

# Human DLL4 / Delta4 Protein (His Tag)

Catalog Number: 10171-H08H



Sino Biological  
Biological Solution Specialist

## General Information

### Gene Name Synonym:

Delta-like 4; hdelta2

### Protein Construction:

A DNA sequence encoding the extracellular domain (Met 1-Pro 524) of human DLL4 (NP\_061947.1) pro-protein was expressed, with a polyhistidine tag at the C-terminus.

**Source:** Human

**Expression Host:** HEK293 Cells

## QC Testing

**Purity:** ≥ 98 % as determined by SDS-PAGE. ≥ 95 % as determined by SEC-HPLC.

### Bio Activity:

1. Immobilized Recombinant Human NOTCH1 Protein (Fc Tag) (Cat: 10954-H02H) at 2 µg/mL (100 µL/well) can bind Recombinant Human DLL4 / Delta4 Protein (His Tag) (Cat: 10171-H08H), the EC<sub>50</sub> is 100-300 ng/mL.
2. Measured by the ability of the immobilized protein to enhance BMP2-induced alkaline phosphatase activity in C3H10T1/2 mouse embryonic fibroblast cells. The ED<sub>50</sub> for this effect is typically 2-10 µg/mL in the presence of 500 ng/mL recombinant human BMP2.
3. Loaded Recombinant Human NOTCH1 Protein, hFc Tag (Cat. No. 10954-H02H) on ProA Biosensor, can bind Recombinant Human DLL4 Protein, His Tag (Cat. No. 10171-H08H) with an affinity constant of 51.7nM as determined in BLI assay (Sartorius Octet RED384) (Routinely tested).
4. Recombinant Human NOTCH1 Protein, hFc Tag (Cat. No. 10954-H02H) captured on Protein A chip can bind Recombinant Human DLL4 Protein, His Tag (Cat. No. 10171-H08H) with an affinity constant of 1.356 µM as determined in an SPR assay (Biacore T200) (Routinely tested).

### Endotoxin:

< 1.0 EU per µg of the protein as determined by the LAL method

### Stability:

Samples are stable for up to twelve months from date of receipt at -70 °C

**Predicted N terminal:** Ser 27

### Molecular Mass:

The recombinant human DLL4 consists of 509 amino acids and predicts a molecular mass of 55.7 kDa. As a result of glycosylation, the apparent molecular mass of rhDLL4 is approximately 55-70 kDa band in SDS-PAGE under reducing conditions.

### Formulation:

Lyophilized from sterile PBS, pH 7.4

Normally 5 % - 8 % trehalose, mannitol and 0.01% Tween80 are added as protectants before lyophilization. Specific concentrations are included in the hardcopy of COA. Please contact us for any concerns or special requirements.

## Usage Guide

### Storage:

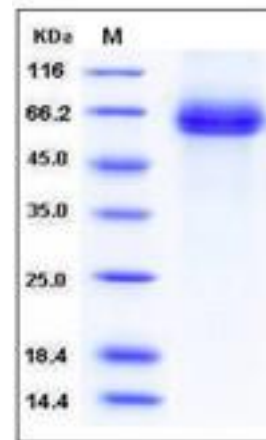
Store it under sterile conditions at -20°C to -80°C upon receiving. Recommend to aliquot the protein into smaller quantities for optimal storage.

**Avoid repeated freeze-thaw cycles.**

### Reconstitution:

Detailed reconstitution instructions are sent along with the products.

### SDS-PAGE:



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### Protein Description

Delta-like protein 4 (DLL4, Delta4), a type I membrane-bound Notch ligand, is one of five known Notch ligands in mammals and interacts predominantly with Notch 1, which has a key role in vascular development. Recent studies yield substantial insights into the role of DLL4 in angiogenesis. DLL4 is induced by vascular endothelial growth factor (VEGF) and acts downstream of VEGF as a 'brake' on VEGF-induced vessel growth, forming an autoregulatory negative feedback loop inactivating VEGF. DLL4 is downstream of VEGF signaling and its activation triggers a negative feedback that restrains the effects of VEGF. Attenuation of DLL4/Notch signaling results in chaotic vascular network with excessive branching and sprouting. DLL4 is widely distributed in tissues other than vessels including many malignancies. Furthermore, the molecule is internalized on binding its receptor and often transported to the nucleus. In pathological conditions, such as cancer, DLL4 is up-regulated strongly in the tumour vasculature. Blockade of DLL4-mediated Notch signaling strikingly increases nonproductive angiogenesis, but significantly inhibits tumor growth in preclinical mouse models. In preclinical studies, blocking of DLL4/Notch signaling is associated with a paradoxical increase in tumor vessel density, yet causes marked growth inhibition due to functionally defective vasculature. Thus, DLL4 blockade holds promise as an additional strategy for angiogenesis-based cancer therapy.

### References

1. Yan M, *et al.* (2007) Delta-like 4/Notch signaling and its therapeutic implications. *Clin Cancer Res.* 13(24): 7243-6. 2. Sainson RC, *et al.* (2007) Anti-Dll4 therapy: can we block tumour growth by increasing angiogenesis? *Trends Mol Med.* 13(9): 389-95. 3. Martinez JC, *et al.* (2009) Nuclear and membrane expression of the angiogenesis regulator delta-like ligand 4 (DLL4) in normal and malignant human tissues. *Histopathology.* 54(5): 598-606.

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