

Product Specification Sheet

Product Name: ZD6474 (Vandetanib)

Catalog Number: C9364

Technical information:

Chemical Formula: $C_{22}H_{24}BrFN_4O_2$

CAS #: 443913-73-3

Molecular Weight: 475.35

Purity: > 98%

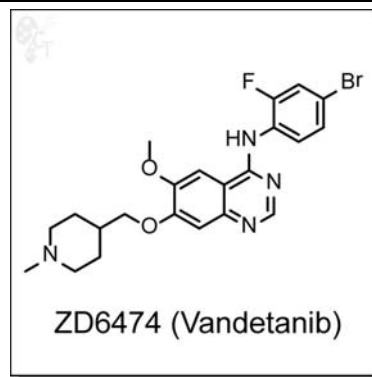
Appearance: White solid

Solubility: Soluble in DMSO up to 40 mM

Chemical Name: N-(4-bromo-2-fluorophenyl)-6-methoxy-7-((1-methylpiperidin-4-yl)methoxy)quinazolin-4-amine

Storage: Store solid powder at 4°C desiccated; Store DMSO solution at -20°C.

Shelf Life: In the unopened package, powder is stable for 1 year and DMSO solution is stable for 6 months under proper storage condition.



Handling: • To make 10 mM stock solution, add 0.21mL of DMSO for each mg of ZD6474 (Vandetanib).

- For DMSO solution, briefly spin the vial at 500 rpm in a 50 mL conical tube to ensure maximum sample recovery.

Biological Activity: Vandetanib (ZD6474) is an orally-bioavailable, ATP-competitive, quinazoline-based inhibitor of VEGFR2 that has been shown to inhibit both VEGF-induced signalling in endothelial cells and tumor-induced angiogenesis. [1] Vandetanib inhibits VEGFR2, VEGFR3, EGFR, and RET at IC50s of 40 nM, 110 nM, 500 nM, and 130 nM, respectively. It has been found to inhibit cell proliferation of VEGFR-stimulated cells (IC50 60 nM) and EGFR-stimulated HUVEC proliferation (IC50 170 nM). [2]

Vandetanib shows robust inhibition of VEGF-stimulated VEGFR2 phosphorylation, Erk-1/2 phosphorylation, as well as endothelial cell proliferation at doses less than 100 nM. [3]

More recently, Vandetanib has been shown to antagonize ABCC1- and ABCG2-mediated MDR by inhibition of transport function. ABCG2, expressed in a wide variety of cancer stem cells, is inhibited by vandetanib, potentially reversing MDR at low concentrations. [4]

- Reference:**
1. Ryan et al., Clin. Cancer 2005, 92, S6-S13.
 2. Flanigan et al., Current status of vandetanib (ZD6474) in the treatment of non-small cell lung cancer. *Biologics: Targets and Therapy*, 2010, 4, 237-243. Pubmed ID: 20859451
 3. McCarty et al., ZD6474, a vascular endothelial growth factor receptor tyrosine kinase inhibitor with additional activity against epidermal growth factor receptor tyrosine kinase, inhibits orthotopic growth and angiogenesis of gastric cancer. *Mol. Cancer Ther.* 2004, 3(9), 1041-1048. Pubmed ID: 15367698
 4. Zheng et al., Vandetanib (Zactima, ZD6474) antagonizes ABCC1- and ABCG2-mediated multidrug resistance by inhibition of their transport function. *PLoS ONE*, 2009, 4(4), e5172. Pubmed ID: 19390592

To reorder: <http://www.cellagentech.com/ZD6474-Vandetanib/>

For Technical Support: technical@cellagentech.com

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