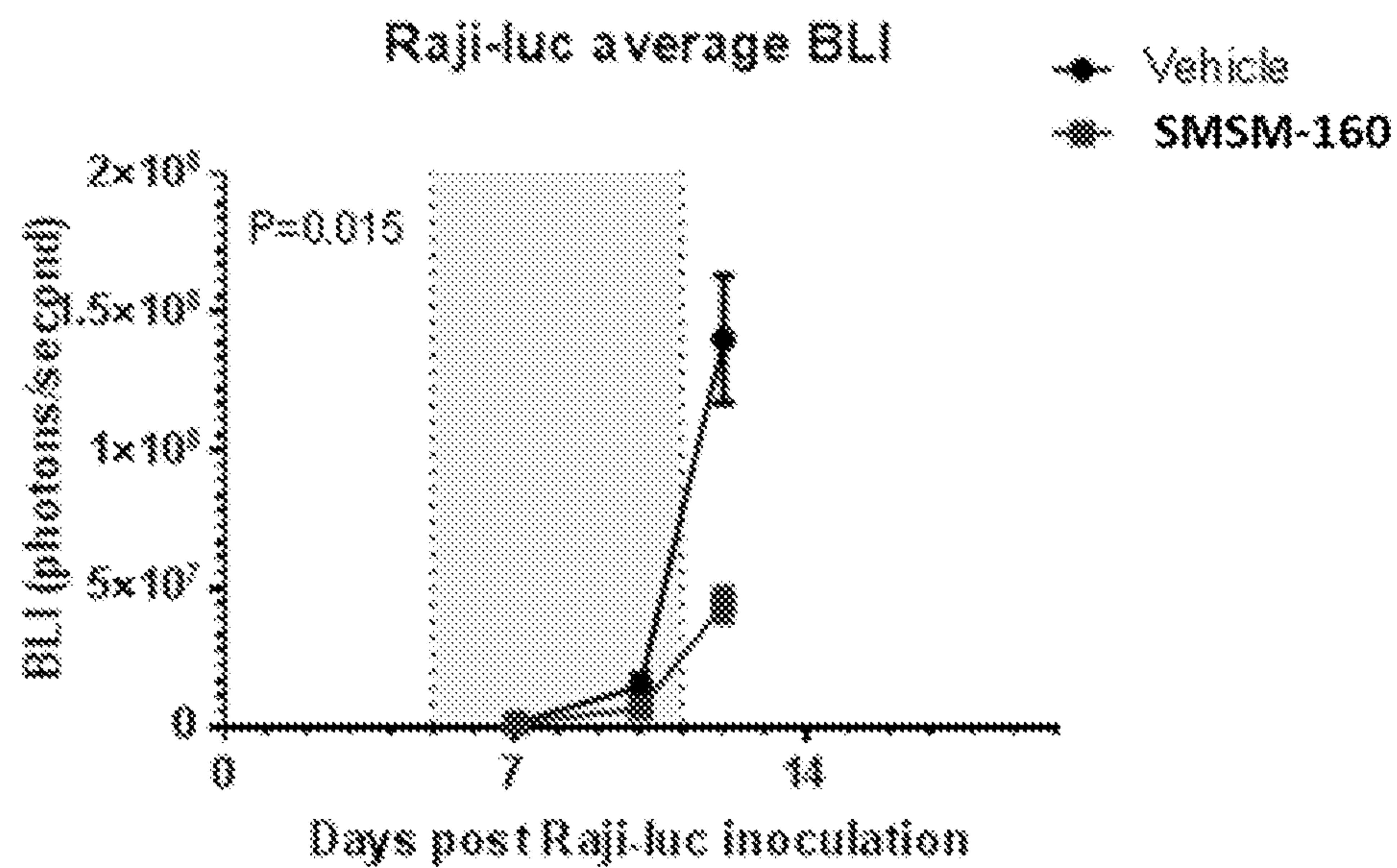


FIG. 19B

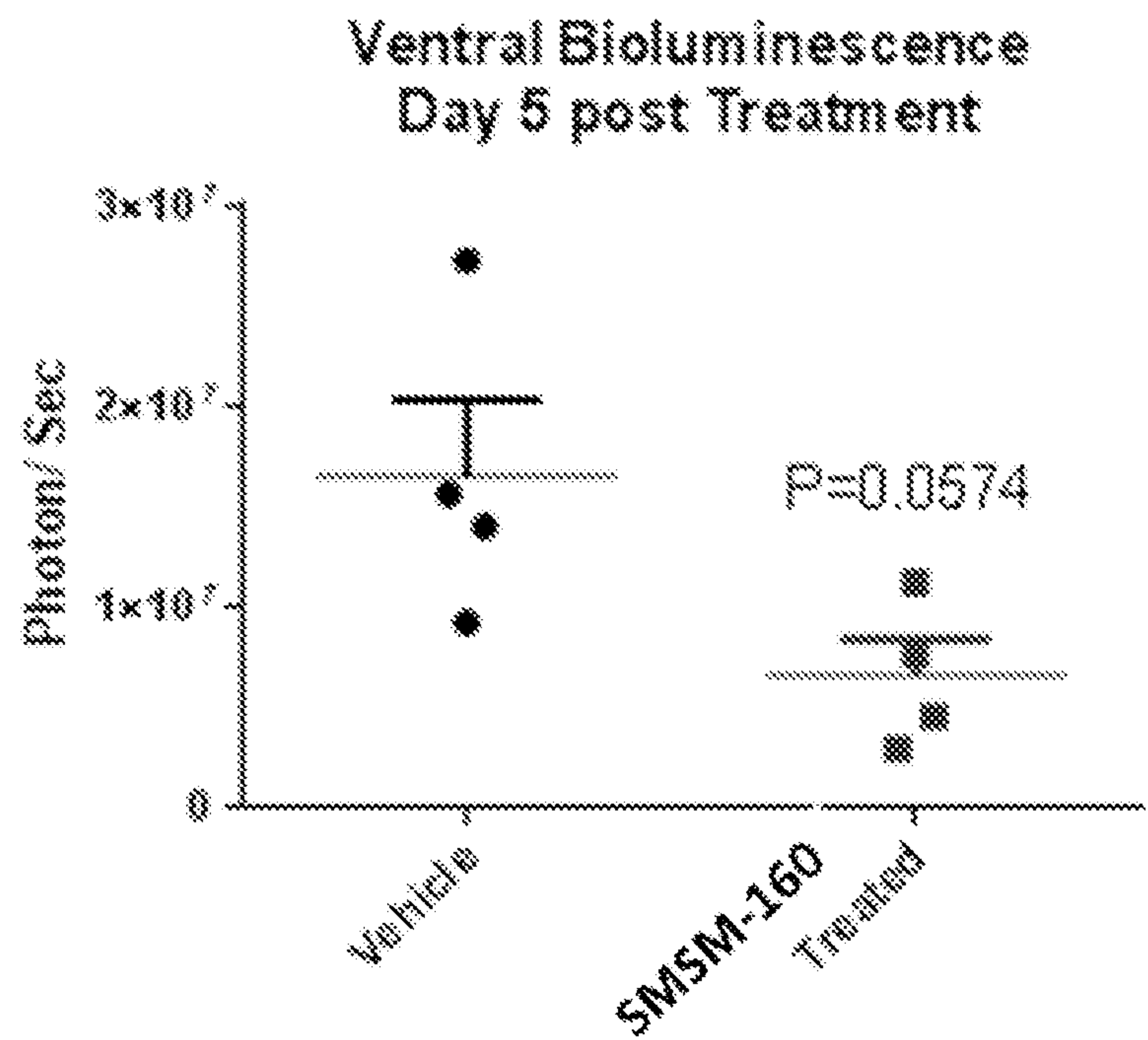


FIG. 19C

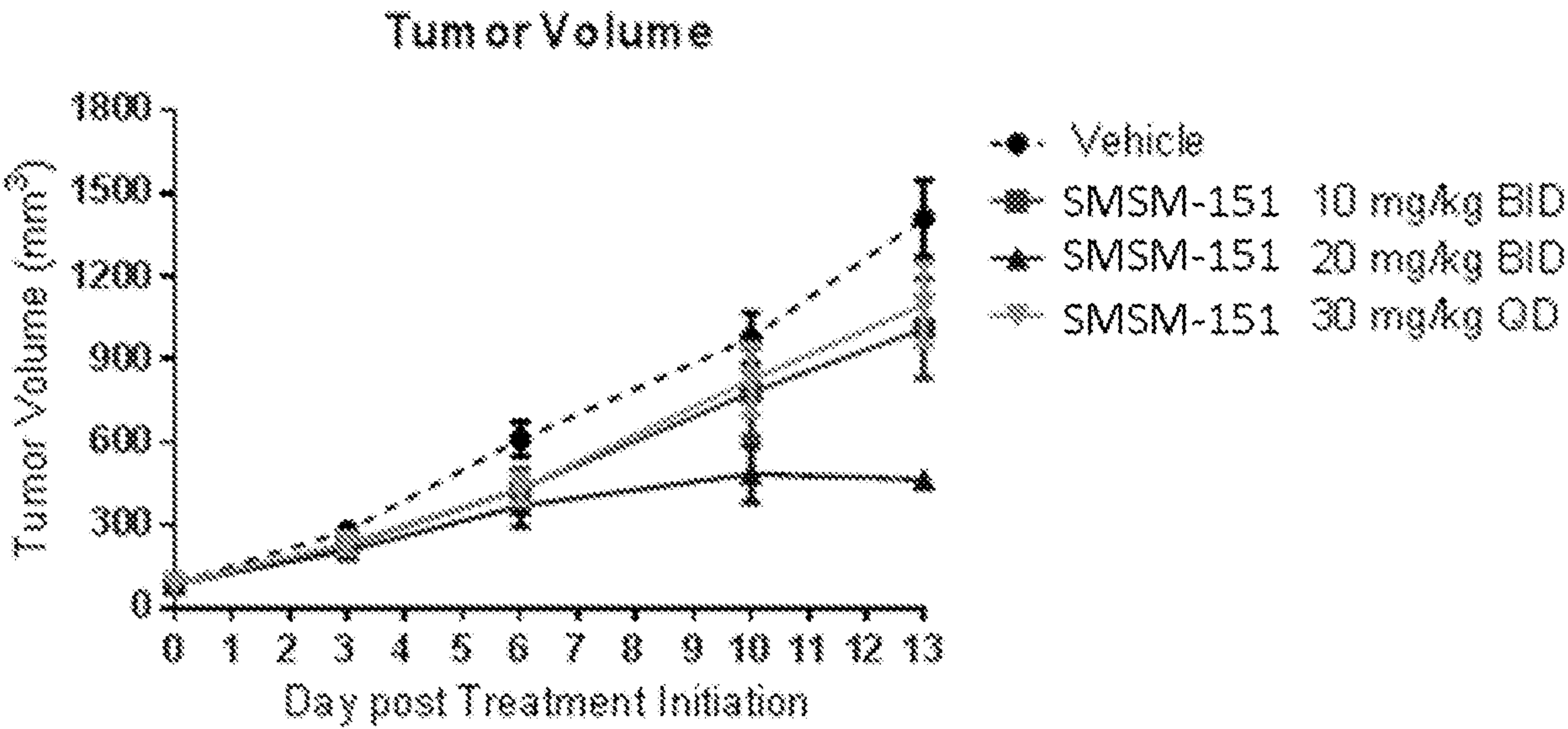


FIG. 20A

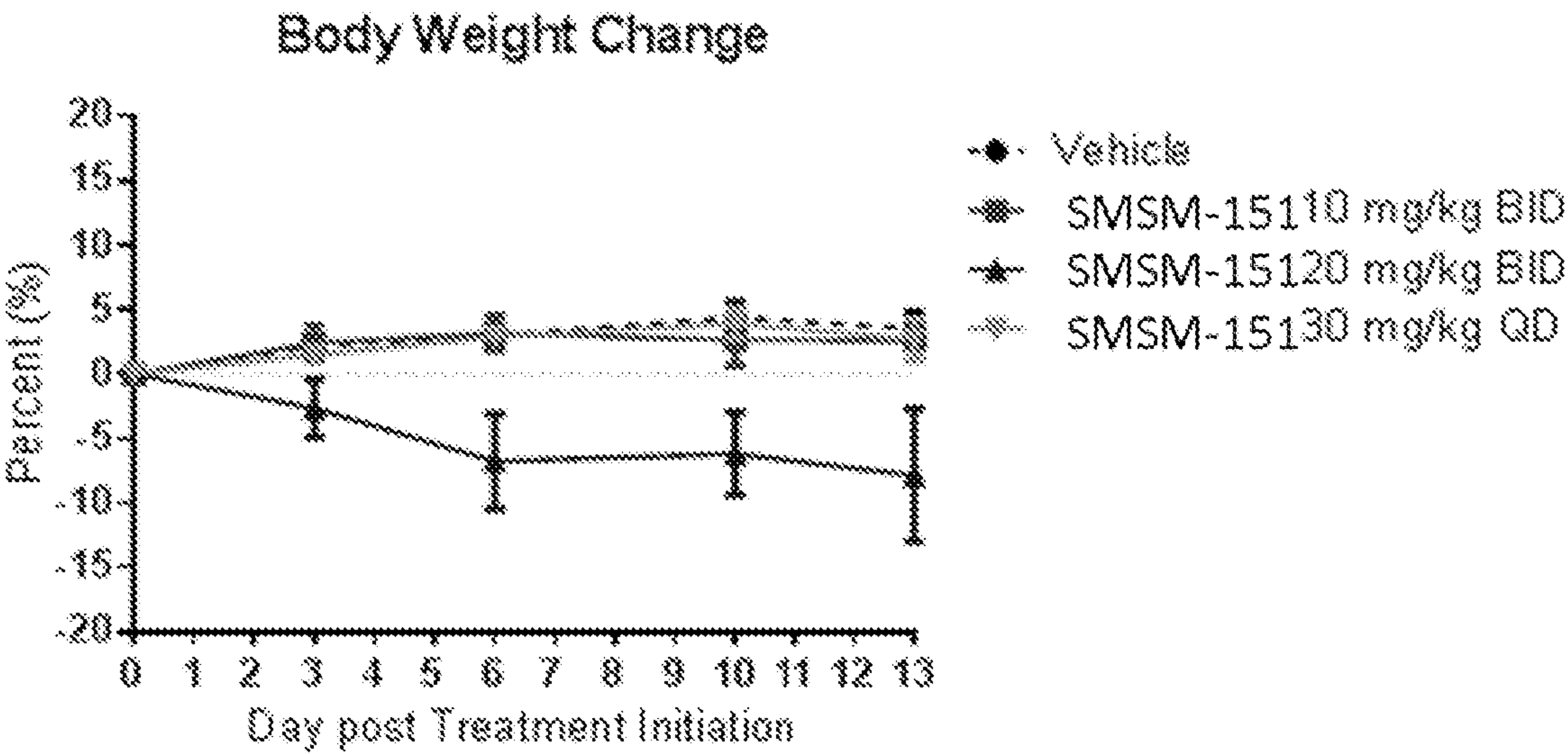


FIG. 20B

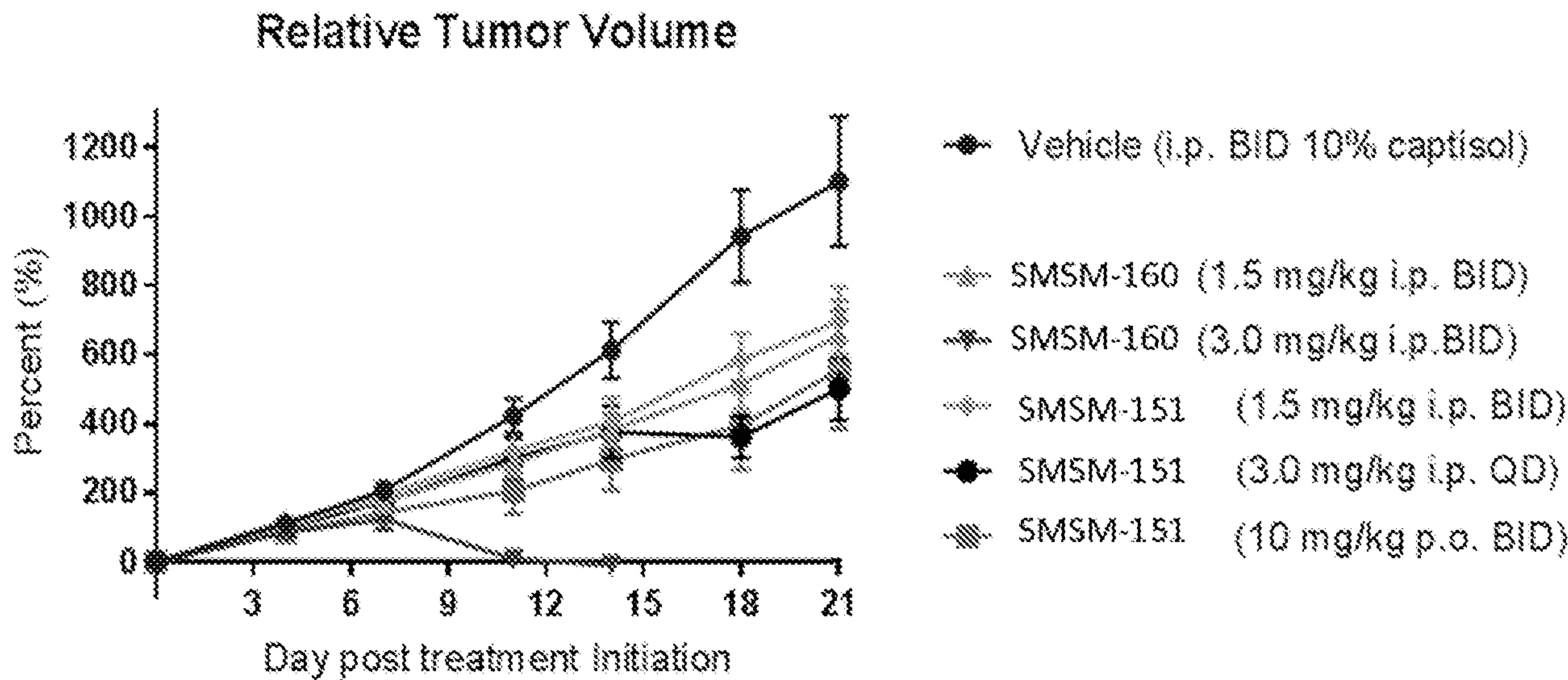


FIG. 21A

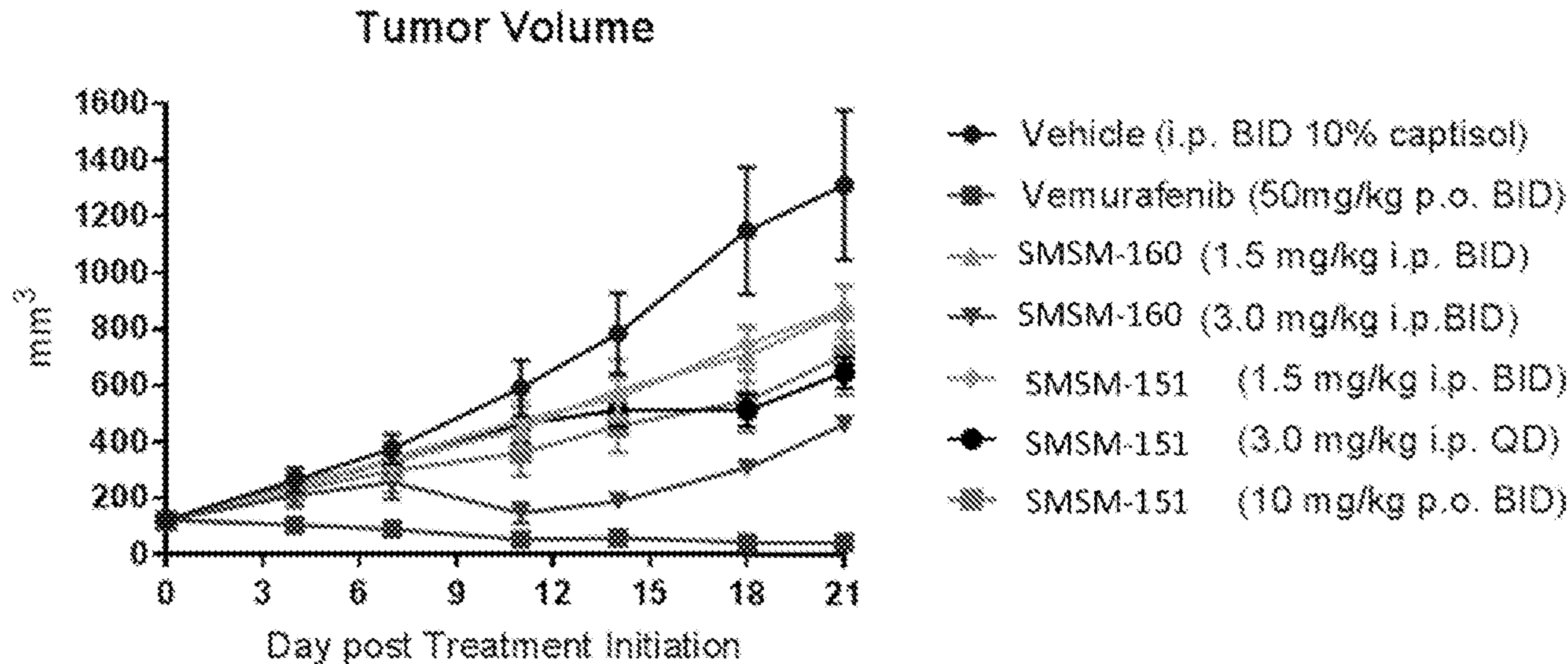


FIG. 21B

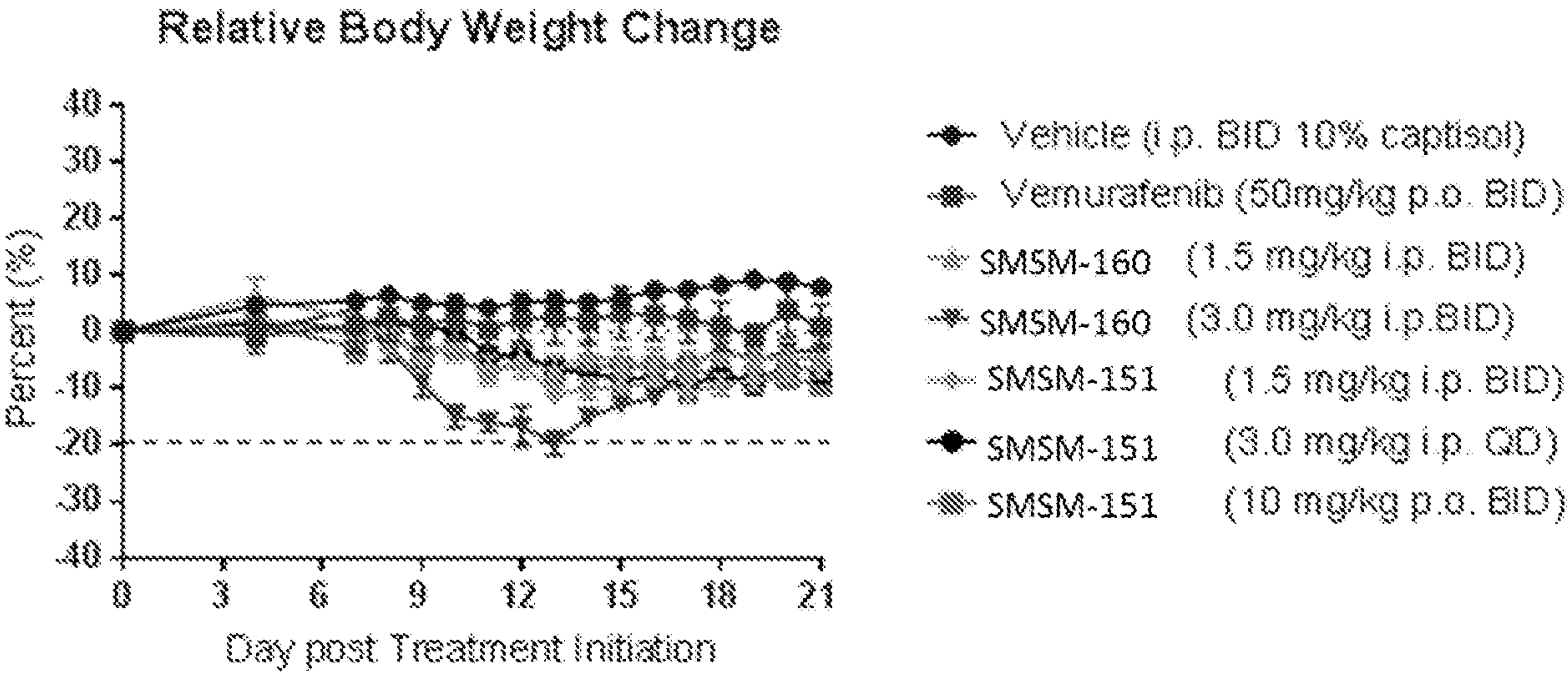
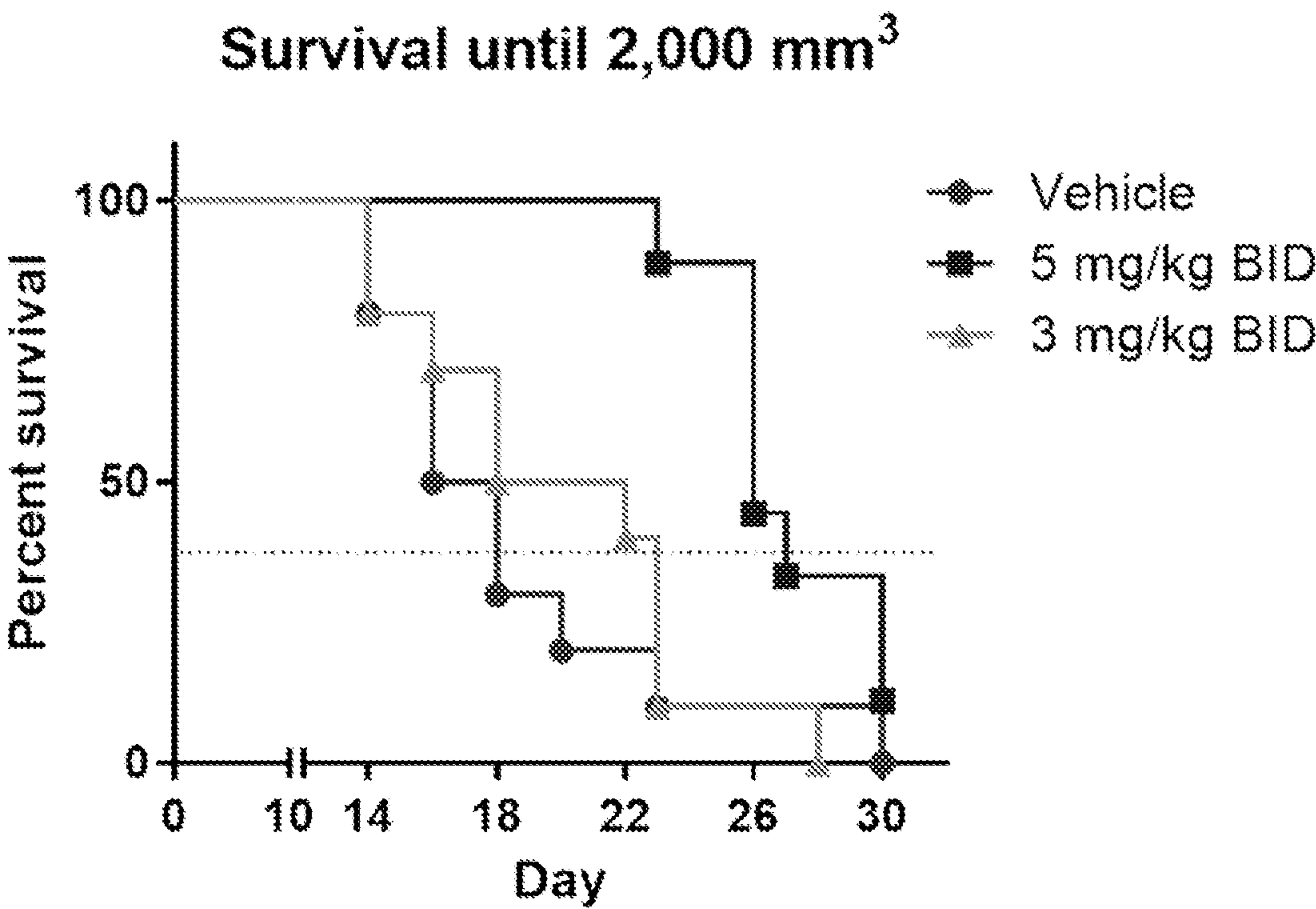


FIG. 21C



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

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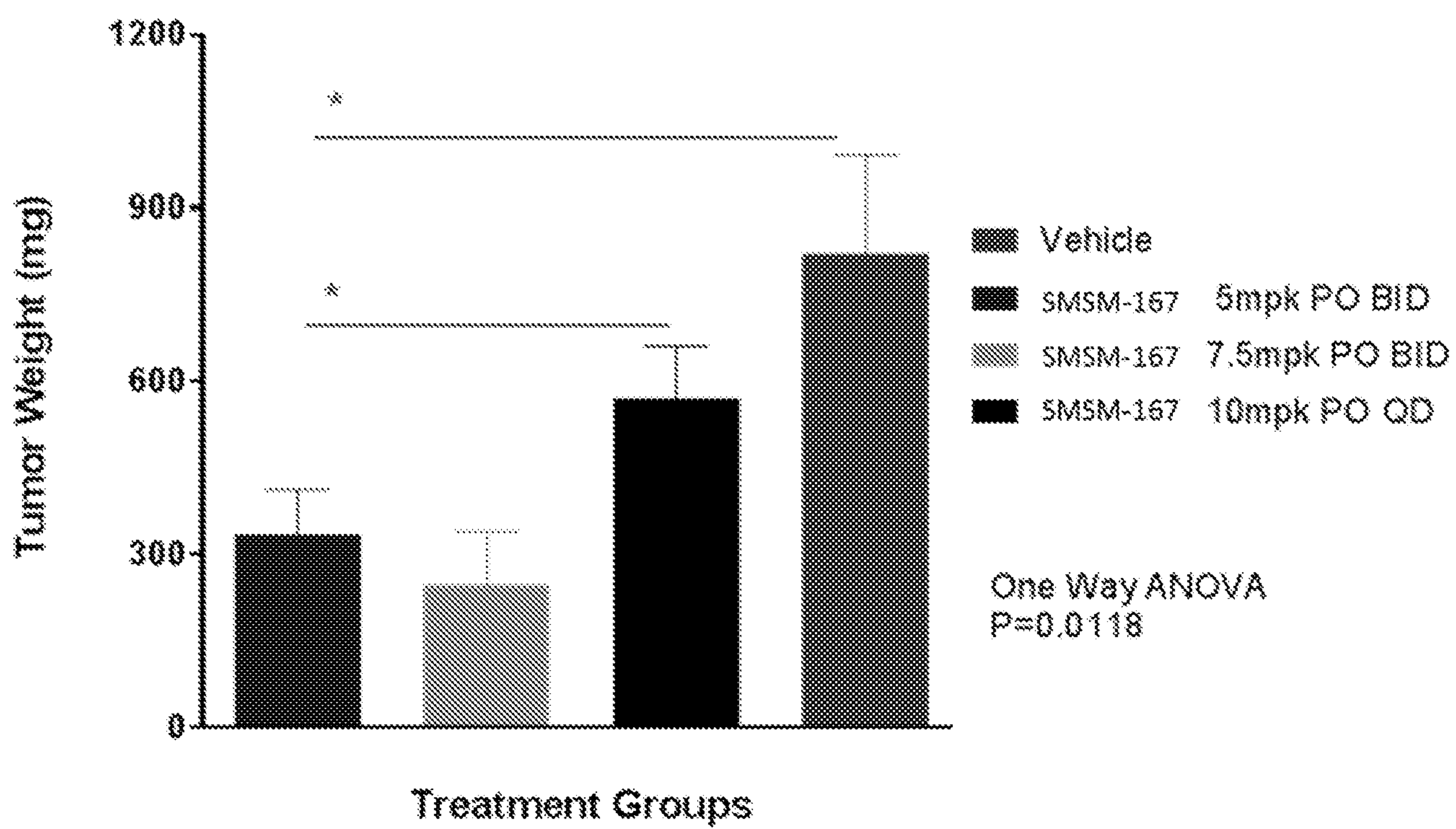


FIG. 23

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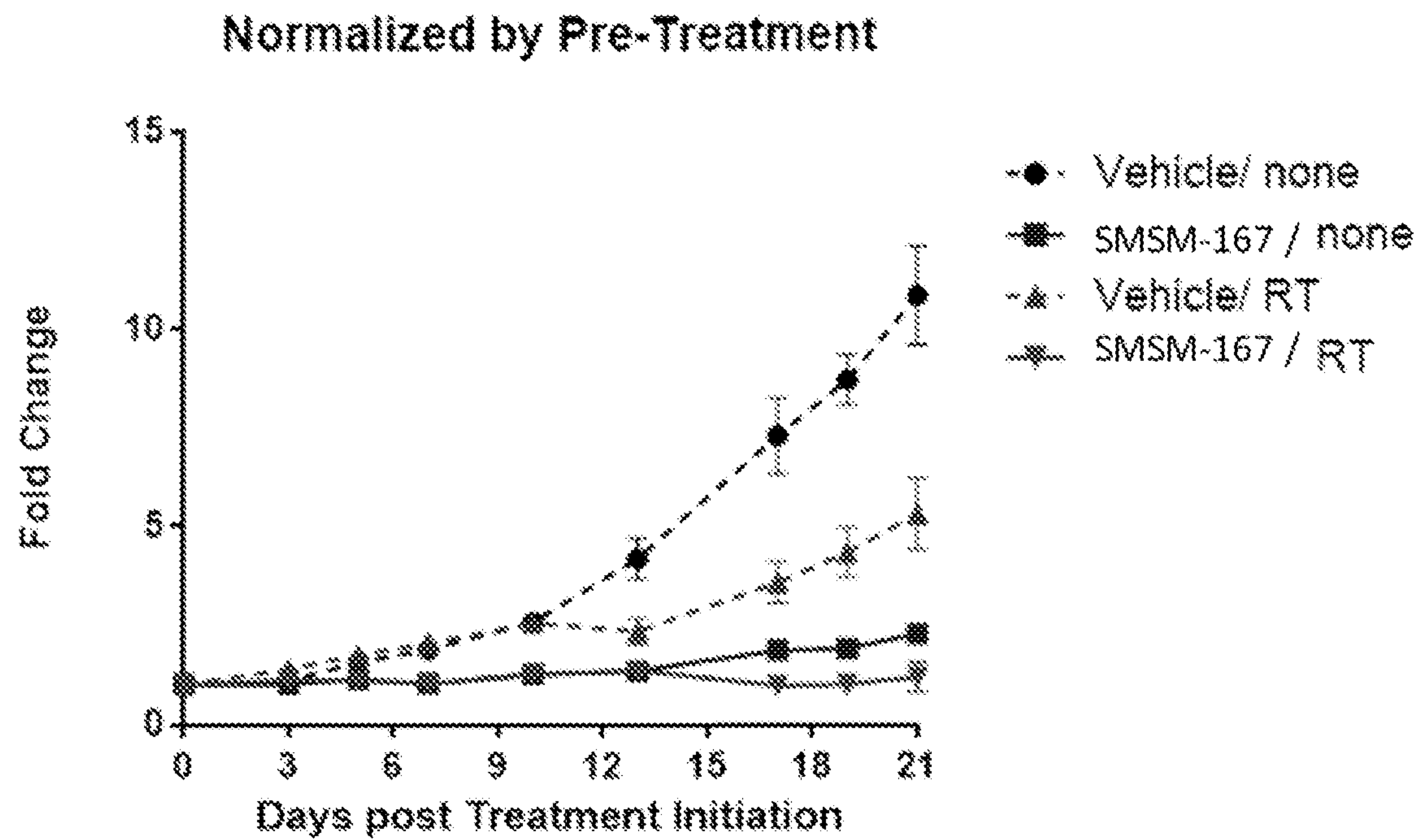


FIG. 24A

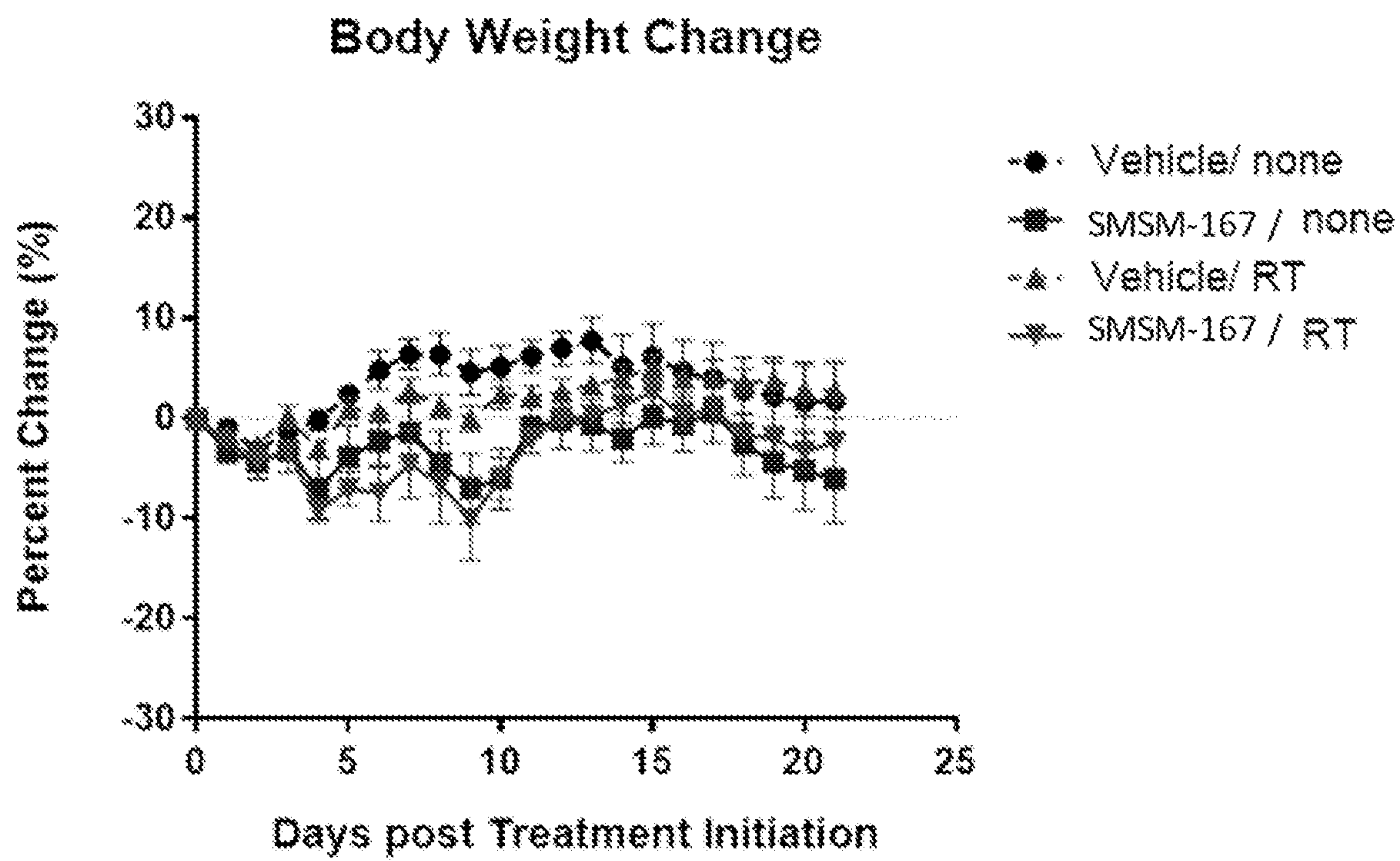


FIG. 24B