

DESCRIPTION

Source	Mouse myeloma cell line, NS0-derived		
	Mouse VE-Cadherin Asp46 - Gln592 Accession # 2208309A	IEGRMD	Human IgG ₁ (Pro100 - Lys330)
	N-terminus		C-terminus
N-terminal Sequence Analysis	Asp46		
Structure / Form	Disulfide-linked homodimer		
Predicted Molecular Mass	88.7 kDa (monomer)		

SPECIFICATIONS

SDS-PAGE	100-120 kDa, reducing conditions
Activity	Measured by its ability to inhibit proliferation of LL/2 mouse Lewis lung carcinoma cells. Immobilized rmVE-Cadherin/Fc Chimera inhibits LL/2 cell growth by 35-50%. The ED ₅₀ for this effect is 1.5-6.0 µg/mL.
Endotoxin Level	<1.0 EU per 1 µg of the protein by the LAL method.
Purity	>80%, by SDS-PAGE under reducing conditions and visualized by silver stain.
Formulation	Lyophilized from a 0.2 µm filtered solution in Tris-Citrate. See Certificate of Analysis for details.

PREPARATION AND STORAGE

Reconstitution	Reconstitute at 100 µg/mL in sterile PBS.
Shipping	The product is shipped at ambient temperature. Upon receipt, store it immediately at the temperature recommended below.
Stability & Storage	Use a manual defrost freezer and avoid repeated freeze-thaw cycles. <ul style="list-style-type: none"> ● 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

The cadherin (Ca⁺⁺-dependent adherence) superfamily is a large group of membrane-associated glycoproteins that engage in homotypic, calcium-dependent, cell-cell adhesion events. The superfamily can be divided into at least five major subfamilies based on molecule gene structure, and/or extracellular (EC) and intracellular domains (1, 2, 3, 4). Subfamilies include classical/type I, atypical/type II, and desmosomal-related cadherins (1, 2, 3). VE-Cadherin (vascular endothelial cadherin; also cadherin-5 and CD144) is a 125 kDa atypical/type II subfamily cadherin. Its subfamily classification is based principally on its genomic structure, as its physical structure is notably divergent from other type II subfamily members (2, 3). Mouse VE-Cadherin is synthesized as a 784 amino acid (aa) type I transmembrane (TM) preproprotein that contains a 24 aa signal peptide, a 21 aa prosequence, a 554 aa extracellular region (ECR), a 21 aa TM segment, and a 164 aa cytoplasmic domain (5, 6). The ECR contains five Ca⁺⁺-binding cadherin domains that are approximately 105 aa in length. Cadherin domains are comprised of two β-sheets that are oriented like bread in a sandwich. Although complex, the N-terminal cadherin domain mediates *trans* interactions, while the internal domains contribute to *cis* multimerizations (7). Mouse VE-Cadherin ECR is 92%, 77%, and 73% aa identical to rat, human and porcine VE-Cadherin ECR, respectively. VE-Cadherin is involved in the maintenance of endothelial permeability. In this regard, VE-Cadherin does not initiate new blood vessel formation; it maintains it once formed. Thus, when VE-Cadherin is downregulated, cells part and permeability increases (8). Notably, VEGF is known to promote vascular leakage, and apparently does so by inducing a β-arrestin-dependent endocytosis of VE-Cadherin (9). Part of this effect may be mediated by VE-Cadherin itself which is reported to increase the membrane half-life of VEGFR2 (10). VE-Cadherin acts homotypically at sites of zonula adherens. On each expressing cell, it is proposed that VE-Cadherin first forms a trimer, which then dimerizes with a trimeric counterpart *in-trans*. Alternatively, two *cis*-dimers could act *in-trans* to generate homotypic binding (11). In addition to cell adhesion, VE-Cadherin also is reported to mediate TGF-β receptor assembly. When clustered, VE-Cadherin enhances TβRII/TβRI assembly into an active receptor complex on endothelial cells (12). VE-Cadherin is expressed on endothelial cells, trophoblast cells, endothelial progenitor cells and embryonic hematopoietic cells (5, 8, 13, 14).

References:

1. Patel, S.D. *et al.* (2007) *Curr. Opin. Struct. Biol.* **13**:690.
2. Vestweber, D. (2008) *Arterioscler. Thromb. Vasc. Biol.* **28**:223.
3. Vincent, P.A. *et al.* (2004) *Am. J. Physiol. Cell. Physiol.* **286**:C987.
4. Cavallaro, U. *et al.* (2006) *Exp. Cell Res.* **312**:659.
5. Breier, G. *et al.* (1996) *Blood* **87**:630.
6. Huber, P. *et al.* (1996) *Genomics* **32**:21.
7. Pokutta, S. and W.I. Weis (2007) *Annu. Rev. Cell Dev. Biol.* **23**:237.
8. Crosby, C.V. *et al.* (2005) *Blood* **105**:2771.
9. Gavard, J. and J.S. Gutkind (2006) *Nat. Cell Biol.* **8**:1223.
10. Calera, M.R. *et al.* (2004) *Exp. Cell Res.* **300**:248.
11. Hewat, E.A. *et al.* (2007) *J. Mol. Biol.* **365**:744.
12. Rudini, N. *et al.* (2008) *EMBO J.* **27**:993.
13. Kogata, N. *et al.* (2006) *Circ. Res.* **98**:897.
14. Ema, M. *et al.* (2006) *Blood* **108**:4018.