**ZSET1446**

**Cat. No.:** HY-11013  
**CAS No.:** 887603-94-3  
**Molecular Formula:** C₁₅H₁₂N₂O  
**Molecular Weight:** 236.27  
**Target:** Calcium Channel; nAChR  
**Pathway:** Membrane Transporter/Ion Channel; Neuronal Signaling  
**Storage:**  
- Powder  
  -20°C  3 years  
  4°C  2 years  
- In solvent  
  -80°C  6 months  
  -20°C  1 month

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**SOLVENT & SOLUBILITY**

**In Vitro**  
DMSO : ≥ 34 mg/mL (143.90 mM)  
*“≥” means soluble, but saturation unknown.*

<table>
<thead>
<tr>
<th>Solvent Concentration</th>
<th>Mass for 1 mg</th>
<th>Mass for 5 mg</th>
<th>Mass for 10 mg</th>
</tr>
</thead>
<tbody>
<tr>
<td>1 mM</td>
<td>4.2324 mL</td>
<td>21.1622 mL</td>
<td>42.3245 mL</td>
</tr>
<tr>
<td>5 mM</td>
<td>0.8465 mL</td>
<td>4.2324 mL</td>
<td>8.4649 mL</td>
</tr>
<tr>
<td>10 mM</td>
<td>0.4232 mL</td>
<td>2.1162 mL</td>
<td>4.2324 mL</td>
</tr>
</tbody>
</table>

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

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**BIOLOGICAL ACTIVITY**

**Description**  
ZSET1446 is a novel cognitive enhancer that significantly improves learning deficits in various types of Alzheimer disease (AD) models.

**In Vitro**  
ZSET1446 (100 pM-10 nM) exerts limited effects on the basal neuronal excitability and synaptic transmission. ZSET1446 potentiates the facilitatory effect of nAChR stimulation on sPSC frequency. ZSET1446 potentiates the increase in sIPSC frequency by local application of nicotine (5 µM; 2 minutes) without affecting the basal sIPSC frequency at the range of 10 pM to 1 nM[1].

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In Vivo  
ZSET1446 enhances object recognition memory in mice and ameliorates cognitive impairment caused by scopolamine in rats. Concomitant administration of subeffective doses of ZSET1446 and memantine significantly ameliorates the cognitive performance in the novel object recognition task in both mice and rats. Moreover, oral administration of ZSET1446 or memantine increases the extracellular level of ACh in the hippocampus as compared with the control. Further, concomitant administration of subeffective doses of ZSET1446 and memantine significantly
increases the extracellular level of ACh as compared with the group of ZSET1446 or memantine alone[2]. ZSET1446 (0.002, 0.01, and 0.1 mg/kg, p.o.) ameliorates cognitive deficits of SAMP8 after 4, 8, 12, and 16 weeks of treatment in a novel object recognition test. ZSET1446 also reduces grading scores of SAMP8 after 16 weeks of treatment. Further, 8-week treatment of ZSET1446 significantly reduces the total number of Aβ-positive granules in the hippocampus[3].

PROTOCOL

Animal Administration[2]

The experimental apparatus consists of a Plexiglas open-field box [25 cm (width) × 41 cm (length) × 17 cm (depth), model TP-105], the floor of which is covered with sawdust. The apparatus is located in a sound-attenuated room. The procedure for the novel object recognition task consists of three different sessions: habituation, training, and retention sessions. Each mouse is individually habituated to the box, with 10 min of exploration in the absence of objects (day 1: habituation session). ZSET1446 at doses of 0.001, 0.003, 0.01, and 0.03 mg/kg and/or memantine at doses of 3 and 10 mg/kg is orally administered 60 min before the training trial. In the experiment of injection of nicotinic receptor antagonists, oral administration of ZSET1446 and i.p. injection of mechamylamine or DHbE at each dose of 1 mg/kg are given 60 min before the training trial. During the training session, two different novel objects are symmetrically fixed to the floor in the box, and each animal is allowed to explore in the box for 10 min (day 2: training session). These objects are different in shape and color but similar in size. The mice are considered to be exploring the object when the mouse is facing, touching, or sniffing the object. The time spent for exploring each object is manually measured by a stopwatch. In the training session, locomotor activity is simultaneously measured for a period of 10 min automatically, using an Animex Auto placed under the open-field box. After the training trial, mice are immediately returned to their home cages.

REFERENCES


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