

Catalog # 10-4856 PF-4708671 dihydrochloride

CAS# 125517-76-0 (free base)

2-[[4-(5-Ethylpyrimidin-4-yl)piperazin-1-yl]methyl]-5-(trifluoromethyl)-1H-benzimidazole dihydrochloride

Lot # FBA9018

PF-4708671 is a cell permeable inhibitor of p70 ribosomal S6 kinase 1 (Ki = 20 nM; IC₅₀ = 160 nM). Selective against 87 kinases including 13 other AGC family members. It enhanced cell death induced by glucose deprivation in breast cancer cells2 and inhibited cell migration in triple-negative breast cancer cells3. PF-4708671 inhibited cell proliferation and invasiveness in multiple non-small cell lung cancer cell lines and inhibited tumorigenesis in mice. It induced AMPK phosphorylation and activation independently of p70SK1 *via* specific inhibition of mitochondrial respiratory chain complex I. PF-4708671 displayed protective effects in a mouse model of cerebral ischemia-reperfusion injury.

- 1) Pearce et al. (2010) Characterization of PF-4708671, a novel and highly specific inhibitor of p70 ribosomal S6 kinase (S6K1); Biochem. J. **431** 245
- 2) Choi et al. (2013) Inhibition of S6K1 enhances glucose deprivation-induced cell death via downregulation of anti-apoptotic proteins in MCF-7 breast cancer cells; Biochem. Biophys. Res. Commun. **432** 123
- 3) Khotskaya et al. (2014) S6K1 promotes invasiveness of breast cancer cells in a model of metastasis of triple-negative breast cancer; Am. J. Transl. Res. **6** 361
- 4) Qiu et al. (2016) The p70S6K Specific Inhibitor PF-4708671 Impedes Non-Small Cell Lung Cancer Growth; PLoS One, 11 e0147185
- 5) Vainer et al. (2014) PF-4708671 activates AMPK independently of p70S7K1 inhibition; PLoS One, 9 e107364
- 6) Chi et al. (2019) Inhibition of p70 ribosomal S6 kinase 1 (S6K1) by PF-7408671 decreased infarct size in early cerebral ischemia-reperfusion with decreased BBB permeability; PLoS One, **9** e107364

PHYSICAL DATA

Molecular Weight: 463.33

Molecular Formula: $C_{19}H_{21}F_3N_6 \cdot 2HCl$ Purity: >98% by HPLC NMR: (Conforms)

Solubility: DMSO (at least 35 mg/ml); Water (at least 50 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. Hygroscopic solid – protect from moisture and

humidity.

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