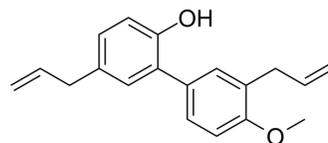


## 4-O-Methyl honokiol

|                           |   |           |       |         |  |     |         |            |       |          |  |       |         |
|---------------------------|---|-----------|-------|---------|--|-----|---------|------------|-------|----------|--|-------|---------|
| <b>Cat. No.:</b>          | HY-U00450   |           |       |         |  |     |         |            |       |          |  |       |         |
| <b>CAS No.:</b>           | 68592-15-4  |           |       |         |  |     |         |            |       |          |  |       |         |
| <b>Molecular Formula:</b> | C <sub>19</sub> H <sub>20</sub> O <sub>2</sub>  |           |       |         |  |     |         |            |       |          |  |       |         |
| <b>Molecular Weight:</b>  | 280.36  |           |       |         |  |     |         |            |       |          |  |       |         |
| <b>Target:</b>            | PPAR $\gamma$ ; NF- $\kappa$ B  |           |       |         |  |     |         |            |       |          |  |       |         |
| <b>Pathway:</b>           | Cell Cycle/DNA Damage; Vitamin D Related/Nuclear Receptor; NF- $\kappa$ B   |           |       |         |  |     |         |            |       |          |  |       |         |
| <b>Storage:</b>           | <table border="0"> <tr> <td>Pure form</td> <td>-20°C</td> <td>3 years</td> </tr> <tr> <td></td> <td>4°C</td> <td>2 years</td> </tr> <tr> <td>In solvent</td> <td>-80°C</td> <td>6 months</td> </tr> <tr> <td></td> <td>-20°C</td> <td>1 month</td> </tr> </table> | Pure form | -20°C | 3 years |  | 4°C | 2 years | In solvent | -80°C | 6 months |  | -20°C | 1 month |
| Pure form                 | -20°C   | 3 years   |       |         |  |     |         |            |       |          |  |       |         |
|                           | 4°C   | 2 years   |       |         |  |     |         |            |       |          |  |       |         |
| In solvent                | -80°C   | 6 months  |       |         |  |     |         |            |       |          |  |       |         |
|                           | -20°C   | 1 month   |       |         |  |     |         |            |       |          |  |       |         |



### SOLVENT & SOLUBILITY

|   |   |                          |              |            |            |
|---|---|--------------------------|--------------|------------|------------|
| <b>In Vitro</b>   | DMSO : 100 mg/mL (356.68 mM; Need ultrasonic)   |                          |              |            |            |
|   |   | Solvent<br>Concentration | Mass<br>1 mg | 5 mg       | 10 mg      |
|   | <b>Preparing Stock Solutions</b>  | 1 mM                     | 3.5668 mL    | 17.8342 mL | 35.6684 mL |
|   |   | 5 mM                     | 0.7134 mL    | 3.5668 mL  | 7.1337 mL  |
| 10 mM   |   | 0.3567 mL                | 1.7834 mL    | 3.5668 mL  |            |
| Please refer to the solubility information to select the appropriate solvent. |   |                          |              |            |            |
| <b>In Vivo</b>  | <ol style="list-style-type: none"> <li>Add each solvent one by one: 10% DMSO &gt;&gt; 90% (20% SBE-<math>\beta</math>-CD in saline)<br/>Solubility: <math>\geq</math> 2.5 mg/mL (8.92 mM); Clear solution</li> <li>Add each solvent one by one: 10% DMSO &gt;&gt; 90% corn oil<br/>Solubility: <math>\geq</math> 2.5 mg/mL (8.92 mM); Clear solution</li> </ol> |                          |              |            |            |

### BIOLOGICAL ACTIVITY

|                                     |  |                |
|-------------------------------------|--|----------------|
| <b>Description</b>                  | 4-O-Methyl honokiol is a natural neolignan isolated from <i>Magnolia officinalis</i> , acts as a PPAR $\gamma$ agonist, and inhibits NF- $\kappa$ B activity, used for cancer and inflammation research.   |                |
| <b>IC<sub>50</sub> &amp; Target</b> | PPAR $\gamma$  | NF- $\kappa$ B |
| <b>In Vitro</b>                     | 4-O-Methyl honokiol is a natural neolignan isolated from <i>Magnolia officinalis</i> , acts as a PPAR $\gamma$ agonist, and inhibits NF- $\kappa$ B activity. 4-O-Methyl honokiol (20 $\mu$ M) increases the expression, transcription and DNA binding activities, and nuclear translocation of PPAR $\gamma$ in both in prostate PC-3 and LNCaP cells. 4-O-Methyl honokiol (0-30 $\mu$ M) inhibits LNCaP and PC-3 cancer cells growth, causes G0/G1 phase arrest and induces apoptotic cell death, and such effects can be reversed by PPAR $\gamma$ antagonist. 4-O-Methyl honokiol inhibits NF- $\kappa$ B activity and cancer cell growth, but such effects as well as its activation of |                |

PPAR $\gamma$  can be abolished by knock-down of p21<sup>[1]</sup>. 4-O-methylhonokiol (0.5, 1 and 2  $\mu$ M) reduces LPS-induced release of NO, PGE2, ROS, TNF- $\alpha$  and IL-1 $\beta$  in cultured astrocytes, and amyloidogenesis in cultured astrocytes and microglial BV-2 cells<sup>[2]</sup>. MCE has not independently confirmed the accuracy of these methods. They are for reference only.

#### In Vivo

4-O-Methyl honokiol (40 or 80 mg/kg, i.p. everyday for 4 weeks) inhibits the growth of SW620 and PC3 tumours in SW620 and PC3 xenograft model. 4-O-Methyl honokiol significantly increases the expression of p21 and PPAR $\gamma$  in the tumour tissues<sup>[1]</sup>. 4-O-Methyl honokiol (0.5 or 1 mg/kg/day daily for 3 weeks) significantly ameliorates LPS-induced memory impairment, and inhibits LPS-induced iNOS and COX-2 expression in mice. 4-O-Methyl honokiol also shows inhibitory activities against the A $\beta$ <sub>1-42</sub> accumulation, and activates astrocytes and microglia in LPS-injected mice brain<sup>[2]</sup>. MCE has not independently confirmed the accuracy of these methods. They are for reference only.

## PROTOCOL

#### Cell Assay <sup>[1]</sup>

Cells ( $5 \times 10^4$  cells per well) are plated onto 24-well plates. The cell growth inhibitory effect of 4-O-Methyl honokiol is evaluated in cells treated with 4-O-Methyl honokiol (0-30  $\mu$ M) for 0-72 h, using an excluded trypan blue assay<sup>[1]</sup>. MCE has not independently confirmed the accuracy of these methods. They are for reference only.

#### Animal Administration <sup>[1]</sup>

Six-week-old male BALB/c athymic nude mice are used in the assay. SW620 and PC3 cells are injected s.c. ( $1 \times 10^7$  cells in 0.1 mL PBS per animal) into the lower right flanks of mice. After 20 days, when the tumours have reached an average volume of 300-400 mm<sup>3</sup> or about 50 mm<sup>3</sup>, the tumour-bearing nude mice are i.p. injected with 4-O-Methyl honokiol (40 and 80 mg/kg dissolved in 0.1% DMSO) twice per week for 3 weeks. Cisplatin (10 mg/kg) is also i.p. injected once a week as a positive control. The group treated with 0.1% DMSO is designated as the control. The tumour volumes are measured with vernier calipers and calculated by the following formula:  $(A \times B^2)/2$ , where A is the larger and B is the smaller of the two dimensions <sup>[1]</sup>.

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

## REFERENCES

[1]. Lee NJ, et al. 4-O-methylhonokiol, a PPAR $\gamma$  agonist, inhibits prostate tumour growth: p21-mediated suppression of NF- $\kappa$ B activity. Br J Pharmacol. 2013 Mar;168(5):1133-45.

[2]. Lee YJ, et al. Inhibitory effect of 4-O-methylhonokiol on lipopolysaccharide-induced neuroinflammation, amyloidogenesis and memory impairment via inhibition of nuclear factor-kappaB in vitro and in vivo models. J Neuroinflammation. 2012 Feb 19;9:35.

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

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