

## Product Information

### Fas Ligand, soluble, human recombinant, expressed in HEK 293 cells

Catalog Number **S8689**

Storage Temperature  $-20\text{ }^{\circ}\text{C}$

Synonym: *SuperFas* Ligand™

#### Product Description

This product is produced in human embryo kidney cells by fusing the extracellular domain of human FasL, fragment 103–281 at the N-terminus to a 26 amino acid linker peptide and a FLAG-tag. The fusion protein migrates on SDS-PAGE under reducing conditions as a 32 kDa protein for the nonglycosylated form and 35 kDa for the glycosylated form of the protein. Glycosylation of recombinant protein is similar or identical to natural human FasL and *SuperFas* Ligand recognizes the Fas receptor (CD95) on human, rat, and mouse cells.

The Fas ligand (FasL) is a cytokine that binds to the Fas antigen, a receptor that transduces the apoptotic signal into cells. FasL is a type II membrane protein belonging to the TNF family and predominantly expressed on activated T cells and NK cells. The Fas/FasL system modulates the immune response by inducing cell apoptosis to maintain homeostasis and in the regulation of immune responses and privilege.<sup>1</sup> The cysteine-rich repeats of the Fas receptor are required for binding by the Fas ligand.<sup>2</sup> Binding of the Fas ligand induces trimerization of Fas in the target cell membrane. Activation of Fas causes the recruitment of Fas-associated protein with death domain (FADD) via interactions between the death domains of Fas and FADD.<sup>3</sup> Pro-caspase 8 binds to Fas-bound FADD via interactions between the death effector domains (DED) of FADD and pro-caspase 8 leading to the activation of caspase 8. Activated caspase 8 cleaves (activates) nine other procaspases, a process that ultimately leads to apoptosis. B cell antigen receptor signaling inhibits Fas-mediated apoptosis via up-regulation of cellular FLICE-inhibitory protein (c-FLIP).<sup>4</sup> The expression of the Fas ligand is regulated by protein phosphatases(s) sensitive to okadaic acid.<sup>5</sup> Serum withdrawal-induced apoptosis is mediated partially by the Fas/FasL interactions.<sup>6</sup>

This product is supplied as a lyophilized powder containing phosphate buffered saline salts.

Purity:  $\geq 95\%$  (SDS-PAGE)

The ability of this product to induce apoptosis of A20B lymphoma cells has been determined ( $\text{ED}_{50} = 1\text{ ng/ml}$ ). It does not require any enhancer since the modified linker sequence facilitates the formation of active FasL complexes.

#### Precautions and Disclaimer

This product is for R&D use only, not for drug, household, or other uses. Please consult the Material Safety Data Sheet for information regarding hazards and safe handling practices.

#### Preparation Instructions

Reconstitute the product with 50  $\mu\text{l}$  of sterile water to a concentration of 0.1 mg/ml. Further dilutions should be made with cell culture medium containing 5% fetal calf serum or phosphate buffered saline containing at least 0.1% human serum albumin or bovine serum albumin.

#### Storage/Stability

Store the product at  $-20\text{ }^{\circ}\text{C}$ .

Upon reconstitution, store under sterile conditions at  $2\text{--}8\text{ }^{\circ}\text{C}$  for one month or at  $-20\text{ }^{\circ}\text{C}$  to  $-70\text{ }^{\circ}\text{C}$  for three months without detectable loss of activity. Avoid repeated freeze-thaw cycles.

## References

1. Starling, G.C. et al., Identification of amino acid residues important for ligand binding to Fas. *J. Exp. Med.*, **185**, 1487-1492 (1997).
2. Orlinick, J.R. et al., Requirement of cysteine-rich repeats of the Fas receptor for binding by the Fas ligand. *J. Biol. Chem.*, **272**, 28889-28894 (1997).
3. Juo, P. et al., FADD is required for multiple signaling events downstream of the receptor Fas. *J. Cell Growth Differ.*, **10**, 797-804 (1999).
4. Wang, J. et al., Inhibition of Fas-mediated apoptosis by the B cell antigen receptor through c-FLIP. *Eur. J. Immunol.*, **30**, 155-163 (2000).
5. Morimoto, Y. et al., Upregulation of the expression of Fas antigen and Fas ligand in a human submandibular gland ductal cell line by okadaic acid. *Arch. Oral Biol.*, **45**, 657-666 (2000).
6. Hu, C.L., Apoptosis of bovine granulosa cells after serum withdrawal is mediated by Fas antigen (CD95) and Fas ligand. *Biol. Reprod.*, **64**, 518-526 (2001).

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