**Proteins** 



# Geranylgeraniol

Cat. No.: HY-W011474 CAS No.: 24034-73-9 Molecular Formula:  $C_{20}H_{34}O$ Molecular Weight: 290.48 Target: NF-κB

Storage: Pure form -20°C 3 years

NF-κΒ

2 years

In solvent -80°C 6 months

> -20°C 1 month

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**Product** Data Sheet

# **SOLVENT & SOLUBILITY**

In Vitro

Pathway:

DMSO: 100 mg/mL (344.26 mM; Need ultrasonic)

	Solvent Mass Concentration	1 mg	5 mg	10 mg
Preparing Stock Solutions	1 mM	3.4426 mL	17.2129 mL	34.4258 mL
	5 mM	0.6885 mL	3.4426 mL	6.8852 mL
	10 mM	0.3443 mL	1.7213 mL	3.4426 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 (8.61 mM); Clear solution
- 2. Add each solvent one by one: 10% DMSO >> 90% (20% SBE- $\beta$ -CD in saline) Solubility: ≥ 2.5 mg/mL (8.61 mM); Suspended solution
- 3. Add each solvent one by one: 10% DMSO >> 90% corn oil Solubility: ≥ 2.5 mg/mL (8.61 mM); Clear solution

# **BIOLOGICAL ACTIVITY**

Description	Geranylgeraniol is an orally acitve vitamin $K_2$ sub-type, an intermediate of the mevalonate pathway. Geranylgeraniol targets NF-kB signaling pathway and could alleviate LPS-induced microglial inflammation in animal model <sup>[1][2][3][4]</sup> .
IC <sub>50</sub> & Target	$NF ext{-}kB^{[1]}$
In Vitro	Geranylgeraniol (0-10 $\mu$ M; 24 h) dose-dependently suppresses the LPS-induced increase in the mRNA levels of Il-1 $\beta$ , Tnf- $\alpha$ , Il-6, and Cox-2 <sup>[1]</sup> .

Geranylgeraniol (10  $\mu$ M; 24 h) inhibits the phosphorylation of TAK1, IKK $\alpha/\beta$ , and NF- $\kappa$ B p65 proteins as well as NF- $\kappa$ B nuclear translocation induced by LPS while maintaining I $\kappa$ B $\alpha$  expression<sup>[1]</sup>.

Geranylgeraniol, (50  $\mu$ M; 24 h) eliminates cell damage caused by <u>Simvastatin</u> (HY-17502) (10  $\mu$ M) and Mevalonat (10 mM), and reduces the inflammatory marker and the damage of the mitochondria, maintaining its shape and component<sup>[2]</sup>. MCE has not independently confirmed the accuracy of these methods. They are for reference only.

Western Blot Analysis  $^{[1]}$ 

Cell Line:	MG6 cell	
Concentration:	0, 1, 10 μΜ	
Incubation Time:	24 hours	
Result:	Suppressed by TAK1, IKKα/β, and NF-κB p65 proteins level at 10 μM.	
RT-PCR <sup>[1]</sup>		
Cell Line:	MG6 cell	
Concentration:	10 μΜ	
Incubation Time:	0, 6, 12, 24 hours	
Result:	Significantly inhibited pro-inflammatory cytokine Il-1 $\beta$ , Tnf- $\alpha$ , Il-6, and Cox-2 mRNA level.	

### In Vivo

Geranylgeraniol (725 mg/kg/d; p.o.; 90 d) is not toxicologically significant with a dose below 725 mg/kg/d in rats<sup>[3]</sup>. Geranylgeraniol (483 mg/kg/d; p.o.; 10 d) suppresses lipopolysaccharide-induced inflammation via inhibition of nuclear factor- $\kappa$ B activation in rats<sup>[4]</sup>.

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

Animal Model:	Han Wistar rats (169-192 g for male; 116-152 g for female) $^{\left[3\right]}$
Dosage:	0, 725, 1450, and 2900 mg/kg
Administration:	Oral gavage; once daily; 90 days
Result:	Showed the lowest observed adverse effect level (LOAEL) for local effects and the no observed adverse effect level (NOAEL) for systemic effects as 725 mg/kg/d.  Reduced body weights by 12.9 and 21.6% in the intermediate- and high-dose group males, respectively, compared to controls.

Animal Model:	Wistar rats (male, 8-week-old, 130-150 g) <sup>[4]</sup>
Dosage:	0, 48.3, 483, 4830 mg/kg
Administration:	Oral gavage; once daily; 10 days; with or not LPS challenge (i.p.; 0.5 mg/kg)
Result:	Suppressed LPS-induced inflammatory cytokines and mRNA expression of LPS-induced inflammatory genes in liver with doses of 483 mg/kg and 4830 mg/kg.  Suppressed protein levels of IRAK1, TRAF6, and TAK1, originating from transcriptional down-regulation with doses of 483 mg/kg and 4830 mg/kg.

## **REFERENCES**

- [1]. Saputra WD, et al. Geranylgeraniol Inhibits Lipopolysaccharide-Induced Inflammation in Mouse-Derived MG6 Microglial Cells via NF-kB Signaling Modulation. Int J Mol Sci. 2021 Sep 29;22(19):10543.
- [2]. Marcuzzi A, et al. Geranylgeraniol and Neurological Impairment: Involvement of Apoptosis and Mitochondrial Morphology. Int J Mol Sci. 2016 Mar 11;17(3):365.
- [3]. Preece K, et al. A toxicological evaluation of geranylgeraniol. Regul Toxicol Pharmacol. 2021 Aug;124:104975.
- [4]. Giriwono PE, et al. Dietary supplementation with geranylgeraniol suppresses lipopolysaccharide-induced inflammation via inhibition of nuclear factor-κB activation in rats. Eur J Nutr. 2013 Apr;52(3):1191-9.

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

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