

Catalog # 10-4839 RK-33

CAS# 1070773-09-9

3,7-Bis(4-methoxybenzyl)-3,7-dihydro-1,3,4,6,7,9-hexaza-2H-cyclopenta[e]azulene-2-one Lot # FBS2061

RK-33 is an inhibitor of the RNA helicase DDX3 – IC_{50} 's = 4.4-8.4 μ M in high DDX3 expressing lung cancer cell lines A549, H1299, H23, and H460.^{1,2} Inhibition of DDX3 led to activation of cell death pathways, inhibition of Wnt pathway signaling, and abrogation of non-homologous end-joining (NHEJ) DNA repair. RK-33 was also active in colorectal cancer ³, prostate cancer⁴, and medulloblastoma cancer⁵ cell lines. RK-33 caused radiosensitization in breast cancer through inhibition of mitochondrial translation.⁶ RK-33 facilitates differentiation in human embryonic stem cells (hESC) and decreases pluripotency markers as well as reducing teratoma formation.⁷

- 1) Kondaskar et al. (2011), Novel, Broad Spectrum Anticancer Agents Containing the Tricyclic 5:7:5-Fused Diimidazodiazepine Ring System; ACS Med. Chem. Lett., 2 252
- 2) Bol et al. (2015), Targeting DDX3 with a small molecule inhibitor for lung cancer therapy; EMBO Mol. Med., 7 648
- 3) Heerma van Voss et al. (2015), Identification of the DEAD box RNA helicase DDX3 as a therapeutic target in colorectal cancer; Oncotarget, 6 28312
- 4) Xie et al. (2016), RK-33 Radiosensitizes Prostate Cancer Cells by Blocking the RNA Helicase DDX3; Cancer Res., 76 6340
- 5) Tantravedi et al. (2019), Targeting DDX3 in Medulloblastoma Using the Small Molecule Inhibitor RK-33; Transl. Oncol., 12 96
- 6) Heerma van Voss et al. (2018), Targeting mitochondrial translation by inhibiting DDX3: a novel radiosensitization strategy for cancer treatment; Oncogene, **37** 63
- 7) Kerr et al. (2019), Targeting RNA helicase DDX3 in stem cell maintenance and teratoma formation; Genes Cancer, 10 11

PHYSICAL DATA

Molecular Weight: 428.44 Molecular Formula: $C_{23}H_{20}N_6O_3$ Purity: 98% by HPLC

NMR: (Conforms)

Solubility: DMSO (20 mg/ml)
Physical Description: Yellow solid

Storage and Stability: Store as supplied desiccated at -20°C for up to 1 year from the date of purchase. Solutions in

DMSO may be stored at -20°C for up to 1 month.

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