

Product Data Sheet

Product Name: KEAP1 Protein, Human (sf9)
Cat. No.: GP27282
Batch No.: 1

Product Data

Purity >98% Source
Physical Appearance solid Shipping Condition

Synonyms KEAP1 []; Kelch-like ECH-associated protein 1; Cytosolic inhibitor of Nrf2; KEAP1; KEAP1

Amino Acid Sequence QPDP RPSGAGACCRFLPLQSQPEGAGDAVMYASTECKAEVTPSQHG NRTFSYTL EDHTKQAFGIMNELRLSQQLCDVTLQVKYQDAPAAQFMAHKV

Formulation Lyophilized from a 0.22 µm filtered solution of 20 mM Tris, 500 mM NaCl, 3 mM DTT, 10% Glycerol, pH 7.4 or 50 mM Tris-HCl, 500 mM NaCl

Introduction

KEAP1 is a substrate-specific adapter within the BCR (BTB-CUL3-RBX1) E3 ubiquitin ligase complex, intricately regulates the cellular response to oxidative stress by orchestrating the ubiquitination of NFE2L2/NRF2. Serving as a crucial sensor for oxidative and electrophilic stress, KEAP1, under normal conditions, facilitates the ubiquitination and subsequent degradation of NFE2L2/NRF2, a transcription factor essential for the expression of numerous cytoprotective genes. When confronted with oxidative stress, distinct electrophile metabolites induce non-enzymatic covalent modifications on highly reactive cysteine residues in KEAP1, effectively dampening the ubiquitin ligase activity of the BCR(KEAP1) complex. This disruption promotes the nuclear accumulation of NFE2L2/NRF2 and triggers the expression of phase II detoxifying enzymes. Furthermore, selective autophagy leads to the sequestration of KEAP1 in inclusion bodies through its interaction with SQSTM1/p62, resulting in the inactivation of the BCR(KEAP1) complex and the activation of NFE2L2/NRF2. Notably, the BCR(KEAP1) complex extends its ubiquitin ligase activity to substrates like SQSTM1/p62, BPTF, and PGAM5, modulating their degradation via the proteasome. The ubiquitin ligase activity of the BCR(KEAP1) complex faces inhibition in response to oxidative stress and electrophile metabolites such as sulforaphane, as these metabolites react with reactive cysteine residues in KEAP1, leading to the non-enzymatic covalent modifications that incapacitate the complex. Moreover, selective autophagy contributes to the inactivation of the BCR(KEAP1) complex through the interaction between KEAP1 and SQSTM1/p62, promoting the sequestration of the complex in inclusion bodies and facilitating its degradation.

Biological Activity

Measured by its ability to inhibit the proliferation of A549 cells. The ED50 for this effect is 1.117-2.321 µg/mL.

Stability

Stored at -20°C for 2 years from date of receipt. After reconstitution, it is stable at 4°C for 1 week or -20°C for longer (with carrier protein). It is recommended to freeze aliquots at -20°C or -80°C for extended storage.

Protocol

KEAP1, as the substrate-specific adapter within the BCR (BTB-CUL3-RBX1) E3 ubiquitin ligase complex, intricately regulates the cellular response to oxidative stress by orchestrating the ubiquitination of NFE2L2/NRF2. Serving as a crucial sensor for oxidative and electrophilic stress, KEAP1, under normal conditions, facilitates the ubiquitination and subsequent degradation of NFE2L2/NRF2, a transcription factor essential for the expression of numerous cytoprotective genes. When confronted with oxidative stress, distinct electrophile metabolites induce non-enzymatic covalent modifications on highly reactive cysteine residues in KEAP1, effectively dampening the ubiquitin ligase activity of the BCR(KEAP1) complex. This disruption promotes the nuclear accumulation of NFE2L2/NRF2 and triggers the expression of phase II detoxifying enzymes. Furthermore, selective autophagy leads to the sequestration of KEAP1 in inclusion bodies through its interaction with SQSTM1/p62, resulting in the inactivation of the BCR(KEAP1) complex and the activation of NFE2L2/NRF2. Notably, the BCR(KEAP1) complex extends its ubiquitin ligase activity to substrates like SQSTM1/p62, BPTF, and PGAM5, modulating their degradation via the proteasome. The ubiquitin ligase activity of the BCR(KEAP1) complex faces inhibition in response to oxidative stress and electrophile metabolites such as sulforaphane, as these metabolites react with reactive cysteine residues in KEAP1, leading to the non-enzymatic covalent modifications that incapacitate the complex. Moreover, selective autophagy contributes to the inactivation of the BCR(KEAP1) complex through the interaction between KEAP1 and SQSTM1/p62, promoting the sequestration of the complex in inclusion bodies and facilitating its degradation.

Background

KEAP1 in the BCR E3 ubiquitin ligase complex complex regulates cellular responses to oxidative stress by ubiquitinating NFE2L2/NRF2. As an oxidative stress sensor, KEAP1 normally promotes NFE2L2/NRF2 degradation. KEAP1 Protein, Human (sf9) is the recombinant human-derived KEAP1 protein, expressed by Sf9 insect cells, with tag free.

Caution: Product has not been fully validated for medical applications. For research use only.
Tel: (909) 407-4943 Fax: (626) 353-8530 E-mail: tech@glpbio.com
Address: 10292 Central Ave. #205, Montclair, CA, USA