

Product Information

MONOCLONAL ANTI-MATRIX METALLOPROTEINASE-3 (MMP-3), CLONE 4B7.3 Purified Mouse Immunoglobulin Fraction

Product Number **M 6552**

Product Description

Monoclonal Anti-Matrix Metalloproteinase-3 (MMP-3) (mouse IgG1 isotype) is derived from the hybridoma produced by the fusion of mouse myeloma cells and splenocytes from mice immunized with native human matrix metalloproteinase-3 (stromelysin-1) as immunogen. The immunoglobulin fraction of antibody to MMP-1 is purified from ascites fluid using protein G affinity chromatography.

Monoclonal Anti-MMP-3 specifically binds to stromelysin-1 and does not cross-react with stromelysin-2 or stromelysin-3 (MMP-10 or MMP-11), nor with other MMP family members (MMP-1, MMP-2, MMP-9, etc). By immunoblotting against the reduced protein, the antibody reacts with bands at 59 kDa and 57 kDa (the pro-form), as well as the active forms. It also reacts with non-reduced MMP-3. Higher antibody concentrations may be necessary for non-human samples.

Monoclonal Anti-MMP-3 may be used for the detection and localization of MMP-3 by various immunochemical techniques such as immunoblotting, immunoprecipitation, immunohistochemistry and ELISA.

The matrix metalloproteinases (MMPs) are a family of at least eighteen secreted and membrane-bound zinc-endopeptidases. Collectively, these enzymes can degrade all the components of the extracellular matrix, including fibrillar and non-fibrillar collagens, fibronectin, laminin and basement membrane glycoproteins. In general, the structure of MMPs is characterized by a signal peptide, a propeptide, and a catalytic domain containing the highly conserved zinc-binding site. In addition, fibronectin-like repeats, a hinge region, and a C-terminal hemopexin-like domain allow categorization of MMPs into the collagenase, gelatinase, stromelysin and membrane-type MMP subfamilies.^{1,2,3} MMPs contain the motif His-Glu-Xaa-His that binds zinc in the catalytic site, as well as another zinc molecule and two calcium molecules structurally. They fall within the matrixin subfamily, and are EC designated 3.4.24.x.

This group also contains astacin, reprotysin, and serralyisin, as well as other more divergent metalloproteinases. All MMPs are synthesized as proenzymes, and most of them are secreted from the cells as proenzymes. Thus, the activation of these proenzymes is a critical step that leads to extracellular matrix breakdown.

MMPs are considered to play an important role in wound healing, apoptosis, bone elongation, embryo development, uterine involution, angiogenesis,⁴ and tissue remodeling, and in diseases such as multiple sclerosis,^{2,5} Alzheimer's,² malignant gliomas,² lupus, arthritis, periodontitis, glomerulonephritis, atherosclerosis, tissue ulceration, and in cancer cell invasion and metastasis.⁶ Numerous studies have shown that there is a close association between expression of various members of the MMP family by tumors and their proliferative and invasive behavior and metastatic potential.

The tissue inhibitors of metalloproteinases (TIMPs) are naturally occurring proteins that specifically inhibit matrix metalloproteinases and regulate extracellular matrix turnover and tissue remodeling by forming tight-binding inhibitory complexes with the MMPs. Thus, TIMPs maintain the balance between matrix destruction and formation. An imbalance between MMPs and the associated TIMPs may play a significant role in the invasive phenotype of malignant tumors.

MMPs and TIMPs can be divided into two groups with respect to gene expression: the majority exhibit inducible expression, and a small number are produced constitutively or are expressed at very low levels and are not inducible. Among agents that induce MMP and TIMP production are the inflammatory cytokines TNF α and IL-1 β . A marked cell type specificity is a hallmark of both MMP and TIMP gene expression (i.e., a limited number of cell types can be induced to make these proteins).

Matrix Metalloproteinase-3 (MMP-3) is also termed stromelysin-1. MMP-3 and MMP-10 (stromelysin-2) are both expressed by keratinocytes and fibroblasts and they are able to degrade a wide range of substrates. MMP-3 degrades gelatin, type IV, V, IX and X collagens, elastin, laminin, vitronectin, and proteoglycans.^{8,9} MMP-3 can be induced by cytokines IL-1 β and TNF α , by growth factors EGF and PDGF, and by the tumor promotor PMA; and expression is inhibited by TGF β and by all-trans retinoic acid (RA). The human MMP-3 gene has the chromosomal location of 11q22.2-22.3.

Reagents

Monoclonal Anti-MMP-3 is supplied in M phosphate buffered saline, pH 7.4, containing 50% glycerol and 15 mM sodium azide as preservative.

Precautions and Disclaimer

Due to the sodium azide content a material safety data sheet (MSDS) for this product has been sent to the attention of the safety officer of your institution. Consult the MSDS for information regarding hazards and safe handling practices.

Storage/Stability

For continuous use, store at 2-8 °C for up to six months. For extended storage, the solution may be stored 0 to !20 °C. The antibody is supplied with 50% glycerol to prevent freezing. If slight turbidity occurs upon pro-longed storage, clarify the solution by centrifugation before use.

Product Profile

A working dilution of 1:1,000 is determined by immunoblotting using a concentrated cell culture media from a stimulated human cell line. (Substrate: BCIP/NBT).

Control: MMP Control-1, Product No. M 2928.

Note: MMP-3 levels in quiescent cells and tissue are minimal, and stimulation of protein concentration is often needed to visualize the bands by immunoblotting. In addition, cell types differ greatly in the quantity of MMP-3 produced.

In order to obtain best results and assay sensitivity in different techniques and preparations we recommend determining optimum working dilutions by titration assay.

References

1. Borkakoti, N., Prog. Biophy. Mol. Biol., **70**, 73 (1998).
2. Yong, V. W., et al., Trends Neurosci., **21**, 75 (1998).
3. K \neq h \neq ri, V. M., and Saarialho-Kere, U., Exp. Dermatol., **6**, 199 (1997).
4. Halbert, I., et al., Proc. Natl. Acad. Sci., USA, **93**, 9748 (1996).
5. Chandler, S., et al., J. Neuroimmunol., **72**, 155 (1997).
6. Birkedal-Hansen, H., et al., Crit. Rev. Oral Biol. Med., **4**, 197 (1993).
7. Hasty, K. A., et al., J. Exp. Med., **159**, 1455 (1984).
8. Murphy, G., et al., Biochem. J., **277**, 277 (1991).
9. Windsor, L. J., J. Biol. Chem., **268**, 17341 (1993).

JWM/DAA 12/98

Sigma brand products are sold through Sigma-Aldrich, Inc.

Sigma-Aldrich, Inc. warrants that its products conform to the information contained in this and other Sigma-Aldrich publications. Purchaser must determine the suitability of the product(s) for their particular use. Additional terms and conditions may apply. Please see reverse side of the invoice or packing slip.