

## CHIC35

Cat. No.: HY-111303

CAS No.: 848193-72-6

Molecular Formula: C<sub>14</sub>H<sub>15</sub>ClN<sub>2</sub>O

Molecular Weight: 262.73

Target: Sirtuin

Pathway: Cell Cycle/DNA Damage; Epigenetics

Storage: Please store the product under the recommended conditions in the Certificate of

Analysis.

**Product** Data Sheet

## **BIOLOGICAL ACTIVITY**

Description CHIC35, an analog of EX-527, is a potent and selective inhibitor of SIRT1 (IC $_{50}$ =0.124  $\mu$ M). CHIC35 shows potential selective inhibition against SIRT1 over SIRT2 (IC $_{50}$ =2.8  $\mu$ M) or SIRT3 (IC $_{50}$ >100  $\mu$ M)<sup>[1]</sup>. CHIC35 has anti-inflammatory effects and can be used for CHARGE syndrome research<sup>[1][2]</sup>.

CHIC-35 (5 µM; 72 hours) exhibits no significant difference in the survival of embryos at early stages<sup>[2]</sup>.

IC<sub>50</sub> & Target SIRT1 SIRT2 SIRT3

0.124  $\mu$ M (IC<sub>50</sub>) 2.8  $\mu$ M (IC<sub>50</sub>) >100  $\mu$ M (IC<sub>50</sub>)

In Vitro CHIC-35 (0.5  $\mu$ M; 16 hours) increases acetylation of histone H4 in BMDMs similar to Cambinol (200  $\mu$ M)<sup>[1]</sup>.

Zebrafish embryos are microinjected with 2.4 ng of chd7 MO to develop to different stages of development. chd7 morphant embryos are treated with CHIC-35 from 8hpf to 24hpf. CHIC-35 ( $5\,\mu\text{M}$ ) is removed at 24hpf and embryos are incubated in fresh egg water until 4dpf. The chd7 morphant larvae has a severely reduced and disrupted pattern of cartilage elements in comparison to the control, CHIC-35 shows partial recovery in craniofacial cartilage elements<sup>[2]</sup>.

At 4dpf, zebrafish embryos show a well-formed lower jaw in controls, while chd7 morphants exhibits reduced lower jaw. Treatment with CHIC-35 (5  $\mu$ M) rescues the expression of sox9a inchd7 morphants<sup>[2]</sup>.

Nearly 30% of chd7 morphant embryos (24hpf to 72hpf) shows a near complete loss of isl2a expression in the cranial region compared to 10% of the wildtype controls. CHIC-35 reduces this to 7.5% significantly. However, CHIC-35 shows no discernible effect on the enteric neurons marked by  $Tg^{[2]}$ .

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

## **REFERENCES**

[1]. Jérôme Lugrin, et al. The sirtuin inhibitor cambinol impairs MAPK signaling, inhibits inflammatory and innate immune responses and protects from septic shock. Biochim Biophys Acta. 2013 Jun;1833(6):1498-510

[2]. Zainab Asad, et al. Chemical screens in a zebrafish model of CHARGE syndrome identifies small molecules that ameliorate disease-like phenotypes in embryo. Eur J Med Genet. 2020 Feb;63(2):103661.

 $\label{lem:caution:Product} \textbf{Caution: Product has not been fully validated for medical applications. For research use only.}$ 

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