

Catalog #10-4108 AZD7648

CAS# 2230820-11-6

7-Methyl-2-[(7-methyl-[1,2,4]triazolo[1,5-a]pyridine-6-yl)amino]-9-(oxan-4-yl)purin-8-one Lot # FBA94036

AZD7648 is a very potent and selective (against 397 kinases) DNA-PK inhibitor (pIC₅₀ = 9.2; cellular pIC50 = 7.0). It is a sensitizer of radiation- and doxorubicin-induced DNA damage and sustained tumor regression in xenograft and patient-derived xenograft models. AZD7648 significantly enhances precise CRISPR/Cas9 gene editing alone 3,4,5 or in combined treatment with a DNA polymerase theta inhibitor (2iHDR cocktail) .

- 1) Goldberg et al. (2020), The Discovery of 7-Methyl-2-[(7-methyl[1,2,4]triazolo[1,5-a]pyridine-6-yl)amino]-9-(tetrahydro-2H-pyran-4-yl)-7,9-dihydro-8H-purin-8-one (AZD7648), a Potent and Selective DNA-Dependent Protein Kinase (DNA-PK) Inhibitor, J. Med. Chem. **63** 3461
- 2) Fok et al. (2019), AZD7648 is a potent and selective DNA-PK inhibitor that enhances radiation, chemotherapy and Olaparib activity; Nat. Commun. **10** 5065
- 3) Cloarec-Ung et al. (2024), Near-perfect precise on-target editing of human hematopoietic stem and progenitor cells; Elife **12** RP91288
- 4) Selvaraj et al. (2024), High-efficiency transgene integration by homology-directed repair in human primary cells using DNA-PKcs inhibition; Nat. Biotech. **42** 731
- 5) Wimberger et al. (2023), Simultaneous inhibition of DNA-PK and PolΘ improves integration efficiency and precision of genome editing; Nat. Commun. **14** 4761

PHYSICAL DATA

 $\begin{tabular}{lll} Molecular Weight: & 380.41 \\ Molecular Formula: & $C_{18}H_{20}N_8O_2$ \\ Purity: & $>98\% \ (HPLC) \end{tabular}$

NMR: (Conforms)

Solubility: DMSO (5 mg/mL with warming)

Physical Description: White solid

Storage and Stability: Store as supplied at -20°C for up to 2 years from the date of purchase. Solutions in

DMSO may be stored at -20°C for up to 3 months.