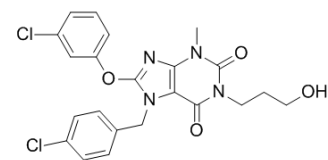


HC-070

Cat. No.:	HY-112302		
CAS No.:	1628291-95-1		
Molecular Formula:	C ₂₂ H ₂₀ Cl ₂ N ₄ O ₄		
Molecular Weight:	475.32		
Target:	TRP Channel		
Pathway:	Membrane Transporter/Ion Channel; Neuronal Signaling		
Storage:	Powder	-20°C	3 years
		4°C	2 years
	In solvent	-80°C	6 months
		-20°C	1 month



SOLVENT & SOLUBILITY

In Vitro

DMSO : ≥ 62.5 mg/mL (131.49 mM)
 * "≥" means soluble, but saturation unknown.

Preparing Stock Solutions	Solvent Concentration	Mass		
		1 mg	5 mg	10 mg
	1 mM	2.1038 mL	10.5192 mL	21.0385 mL
	5 mM	0.4208 mL	2.1038 mL	4.2077 mL
	10 mM	0.2104 mL	1.0519 mL	2.1038 mL

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

In Vivo

- Add each solvent one by one: 10% DMSO >> 40% PEG300 >> 5% Tween-80 >> 45% saline
 Solubility: ≥ 2.5 mg/mL (5.26 mM); Clear solution
- Add each solvent one by one: 10% DMSO >> 90% corn oil
 Solubility: ≥ 2.5 mg/mL (5.26 mM); Clear solution

BIOLOGICAL ACTIVITY

Description

HC-070 is an antagonist of TRPC4/TRPC5, with IC₅₀s of 9.3 nM and 46 nM for hTRPC5 and hTRPC4 in cells, respectively.

IC₅₀ & Target

IC₅₀: 9.3 nM (hTRPC5, cell assay), 46 nM (hTRPC4, cell assay)^[1]

In Vitro

HC-070 is an antagonist of TRPC4/TRPC5, with IC₅₀s of 9.3 nM and 46 nM for hTRPC5 and hTRPC4, respectively. HC-070 weakly inhibits TRPC3 (IC₅₀, 1 μM), and is at least 400-fold selective for human TRPC4 and TRPC5-containing channels versus the other channels examined. HC-070 inhibits lanthanum-activated hTRPC5-, mTRPC5-, rTRPC5-mediated currents with IC₅₀s of 0.52 nM, 0.55 nM, and 0.32 nM in whole-cell manual patch clamp. Furthermore, HC-070 blocks M2R-activated human TRPC1/TRPC4 channels with an IC₅₀ of 1.3 nM and La³⁺- and M1R-activated human TRPC1/5 channels with IC₅₀s of 1.4 nM

and 4.4 nM. HC-070 inhibits human TRPC5 currents activated via muscarinic type 1 (M1R) with an IC₅₀ of 2.0 nM. HC-070 also suppresses hTRPC4 currents via M2R with an IC₅₀ of 0.49 nM. HC-070 (20 nM) reduces CCK-4 evoked neuronal activity in the amygdala slices^[1].

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

In Vivo

HC-070 (1 mg/kg, p.o.) affects mice with increased evoked anxiety (CCK-4), but shows no effects in the absence of CCK-4. HC-070 (0.3, 1 or 3 mg/kg, p.o.) decreases anxiety in a standard EPM (more light/high anxiety). HC-070 (1 mg/kg) reduces the increased capacity for fear memory in mice subjected to chronic social stress on days 1-15. In addition, HC-070 (1, 3, 10 mg/kg, p.o.) causes reduction in marble burying behavior. HC-070 (0.3, 1, 3, 10 mg/kg, p.o.) also reduces time of immobility in a tail suspension test but does not impact locomotor activity in mice^[1].

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PROTOCOL

Animal Administration ^[1]

Mice^[1]

The room is illuminated with fluorescent lighting on a 12-hour light/dark cycle. The light cycle is reversed, so that the dark cycle is from 6 am-6 pm daily and studies are performed when animals are more active. Groups of male C57/BL6 mice (10 weeks old) are dosed PO with 0.5% methyl cellulose or HC-070 at 0.3, 1 or 3 mg/kg (n = 10). The positive control, 1.5 mg/kg diazepam, is administered IP 30 minutes prior to testing (n = 10). Immediately following dosing, mice are returned to their home cage. At 60 minutes post vehicle or HC-070 administration, and 30 minutes post diazepam administration, mice are placed onto the elevated plus maze, one at a time, and their session recorded for 5 minutes. Videos are manually scored for number of open arm entries by a scorer blinded to treatment. All animals that fall off the maze during the test are removed from analysis^[1].

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

CUSTOMER VALIDATION

- Sci Transl Med. 2021 May 26;13(595):eabd7702.
- bioRxiv. 2020 Jul.

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REFERENCES

[1]. Just S, et al. Treatment with HC-070, a potent inhibitor of TRPC4 and TRPC5, leads to anxiolytic and antidepressant effects in mice. PLoS One. 2018 Jan 31;13(1):e0191225.

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

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