abcam

Product datasheet

Recombinant human ALK (mutated R1275Q) protein ab186455

5 Images

Description

Product name Recombinant human ALK (mutated R1275Q) protein

Biological activity The specific activity of ab186455 was determined to be 19 nmol /min/mg as per activity assay

protocol.

Purity > 90 % Densitometry.

Affinity purified.

Expression system Baculovirus infected Sf9 cells

Accession Q9UM73

Protein length Protein fragment

Animal free No

Nature Recombinant

Species Human

Sequence RRKHQELQAMQMELQSPEYKLSKLRTSTIMTDYNPNYCFA

GKTSSISDLK

EVPRKNITLIRGLGHGAFGEVYEGQVSGMPNDPSPLQVAV

KTLPEVCSEQ

DELDFLMEALIISKFNHQNIVRCIGVSLQSLPRFILLELMAG

GDLKSFLR

ETRPRPSQPSSLAMLDLLHVARDIACGCQYLEENHFIHRDI

AARNCLLTC

PGPGRVAKIGDFGMAQDIYRASYYRKGGCAMLPVKWMPP

EAFMEGIFTSK

TDTWSFGVLLWEIFSLGYMPYPSKSNQEVLEFVTSGGRM

DPPKNCPGPVY

RIMTQCWQHQPEDRPNFAIILERIEYCTQDPDVINTALPIEY

GPLVEEEE

KVPVRPKDPEGVPPLLVSQQAKREEERSPAAPPPLPTTS

SGKAAKKPTAA

EISVRVPRGPAVEGGHVNMAFSQSNPPSELHKVHGSRNK

PTSLWNPTYGS

WFTEKPTKKNNPIAKKEPHDRGNLGLEGSCTVPPNVATG

RLPGASLLLEP

SSLTANMKEVPLFRLRHFPCGNVNYGYQQQGLPLEAATA

1

PGAGHYEDTIL KSKNSMNQPGP

Predicted molecular weight 90 kDa

Amino acids 1060 to 1620

Modifications mutated R1275Q

Tags proprietary tag N-Terminus

Specifications

Our **Abpromise guarantee** covers the use of **ab186455** in the following tested applications.

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

Applications SDS-PAGE

Functional Studies

Form Liquid

Preparation and Storage

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

pH: 7.5

Preservative: 1.02% Imidazole

Constituents: 0.81% Sodium phosphate, 1.75% Sodium chloride, 0.002% PMSF, 0.005% DTT,

25% Glycerol (glycerin, glycerine)

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

General Info

Function

Neuronal receptor tyrosine kinase that is essentially and transiently expressed in specific regions of the central and peripheral nervous systems and plays an important role in the genesis and differentiation of the nervous system. Transduces signals from ligands at the cell surface, through specific activation of the mitogen-activated protein kinase (MAPK) pathway. Phosphorylates almost exclusively at the first tyrosine of the Y-x-x-Y-Y motif. Following activation by ligand, ALK induces tyrosine phosphorylation of CBL, FRS2, IRS1 and SHC1, as well as of the MAP kinases MAPK1/ERK2 and MAPK3/ERK1. Acts as a receptor for ligands pleiotrophin (PTN), a secreted growth factor, and midkine (MDK), a PTN-related factor, thus participating in PTN and MDK signal transduction. PTN-binding induces MAPK pathway activation, which is important for the anti-apoptotic signaling of PTN and regulation of cell proliferation. MDK-binding induces phosphorylation of the ALK target insulin receptor substrate (IRS1), activates mitogen-activated protein kinases (MAPKs) and Pl3-kinase, resulting also in cell proliferation induction. Drives NF-kappa-B activation, probably through IRS1 and the activation of NF-kappa-B are essential for the autocrine growth and survival signaling of MDK.

Tissue specificity

Expressed in brain and CNS. Also expressed in the small intestine and testis, but not in normal lymphoid cells.

Involvement in disease

A chromosomal aberration involving ALK is found in a form of non-Hodgkin lymphoma. Translocation t(2;5)(p23;q35) with NPM1. The resulting chimeric NPM1-ALK protein homodimerize and the kinase becomes constitutively activated. The constitutively active fusion proteins are responsible for 5-10% of non-Hodgkin lymphomas.

A chromosomal aberration involving ALK is associated with inflammatory myofibroblastic tumors (IMTs). Translocation t(2;11)(p23;p15) with CARS; translocation t(2;4)(p23;q21) with SEC31A. A chromosomal aberration involving ALK is associated with anaplastic large-cell lymphoma (ALCL). Translocation t(2;17)(p23;q25) with ALO17.

Neuroblastoma 3

The ALK signaling pathway plays an important role in glioblastoma, the most common malignant brain tumor of adults and one of the most lethal cancers. It regulates both glioblastoma migration and growth.

A chromosomal aberration involving ALK is found in one subject with colorectal cancer.

Translocation t(2;2)(p23.1;p23.3). A 5 million base pair tandem duplication generates an in-frame WDCP-ALK gene fusion.

Sequence similarities

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

Contains 1 LDL-receptor class A domain.

Contains 2 MAM domains.

Contains 1 protein kinase domain.

Post-translational modifications

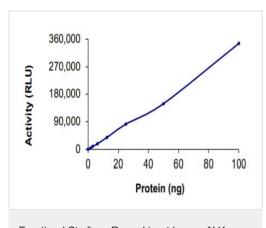
Phosphorylated at tyrosine residues by autocatalysis, which activates kinase activity. In cells not stimulated by a ligand, receptor protein tyrosine phosphatase beta and zeta complex (PTPRB/PTPRZ1) dephosphorylates ALK at the sites in ALK that are undergoing autophosphorylation through autoactivation. Phosphorylation at Tyr-1507 is critical for SHC1 association.

N-glycosylated.

Cellular localization

Cell membrane. Membrane attachment was crucial for promotion of neuron-like differentiation and cell proliferation arrest through specific activation of the MAP kinase pathway.

Images



Functional Studies - Recombinant human ALK (mutated R1275Q) protein (ab186455)

The specific activity of ALK (ab186455) was determined to be 10 nmol/min/mg as per activity assay protocol and was equivalent to 21 nmol/min/mg as per radiometric assay



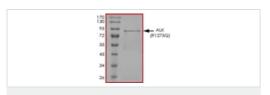
SDS PAGE analysis of ab186455

SDS-PAGE - Recombinant human ALK (mutated R1275Q) protein (ab186455)



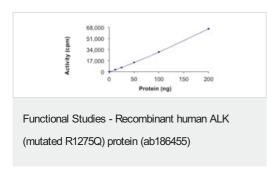
SDS PAGE analysis of ab186455

SDS-PAGE - Recombinant human ALK (mutated R1275Q) protein (ab186455)



SDS-PAGE analysis of ab186455.

SDS-PAGE - Recombinant human ALK (mutated R1275Q) protein (ab186455)



Kinase Assay demonstrating the specific activity of ab186455.

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