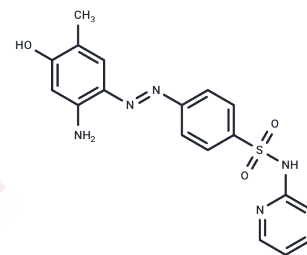


MS436

Chemical Properties

CAS No. : 1395084-25-9
 Formula: C₁₈H₁₇N₅O₃S
 Molecular Weight: 383.42
 Storage: Powder: -20°C for 3 years | In solvent: -80°C for 1 year
Actual storage temperature shall be subject to the COA.



Biological Description

Description	MS436 is a selective, small-molecule inhibitor for the BRD4 bromodomains.
Targets(IC50)	Epigenetic Reader Domain
In vivo	MS436 exhibits low nanomolar affinity (K _i =30-50 nM) for the first bromodomain of BRD4. In mouse macrophages with NF-κB-directed expression, MS436 inhibits BRD4 activity, reducing the levels of NO and IL-6.
Kinase Assay	Fluorescence Anisotropy Binding Assay: Binding affinity of the newly synthesized diazobenzene compounds for various bromodomains is assessed in a fluorescence anisotropy competition assay using a fluorescein isothiocyanate (FITC)-labeled MS417 as an assay probe. Competition experiments are performed with a BrD protein (0.25-1 μM) and the fluorescent probe (80 nM), and increasing concentration of unlabeled competing ligand in a PBS buffer (pH 7.4) in total volume of 80 μL. Measurements are obtained after a 1 hour incubation of the fluorescent ligand and the protein at 25°C with Safire 2 microplate reader. In a competition-binding assay, fluorescent ligand concentration is ≤ 2Kd, and protein concentration was set at which 50-80% of fluorescent ligand is bound. Dissociation constant of a competing ligand is calculated with the correction to Cheng-Prussoff equation introduced by Nicolovska-Coleska and colleagues. Assuming one-site competitive binding model, the equation used to calculate K _i 's from IC ₅₀ values recovered from fitting data using Prism.
Cell Research	Murine macrophage RAW264.7 cells are plated at a density of 10000 cells per well in a 96-well plate and incubated at 37 °C for 18 h. The cells are then treated with the diazobenzene bromodomain inhibitors up to 100 μL for 24 hours. At the end of the 24 hr incubation, 10 μL of the MTT solution (4 mg/ml) is added to each well and incubated at 37°C for 4 h. The supernatants are then removed and the cells were solubilized in 100 μL of 100% DMSO. The diazobenzene compounds are first dissolved in DMSO then diluted with culture medium to concentrations that ranged from 0.28 to 50000 nM. The final concentration of DMSO is adjusted to 0.05% (v/v). The extent of the reduction is measured by the absorbance at 570/630 nm using EnVison 2104 Multilabel Reader.(Only for Reference)

Solubility Information

A DRUG SCREENING EXPERT

Solubility	DMSO: 40 mg/mL (104.32 mM),Sonication is recommended. (< 1 mg/ml refers to the product slightly soluble or insoluble)
In vivo Formulation	10% DMSO+40% PEG300+5% Tween 80+45% Saline: 1 mg/mL (2.61 mM),Sonication is recommended. <i>Please add the solvents sequentially, clarifying the solution as much as possible before adding the next one. Dissolve by heating and/or sonication if necessary. Working solution is recommended to be prepared and used immediately. The formulation provided above is for reference purposes only. In vivo formulations may vary and should be modified based on specific experimental conditions.</i>

Preparing Stock Solutions

	1mg	5mg	10mg
1 mM	2.6081 mL	13.0405 mL	26.0811 mL
5 mM	0.5216 mL	2.6081 mL	5.2162 mL
10 mM	0.2608 mL	1.3041 mL	2.6081 mL
50 mM	0.0522 mL	0.2608 mL	0.5216 mL

Please select the appropriate solvent to prepare the stock solution, according to the solubility of the product in different solvents. Please use it as soon as possible.

Note: The dilution table applies only to solid products. For liquid products, please calculate the stock solution based on the stated concentration and/or density.

Reference

Zhang G, et al. J Med Chem. 2013, 56(22), 9251-9264.

Inhibitor · Natural Compounds · Compound Libraries · Recombinant Proteins

This product is for Research Use Only· Not for Human or Veterinary or Therapeutic Use

Tel:781-999-4286 E_mail:info@targetmol.com Address:34 Washington Street,Wellesley Hills,MA 02481