

**Recombinant Human TNFSF9 (4-1BBL), Animal Component-Free**

<b>Cat. No. :</b>	H064H
<b>Alternative Names:</b>	TNFSF9; 4-1BB ligand; 4-1BBL; TNFL9; Tumor necrosis factor ligand superfamily member 9; TNF Superfamily Member 9
<b>Species:</b>	Human
<b>Accession No.:</b>	P41273
<b>Expression System:</b>	HEK293
<b>Protein Sequence:</b>	Arg71-Glu254
<b>Theoretical MW:</b>	19.44 kDa
<b>Theoretical pI:</b>	6.22
<b>Tag:</b>	Tag-Free.
<b>Formulation buffer:</b>	PBS, 5% Mannitol and 0.01% Tween 80, pH7.4.
<b>Appearance:</b>	Lyophilized Powder.
<b>Purity:</b>	≥95% as determined by SDS-PAGE.
<b>Bioactivity:</b>	The activity was determined by its ability to induce IL-8 secretion in peripheral blood mononuclear cells (PBMCs).
<b>Endotoxin Level:</b>	≤0.05 EU/μg, as determined by the LAL assay.
<b>Application:</b>	Cell Culture; Activity Assays.

**Preparation & Storage**

<b>Reconstitution:</b>	<p>Reconstitute with sterile double-distilled water (ddH<sub>2</sub>O).</p> <p>⚠ Centrifuge the vial briefly before opening to ensure full recovery of the solution. Avoid vortexing and minimize vigorous pipetting to maintain protein stability.</p> <p>❄ Immediately aliquot the reconstituted protein solution and store under recommended conditions. Avoid repeated freeze-thaw cycles.</p>
<b>Shipping:</b>	Shipped on dry ice. Short-term transit on cold packs (2-8°C) is acceptable.
<b>Storage:</b>	<p>Use a manual defrost freezer and avoid repeated freeze-thaw cycles.</p> <ul style="list-style-type: none"> <li>● 12 months from date of receipt, -20 to -80°C as supplied.</li> <li>● 2-7 days at 2 to 8°C under sterile conditions after reconstitution.</li> <li>● 3-6 months at -20 to -80°C under sterile conditions after reconstitution.</li> </ul>

**Protein Description**

**Background:** TNFSF9 (tumor necrosis factor superfamily member 9), commonly designated 4-1BB ligand (4-1BBL) or CD137L, is a type II transmembrane glycoprotein critical for adaptive immune regulation. Synthesized as a 254-amino acid precursor, it forms homotrimers on the surface of activated antigen-presenting cells (dendritic cells, macrophages, B cells), endothelial cells, and inflamed tissues; proteolytic cleavage releases a soluble trimeric form. Binding to its receptor 4-1BB (TNFRSF9/CD137) on activated CD8<sup>+</sup> T cells, CD4<sup>+</sup> T cells, NK cells, and regulatory T cells triggers TRAF1/2-dependent NF-κB and MAPK signaling, enhancing T cell proliferation, survival (via Bcl-xL/Bfl-1 upregulation), IFN-γ/IL-2 production, cytotoxic activity, and memory differentiation. The axis also augments NK cell function and can modulate regulatory T cell activity context-dependently. Physiologically vital for antiviral and antitumor immunity, dysregulation contributes to autoimmune disorders (e.g., rheumatoid arthritis, lupus), graft-versus-host disease, and tumor immune evasion. Clinically, agonistic anti-4-1BB antibodies (urelumab, utomilumab) are under active investigation in oncology—often combined with PD-1/PD-L1 inhibitors—while recombinant 4-1BBL fusion proteins and bispecific engagers targeting TNFSF9 are emerging strategies. Hepatotoxicity with early systemic agonists has spurred development of tumor-localized delivery platforms to improve therapeutic windows.

**References:**

1. Chalupny NJ, Smith CA, Armitage RJ, et al. Inducible expression of the gp39 homologue 4-1BBL in human T cells. Proc Natl Acad Sci U S A. 1992;89(22):10360-10364.
2. Shuford WW, Klussman K, Tritchler DD, et al. 4-1BB costimulatory signals preferentially induce CD8<sup>+</sup> T cell proliferation and lead to the amplification in vivo of cytotoxic T cell responses. J Exp Med. 1997;186(1):47-55.
3. Watts TH. TNF/TNFR family members in costimulation of T cell responses. Annu Rev Immunol. 2005;23:23-68.
4. Chester C, Marabelle A, Houot R, et al. Immunotherapy targeting 4-1BB: mechanistic rationale, clinical results, and future strategies. Blood. 2018;131(1):49-57.

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