Semapimod tetrahydrochloride

Cat. No.: HY-15509A

CAS No.: 164301-51-3

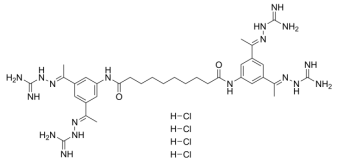
Molecular Formula: $C_{34}H_{56}Cl_4N_{18}O_2$

Molecular Weight: 890.74

Target: p38 MAPK; Interleukin Related; TNF Receptor

Pathway: MAPK/ERK Pathway; Immunology/Inflammation; Apoptosis

Storage: -20°C, sealed storage, away from moisture and light
* In solvent : -80°C, 6 months; -20°C, 1 month (sealed storage, away from moisture and light)



SOLVENT & SOLUBILITY

In Vitro

H_2O : 2.17 mg/mL (2.44 mM; ultrasonic and warming and heat to 60°C)

Preparing Stock Solutions	Mass / Solvent / Concentration	1 mg	5 mg	10 mg
	1 mM	1.1227 mL	5.6133 mL	11.2266 mL
	5 mM	---	---	---
	10 mM	---	---	---

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

BIOLOGICAL ACTIVITY

Description

Semapimod tetrahydrochloride (CNI-1493), an inhibitor of proinflammatory cytokine production, can inhibit TNF-α, IL-1β, and IL-6. Semapimod tetrahydrochloride inhibits TLR4 signaling (IC_{50}≈0.3 μM). Semapimod tetrahydrochloride inhibits p38 MAPK and nitric oxide production in macrophages. Semapimod tetrahydrochloride has potential in a variety of inflammatory and autoimmune disorders[1][2][3].

IC_{50} & Target	IL-1β	IL-6	p38 MAPK

In Vitro

Semapimod tetrahydrochloride leads to a significant decrease of p38-MAPK phosphorylation in macrophages, proinflammatory gene expression of macrophage inflammatory protein-1alpha, interleukin-6, monocyte chemoattractant protein-1, and intercellular adhesion molecule-1, and neutrophil infiltration. Semapimod tetrahydrochloride completely abrogated nitric oxide production within the tunica muscularis[2].

?Semapimod tetrahydrochloride desensitizes TLR signaling via its effect on the TLR chaperone gp96. Semapimod tetrahydrochloride inhibits ATP-binding and ATPase activities of gp96 in vitro (IC_{50}≈0.2-0.4 μM). Semapimod tetrahydrochloride desensitizes TLR signaling via its effect on the TLR chaperone gp96[3].

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

In Vivo

Semapimod tetrahydrochloride (5 mg/kg; i.p; daily for 2 weeks) ameliorates endothelial dysfunction in Obese Zucker (OZ) rats[1].

?Semapimod tetrahydrochloride restores AM-induced akt phosphorylation and cGMP production in OZ rats[1].

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

Animal Model:	Male OZ rats[1]
Dosage:	5 mg/kg
Administration:	I.p; daily for 2 weeks
Result:	Restored endothelium-dependent vasorelaxation in OZ rats.

REFERENCES

[1]. Wehner S, Set al. Inhibition of p38 mitogen-activated protein kinase pathway as prophylaxis of postoperative ileus in mice. Gastroenterology. 2009;136(2):619-629.

[2]. Nishimatsu H, et al. Blockade of endogenous proinflammatory cytokines ameliorates endothelial dysfunction in obese Zucker rats. Hypertens Res. 2008;31(4):737-743.

[3]. Wang J, et al. Experimental Anti-Inflammatory Drug Semapimod Inhibits TLR Signaling by Targeting the TLR Chaperone gp96. J Immunol. 2016;196(12):5130-5137.

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