EX527

1 mg

Small Molecules

Epigenetic modifier; Inhibits SIRT1

histone deacetylase

Catalog # 73652

73654 10 mg



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Product Description

EX527 is a cell-permeable, selective inhibitor of mammalian sirtuin 1 (SIRT1; $IC_{50} = 98 \text{ nM}$) over SIRT2 and SIRT3 and has no effect on other histone deacetylases (HDACs; Nayagam et al.). SIRT1 is a nicotinamide adenine dinucleotide (NAD)-dependent deacetylase with roles in energy metabolism and inflammation. Studies have shown that EX527 inhibits sirtuins by forming a trimeric sirtuin complex with an NAD+-derived coproduct (Gertz et al.).

Chemical Name: 6-chloro-2,3,4,9-tetrahydro-1H-carbazole-1-carboxamide

Structure:

Properties

Physical Appearance: A crystalline solid

Storage: Product stable at -20°C as supplied. Protect product from prolonged exposure to light. For long-term storage

store with a desiccant.

Stable as supplied for 12 months from date of receipt.

Solubility: \cdot DMSO \leq 80 mM

· Ethanol ≤ 20 mM

For example, to prepare a 10 mM stock solution in DMSO, resuspend 1 mg in 402 µL of DMSO.

Prepare stock solution fresh before use. Information regarding stability of small molecules in solution has rarely been reported, however, as a general guide we recommend storage in DMSO at -20°C. Aliquot into working volumes to avoid repeated freeze-thaw cycles. The effect of storage of stock solution on compound performance should be tested for each application.

Compound has low solubility in aqueous media. For use as a cell culture supplement, stock solution should be diluted into culture medium immediately before use. Avoid final DMSO concentration above 0.1% due to potential cell toxicity.

Small Molecules EX527



Published Applications

DIFFERENTIATION

- · Increases the production of oligodendrocytes from differentiating neural stem cells and neural progenitor cells in vitro (Rafalski et al.). IMMUNOLOGY
- · Restores the microvascular response during the hypoinflammatory phase in a mouse model of sepsis, and enhances the systemic innate immune response (Vachharajani et al.).

DISEASE MODELING

· Delays cyst growth in kidneys of PKD1 knockout mouse models (Zhou et al.).

References

Gertz M et al. (2013) Ex-527 inhibits Sirtuins by exploiting their unique NAD+-dependent deacetylation mechanism. Proc Natl Acad Sci USA 110(30): E2772–81.

Nayagam VM et al. (2006) SIRT1 modulating compounds from high-throughput screening as anti-inflammatory and insulin-sensitizing agents. J Biomol Screen 11(8): 959–67.

Rafalski VA et al. (2013) Expansion of oligodendrocyte progenitor cells following SIRT1 inactivation in the adult brain. Nat Cell Biol 15(6): 614–24.

Vachharajani VT et al. (2014) SIRT1 inhibition during the hypoinflammatory phenotype of sepsis enhances immunity and improves outcome. J Leukoc Biol 96(5): 785–96.

Zhou X et al. (2013) Sirtuin 1 inhibition delays cyst formation in autosomal-dominant polycystic kidney disease. J Clin Invest 123(7): 3084–98.

Related Small Molecules

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This product is hazardous. Please refer to the Safety Data Sheet (SDS).

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