

Recombinant Rhesus Macaque IL-18/IL-1F4

Catalog Number: 2548-RM

DESCRIPTION	
Source	E. coli-derived Tyr37-Asp193 Accession # AAK13416
N-terminal Sequence Analysis	Tyr37
Predicted Molecular Mass	18.2 kDa
SPECIFICATIONS	
Activity	Measured by its ability to induce IFN γ secretion by KG-1 human acute myelogenous leukemia cells in the presence of TNF- α . The ED ₅₀ for this effect is typically 1-4 ng/mL in the presence of 20 ng/mL rhTNF- α .
Endotoxin Level	<1.0 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 μ m filtered solution in MOPS, Na $_2$ SO $_4$, EDTA and DTT with BSA as a carrier protein. See Certificate of Analysis for details.

PREPARATION AND STORAGE	
Reconstitution	Reconstitute at 10 µg/mL in sterile PBS containing at least 0.1% human or bovine serum albumin.
Shipping	The product is shipped with polar packs. 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

Interleukin-18 (IL-18), also known as IL-1F4 and IFN γ inducing factor (IGIF), is a member of the IL-1 family of cytokines and is a key molecule in the innate immune response (1). Rhesus IL-18 is synthesized as a 24 kDa proprotein that contains a 36 amino acid (aa) propeptide and a 157 aa mature region (2). Under inflammatory conditions, the propeptide is cleaved by Caspase-1 in the cytoplasm to liberate the mature nonglycosylated 18 kDa monomeric IL-18 (3, 4). Mature rhesus IL-18 shares 96% aa sequence identity with human IL-18 and 60% - 76% with mouse, rat, canine, feline, and porcine IL-18. IL-18 is secreted by a variety of cell types including macrophages, dendritic cells, and epithelial cells (1, 5). Circulating mature IL-18 is sequestered by soluble IL-18 binding proteins (IL-18 BP) that inhibit IL-18 bioactivity (6). IL-18 interacts with the widely expressed IL-18 Ra which then recruits the signaling subunit IL-18 R β (7, 8). The IL-1 family member IL-1F7 also binds to IL-18 Ra but does not recruit IL-18 R β or induce signaling (9). IL-1F7 binds IL-18 BP and enhances its neutralizing effect on IL-18 activity (9). IL-18 synergizes with other cytokines to activate NK, Th1, and Th17 cells and to increase the production of IFN γ (1, 5, 10, 11, 12). IL-18 can also promote Th2 cytokine release which reduces the effectiveness of antiviral responses (13, 14). Increased levels of active IL-18 contribute to the severity of autoimmunity and hypertension, while deficiency of IL-18 results in symptoms of metabolic syndrome (1, 5, 15, 16). In cancer, IL-18 stimulates Th1 and NK cells to target tumor cells, but it can also promote angiogenesis, metastasis, and tumor cell immune evasion (11).

References:

- 1. Arend, W.P. et al. (2008) Immunol. Rev. 223:20.
- 2. Giavedoni, L.D. et al. (2001) J. Interferon Cytokine Res. 21:173.
- 3. Ghayur, T. *et al.* (1997) Nature **386**:619.
- 4. Gu, Y. et al. (1997) Science **275**:206.
- 5. Boraschi, D. and C.A. Dinarello (2006) Eur. Cytokine Netw. 17:224.
- 6. Novick, D. et al. (1999) Immunity 10:127.
- 7. Torigoe, K. et al. (1997) J. Biol. Chem. 272:25737.
- 8. Born, T.L. et al. (1998) J. Biol. Chem. 273:29445.
- 9. Bufler, P. et al. (2002) Proc. Natl. Acad. Sci. 99:13723.
- 10. Takeda, K. et al. (1998) Immunity 8:383.
- 11. Park, S. et al. (2007) Cell. Mol. Immunol. 4:329.
- 12. Yoshimoto, T. et al. (1998) J. Immunol. 161:3400.
- 13. Hoshino, T. et al. (2001) J. Immunol. 166:7014.
- 14. Iannello, A. et al. (2009) AIDS Rev. 11:115.
- 15. Rabkin, S.W. (2009) Nat. Clin. Pract. Cardiovasc. Med. 6:192.
- Netea, M.G. et al. (2006) Nat. Med. 12:650.

