**Proteins** 

# **Screening Libraries**

# CR-1-31-B

Cat. No.: HY-136453 CAS No.: 1352914-52-3 Molecular Formula:  $C_{28}H_{29}NO_{8}$ Molecular Weight: 508

Target: Eukaryotic Initiation Factor (eIF); Apoptosis

Pathway: Cell Cycle/DNA Damage; Apoptosis

In solvent

Storage: Powder -20°C 3 years

4°C 2 years -80°C 6 months

-20°C 1 month

**Product** Data Sheet

# **SOLVENT & SOLUBILITY**

In Vitro

DMSO: 230 mg/mL (452.76 mM; Need ultrasonic)

	Solvent Mass Concentration	1 mg	5 mg	10 mg
Preparing Stock Solutions	1 mM	1.9685 mL	9.8425 mL	19.6850 mL
	5 mM	0.3937 mL	1.9685 mL	3.9370 mL
	10 mM	0.1969 mL	0.9843 mL	1.9685 mL

Please refer to the solubility information to select the appropriate solvent.

In Vivo

- 1. Add each solvent one by one: 10% DMSO >> 40% PEG300 >> 5% Tween-80 >> 45% saline Solubility: ≥ 5.75 mg/mL (11.32 mM); Clear solution
- 2. Add each solvent one by one: 10% DMSO >> 90% (20% SBE-β-CD in saline) Solubility: ≥ 5.75 mg/mL (11.32 mM); Clear solution
- 3. Add each solvent one by one: 10% DMSO >> 90% corn oil Solubility: ≥ 5.75 mg/mL (11.32 mM); Clear solution

# **BIOLOGICAL ACTIVITY**

Description

CR-1-31-B is a synthetic rocaglate and a potent eIF4A inhibitor. CR-1-31-B exhibits powerful inhibitory effects over eIF4A by perturbing the interaction between eIF4A and RNA, sequentially impeding initiation during protein synthesis. CR-1-31-B perturbs association of Plasmodium falciparum eIF4A (PfeIF4A) with RNA. CR-1-31-B induces apoptosis of neuroblastoma and gallbladder cancer cells<sup>[1][2][3][4]</sup>.

IC<sub>50</sub> & Target

eIF4

#### In Vitro

CR-1-31-B (100 nM; 24 hours) inhibits MUC1-C translation in MCF-10A cells (EGF-stimulated)<sup>[1]</sup>.

CR-1-31-B (10 and 100 nM) decreases MUC1-C abundance in MDA-MB-468 breast cancer cells<sup>[1]</sup>.

CR-1-31B sensitizes gallbladder cancer cells to TRAIL-mediated apoptosis through the translational downregulation of c- $\text{FLIP}^{[2]}$ .

Neuroblastoma (NB) cell lines exhibit decreased viability, increased apoptosis rates as well as changes in cell cycle distribution when treated with the synthetic rocaglate CR-1-31-B (24-72 hours), which clamps eIF4A and eIF4F onto mRNA, resulting in a translational block<sup>[4]</sup>.

CR-1-31-B (100 nM; 5 hours) treatment increases reverse glutamine metabolism in pancreatic cancer cells<sup>[5]</sup>.

MCE has not independently confirmed the accuracy of these methods. They are for reference only.

# Western Blot Analysis $^{[1]}$

Cell Line:	MCF-10A cells (EGF-stimulated)
Concentration:	100 nM
Incubation Time:	24 hours
Result:	Blocked increases in MUC1-C abundance.

#### Cell Viability Assay<sup>[4]</sup>

Cell Line:	SH-SY5Y cells and Kelly cells	
Concentration:	0.1-100 nM	
Incubation Time:	24-72 hours	
Result:	A significant decrease in SH-SY5Y viability was observed at 10 nM for all time points. Significantly decreased the viability of Kelly cells at 5 nM. The calculated IC <sub>50</sub> at 48 h was 20 nM for SH-SY5Y and 4 nM for Kelly cells.	

#### Apoptosis Analysis<sup>[4]</sup>

Cell Line:	SH-SY5Y and Kelly cells
Concentration:	SH-SY5Y cells were treated with 10, 20, and 50 nM and Kelly cells with 1, 5, and 10 nM
Incubation Time:	24-72 hours
Result:	Triggered apoptosis.

### In Vivo

CR-1-31-B (2 mg/kg in 60  $\mu$ L olive oil; IP; once every 2 days for 28 days) reduces the growth and initiates TRAIL-induced apoptosis in a BALB/c nude mice model of gallbladder cancer cells (GBC)<sup>[2]</sup>.

CR-1-31-B (0.2 mg/kg; IP; daily for 7 days; murine orthotopic transplant model of pancreatic ductal adenocarcinoma) effectively inhibits protein synthesis and growth of pancreatic tumours<sup>[5]</sup>.

 $\label{eq:mce} \mbox{MCE has not independently confirmed the accuracy of these methods. They are for reference only.}$ 

## **CUSTOMER VALIDATION**

- J Exp Clin Cancer Res. 2022 Dec 9;41(1):340.
- iScience, 2023 Nov 16.
- Int J Mol Sci. 2023, 24(3), 2055.
- Research Square Preprint. 2022 Jun.

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#### **REFERENCES**

- [1]. Jin C, Rajabi H, Rodrigo CM, Porco JA Jr, Kufe D. Targeting the eIF4A RNA helicase blocks translation of the MUC1-C oncoprotein. Oncogene. 2013;32(17):2179-2188.
- [2]. Cao Y, et al. Targeting eIF4A using rocaglate CR 1 31B sensitizes gallbladder cancer cells to TRAIL mediated apoptosis through the translational downregulation of c FLIP. Oncol Rep. 2021;45(1):230-238.
- [3]. Langlais D, et al. Rocaglates as dual-targeting agents for experimental cerebral malaria. Proc Natl Acad Sci U S A. 2018;115(10):E2366-E2375.
- [4]. Skofler C, et al. Eukaryotic Translation Initiation Factor 4Al: A Potential Novel Target in Neuroblastoma. Cells. 2021;10(2):301. Published 2021 Feb 2.
- [5]. Chan K, et al. eIF4A supports an oncogenic translation program in pancreatic ductal adenocarcinoma. Nat Commun. 2019;10(1):5151. Published 2019 Nov 13.

Caution: Product has not been fully validated for medical applications. For research use only.

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