**Proteins** 

# **Product** Data Sheet

# AZD1208 hydrochloride

Cat. No.: HY-15604A CAS No.: 1621866-96-3

Molecular Formula:  $\mathsf{C}_{21}\mathsf{H}_{22}\mathsf{CIN}_3\mathsf{O}_2\mathsf{S}$ 

Target: Pim; Autophagy; Apoptosis

415.94

Pathway: JAK/STAT Signaling; Autophagy; Apoptosis

Storage: Please store the product under the recommended conditions in the Certificate of

Analysis.

## **SOLVENT & SOLUBILITY**

In Vitro

Molecular Weight:

DMSO: 83.3 mg/mL (200.27 mM; Need ultrasonic and warming)

Preparing Stock Solutions	Solvent Mass Concentration	1 mg	5 mg	10 mg
	1 mM	2.4042 mL	12.0210 mL	24.0419 mL
	5 mM	0.4808 mL	2.4042 mL	4.8084 mL
	10 mM	0.2404 mL	1.2021 mL	2.4042 mL

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

## **BIOLOGICAL ACTIVITY**

Description	${\sf AZD1208}\ hydrochloride\ is\ an\ orally\ bioavailable,\ highly\ selective\ PIM\ kinases\ inhibitor^{[1]}.$
IC <sub>50</sub> & Target	PIM <sup>[1]</sup>
In Vitro	AZD1208 hydrochloride shows good antiproliferative activity in a megakaryoblastic leukemia cell line, MOLM-16, with ${\rm GI}_{50}$ values less than 100 nM $^{[1]}$ . AZD1208 hydrochloride (10 $\mu$ M) inhibits the growth of Ramos cells, and at 1 $\mu$ M, strongly inhibits PIM kinases in all cells at 1 $\mu$ M. AZD1208 hydrochloride induces apoptosis, and PIM2 knockdown is mainly associated with an alteration of the cell cycle $^{[2]}$ . The combination of AZD1208 hydrochloride and AZD2014 rapidly activates AMPK $\alpha$ , a negative regulator of translation machinery through mTORC1/2 signaling in AML cells; profoundly inhibits AKT and 4EBP1 activation; and suppresses polysome formation $^{[3]}$ .

## **PROTOCOL**

Cell Assay [1]

MOLM-16 cells, purchased from DSMZ and cultured in RPMI containing 10% fetal bovine serum (FBS) and 1% L-glutamine,

are plated at 20,000 cells per well in 96 well plates overnight. Cells are treated for 72 hours with compound (including AZD1208 hydrochloride) or control vehicle (dimethyl sulfoxide) and cell viability is measured after the addition of Cell Titer-Blue for 4 hours at 37 $^{\circ}$ C and reading of fluorescence on a Tecan Infinite $^{\circ}$ 200. The Gl<sub>50</sub> is determined by calculating growth at each dose relative to vehicle treated cells and cell viability at the time of treatment<sup>[1]</sup>.

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

#### **CUSTOMER VALIDATION**

- Science. 2017 Dec 1;358(6367):eaan4368.
- Sci Transl Med. 2018 Jul 18;10(450):eaaq1093.
- Nat Commun. 2019 Apr 23;10(1):1844.
- eNeuro. 2019 Aug 22;6(4):ENEURO.0003-19.2019.
- Exp Ther Med. November 25, 2021.

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#### **REFERENCES**

[1]. Dakin LA, et al. Discovery of novel benzylidene-1,3-thiazolidine-2,4-diones as potent and selective inhibitors of the PIM-1, PIM-2, and PIM-3 protein kinases. Bioorg Med Chem Lett. 2012 Jul 15;22(14):4599-604.

[2]. Kreuz S, et al. Loss of PIM2 enhances the anti-proliferative effect of the pan-PIM kinase inhibitor AZD1208 in non-Hodgkin lymphomas. Mol Cancer. 2015 Dec 8;14:205.

[3]. Harada M, et al. The novel combination of dual mTOR inhibitor AZD2014 and pan-PIM inhibitor AZD1208 inhibits growth in acute myeloid leukemia via HSF pathway suppression. Oncotarget. 2015 Nov 10;6(35):37930-47.

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

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