PRODUCT INFORMATION



L-778,123

Item No. 22940

CAS Registry No.: 253863-00-2

Formal Name: 4-[[5-[[4-(3-chlorophenyl)-

3-oxo-1-piperazinyl]methyl]-1H-imidazol-1-yl]methyl]-

benzonitrile, monohydrochloride

C₂₂H₂₀CIN₅O • HCI MF:

442.3 FW: **Purity:** ≥98%

Supplied as: A crystalline solid

Storage: -20°C Stability: ≥4 years

Information represents the product specifications. Batch specific analytical results are provided on each certificate of analysis.

• HCI

Laboratory Procedures

L-778,123 is supplied as a crystalline solid. A stock solution may be made by dissolving the L-778,123 in the solvent of choice, which should be purged with an inert gas. L-778,123 is soluble in organic solvents such as ethanol, DMSO, and dimethyl formamide (DMF). The solubility of L-778,123 is approximately 10 mg/ml in ethanol and approximately 12.5 mg/ml in DMSO and DMF.

Description

L-778,123 is a dual inhibitor of farnesyl transferase (FTase; IC_{50} = 2 nM) and geranylgeranyl transferase type I (GGTase I; IC_{50} = 98 nM).¹ It inhibits prenylation of the FTase and GGTase I substrates HDJ2 and RAP1A in PSN-1 pancreatic tumor cells (EC₅₀s = 92 and 6,760 nM, respectively). L-778,123 (1-300 μM) also inhibits prenylation of the oncogenic protein KI-RAS in PSN-1 cells in a concentration-dependent manner. Ex vivo, L-778,123 (35-50 mg/kg per day) reduces HDJ2 and RAP1A prenylation in dog peripheral blood mononuclear cells (PBMCs) but has no effect on KI-RAS prenylation in patient-derived PBMCs. L-778,123 inhibits lectin-induced expression of the T cell activation markers CD71 and CD25 on human PMBCs (IC₅₀s = 6.48 and 84.1 μM, respectively) and inhibits IL-2-induced proliferation of CTLL-2 cells (IC₅₀ = $0.81 \mu M$).²

References

- 1. Lobell, R.B., Liu, D., Buser, C.A., et al. Preclinical and clinical pharmacodynamic assessment of L-778,123, a dual inhibitor of farnesyl:protein transferase and geranylgeranyl:protein transferase type-I. Mol. Cancer Ther. 1(9), 747-758 (2002).
- 2. Si, M.-S., Reitz, B.A., and Borie, D.C. Inhibition of lymphocyte activation and function by the prenylation inhibitor L-778,123. Invest New Drugs 23(1), 21-29 (2005).

WARNING
THIS PRODUCT IS FOR RESEARCH ONLY - NOT FOR HUMAN OR VETERINARY DIAGNOSTIC OR THERAPEUTIC USE.

This material should be considered hazardous until further information becomes available. Do not ingest, inhale, get in eyes, on skin, or on clothing. Wash thoroughly after handling. Before use, the user must review the complete Safety Data Sheet, which has been sent via email to your institution.

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CAYMAN CHEMICAL

1180 EAST ELLSWORTH RD ANN ARBOR, MI 48108 · USA PHONE: [800] 364-9897

[734] 971-3335

FAX: [734] 971-3640 CUSTSERV@CAYMANCHEM.COM WWW.**CAYMANCHEM**.COM