ANA-12

Cat. No.: HY-12497 CAS No.: 219766-25-3 Molecular Formula: $C_{22}H_{21}N_3O_3S$ Molecular Weight: 407.49 Target: Trk Receptor

Pathway: Neuronal Signaling; Protein Tyrosine Kinase/RTK

-20°C Storage: Powder 3 years

4°C 2 years -80°C In solvent 1 year

> -20°C 6 months

Product Data Sheet

SOLVENT & SOLUBILITY

In Vitro

DMSO: 14.29 mg/mL (35.07 mM; ultrasonic and warming and heat to 60°C)

Preparing Stock Solutions	Solvent Mass Concentration	1 mg	5 mg	10 mg
	1 mM	2.4540 mL	12.2702 mL	24.5405 mL
	5 mM	0.4908 mL	2.4540 mL	4.9081 mL
	10 mM	0.2454 mL	1.2270 mL	2.4540 mL

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

In Vivo

- 1. Add each solvent one by one: 10% DMSO >> 40% PEG300 >> 5% Tween-80 >> 45% saline Solubility: 1.43 mg/mL (3.51 mM); Suspended solution; Need ultrasonic
- 2. Add each solvent one by one: 10% DMSO >> 90% corn oil Solubility: ≥ 1.43 mg/mL (3.51 mM); Clear solution
- 3. Add each solvent one by one: 5% DMSO >> 40% PEG300 >> 5% Tween-80 >> 50% saline Solubility: 1 mg/mL (2.45 mM); Suspended solution; Need ultrasonic
- 4. Add each solvent one by one: 5% DMSO >> 95% (20% SBE-β-CD in saline) Solubility: ≥ 0.45 mg/mL (1.10 mM); Clear solution
- 5. Add each solvent one by one: 5% DMSO >> 95% corn oil Solubility: ≥ 0.45 mg/mL (1.10 mM); Clear solution

BIOLOGICAL ACTIVITY

Description

ANA-12 is a potent and selective TrkB antagonist with IC₅₀s of 45.6 nM and 41.1 μM for the high and low affinity sites, respectively.

IC ₅₀ & Target	TrkB
In Vitro	ANA-12 (10 nM) prevents BDNF-induced neurite outgrowth in the TrkB-expressing cells, and completely abolishes the effects of BDNF at concentrations up to 10-100 μ M[1]. MCE has not independently confirmed the accuracy of these methods. They are for reference only.
In Vivo	ANA-12 (0.5 mg/kg, i.p.) partially inhibits the total endogenous TrkB activity in the whole brain of mice. ANA-12, injected in mice, demonstrates anxiolytic and antidepressive activities at 0.5 mg/kg. ANA-12 (0.5, 1.0, and 2.0 mg/kg) does not affect neuron survival ^[1] . ANA-12 (0.5 mg/kg) shows antidepressant effects in lipopolysaccharide (LPS)-induced depression-like behavior. ANA-12 (0.5 mg/kg) significantly attenuates an increased immobility time in depressed mice. In the TST, FST, and SPT, ANA-12 (0.5 mg/kg) does not show antidepressant-like effects in the control mice ^[2] . ANA-12 (0.5 mg/kg, i.p.) reverses the diminished self-administration of cocaine in CocSired rats ^[3] . MCE has not independently confirmed the accuracy of these methods. They are for reference only.

PROTOCOL

Animal
Administration [2]

On the day of injection, ketamine (ketamine hydrochloride, 10 mg/kg), 7,8-dihydroxyflavone (7,8-DHF; 10 mg/kg), and ANA-12, N2-(2-{[(2-oxoazepan-3-yl) amino]carbonyl}phenyl)benzo[b]thiophene-2-carboxamide (0.5 mg/kg) are prepared in vehicle of 17 % dimethyl sulfoxide (DMSO) in phosphate-buffered saline. The doses of ketamine (10 mg/kg), 7,8-DHF (10 mg/kg), and ANA-12 (0.5 mg/kg) are selected. All compounds are administered intraperitoneally (i.p.) to mice.

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

CUSTOMER VALIDATION

- Cell Metab. 2022 Nov 11;S1550-4131(22)00490-9.
- Nat Commun. 2023 Nov 16;14(1):7406.
- J Neuroinflammation. 2021 Aug 23;18(1):184.
- J Neuroinflammation. 2020 Jan 13;17(1):19.
- Transl Psychiatry. 2023 May 24;13(1):173.

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REFERENCES

- $[1]. \ Cazorla\ M,\ et\ al.\ Identification\ of\ a\ low-molecular\ weight\ TrkB\ antagonist\ with\ anxiolytic\ and\ antidepressant\ activity\ in\ mice.\ J\ Clin\ Invest.\ 2011\ May; 121(5):1846-57.$
- [2]. Zhang JC, et al. Comparison of ketamine, 7,8-dihydroxyflavone, and ANA-12 antidepressant effects in the social defeat stress model of depression. Psychopharmacology (Berl). 2015 Dec;232(23):4325-35.
- [3]. Vassoler FM, et al. Epigenetic inheritance of a cocaine-resistance phenotype. Nat Neurosci. 2013 Jan;16(1):42-7.
- [4]. Fang X, et al. Brain-derived neurotrophic factor-TrkB signaling in the medial prefrontal cortex plays a role in the anhedonia-like phenotype after spared nerve injury. Eur Arch Psychiatry Clin Neurosci. 2018 Jun 7.

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 $\label{lem:caution:Product} \textbf{Caution: Product has not been fully validated for medical applications. For research use only.}$

Tel: 609-228-6898 Fax: 609-228-5909

E-mail: tech@MedChemExpress.com

Address: 1 Deer Park Dr, Suite Q, Monmouth Junction, NJ 08852, USA

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