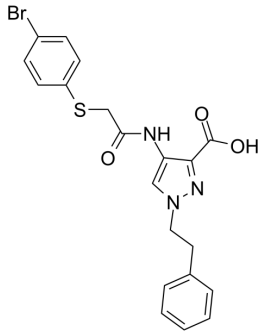
KR-33493

Cat. No.:	HY-100755
CAS No.:	1021497-97-1
Molecular Formula:	$C_{20}H_{18}BrN_3O_3S$
Molecular Weight:	460.34
Target:	TNF Receptor
Pathway:	Apoptosis
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 : ≥ 31 mg/mL (67.34 mM)

* "≥" means soluble, but saturation unknown.

Preparing Stock Solutions	Solvent / Mass / Concentration	1 mg	5 mg	10 mg
	1 mM	2.1723 mL	10.8615 mL	21.7231 mL
	5 mM	0.4345 mL	2.1723 mL	4.3446 mL
	10 mM	0.2172 mL	1.0862 mL	2.1723 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 (5.43 mM); Clear solution
2. Add each solvent one by one: 10% DMSO >> 90% corn oil
   Solubility: ≥ 2.5 mg/mL (5.43 mM); Clear solution

BIOLOGICAL ACTIVITY

Description	KR-33493 is a potent inhibitor of Fas-mediated cell death (FAF1).
IC_{50} & Target	Fas
In Vivo	Body weight changes of both sexes are not related to KR-33493 in all doses. In rats administrated KR-33493 for 4 weeks, no test article-related changes in any treated groups of either sex are found in hematology, serum biochemistry, and urinalysis. In dogs administrated KR-33493 for 2 weeks, red blood cell count (RBC) value in males is significantly higher at the 1000 mg/kg/day dose than that of the control group (i.e., 6.96±0.323 vs. 6.12±0.418). However, the change of RBC is recovered after the end of the administration period. The dose-normalized AUC_{last} is not significantly different between the groups,

suggesting that KR-33493 is governed by linear kinetics[1].

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

PROTOCOL

Animal Administration [1]

A total of 93 male and 93 female specific pathogen-free rats (6 weeks of age), and 16 male and 16 female beagle dogs (8 months of age) are used in this study. In a toxicokinetic study, rat blood samples (approximately 0.6 mL) are collected into tubes containing heparin from the lateral tail vein at 0, 0.5, 1, 2, 4, 8, 12, and 24 h after dosing with KR-33493 at doses of 50, 150, and 500 mg/kg/day on Day 1 and Week 4. Dog blood samples (approximately 0.6 mL) are collected into tubes containing EDTA-2K from the cephalic vein at 0, 0.5, 1, 2, 4, 6, 8, and 24 h after dosing at KR-33493 doses of 50, 250, and 1000 mg/kg/day on Day 1 and Week 2. The plasma is separated by centrifugation (approximately 132,000 g, 3 min, 4°C) and stored at approximately -80°C until analysis. The KR-33493 concentration in plasma is quantified[1].

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

REFERENCES

[1]. Jeong JW, et al. Subacute toxicity evaluation of KR-33493, FAF1 inhibitor for a new anti-parkinson's disease agent, after oral administration in rats and dogs. Regul Toxicol Pharmacol. 2016 Nov;81:387-396.

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