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Insulin Receptor Protein (INSR) (AA 763-1382) (rho-1D4 tag)



Image



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Overview

Quantity:	1 mg
Target:	Insulin Receptor (INSR)
Protein Characteristics:	AA 763-1382
Origin:	Human
Source:	Insect Cells
Protein Type:	Recombinant
Purification tag / Conjugate:	This Insulin Receptor protein is labelled with rho-1D4 tag.
Application:	Western Blotting (WB), SDS-PAGE (SDS), ELISA, Crystallization (Crys)

Product Details

Sequence:

SLGDVGNVTV AVPTVAAFPN TSSTSVPTSP EEHRPFEKVV NKESLVISGL RHFTGYRIEL

QACNQDTPEE RCSVAAYVSA RTMPEAKADD IVGPVTHEIF ENNVVHLMWQ EPKEPNGLIV

LYEVSYRRYG DEELHLCVSR KHFALERGCR LRGLSPGNYS VRIRATSLAG NGSWTEPTYF

YVTDYLDVPS NIAKIIIGPL IFVFLFSVVI GSIYLFLRKR QPDGPLGPLY ASSNPEYLSA SDVFPCSVYV

PDEWEVSREK ITLLRELGQG SFGMVYEGNA RDIIKGEAET RVAVKTVNES ASLRERIEFL

NEASVMKGFT CHHVVRLLGV VSKGQPTLVV MELMAHGDLK SYLRSLRPEA ENNPGRPPPT

LQEMIQMAAE IADGMAYLNA KKFVHRDLAA RNCMVAHDFT VKIGDFGMTR DIYETDYYRK

GGKGLLPVRW MAPESLKDGV FTTSSDMWSF GVVLWEITSL AEQPYQGLSN EQVLKFVMDG

GYLDQPDNCP ERVTDLMRMC WQFNPKMRPT FLEIVNLLKD DLHPSFPEVS FFHSEENKAP

ESEELEMEFE DMENVPLDRS SHCQREEAGG RDGGSSLGFK RSYEEHIPYT HMNGGKKNGR

ILTLPRSNPS

Sequence without tag. Tag location is at the discretion of the manufacturer. If you have a

special request, please contact us.

Characteristics:

- Made in Germany from design to production by highly experienced protein experts.
- Human INSR Protein (raised in Insect Cells) purified by multi-step, protein-specific process to ensure crystallization grade.
- State-of-the-art algorithm used for plasmid design (Gene synthesis).

This protein is a made to order protein and will be made for the first time for your order. Our experts in the lab will ensure that you receive a correctly folded protein.

The big advantage of ordering our made-to-order proteins in comparison to ordering custom made proteins from other companies is that there is no financial obligation in case the protein cannot be expressed or purified.

In the unlikely event that the protein cannot be expressed or purified we do not charge anything (other companies might charge you for any performed steps in the expression process for custom-made proteins, e.g. fees might apply for the expression plasmid, the first expression experiments or purification optimization).

When you order this made-to-order protein you will only pay upon receival of the correctly folded protein. With no financial risk on your end you can rest assured that our experienced protein experts will do everything to make sure that you receive the protein you ordered.

The concentration of our recombinant proteins is measured using the absorbance at 280nm.

The protein's absorbance will be measured in several dilutions and is measured against its specific reference buffer.

The concentration of the protein is calculated using its specific absorption coefficient. We use the Expasy's protparam tool to determine the absorption coefficient of each protein.

Purification:

Three step purification of membrane proteins expressed in baculovirus infected SF9 insect cells:

- 1. Membrane proteins are fractioned by ultracentrifugation and subsequently solubilized with different detergents (detergent screen). Samples are analyzed by Western blot.
- 2. The best performing detergent is used for solubilization and the proteins are purified via their rho1D4 tag via two rho1D4 antibody columns: one DTT resistant, the other one not. Eluate fractions are analyzed by Western blot.
- Protein containing fractions of the best purification are subjected to second purification step through size exclusion chromatograph. Eluate fractions are analyzed by SDS-PAGE and Western blot.

Purity:

>95 % as determined by SDS PAGE, Size Exclusion Chromatography and Western Blot.

Sterility:

0.22 µm filtered

Endotoxin Level:

Protein is endotoxin-free.

Grade:

Crystallography grade

Target Details

Target:

Insulin Receptor (INSR)

Alternative Name:

INSR (INSR Products)

Background:

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 Srchomology-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 PI3K to phosphotyrosines on IRS1 leads to the activation of PI3K 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: anti-apoptotic 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

Target Details	
	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. {ECO:0000269 PubMed:12138094, ECO:0000269 PubMed:16314505,
	ECO:0000269 PubMed:16831875, ECO:0000269 PubMed:8257688,
	ECO:0000269 PubMed:8276809, ECO:0000269 PubMed:8452530,
	ECO:0000269 PubMed:9428692}.
Molecular Weight:	70.9 kDa Including tag.
UniProt:	P06213
Pathways:	NF-kappaB Signaling, RTK Signaling, AMPK Signaling, Carbohydrate Homeostasis, Regulation
	of Cell Size, Regulation of Carbohydrate Metabolic Process, Growth Factor Binding, Negative
	Regulation of Transporter Activity
Application Details	
Application Notes:	In addition to the applications listed above we expect the protein to work for functional studies
	as well. As the protein has not been tested for functional studies yet we cannot offer a gurantee
	though.
Comment:	In cases in which it is highly likely that the recombinant protein with the default tag will be
	insoluble our protein lab may suggest a higher molecular weight tag (e.g. GST-tag) instead to
	increase solubility. We will discuss all possible options with you in detail to assure that you
	receive your protein of interest.
Restrictions:	For Research Use only
Handling	
Format:	Liquid
Buffer:	100 mM NaCL, 20 mM Hepes, 10% glycerol. pH value is at the discretion of the manufacturer.
Handling Advice:	Avoid repeated freeze-thaw cycles.
Storage:	-80 °C
Storage Comment:	Store at -80°C.
Expiry Date:	Unlimited (if stored properly)

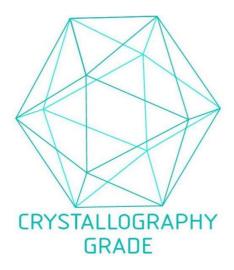


Image 1. "Crystallography Grade" protein due to multi-step, protein-specific purification process