

DESCRIPTION

Source	Human embryonic kidney cell, HEK293-derived cynomolgus monkey TNF RI/TNFRSF1A protein		
	Cynomolgus Monkey TNF RI/TNFRSF1A (Leu30-Thr211) Accession # NP_001306550	IEGRMD	Human IgG ₁ (Pro100-Lys330)
	N-terminus		C-terminus
N-terminal Sequence Analysis	Leu30		
Structure / Form	Disulfide-linked homodimer		
Predicted Molecular Mass	47 kDa		

SPECIFICATIONS

SDS-PAGE	55-66 kDa, reducing conditions
Activity	Measured by its ability to inhibit the TNF- α mediated cytotoxicity in the L-929 mouse fibroblast cells in the presence of the metabolic inhibitor actinomycin D. Matthews, N. and M.L. Neale (1987) in <i>Lymphokines and Interferons, A Practical Approach</i> . Clemens, M.J. <i>et al.</i> (eds): IRL Press. 221. The ED ₅₀ for this effect is 0.6-3.6 ng/mL.
Endotoxin Level	<0.10 EU per 1 μ g of the protein by the LAL method.
Purity	>95%, by SDS-PAGE visualized with Silver Staining and quantitative densitometry by Coomassie® Blue Staining.
Formulation	Lyophilized from a 0.2 μ m filtered solution in PBS. See Certificate of Analysis for details.

PREPARATION AND STORAGE

Reconstitution	Reconstitute at 500 μ g/mL in PBS.
Shipping	The product is shipped at ambient temperature. Upon receipt, store it immediately at the temperature recommended below.
Stability & Storage	<ul style="list-style-type: none"> • 12 months from date of receipt, \leq -20 °C as supplied. • 1 month, 2 to 8 °C under sterile conditions after reconstitution. • 3 months, \leq -20 °C under sterile conditions after reconstitution.

DATA

<p>Inhibition Activity</p> <p>Recombinant Cynomolgus TNF RI (ng/mL)</p>	<p>Recombinant Cynomolgus Monkey TNF RI/TNFRSF1A Fc Chimera (Catalog # 9884-TR) inhibits Recombinant Human TNF-α (Catalog # 210-TA) mediated cytotoxicity in the L-929 mouse fibroblast cells. The ED₅₀ for this effect is 0.6-3.6 ng/mL.</p>	<p>SDS-PAGE</p> <p>2 μg/lane of Recombinant Cynomolgus Monkey TNF RI/TNFRSF1A was resolved with SDS-PAGE under reducing (R) and non-reducing (NR) conditions and visualized by Coomassie® Blue staining, showing bands at 55-66 kDa and 110-130 kDa, respectively.</p>
---	--	--

BACKGROUND

TNF receptor 1 is a 55 kDa type I transmembrane protein member of the TNF receptor superfamily, designated TNFRSF1A (1, 2). TNF RI is a 455 amino acid (aa) protein that contains a signal sequence and ECD with a PLAD (pre-ligand assembly domain) that mediates constitutive dimer/trimer formation, followed by four CRD (cysteine-rich domains), a transmembrane domain, and a cytoplasmic domain that contains a neutral sphingomyelinase activation domain and a death domain (3, 4). The ECD of cynomolgus TNF RI shows 97%, 69%, and 68% aa identity with human, mouse and rat TNF RI, respectively. Both TNF RI and TNF RII (TNFRSF1B) are widely expressed and contain four TNF-alpha trimer-binding CRD in their ECD. However, TNF RI is thought to mediate most of the cellular effects of TNF-alpha (3). TNF RI is essential for proper development of lymph node germinal centers and Peyer's patches, and for combating intracellular pathogens such as *Listeria* (1, 2, 5). TNF RI is also a receptor for TNF-beta/TNFSF1B (lymphotoxin-alpha) (6). TNF RI is stored in the Golgi and translocates to the cell surface following pro-inflammatory stimuli (7). TNF-alpha stabilizes TNF RI and induces its sequestering in lipid rafts, where it activates NF kappa B and is cleaved by ADAM-17/TACE (8, 9, 16). Release of the 28-34 kDa TNF RI ECD also occurs constitutively and in response to products of pathogens such as LPS, CpG DNA or *S. aureus* protein A (1, 10-12). Full-length TNF RI may also be released in exosome-like vesicles (13). Release helps to resolve inflammatory reactions, since it down-regulates cell surface TNF RI and provides soluble TNF RI to bind TNF-alpha (10, 14-15). Exclusion from lipid rafts causes endocytosis of TNF RI complexes and induces apoptosis (1). Mutations of human TNF R1 can result in inflammatory episodes known as TRAPS (TNFR-associated periodic syndrome) (7).

References:

1. Pfeffer, K. (2003) Cytokine Growth Factor Rev. **14**:185.
2. Hehlgans, T. and K. Pfeffer (2005) Immunology **115**:1.
3. Chan, F.K. *et al.* (2000) Science **288**:2351.
4. Schall, T.J. *et al.* (1990) Cell **61**:361.
5. Peschon, J.J. *et al.* (1998) J. Immunol. **160**:943.
6. Banner, D.W. *et al.* (1993) Cell **73**:431.
7. Turner, M.D. *et al.* (2012) Biosci. Rep. **32**:105.
8. Legler, D.F. *et al.* (2003) Immunity **18**:655.
9. Tellier, E. *et al.* (2006) Exp. Cell Res. **312**:3969.
10. Xanthoulea, S. *et al.* (2004) J. Exp. Med. **200**:367.
11. Jin, L. *et al.* (2000) J. Immunol. **165**:5153.
12. Gomez, M.I. *et al.* (2006) J. Biol. Chem. **281**:20190.
13. Islam, A. *et al.* (2006) J. Biol. Chem. **281**:6860.
14. Garton, K.J. *et al.* (2006) J. Leukoc. Biol. **79**:1105.
15. McDermott, M.F. *et al.* (1999) Cell **97**:133.