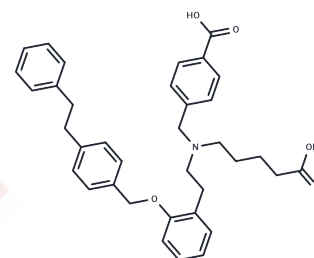


Cinaciguat

Chemical Properties

CAS No. :	329773-35-5
Formula:	C36H39NO5
Molecular Weight:	565.7
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	Cinaciguat (BAY 58-2667) (BAY 58-2667) is the first of a new class of soluble guanylate cyclase (sGC) activators in Clinicalal development for acute decompensated heart failure.
Targets(IC50)	Guanylate cyclase
In vitro	Cinaciguat (10 μM) significantly enhances intracellular cGMP generation without exhibiting dose-dependent effects on cell contraction and calcium transients[2].
In vivo	Cinaciguat (10 mg/kg/day, p.o.) treatment in diabetic rats does not affect blood glucose levels but reduces water intake, alleviates diabetes mellitus-related oxidative stress, protects against alterations in the NO-sGC-cGMP-PKG signaling pathway, and reduces myocardium hypertrophy and apoptosis[1]. Cinaciguat (1-10-100 nM) induces concentration-dependent relaxations in strips from both WT and apo-sGC mice, but does not influence phasic activity induced by PGF2α in WT or apo-sGC strips[3].
Kinase Assay	HDAC Inhibition Assays: Purified HDACs are incubated with 1 mM carboxyfluorescein (FAM)-labeled acetylated peptide substrate and test compound for 17 h at 25 °C in HDAC assay buffer containing 100 mM HEPES (pH 7.5), 25 mM KCl, 0.1% BSA, and 0.01% Triton X-100. Reactions are terminated by the addition of buffer containing 0.078% SDS for a final SDS concentration of 0.05%. Substrate and product are separated electrophoretically using a Caliper LabChip 3000 system with blue laser excitation and green fluorescence detection (CCD2). The fluorescence intensity in the substrate and product peaks is determined using the Well Analyzer software on the Caliper system. The reactions are performed in duplicate for each sample. IC50 values are automatically calculated using the IDBS XLFit version 4.2.1 plug-in for Microsoft Excel and the XLFit 4-Parameter Logistic Model: $((A+((B-A)/(1+((C/x)^D)))))$, in which x is compound concentration, A and B are respectively the estimated minimum and maximum of percent inhibition, C is the inflection point, and D is the Hill slope of the sigmoidal curve. The standard errors of the IC50 values are automatically calculated using the IDBS XLFit version 4.2.1 plug-in for Microsoft Excel and the formula $xf4_FitResultStdError$.

Solubility Information

A DRUG SCREENING EXPERT

Solubility	DMSO: 62.5 mg/mL (110.48 mM),Sonication is recommended. (< 1 mg/ml refers to the product slightly soluble or insoluble)
In vivo Formulation	10% DMSO+90% Saline: 6.25 mg/mL (11.05 mM),Suspension. 10% DMSO+90% Corn Oil: 2 mg/mL (3.54 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	1.7677 mL	8.8386 mL	17.6772 mL
5 mM	0.3535 mL	1.7677 mL	3.5354 mL
10 mM	0.1768 mL	0.8839 mL	1.7677 mL
50 mM	0.0354 mL	0.1768 mL	0.3535 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

The soluble guanylate cyclase activator cinaciguat prevents cardiac dysfunction in a rat model of type-1 diabetes mellitus[J]. Cardiovascular Diabetology, 2015, 14(1):145.

Liu R, Kang Y, Chen L. Activation mechanism of human soluble guanylate cyclase by stimulators and activators. Nature Communications. 2021, 12(1): 1-10.

Reinke Y, et al. The soluble guanylate cyclase stimulator riociguat and the soluble guanylate cyclase activator cinaciguat exert no direct effects on contractility and relaxation of cardiac myocytes from normal rats. Format: AbstractSend to Eur J Pharmaco

Cosyns SM, et al. Influence of cinaciguat on gastrointestinal motility in apo-sGC mice. Neurogastroenterol Motil. 2014 Nov;26(11):1573-85.

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