

3050 Spruce Street Saint Louis, Missouri 63103 USA Telephone (800) 325-5832 (314) 771-5765 Fax (314) 286-7828 email: techserv@sial.com sigma-aldrich.com

# **ProductInformation**

ACTIVIN RECEPTOR IIB/Fc CHIMERA Human, Recombinant Expressed in insect Sf 21 cells

Product Number A 9579

## **Product Description**

Recombinant human Activin Receptor IIB is a chimeric protein expressed in insect Sf 21 cells. A DNA sequence encoding the signal peptide from CD33 and Ser19 - Thr134 from the extracellular domain of the human activin receptor IIB protein' is fused to the carboxy-terminal 6X histidine-tagged Fc region of human IgG1 via a polypeptide linker. Recombinant human activin receptor IIB/Fc chimera, generated after removal of amino acid residues (Met1 to Ala16) of the CD33 signal peptide, is a disulfide-linked homodimeric protein. This protein contains 360 amino acid residues with a calculated molecular mass of approximately 41 kDa. Due to glycosylation, the monomer migrates as a 55 kDa to 60 kDa protein in SDS-PAGE under reducing conditions. Human, mouse, and rat type II activin receptors share greater than 98 % homology.

Activin is secreted by Sertoli<sup>2</sup> cells in the testis and grandulosa cell in the ovary. In early studies, this peptide was thought to be an inhibin and not recognized as a unique compound.<sup>3,4</sup> Activins and inhibins are members of the TGF- $\beta$  superfamily due to amino acid homology with respect to the conservation of 7 of the 9 cysteine residues common to all TGF- $\beta$  forms.<sup>4</sup> Activins are homodimers or heterodimers of the various  $\beta$  subunit isoforms, while inhibins are heterodimers of a unique  $\alpha$  subunit and one of the various  $\beta$  subunits.<sup>5</sup> Five  $\beta$  subunits have been cloned (mammalian  $\beta_{A}$ ,  $\beta_{B}$ ,  $\beta_{C}$ ,  $\beta_{E}$ , and Xenopus  $\beta_{D}$ ).<sup>4</sup> The activin/inhibin nomenclature reflects the subunit composition of the proteins: activin A ( $\beta_{A}$ - $\beta_{A}$ ), activin B ( $\beta_{B}$ - $\beta_{B}$ ), activin AB ( $\beta_{B}$ - $\beta_{A}$ ), inhibin A ( $\alpha$ - $\beta_{A}$ ), and inhibin B ( $\alpha$ - $\beta_{B}$ ).

Activins have a wide range of biological activities including mesoderm induction<sup>6,7</sup>, neural cell differentiation, bone remodeling, hematopoiesis, and reproductive physiology. Activins are involved in growth and differentiation of several tissues from different species.<sup>2,3,7,8</sup> This protein also plays a key role in the

production and regulation of hormones such as FSH, LH, GnRH, and ACTH. Activin influences erythropoiesis and the potentiation of erythroid colony formation, oxytocin secretion, paracrine, and autocrine regulation.<sup>3</sup>

Similar to other TGF- $\beta$  family members, activins exert their biological activities through the effects of the heterodimeric complex composed of two membrane spanning serine-threonine kinases designated type I and type II receptors.<sup>9</sup> Activin type I and type II receptors are distinguished by the level of sequence homology of their kinase domains and other structural and functional features. To date, seven type I and five type II activin receptors have been cloned from mammals, including activin receptor IA, activin receptor IIA, activin receptor IIA, and five splice variants of activin receptor IIB have been reported.<sup>10</sup>

Type I activin receptors do not bind directly to activin but will associate with the type II receptor-activin complex and initiate signal transduction.<sup>11</sup> Recombinant soluble activin type II receptors are highly conserved, bind activin with high affinity, and are potent activin antagonists. Activin receptor IIB will also bind inhibin, BMP-2, and BMP-7 with lower affinities.

## Reagent

Activin Receptor IIB is supplied as approximately 100  $\mu$ g of protein lyophilized from a 0.2  $\mu$ m filtered solution in phosphate buffered saline (PBS).

#### **Preparation Instructions**

Reconstitute the contents of the vial using sterile phosphate-buffered saline (PBS) containing 0.1 % human serum albumin or bovine serum albumin. Prepare a stock solution of no less than 50 µg/ml.

#### Storage/Stability

Store at -20 °C. Upon reconstitution, store at 2 °C to 8 °C for one month. For extended storage, freeze in working aliquots at. Repeated freezing and thawing is not recommended.

## **Product Profile**

Activin Receptor IIB/Fc Chimera is measured by its ability to inhibit hemoglobin expression induced by activin in K562 erythroleukemia cells.<sup>1</sup> Approximately 0.02  $\mu$ g/ml to 0.06  $\mu$ g/ml of recombinant human activin receptor IIB/Fc will inhibit 50 % of the biological response due to 3 ng/ml of recombinant human activin A.

Purity: >95 % as determined by SDS-PAGE, visualized by silver stain.

Endotoxin: <0.1 ng/ $\mu$ g determined by the LAL method.

## References

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