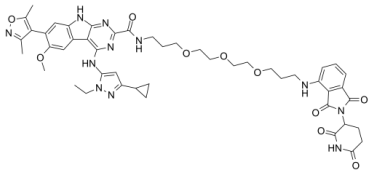
Data Sheet

Product Name:	BETd-246
Cat. No.:	CS-0087862
CAS No.:	2140289-17-2
Molecular Formula:	C48H55N11O10
Molecular Weight:	946.02
Target:	Epigenetic Reader Domain; PROTAC
Pathway:	Epigenetics; PROTAC
Solubility:	DMSO : 200 mg/mL (211.41 mM; Need ultrasonic)



BIOLOGICAL ACTIVITY:

BETd-246 is a second-generation **BET bromodomain (BRD)** inhibitor, exhibiting superior selectivity, potency and antitumor activity[1]. IC50 & Target: BET BRD[1]. **In Vitro:** BETd-246 treatment (0-100 nM, 1-3 h) causes a dose-dependent depletion of BRD2, BRD3 and BRD4 in representative TNBC cell lines with 30-100 nM for 1 h or with 10-30 nM for 3 h incubation.

BETd-246 (100 nM, 24/48 hours) displays strong growth inhibition and apoptosis induction activity in MDA-MB-468 cell lines. BETd-246 induces a rapid and time-dependent downregulation of MCL1 protein in all the TNBC cell lines evaluated. BETd-246 induces much stronger apoptosis than BETi-211.

BETd-246 (100 nM, 24 hours) induces pronounced cell cycle arrest and apoptosis in TNBC cell lines[1]. **In Vivo:** BETd-246 (5 mg/kg, IV, 3 times per week for 3 weeks) treatment effectively inhibits WHIM24 tumor growth, similar to the antitumor activity of BETi-211 with higher dosage and more frequently administration. The treatment of 10 mg/kg induces partial tumor regression during treatment without apparent toxicity. BETd-246 has very limited drug exposure in the xenograft tumor tissue in MDA-M-231and MDA-MB-468 models[1].

References:

[1]. Bai L, et al. Targeted Degradation of BET Proteins in Triple-Negative Breast Cancer. Cancer Res. 2017 May 1;77(9):2476-2487.

CAIndexNames:

9H-Pyrimido[4,5-b]indole-2-carboxamide, 4-[(3-cyclopropyl-1-ethyl-1H-pyrazol-5-yl)amino]-7-(3,5-dimethyl-4-isoxazolyl)-N-[3-[2-[2-[3-[[2-(2,6-dioxo-3-piperidinyl)-2,3-dihydro-1,3-dioxo-1H-isoindol-4-yl]amino]propoxy]ethoxy]ethoxy]propyl]-6-methoxy-

SMILES:

O=C(C1=NC(NC2=CC(C3CC3)=NN2CC)=C4C(NC5=C4C=C(OC)C(C6=C(C)ON=C6C)=C5)=N1)NCCCOCCOCCOCCCNC7=CC=CC(C(N8C(CC9)C(NC9=O)=O)=O)=C7C8=O

Tel: 732-484-9848 Fax: 888-484-5008 E-mail: sales@ChemScene.com

Address: 1 Deer Park Dr, Suite Q, Monmouth Junction, NJ 08852, USA