# abcam

## Product datasheet

## Recombinant Human Smad3 protein ab151882

1 References 1 Image

**Description** 

Product name Recombinant Human Smad3 protein

**Purity** > 80 % Densitometry.

**Expression system** Escherichia coli

Accession P84022

Protein length Full length protein

Animal free No

Nature Recombinant

**Species** Human

**Sequence** MSSILPFTPPIVKRLLGWKKGEQNGQEEKWCEKAVKSLV

KKLKKTGQLDE LEKAITTQNV

NTKCITIPRSLDGRLQVSHRKGLPHVIYCRLWRWPDLH

SHHELRAMELCEFAFNMKKDEV CVNPYHYQRVETPVLPPVLVPRHTEI

PAEFPPLDDYSHSIPENTNFPAGIEPQSNIPETP

**PPGYLSEDGETSDH** 

QMNHSMDAGSPNLSPNPMSPAHNNLDLQPVTYCEPAFW

CSISYYELNQRV

GETFHASQPSMTVDGFTDPSNSERFCLGLLSNVNRNAAV

**ELTRRHIGRGV** 

RLYYIGGEVFAECLSDSAIFVQSPNCNQRYGWHPATVCKI

**PPGCNLKIFN** 

NQEFAALLAQSVNQGFEAVYQLTRMCTIRMSFVKGWGAE

YRRQTVTSTPC

WIELHLNGPLQWLDKVLTQMGSPSIRCSSVS

Predicted molecular weight 77 kDa including tags

Amino acids 1 to 425

Tags GST tag N-Terminus

**Specifications** 

Our Abpromise guarantee covers the use of ab151882 in the following tested applications.

The application notes include recommended starting dilutions; optimal dilutions/concentrations should be determined by the end user.

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**Applications** Western blot

Form Liquid

Additional notes ab64311 (Myelin Basic Protein protein) can be utilized as a substrate for assessing kinase

activity

#### **Preparation and Storage**

Stability and Storage Shipped on dry ice. Upon delivery aliquot and store at -80°C. Avoid freeze / thaw cycles.

pH: 7.50

Constituents: 0.002% PMSF, 0.004% DTT, 0.79% Tris HCl, 25% Glycerol (glycerin, glycerine),

0.88% Sodium chloride

#### General Info

#### **Function**

Receptor-regulated SMAD (R-SMAD) that is an intracellular signal transducer and transcriptional modulator activated by TGF-beta (transforming growth factor) and activin type 1 receptor kinases. Binds the TRE element in the promoter region of many genes that are regulated by TGF-beta and, on formation of the SMAD3/SMAD4 complex, activates transcription. Also can form a SMAD3/SMAD4/JUN/FOS complex at the AP-1/SMAD site to regulate TGF-beta-mediated transcription. Has an inhibitory effect on wound healing probably by modulating both growth and migration of primary keratinocytes and by altering the TGF-mediated chemotaxis of monocytes. This effect on wound healing appears to be hormone-sensitive. Regulator of chondrogenesis and osteogenesis and inhibits early healing of bone fractures. Positively regulates PDPK1 kinase activity by stimulating its dissociation from the 14-3-3 protein YWHAQ which acts as a negative regulator.

Involvement in disease

Colorectal cancer

Loeys-Dietz syndrome 3

Sequence similarities

Belongs to the dwarfin/SMAD family.

Contains 1 MH1 (MAD homology 1) domain. Contains 1 MH2 (MAD homology 2) domain.

**Domain** 

The MH1 domain is required for DNA binding. Also binds zinc ions which are necessary for the  $\,$ 

DNA binding.

The MH2 domain is required for both homomeric and heteromeric interactions and for transcriptional regulation. Sufficient for nuclear import.

The linker region is required for the TGFbeta-mediated transcriptional activity and acts synergistically with the MH2 domain.

Post-translational modifications

Phosphorylated on serine and threonine residues. Enhanced phosphorylation in the linker region on Thr-179, Ser-204 and Ser-208 on EGF and TGF-beta treatment. Ser-208 is the main site of MAPK-mediated phosphorylation. CDK-mediated phosphorylation occurs in a cell-cycle dependent manner and inhibits both the transcriptional activity and antiproliferative functions of SMAD3. This phosphorylation is inhibited by flavopiridol. Maximum phosphorylation at the G(1)/S junction. Also phosphorylated on serine residues in the C-terminal SXS motif by TGFBR1 and ACVR1. TGFBR1-mediated phosphorylation at these C-terminal sites is required for interaction with SMAD4, nuclear location and transactivational activity, and appears to be a prerequisite for the TGF-beta mediated phosphorylation in the linker region. Dephosphorylated in the C-terminal SXS motif by PPM1A. This dephosphorylation disrupts the interaction with SMAD4, promotes nuclear export and terminates TGF-beta-mediated signaling. Phosphorylation at Ser-418 by

CSNK1G2/CK1 promotes ligand-dependent ubiquitination and subsequent proteasome degradation, thus inhibiting SMAD3-mediated TGF-beta responses. Phosphorylated by PDPK1. Acetylation in the nucleus by EP300 in the MH2 domain regulates positively its transcriptional activity and is enhanced by TGF-beta.

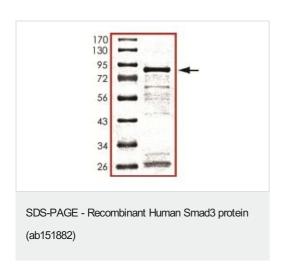
Ubiquitinated. Monoubiquitinated, leading to prevent DNA-binding. Deubiquitination by USP15 alleviates inhibition and promotes activation of TGF-beta target genes.

Poly-ADP-ribosylated by PARP1 and PARP2. ADP-ribosylation negatively regulates SMAD3 transcriptional responses during the course of TGF-beta signaling.

#### **Cellular localization**

Cytoplasm. Nucleus. Cytoplasmic and nuclear in the absence of TGF-beta. On TGF-beta stimulation, migrates to the nucleus when complexed with SMAD4 (PubMed:15799969). Through the action of the phosphatase PPM1A, released from the SMAD2/SMAD4 complex, and exported out of the nucleus by interaction with RANBP1 (PubMed:16751101, PubMed:19289081). Co-localizes with LEMD3 at the nucleus inner membrane (PubMed:15601644). MAPK-mediated phosphorylation appears to have no effect on nuclear import (PubMed:19218245). PDPK1 prevents its nuclear translocation in response to TGF-beta (PubMed:17327236).

#### **Images**



SDS-PAGE analysis of ab151882 which was determined to be >80% pure by densitometry.

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