

## ML-323

Catalog No: tcsc2274

Available Sizes

Size: 5mg

Size: 10mg

Size: 50mg

**Size:** 100mg

**Specifications** 

CAS No:

1572414-83-5

Formula:

 $C_{23}H_{24}N_{6}$ 

Pathway: Cell Cycle/DNA Damage

**Target:** 

Deubiquitinase

Purity / Grade:

>98%

## Solubility:

DMSO : ≥ 49 mg/mL (127.44 mM)

## **Observed Molecular Weight:**

384.48

## **Product Description**

ML-323 is a reversible, potent **USP1-UAF1** inhibitor with **IC**<sub>50</sub> of 76 nM in a Ub-Rho assay. The measured inhibition constants of ML-

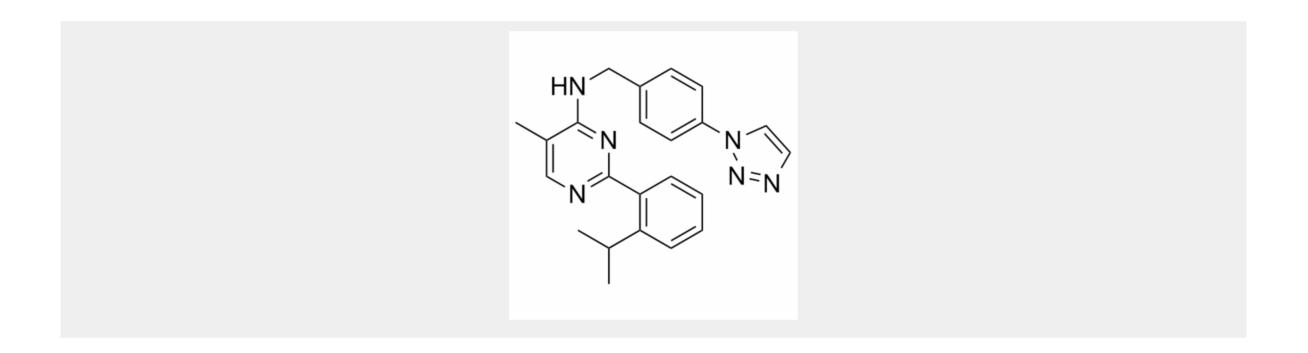


323 for the free enzyme ( $\mathbf{K}_{\mathbf{i}}$ ) is 68 nM.

IC50 & Target: IC50: 76 nM (USP1-UAF1, in Ub-Rho assay)<sup>[1]</sup>

Ki: 68 nM (USP1-UAF1)<sup>[1]</sup>

*In Vitro:* ML-323 (ML323) is a highly potent inhibitor of the USP1-UAF1 deubiquitinase complex with excellent selectivity against human DUBs, deSUMOylase, deneddylase and unrelated proteases. ML-323 is a potent USP1-UAF1 inhibitor with IC50 values of 76 nM in a ubiquitin-rhodamine (Ub-Rho) assay and 174 nM and 820 nM in orthogonal gel-based assays using K63-linked diubiquitin (di-Ub) and monoubiquitinated PCNA (Ub-PCNA) as substrates, respectively. ML-323 probably exerts its inhibitory effect through an allosteric mechanism. The measured inhibition constants of ML-323 for the free enzyme (K<sub>i</sub>) and the enzyme-substrate complex (K'<sub>i</sub>) are 68 nM and 183 nM. Besides, ML-323 potentiates Cisplatin cytotoxicity in non-small cell lung cancer and osteosarcoma cells<sup>[1]</sup>. ML-323 (ML323), a probe molecule that displays reversible, nanomolar inhibitory activity and excellent selectivity toward USP1/UAF1. In addition, ML-323 potentiates the cytotoxicity of Cisplatin and increases endogenous monoubiquitination levels of both PCNA and FANCD2, two known cellular targets of USP1/UAF1<sup>[2]</sup>.



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