

#### DESCRIPTION

**Source** *E. coli*-derived  
Leu79-Leu234, with an N-terminal Met  
Accession # NP\_001075732

**N-terminal Sequence Analysis** Met - Leu79

**Predicted Molecular Mass** 17.4 kDa

#### SPECIFICATIONS

**SDS-PAGE** 19 kDa, reducing conditions

**Activity** Measured in a cytotoxicity assay using L-929 mouse fibroblast cells in the presence of the metabolic inhibitor actinomycin D. Matthews, N. and M.L. Neale (1987) in *Lymphokines and Interferons, A Practical Approach*. Clemens, M.J. *et al.* (eds): IRL Press. 221.  
The ED<sub>50</sub> for this effect is typically 0.01-0.04 ng/mL.

**Endotoxin Level** <0.10 EU per 1  $\mu$ g of the protein by the LAL method.

**Purity** >95%, by SDS-PAGE under reducing conditions and visualized by silver stain.

**Formulation** Lyophilized from a 0.2  $\mu$ m filtered solution in PBS with BSA as a carrier protein. See Certificate of Analysis for details.

#### PREPARATION AND STORAGE

**Reconstitution** Reconstitute at 100  $\mu$ g/mL in PBS containing at least 0.1% human or bovine serum albumin.

**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.

#### BACKGROUND

Tumor necrosis factor alpha (TNF- $\alpha$ , also known as cachectin and TNFSF1A), is the prototypic ligand of the TNF superfamily. It is a pleiotropic molecule that plays a central role in inflammation, apoptosis, and immune system development. TNF- $\alpha$  is produced by a wide variety of immune and epithelial cell types (1, 2). Rabbit TNF- $\alpha$  precursor is a type II transmembrane protein that consists of a 35 amino acid (aa) region, a 21 aa transmembrane segment, and a 179 aa extracellular domain (ECD) (3). This 26 kDa protein is assembled intracellularly to form a noncovalently linked homotrimer (4). Cleavage of membrane bound TNF- $\alpha$  by TACE/ADAM17 releases a 55 kDa soluble trimeric form of TNF- $\alpha$  (5, 6). Within the cleaved ECD, rabbit TNF- $\alpha$  shares 76% - 83% with bovine, canine, cotton rat, equine, feline, human, mouse, porcine, rat, and rhesus TNF- $\alpha$ . Ligation of the membrane bound trimeric complex induces reverse signaling that promotes lymphocyte costimulation but diminishes monocyte responsiveness (7). TNF- $\alpha$  trimers bind a ubiquitous TNF RI as well as a hematopoietic cell-restricted TNF RII, both of which are also expressed as preformed homotrimers (1, 8). TNF- $\alpha$  regulates lymphoid tissue development through control of apoptosis (2). It also promotes inflammatory responses by inducing the activation of vascular endothelial cells and macrophages (2). TNF- $\alpha$  is a key cytokine in several inflammatory disorders (9). It contributes to the development of type 2 diabetes through its effects on insulin resistance and fatty acid metabolism (10, 11).

#### References:

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