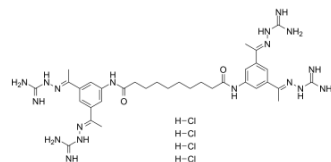


Semapimod tetrahydrochloride

Cat. No.:	HY-15509A
CAS No.:	164301-51-3
Molecular Formula:	C ₃₄ H ₅₆ Cl ₄ N ₁₈ O ₂
Molecular Weight:	890.74
Target:	TNF Receptor; Interleukin Related; p38 MAPK
Pathway:	Apoptosis; Immunology/Inflammation; MAPK/ERK Pathway
Storage:	-20°C, protect from light * In solvent : -80°C, 6 months; -20°C, 1 month (protect from light)



BIOLOGICAL ACTIVITY

Description	Semapimod tetrahydrochloride (CNI-1493), an inhibitor of proinflammatory cytokine production, can inhibit TNF- α , IL-1 β , and IL-6. Semapimod tetrahydrochloride inhibits TLR4 signaling (IC ₅₀ ≈0.3 μ M). 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]} .											
IC₅₀ & Target	TNF- α	IL-1 β	IL-6	p38 MAPK								
In Vitro	<p>Semapimod tetrahydrochloride leads to a significant decrease of p38-MAPK phosphorylation in macrophages, proinflammatory gene expression of macrophage inflammatory protein-1α, 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].</p> <p>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 (IC₅₀≈0.2-0.4 μM). Semapimod tetrahydrochloride desensitizes TLR signaling via its effect on the TLR chaperone gp96^[3].</p> <p>MCE has not independently confirmed the accuracy of these methods. They are for reference only.</p>											
In Vivo	<p>Semapimod tetrahydrochloride (5 mg/kg; i.p; daily for 2 weeks) ameliorates endothelial dysfunction in Obese Zucker (OZ) rats^[1].</p> <p>Semapimod tetrahydrochloride restores AM-induced akt phosphorylation and cGMP production in OZ rats^[1].</p> <p>MCE has not independently confirmed the accuracy of these methods. They are for reference only.</p> <table border="1" style="width: 100%; border-collapse: collapse;"> <tr> <td style="width: 30%;">Animal Model:</td> <td>Male OZ rats^[1]</td> </tr> <tr> <td>Dosage:</td> <td>5 mg/kg</td> </tr> <tr> <td>Administration:</td> <td>i.p; daily for 2 weeks</td> </tr> <tr> <td>Result:</td> <td>Restored endothelium-dependent vasorelaxation in OZ rats.</td> </tr> </table>				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.
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Dosage:	5 mg/kg											
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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.

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

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