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Certificate of Analysis

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Product Name: L-685,458

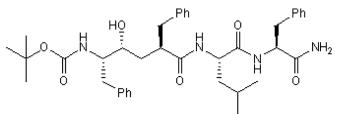
Catalog No.: 2627 Batch No.: 4

CAS Number: IUPAC Name: 292632-98-5

(5S)-(tert-Butoxycarbonylamino)-6-phenyl-(4R)-hydroxy-(2R)-benzylhexanoyl)-L-leucy-L-phenylalaninamide

1. PHYSICAL AND CHEMICAL PROPERTIES

Batch Molecular Formula: Batch Molecular Weight: Physical Appearance: Solubility: Storage: Batch Molecular Structure: $C_{39}H_{52}N_4O_6$ 672.85 White solid DMSO to 15 mM Store at -20°C



8.39

2. ANALYTICAL DATA

TLC:	R _f = 0.3 (Dichloromethane:Methanol [95:5])			
HPLC:	Shows 98.2% purity			
¹ H NMR:	Consistent with structure			
Mass Spectrum:	Consistent with structure			
Microanalysis:	Carbon Hydrogen Nitrogen			
	Theoretical 69.62 7.79 8.33			

Found

Caution - Not Fully Tested • Research Use Only • Not For Human or Veterinary Use

69.31

7.84

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Print Date: Feb 28th 2024

4

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CAS Number: 292632-98-5

IUPAC Name:

(5S)-(tert-Butoxycarbonylamino)-6-phenyl-(4R)-hydroxy-(2R)-benzylhexanoyl)-L-leucy-L-phenylalaninamide

Description:

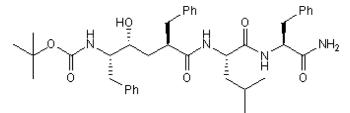
L-685,458 is a potent and selective γ -secretase inhibitor (IC₅₀ = 17 nM) that displays > 50-fold selectivity over a range of aspartyl, serine and cysteine proteases. L-685,458 binds with high affinity to the related aspartyl protease, signal peptide peptidase (SPP; K_D = 5.1 nM), and inhibits SPP expressed in HEK293 cells (IC₅₀ = 10 μ M). L-685,458 exhibits equal potency for inhibition of Aβ40 and Aβ42 peptides (IC₅₀ values are 48 and 67 nM respectively in human neuroblastoma cells). It also regulates CXCR4 and VEGFR2 expression through inhibition of Notch signaling in vitro.

Physical and Chemical Properties:

Batch Molecular Formula: $C_{39}H_{52}N_4O_6$ Batch Molecular Weight: 672.85 Physical Appearance: White solid

Minimum Purity: ≥97%

Batch Molecular Structure:



References:

Storage: Store at -20°C

Solubility & Usage Info:

DMSO to 15 mM

This product is supplied in lyophilized form. It may appear as a solid, gel or film and be very hard to visualize. Solutions should be made by adding solvent directly to the vial. The vial should then be vortexed vigorously to ensure the product has completely dissolved.

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Stability and Solubility Advice:

Some solutions can be difficult to obtain and can be encouraged by rapid stirring, sonication or gentle warming (in a 45-60°C water bath).

Information concerning product stability, particularly in solution, has rarely been reported and in most cases we can only offer a general guide. *Unless contradicted by product-specific protocols or instructions, our standard recommendations apply:

SOLIDS: Provided storage is as stated on the product label and the vial is kept tightly sealed, the product can be stored for up to 6 months from date of receipt.

SOLUTIONS: We recommend that stock solutions, once prepared, are stored aliquoted in tightly sealed vials at -20°C or below and used within 1 month. Wherever possible solutions should be made up and used on the same day.

Williams *et al* (2008) Regulation of CXCR4 by the Notch ligand delta-like 4 in endothelial cells. Cancer Res. **68** 1889. PMID: 18339870. **Iben** *et al* (2007) Signal peptide peptidase and gamma-secretase share equivalent inhibitor binding pharmacology. J.Biol.Chem. **282** 36829. PMID: 17932033.

Williams et al (2006) Up-regulation of the Notch ligand delta-like 4 inhibits VEGF-induced endothelial cell function. Blood **107** 931. PMID: 16219802.

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