BIO

Small Molecules

WNT pathway activator; Inhibits GSK3



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TOLL FREE PHONE 1 800 667 0322 • PHONE +1 604 877 0713 INFO@STEMCELL.COM • TECHSUPPORT@STEMCELL.COM FOR GLOBAL CONTACT DETAILS VISIT OUR WEBSITE

Catalog # 72032 1 mg 72034 5 mg

Product Description

BIO is an indirubin compound that is cell-permeable and is a selective, potent, and reversible inhibitor of glycogen synthase kinase (GSK) 3α and GSK3 β (IC₅₀ = 5 nM) which acts by competing for the ATP binding site of GSK3. GSK3 is a serine/threonine kinase that is a key inhibitor of the WNT pathway; therefore BIO functions as a WNT activator. It shows some inhibitory activity against cyclin-dependent kinase (CDK) 5 (IC₅₀ = 83 nM), CDK2 (IC₅₀ = 300 nM), and CDK1 (IC₅₀ = 320 nM), and little activity against other common kinases including MAPK, PKA, PKC, and PKG (Meijer et al.).

Molecular Name: BIO

Alternative Names: GSK 3 IX; MLS 2052; 6-Bromoindirubin-3'-oxime

CAS Number: 667463-62-9 Chemical Formula: $C_{16}H_{10}BrN_3O_2$ Molecular Weight: 356.2 g/mol Purity: $\geq 98\%$

Chemical Name: 6-bromo-3-[(3E)-1,3-dihydro-3-(hydroxyimino)-2H-indol-2-ylidene]-1,3-dihydro-(3Z)-2H-indol-2-one

Structure:

Properties

Physical Appearance: A crystalline solid

Storage: Product stable at -20°C as supplied. Protect from prolonged exposure to light. For product expiry date, please

contact techsupport@stemcell.com.

Solubility: \cdot DMSO \leq 25 mM

· Absolute ethanol ≤ 1.5 mM

For example, to prepare a 10 mM stock solution in DMSO, resuspend 1 mg in 281 µL of fresh 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.

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Published Applications

MAINTENANCE AND SELF-RENEWAL

- · Maintains self-renewal of mouse embryonic stem (ES) cells in the absence of LIF (Sato et al.).
- · Maintains self-renewal of human ES cells in unconditioned medium through a SMAD 2/3 dependent mechanism (James et al.; Sato et al.).
- · Induces proliferation of neonatal and adult rat cardiomyocytes (Tseng et al.).

REPROGRAMMING

- · Enhances reprogramming of mouse fibroblasts, neural stem cells, and thymocytes to induced pluripotent stem (iPS) cells (Lluis et al.). DIFFERENTIATION
- · Promotes differentiation of cardiomyocytes from human ES and iPS cells (Lian et al.).

References

James D et al. (2005) TGFbeta/activin/nodal signaling is necessary for the maintenance of pluripotency in human embryonic stem cells. Development 132(6): 1273–82.

Lian X et al. (2012) Robust cardiomyocyte differentiation from human pluripotent stem cells via temporal modulation of canonical Wnt signaling. Proc Natl Acad Sci U S A 109(27): E1848–57.

Lluis F et al. (2008) Periodic activation of Wnt/beta-catenin signaling enhances somatic cell reprogramming mediated by cell fusion. Cell Stem Cell 3(5): 493–507.

Meijer L et al. (2003) GSK-3-selective inhibitors derived from Tyrian purple indirubins. Chem Biol 10(12): 1255-66.

Sato N et al. (2004) Maintenance of pluripotency in human and mouse embryonic stem cells through activation of Wnt signaling by a pharmacological GSK-3-specific inhibitor. Nat Med 10(1): 55–63.

Tseng A-S et al. (2006) The GSK-3 inhibitor BIO promotes proliferation in mammalian cardiomyocytes. Chem Biol 13(9): 957-63.

Related Small Molecules

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