



3050 Spruce Street
Saint Louis, Missouri 63103 USA
Telephone 800-325-5832 • (314) 771-5765
Fax (314) 286-7828
email: techserv@sial.com
sigma-aldrich.com

Product Information

Mu-Phe-hPhe-FMK

Catalog Number **M4070**
Storage Temperature $-20\text{ }^{\circ}\text{C}$

Synonym: N-Morpholineurea-phenylalanyl-homophenylalanylfluoromethyl ketone

Product Description

Molecular formula: $\text{C}_{25}\text{H}_{30}\text{FN}_3\text{O}_4$
Molecular weight: 455.5

Mu-Phe-hPhe-FMK is a cell-permeable inhibitor of cathepsins B and L.

Among the intracellular proteinases, thiol proteinases such as cathepsin B and cathepsin L, active at slightly acidic pH, play a role in lysosomal protein catabolism. Both have identical substrate specificity, but 4 M urea at pH 5.0 will inactivate cathepsin B, while cathepsin L retains its activity under these conditions.³

Cathepsin B is a lysosomal cysteine protease involved in cellular protein turnover.⁴ Cathepsin B has a high abundance and exhibits both endopeptidase and peptidyl dipeptidase activity.³ This enzyme has been implicated in several pathological conditions including arthritis and tumor metastasis. Cathepsin L is the most powerful of the lysosomal proteinases and has a higher specific activity than cathepsin B in the degradation of physiological protein substrates. Cathepsin L, but not cathepsin B, can also generate kinins from high and low molecular weight kininogens in vitro.⁴

FMK is a trapping group responsible for irreversible inhibition, but is non-cytotoxic. Inhibition occurs when the FMK group covalently bonds to the $-\text{SH}$ of an adjacent cysteine residue on the target protein.

Precautions and Disclaimer

This product is for R&D use only, not for drug, household, or other uses. Please consult the Material Safety Data Sheet for information regarding hazards and safe handling practices.

Preparation Instructions

Prepare stock 20 mM solutions in dry DMSO (>99.9%). Suggested working concentration is 40 μM

Storage/Stability

Store desiccated at $-20\text{ }^{\circ}\text{C}$. Stock solutions stored in frozen aliquots at $-20\text{ }^{\circ}\text{C}$ are stable for 6-8 months. Allow containers to equilibrate to room temperature before opening.

References

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2. Harth, G. et al. Peptide-fluoromethyl ketones arrest intracellular replication and intercellular transmission of *Trypanosoma cruzi*. *Mol. Biochem. Parasitol.* **58**, 17-24 (1993).
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4. Kirschke H, Barrett AJ. 1987. Lysosomal cysteine proteases. In: Glaumann H, Ballard FJ, eds. *Lysosomes: Their role in protein breakdown*. London: Academic Press. pp 193-238.
5. Aronson, N. N., Jr., and Barrett, A. J., The specificity of cathepsin B. Hydrolysis of glucagon at the C-terminus by a peptidyl dipeptidase mechanism. *Biochem. J.*, **171**, 759-765 (1978).
6. Desmazes, C., et al., Cathepsin L, but not cathepsin B, is a potential kininogenase. *Biol. Chem.*, **382**, 811-815 (2001).

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