

## Catalog #10-3956 BI-3406

CAS# 2230836-55-0

N-[(1R)-1-[3-Amino-5-(trifluoromethyl)phenyl]ethyl]-7-methoxy-2-methyl-6-[(3S)-oxolan-3-yl]oxyquinazolin-4-amine Lot # FBS4047



BI-3406 is a potent ( $IC_{50} = 5 \text{ nM}$ ), selective, and orally bioavailable SOS1 inhibitor.<sup>1</sup> It blocks SOS1 interaction with KRAS preventing activation of KRAS. BI-3406 was active against multiple cancer cell types and xenograft models harboring various KRAS mutant alleles including G12C,V,S,A,and G13D. It displayed strong synergistic effects with the MEK inhibitor trametinib<sup>1,2</sup> and this combination was able to overcome resistance to the KRAS inhibitors sotorasib and adagrasib<sup>3</sup>. BI-3406 acted synergistically with the KRAS inhibitor adagrasib in KRAS G12C mutant lung and colorectal cancer models.<sup>4</sup>

- 1) Hofmann et al. (2020), BI-3406, a potent and selective SOS1::KRAS interaction inhibitor, is effective in KRAS-driven cancers through combined MEK inhibition; Cancer Discov. 11 142
- 2) Ma et al. (2022), Inhibition of KRAS, MEK and PI3K Demonstrate Synergistic Anti-Tumor Effects in Pancreatic Ductal Adenocarcinoma Cell Lines; Cancers (Basel) 14 4467
- 3) Koga et al. (2021), KRAS Secondary Mutations That Confer Acquired Resistance to KRAS G12C Inhibitors, Sotorasib and adagrasib, and Overcoming Strategies: Insights From In Vitro Experiments; J. Thorac. Oncol. **16** 1321
- 4) Thatikonda et al. (2024), Co-targeting SOS1 enhances the antitumor effects of KRAS<sup>G12C</sup> inhibitors by addressing intrinsic and acquired resistance; Nat. Cancer **5** 1352

## PHYSICAL DATA

Molecular Weight:	462.47
Molecular Formula:	C <sub>23</sub> H <sub>25</sub> F <sub>3</sub> N <sub>4</sub> O <sub>3</sub>
Purity:	>98% (HPLC)
	NMR: (Conforms)
Solubility:	DMSO (>70 mg/mL)
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.
Materials provided by Fo	cus Biomolecules are for laboratory research use only and are not intended for human or veterinary applications.
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