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# **Certificate of Analysis**

### www.tocris.com

#### Product Name: MRT 67307 dihydrochloride

Catalog No.: 5134 Batch No.: 2

CAS Number: IUPAC Name: 1781882-89-0

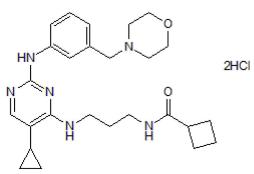
Name: *N*-[3-[[5-Cyclopropyl-2-[[3-(4-morpholinylmethyl)phenyl]amino]-4-pyrimidinyl]amino]propyl]cyclobutanecarboxamide dihydrochloride

## 1. PHYSICAL AND CHEMICAL PROPERTIES

Batch Molecular Formula: Batch Molecular Weight: Physical Appearance: Solubility: C<sub>26</sub>H<sub>36</sub>N<sub>6</sub>O<sub>2</sub>.2HCl.1<sup>3</sup>/<sub>4</sub>H<sub>2</sub>O 569.04 Beige solid water to 20 mM DMSO to 100 mM Desiccate at RT

# Storage:

**Batch Molecular Structure:** 



#### 2. ANALYTICAL DATA

TLC: HPLC: <sup>1</sup>H NMR: Mass Spectrum: Microanalysis: R<sub>f</sub> = 0.12 (Chloroform:Methanol [9:1]) Shows 98.3% purity Consistent with structure Consistent with structure Carbon Hydrogen Nitrogen Theoretical 54.88 7.35 14.77 Found 54.83 7.23 14.67

Caution - Not Fully Tested • Research Use Only • Not For Human or Veterinary Use

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2

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IUPAC Name: N-[3-[[5-Cyclopropyl-2-[[3-(4-morpholinylmethyl)phenyl]amino]-4-pyrimidinyl]amino]propyl]cyclobutanecarboxamide dihydrochloride

#### **Description:**

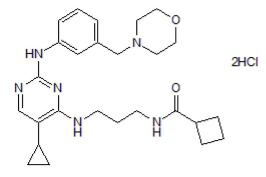
MRT 67307 dihydrochloride is a salt inducible kinase (SIK) inhibitor (IC<sub>50</sub> values are 67, 250 and 430 nM for SIK2, SIK1 and SIK3 respectively) and a potent inhibitor of ULK1 and 2 (IC<sub>50</sub> values of 45 and 38 nM, respectively). MRT 67307 dihydrochloride also inhibits TBK1, MARK1-4, IKK $\epsilon$  and NUAK1 (IC<sub>50</sub> values are 19, 27-52, 160 and 230 nM respectively). Has no effect on IKK $\alpha$  or IKK $\beta$ . Induces IL-10 secretion and inhibits TNF- $\alpha$  and IL-6 secretion in bacterial LPS-stimulated macrophages. Also enhances IL-1-induced activation of NF $\kappa$ B-dependent gene transcription in mouse embryonic fibroblast (MEF) cells. Inhibits autophagy.... Please see product specific page on www.tocris.com for full description.

#### **Physical and Chemical Properties:**

Batch Molecular Formula:  $C_{26}H_{36}N_6O_2.2HCI.1^3/_4H_2O$ Batch Molecular Weight: 569.04 Physical Appearance: Beige solid

#### Minimum Purity: ≥98%

#### **Batch Molecular Structure:**



**Storage:** Desiccate at RT. This product is packaged under an inert atmosphere.

Catalog No.: 5134

#### Solubility & Usage Info:

water to 20 mM DMSO to 100 mM

This product is supplied in lyophilized form. It may appear as a solid, gel or film and be very hard to visualize. Solutions should be made by adding solvent directly to the vial. The vial should then be vortexed vigorously to ensure the product has completely dissolved.

#### Stability and Solubility Advice:

Some solutions can be difficult to obtain and can be encouraged by rapid stirring, sonication or gentle warming (in a  $45-60^{\circ}C$  water bath).

Information concerning product stability, particularly in solution, has rarely been reported and in most cases we can only offer a general guide. \*Unless contradicted by product-specific protocols or instructions, our standard recommendations apply:

SOLIDS: Provided storage is as stated on the product label and the vial is kept tightly sealed, the product can be stored for up to 6 months from date of receipt.

SOLUTIONS: We recommend that stock solutions, once prepared, are stored aliquoted in tightly sealed vials at -20°C or below and used within 1 month. Wherever possible solutions should be made up and used on the same day.

#### **References:**

**Galluzzi** *et al* (2017) Pharmacological modulation of autophagy: therapeutic potential and persisting obstacles. Nat.Rev.Drug.Discov. **16** 487. PMID: 28529316 .

**Petherick** *et al* (2015) Pharmacological inhibition of ULK1 kinase blocks mammalian target of rapamycin (mTOR)-dependent autophagy. J.Biol.Chem. **290** 28726. PMID: 26614783.

**Clark** *et al* (2012) Phosphorylation of CRTC3 by the salt-inducible kinases controls the interconversion of classically activated and regulatory macrophages. Proc.Natl.Acad.Sci.U.S.A. **109** 16986. PMID: 23033494.

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