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Zuschläge

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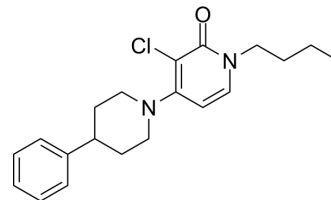
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JNJ-40411813

Cat. No.:	HY-15748
CAS No.:	1127498-03-6
Molecular Formula:	C ₂₀ H ₂₅ ClN ₂ O
Molecular Weight:	344.88
Target:	mGluR
Pathway:	GPCR/G Protein; Neuronal Signaling
Storage:	Powder -20°C 3 years 4°C 2 years In solvent -80°C 2 years -20°C 1 year



SOLVENT & SOLUBILITY

In Vitro	DMSO : 25 mg/mL (72.49 mM; Need ultrasonic)					
	Preparing Stock Solutions	Solvent Concentration	Mass	1 mg	5 mg	10 mg
		1 mM		2.8996 mL	14.4978 mL	28.9956 mL
		5 mM		0.5799 mL	2.8996 mL	5.7991 mL
		10 mM		0.2900 mL	1.4498 mL	2.8996 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: 2.5 mg/mL (7.25 mM); Suspended solution; Need ultrasonic and warming					
	2. Add each solvent one by one: 10% DMSO >> 90% (20% SBE-β-CD in saline) Solubility: ≥ 2.5 mg/mL (7.25 mM); Clear solution					
	3. Add each solvent one by one: 10% DMSO >> 90% corn oil Solubility: 2.5 mg/mL (7.25 mM); Clear solution; Need ultrasonic					

BIOLOGICAL ACTIVITY

Description	JNJ-40411813 (ADX-71149) is a novel positive allosteric modulator of the metabotropic Glutamate 2 receptor (mGlu2R) with EC ₅₀ of 147 nM. JNJ-40411813 has orally bioactivity and penetrate the blood-brain barriers. JNJ-40411813 has the potential property of anti-depression ^{[1][2][3]} .
IC ₅₀ & Target	mGlu2 Receptor 147 nM (EC50)

In Vivo

JNJ-40411813 (ADX-71149) (5-20 mg/kg; p.o.; 6 times in 24 h) clearly potentiated glutamate response in brain regions, more specifically in the cortical regions, striatum, and the hippocampus, coaddition of Glutamate (HY-N0455B)^[2].

Pharmacokinetic Analysis in SD rat^[2]

Route	Dose (mg/kg)	AUC _{INF_obs} (ng·h/mL)	t _{1/2} (h)	T _{max} (h)	C _{max} (ng/mL)	Cl _{obs} (L·h/kg)	V _{ss_obs} (L/kg)
i.v.	2.5	1833±90	/	/	/	1.4±0.1	2.31.4±0.10.2
p.o.	10	2250±417	2.31±0.5	7	938	/	/

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

Animal Model:	Male Wistar Rats (150 mg) ^[2]
Dosage:	5 mg/kg, 20 mg/kg
Administration:	Oral Gavage (p.o.)
Result:	Showed increasing in basal binding with an EC ₅₀ of 48 µM and a Combined with 10 µM glutamate, [³⁵ S]GTPγS binding with an EC ₅₀ of 7.3 µM and a maximal effect of 380%.

REFERENCES

- [1]. Hilde Lavreysen, et al. Pharmacological and pharmacokinetic properties of JNJ-40411813, a positive allosteric modulator of the mGlu2 receptor. *Pharmacol Res Perspect* 2015 Feb;3(1):e00096.
- [2]. Dunlop J, Schizophrenia, et al. Schizophrenia drug discovery and development in an evolving era: are new drug targets fulfilling expectations?. *Journal of Psychopharmacology*. 29 (2): 230–8.
- [3]. Cid JM, et al. Discovery of 1-Butyl-3-chloro-4-(4-phenyl-1-piperidinyl)-(1H)-pyridone (JNJ-40411813): A Novel Positive Allosteric Modulator of the Metabotropic Glutamate 2 Receptor. *J Med Chem*. 2014 Aug 14;57(15):6495-512.

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

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