

DESCRIPTION

Source *E. coli*-derived
Val45-Ala118
Accession # P50228

N-terminal Sequence Analysis Val45

Predicted Molecular Mass 8 kDa

SPECIFICATIONS

Activity Measured by its ability to chemoattract BaF3 mouse pro-B cells transfected with human CXCR2. The ED₅₀ for this effect is typically 0.03-0.1 µg/mL.

Measured by its ability to induce myeloperoxidase release from cytochalasin B-treated human neutrophils. Schröder, J.M. *et al.* (1987) *J. Immunol.* **139**:3474.
The ED₅₀ for this effect is typically 1-3 µg/mL.

Endotoxin Level <0.01 EU per 1 µg of the protein by the LAL method.

Purity >97%, by SDS-PAGE under reducing conditions and visualized by silver stain.

Formulation Lyophilized from a 0.2 µm filtered solution in PBS. See Certificate of Analysis for details.

PREPARATION AND STORAGE

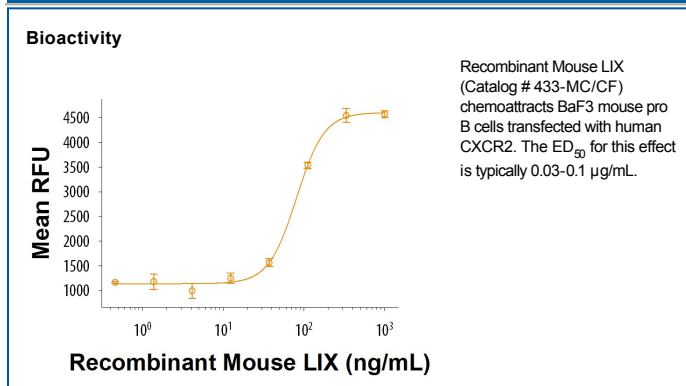
Reconstitution Reconstitute at 100 µg/mL in sterile PBS.

Shipping The product is shipped at ambient temperature. Upon receipt, store it immediately at the temperature recommended below.

Stability & Storage **Use a manual defrost freezer and avoid repeated freeze-thaw cycles.**

- 12 months from date of receipt, -20 to -70 °C as supplied.
- 1 month, 2 to 8 °C under sterile conditions after reconstitution.
- 3 months, -20 to -70 °C under sterile conditions after reconstitution.

DATA



BACKGROUND

LIX (Liposaccharide-Induced CXC chemokine; also GARG-8) is a secreted 8-10 kDa member of the ELR+ class, CXC family, chemokine superfamily of molecules (1-4). It is one of five CXC chemokines in mouse, and serves as a functional equivalent of human ENA-78 and GCP-2. Mouse LIX is synthesized as a 132 amino acid (aa) precursor that contains an extended 40 aa signal sequence plus a 92 aa mature peptide (aa 41-132) (1, 5). No glycosylation has been reported for LIX.

Depending upon the author, the 92 aa full-length isoform is typically referred to as either simply LIX, or LIX (1-92). Shorter isoforms have an accompanying range designation. Although LIX has (marginal) activity, proteolytic processing is necessary for full bioactivity (5, 6). To date, over 25 enzymatically-generated isoforms have been experimentally noted, ranging from 69-92 aa in length (6). These may show only N-terminal, C-terminal, or dual-end processing, with removal of up to 10 N-terminal amino acids (#41-48) and 13 or 14 C-terminal amino acids (#119-132). In vivo, it appears that multiple MMPs act sequentially (and redundantly) to generate a "standard" LIX isoform that consists of aa 5-78 of the mature form. It is suggested that initially, MMP-2 is induced locally, and cleaves LIX 1-92 between Ser4-Val5. This truncation results in an active product (LIX [5-92]) that chemoattracts PMNs. Newly arriving PMNs now secrete MMP-9, which performs the same cleavage as MMP-2, chemoattracting more PMNs. Neutrophil-derived MMP-8 is secreted next, which cleaves both the Ser4-Val5 and Lys79-Arg80 bonds, creating LIX (5-79) (7, 8). Although somewhat unclear, macrophage derived MMP-12 also contributes to LIX activation, and in some circumstances cleaves the ELR motif of activated chemokines (but not LIX), rendering them inactive and downmodulating the inflammatory response (9). Over aa 41-119 of the precursor, mouse LIX shares 73% and 63% aa sequence identity with rat LIX and human GCP-2, respectively. Mouse LIX is known to be active on human cells (6). Cells known to express LIX vary widely and include oligodendroglia (10), adipose tissue stem cells and macrophages (11, 12), platelets and endothelial cells (13), colonic epithelium (14), cardiomyocytes (15), Type II greater alveolar cells and ileal enterocytes (16), hepatocytes (17) and fibroblasts from multiple tissues (18, 19, 20).

LIX is both constitutively expressed (16), and induced in response to a variety of stimuli, including LPS (18, 21), TNF- α (22), mast cell protease-6 (19), leptin (14), and oncostatin M (20). Circulating LIX has two known signaling receptors, CXCR2 and CXCR1. Although the former is considered most important, the latter may prove to have nonredundant functions (6, 21, 23). LIX also binds to DARC, a non-signaling receptor prominently expressed on erythrocytes. Although non-signaling, this receptor serves as a sink or depot for LIX, keeping it from activating CXCR1 and CXCR2 (24). Functionally, LIX is best known as a chemoattractant for neutrophils (5, 9, 15). But it also reportedly chemoattracts macrophages and induces production of NO (14, 25), regulates gut IL-17 and G-CSF secretion (16), promotes TNF- α expression from mast cells and macrophages (26), and induces neurite outgrowth and protects against neuronal apoptosis by serving as an atypical growth factor (10).

References:

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