abcam

Product datasheet

Recombinant human Insulin Receptor protein ab80251

画像数 2

製品の詳細

製品名 Recombinant human Insulin Receptor protein

生理活性 Activity: 752 pmol/min/µg. Assay conditions: 40 mM Tris-HCL pH 7.4, 20 mM MgCl2, 0.1 mg/mL

BSA and 0.2 mM DTT using 0.1 mg/ml Alextide substrate and 20 mM ATP. Reaction was done at

30°C for 40 min.

精製度 > 57 % SDS-PAGE.

Affinity purified.

発現系 Baculovirus infected Sf9 cells

アクセッション番号 <u>P06213</u>

タンパク質長 Protein fragment

Animal free No

由来 Recombinant

生物種 Human

配列 MSPILGYWKIKGLVQPTRLLLEYLEEKYEEHLYERDEGDKWR

NKKFELGL

EFPNLPYYIDGDVKLTQSMAIIRYIADKHNMLGGCPKERAEI

SMLEGAVL

DIRYGVSRIAYSKDFETLKVDFLSKLPEMLKMFKDRLCHKTY

LNGDHVTH

PDFMLYDALDVVLYMDPMCLDAFPKLVCFKKRIEAIPQIDKY

LKSSKYIA

WPLQGWQATFGGGDHPPKSDPAGSAAAVLEENLYFQGSFTMY

VPDEWEVS

REKITLLRELGQGSFGMVYEGNARDIIKGEAETRVAVKTVNE

SASLRERI

EFLNEASVMKGFTCHHVVRLLGVVSKGQPTLVVMELMAHGDL

KSYLRSLR

PEAENNPGRPPPTLQEMIQMAAEIADGMAYLNAKKFVHRDLA

ARNCMVAH

DFTVKIGDFGMTRDIYETDYYRKGGKGLLPVRWMAPESLKDG

VFTTSSDM

WSFGVVLWEITSLAEQPYQGLSNEQVLKFVMDGGYLDQPDNC

PERVTDLM

RMCWQFNPKMRPTFLEIVNLLKDDLHPSFPEVSFFHSEENKA

PESEELEM

1

EFEDMENVPLDRSSHCQREEAGGRDGGSSLGFKRSYEEHIPY THMNGGKK NGRILTLPRSNPS

予測される分子量 70 kDa including tags

領域 999 to 1370

タグ GST tag N-Terminus

配列の追加情報 70 kDa including tag

特性

Our Abpromise quarantee covers the use of ab80251 in the following tested applications.

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

アプリケーション Functional Studies

SDS-PAGE

製品の状態 Liquid

前処理および保存

保存方法および安定性

Shipped on Dry Ice. Upon delivery aliquot. Store at -80°C. Avoid freeze / thaw cycle.

pH: 8.00

Constituents: 0.0462% (R*,R*)-1,4-Dimercaptobutan-2,3-diol, 0.395% Tris HCl, 0.05% Tween,

30% Glycerol (glycerin, glycerine), 0.58% Sodium chloride

This product is an active protein and may elicit a biological response in vivo, handle with caution.

関連情報

機能

Receptor tyrosine kinase which mediates the pleiotropic actions of insulin. Binding of insulin leads to phosphorylation of several intracellular substrates, including, insulin receptor substrates (IRS1, 2, 3, 4), SHC, GAB1, CBL and other signaling intermediates. Each of these phosphorylated proteins serve as docking proteins for other signaling proteins that contain Src-homology-2 domains (SH2 domain) that specifically recognize different phosphotyrosines residues, including the p85 regulatory subunit of PI3K and SHP2. Phosphorylation of IRSs proteins lead to the activation of two main signaling pathways: the PI3K-AKT/PKB pathway, which is responsible for most of the metabolic actions of insulin, and the Ras-MAPK pathway, which regulates expression of some genes and cooperates with the PI3K pathway to control cell growth and differentiation. Binding of the SH2 domains of Pl3K to phosphotyrosines on IRS1 leads to the activation of Pl3K and the generation of phosphatidylinositol-(3, 4, 5)-triphosphate (PIP3), a lipid second messenger, which activates several PIP3-dependent serine/threonine kinases, such as PDPK1 and subsequently AKT/PKB. The net effect of this pathway is to produce a translocation of the glucose transporter SLC2A4/GLUT4 from cytoplasmic vesicles to the cell membrane to facilitate glucose transport. Moreover, upon insulin stimulation, activated AKT/PKB is responsible for: antiapoptotic effect of insulin by inducing phosphorylation of BAD; regulates the expression of gluconeogenic and lipogenic enzymes by controlling the activity of the winged helix or forkhead (FOX) class of transcription factors. Another pathway regulated by PI3K-AKT/PKB activation is mTORC1 signaling pathway which regulates cell growth and metabolism and integrates signals from insulin. AKT mediates insulin-stimulated protein synthesis by phosphorylating TSC2 thereby activating mTORC1 pathway. The Ras/RAF/MAP2K/MAPK pathway is mainly involved in

mediating cell growth, survival and cellular differentiation of insulin. Phosphorylated IRS1 recruits GRB2/SOS complex, which triggers the activation of the Ras/RAF/MAP2K/MAPK pathway. In addition to binding insulin, the insulin receptor can bind insulin-like growth factors (IGFI and IGFII). Isoform Short has a higher affinity for IGFII binding. When present in a hybrid receptor with IGF1R, binds IGF1. PubMed:12138094 shows that hybrid receptors composed of IGF1R and INSR isoform Long are activated with a high affinity by IGF1, with low affinity by IGF2 and not significantly activated by insulin, and that hybrid receptors composed of IGF1R and INSR isoform Short are activated by IGF1, IGF2 and insulin. In contrast, PubMed:16831875 shows that hybrid receptors composed of IGF1R and INSR isoform Long and hybrid receptors composed of IGF1R and INSR isoform Short have similar binding characteristics, both bind IGF1 and have a low affinity for insulin.

組織特異性

Isoform Long and isoform Short are predominantly expressed in tissue targets of insulin metabolic effects: liver, adipose tissue and skeletal muscle but are also expressed in the peripheral nerve, kidney, pulmonary alveoli, pancreatic acini, placenta vascular endothelium, fibroblasts, monocytes, granulocytes, erythrocytes and skin. Isoform Short is preferentially expressed in fetal cells such as fetal fibroblasts, muscle, liver and kidney. Found as a hybrid receptor with IGF1R in muscle, heart, kidney, adipose tissue, skeletal muscle, hepatoma, fibroblasts, spleen and placenta (at protein level). Overexpressed in several tumors, including breast, colon, lung, ovary, and thyroid carcinomas.

関連疾患

Rabson-Mendenhall syndrome

Leprechaunism

Diabetes mellitus, non-insulin-dependent Familial hyperinsulinemic hypoglycemia 5

Insulin-resistant diabetes mellitus with acanthosis nigricans type A

配列類似性

 $Belongs \ to \ the \ protein \ kinase \ superfamily. \ Tyr \ protein \ kinase \ family. \ Insulin \ receptor \ subfamily.$

Contains 3 fibronectin type-III domains. Contains 1 protein kinase domain.

ドメイン

The tetrameric insulin receptor binds insulin via non-identical regions from two alpha chains, primarily via the C-terminal region of the first INSR alpha chain. Residues from the leucine-rich N-terminus of the other INSR alpha chain also contribute to this insulin binding site. A secondary insulin-binding site is formed by residues at the junction of fibronectin type-III domain 1 and 2.

翻訳後修飾

After being transported from the endoplasmic reticulum to the Golgi apparatus, the single glycosylated precursor is further glycosylated and then cleaved, followed by its transport to the plasma membrane.

Autophosphorylated on tyrosine residues in response to insulin. Phosphorylation of Tyr-999 is required for binding to IRS1, SHC1 and STAT5B. Dephosphorylated by PTPRE at Tyr-999, Tyr-1185, Tyr-1189 and Tyr-1190. Dephosphorylated by PTPRF and PTPN1. Dephosphorylated by PTPN2; down-regulates insulin-induced signaling.

細胞内局在

Cell membrane.

画像

3

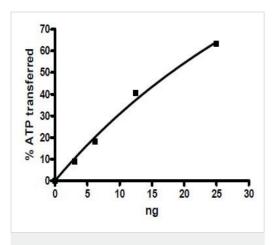
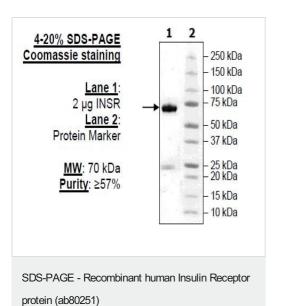


Image showing specific activity of ab80251.





SDS-PAGE anaylsis of ab80251 at approximately 70kDa (2µg).

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