

Catalog # 10-4788 Duvelisib CAS# 1201438-56-3

8-Chloro-2-phenyl-3-[(1S)-1-(7H-purin-6-ylamino)ethyl]isoquinolin-1-one; IPI-145 Lot # FBS1119

Duvelisib is a potent and selective (IC_{50} 's: $PI3K_{\alpha}=1602$ nM, $PI3K_{\beta}=85$ nM, $PI3K_{\delta}=2.5$ nM, $PI3K_{\gamma}=27$ nM) dual $PI3K_{\delta/\gamma}$ inhibitor. It inhibits B and T cell proliferation, blocks neutrophil migration, and inhibits basophil activation. Duvelisib antagonizes B-cell receptor crosslinking activated pro-survival signals in primary chronic lymphocytic leukemia cells. Duvelisib also shows preclinical/clinical activity against other hematologic malignancies such as Non-Hodgkins lymphoma, T-cell lymphoma, and others. Useful clinical agent for the treatment of various blood cancers. Low-dose treatment of T-cell-inflamed tumor models of head and neck cancers with Duvelisib enhanced responses to PD-L1 blockade via suppression of myeloid-derived suppressor cells. Higher doses reversed the effect due to suppression of tumor-infiltrating T lymphocytes.

- 1) Winkler et al. (2013) PI3K- δ and PI3K- γ Inhibition by IPI-145 Abrogates Immune Response and Suppresses Activity in Autoimmune and Inflammatory Disease Models; Chem.Biol. **20** 1309
- 2) Dong et al. (2014) IPI-145 antagonizes intrinsic and extrinsic survival signals in chronic lymphocytic leukemia cells; Blood 124 3583
- 3) Flinn et al. (2018) Duvelisib, a novel dual inhibitor of PI3K-δ,γ, is clinically active in advances hematologic malignancies; Blood 131 877
- 4) Faia et al. (2018) The phosphoinositide-3 kinase (PI3K)-δ,γ inhibitor, duvelisib, shows preclinical synergy with multiple targeted therapies in hematologic malignancies; PLoS One 13 e0200725
- 5) Davis et al. (2017) Anti-PD-L1 Efficacy Can Be Enhanced by Inhibition of Myeloid-Derived Suppressor Cells with a Selective Inhibitor of Pl3Kδ/γ; Cancer Res. **77** 2607

PHYSICAL DATA

Molecular Weight: 416.87

 $\begin{tabular}{lll} Molecular Formula: & $C_{22}H_{17}CIN_6O$ \\ Purity: & $>98\%$ by HPLC \\ \end{tabular}$

NMR: (Conforms)

Solubility: DMSO (>25 mg/ml)

Physical Description: White solid

Storage and Stability: Store as supplied at -20°C for up to 1 year from the date of purchase. Solutions in

DMSO may be stored at -20°C for up to 1 month.

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