

## **Biotinylated Recombinant Human BMP-6**

Catalog Number: BT507/CF

DESCRIPTION	
Source	Mouse myeloma cell line, NS0-derived Gln382-His513 Accession # P22004
N-terminal Sequence Analysis	No results obtained. Gln382 expected
Structure / Form	Disulfide-linked homodimer, biotinylated via sugars
Predicted Molecular Mass	15 kDa (unlabeled)
SPECIFICATIONS	
Activity	Measured by its ability to induce alkaline phosphatase production by ATDC5 mouse chondrogenic cells. Nakamura, K. <i>et al.</i> (1999) Exp. Cell Res. <b>250</b> :351.  The ED <sub>50</sub> for this effect is 0.02-0.15 μg/mL.
Endotoxin Level	<0.10 EU per 1 µg of the protein by the LAL method.
Purity	>95%, by SDS-PAGE visualized with Silver Staining and quantitative densitometry by Coomassie® Blue Staining.
Formulation	Lyophilized from a 0.2 µm filtered solution in HCl. See Certificate of Analysis for details.

PREPARATION AND STORAGE	
Reconstitution	Reconstitute at 100 μg/mL in 4 mM HCl.
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.  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

Bone Morphogenetic Protein 6 (BMP-6), also known as Vgr-1, is a member of the BMP subfamily of TGF-β superfamily proteins. BMPs are involved in a wide range of processes including embryogenesis, tissue morphogenesis, cell differentiation and migration, and tumorigenesis (1). Human BMP-6 is synthesized as a 513 amino acid (aa) precursor protein that is cleaved at the dibasic cleavage site (RxxR) to release the 18 kDa C-terminal mature protein. Biologically active BMP-6 consists of a disulfide-linked homodimer of the mature protein, although it can also form heterodimers with mature BMP-2 (2, 3). Mature human BMP-6 shares 96% and 98% aa sequence identity with mouse and rat BMP-6, respectively. Cellular responses to BMP-6 are mediated by hetero-oligomeric complexes of type I (Activin RIA/ALK-2 and BMPR-IA/ALK-3) and type II (Activin RIIA and BMPR-II) serine/threonine kinase receptors (4, 5). BMP-6 induces the expression of Noggin and is subsequently antagonized by Noggin (6). BMP-6 induces a wide range of cellular responses. It promotes osteoblast differentiation from mesenchymal stem cells (7), chondrocyte maturation (8), Ang II-induced aldosterone production in the adrenal cortex (4), hormone production and responsiveness in ovarian granulosa cells (9), iNOS and TNF-α production in macrophages (5), the cell death of B cells (10), and neurite outgrowth (11). BMP-6 expression is induced in astrocytes surrounding sites of brain injury where it functions as a neuroprotectant (11, 12). It enhances tumor progression by promoting local angiogenesis and differentiation of immune tolerizing M2 macrophages (13-15). Through interactions with the BMP co-receptor RGM-C/Hemojuvelin, BMP-6 plays an important role in iron homeostasis by promoting Hepcidin expression and preventing serum iron overload (16). Heterodimers of BMP-2 and BMP-6 show increased potency at inducing osteoblastic calcium deposition, chondrogenesis, and *in vivo* bone formation compared to either BMP-2 or BMP-6 homodimers (3).

## References:

- 1. Bragdon, B. et al. (2010) Cell Signal. 23:609.
- 2. Celeste, A.J. et al. (1990) Proc. Natl. Acad. Sci. USA 87:9843.
- 3. Israel, D.I. et al. (1996) Growth Factors 13:291.
- 4. Inagaki, K. et al. (2006) Endocrinology 147:2681.
- 5. Hong, J.H. et al. (2008) Immunology **128**:e442.
- 6. Haudenschild, D.R. et al. (2004) Cancer Res. 64:8276.
- 7. Lavery, K. et al. (2008) J. Biol. Chem. 283:20948.
- 8. Grimsrud, C.D. et al. (1999) J. Bone Miner. Res. 14:475.
- 9. Shi, J. et al. (2009) Fertil. Steril. 92:1794.
- 10. Kersten, C. et al. (2005) BMC Immunol. **6**:9
- 11. Yabe, T. et al. (2002) J. Neurosci. Res. 68:161.
- 12. Zhang, Z. et al. (2006) Neuroscience **138**:47.
- 13. Dai, J. et al. (2005) Cancer Res. 65:8274.
- Kwon, S.J. et al. (2014) Prostate 74:121.
- 15. Lee, J.-H. et al. (2013) Cancer Res. 73:3604.
- Andriopoulos, B. Jr. et al. (2009) Nat. Genet. 41:482.

