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# Cleaved-Caspase-4 p20 (Q81) Polyclonal Antibody

Catalog # AP63103

#### **Product Information**

Application WB, IHC-P
Primary Accession P49662
Reactivity Human
Host Rabbit
Clonality Polyclonal
Calculated MW 43262

#### **Additional Information**

Gene ID 837

Other Names CASP4; ICH2; Caspase-4; CASP-4; ICE(rel)-II; Protease ICH-2; Protease TX

**Dilution** WB~~Western Blot: 1/500 - 1/2000. Immunohistochemistry: 1/100 - 1/300.

ELISA: 1/40000. Not yet tested in other applications. IHC-P~~N/A

Format Liquid in PBS containing 50% glycerol, 0.5% BSA and 0.09% (W/V) sodium

azide.

Storage Conditions -20°C

### **Protein Information**

Name CASP4 {ECO:0000303 | PubMed:15123740, ECO:0000312 | HGNC:HGNC:1505}

**Function** Inflammatory caspase that acts as the effector of the non- canonical

inflammasome by mediating lipopolysaccharide (LPS)-induced pyroptosis

(PubMed:<u>25119034</u>, PubMed:<u>26375003</u>, PubMed:<u>32109412</u>,

PubMed:<u>34671164</u>, PubMed:<u>37001519</u>, PubMed:<u>37993712</u>,

PubMed: 37993714). Also indirectly activates the NLRP3 and NLRP6

inflammasomes (PubMed:<u>23516580</u>, PubMed:<u>26375003</u>, PubMed:<u>32109412</u>, PubMed:<u>7797510</u>). Acts as a thiol protease that cleaves a tetrapeptide after an

Asp residue at position P1: catalyzes cleavage of CGAS, GSDMD and IL18

(PubMed: <u>15326478</u>, PubMed: <u>23516580</u>, PubMed: <u>26375003</u>, PubMed: <u>28314500</u>, PubMed: <u>23100413</u>, PubMed: <u>27002713</u>

PubMed:<u>28314590</u>, PubMed:<u>32109412</u>, PubMed:<u>37993712</u>,

PubMed:<u>37993714</u>, PubMed:<u>7797510</u>). Effector of the non-canonical inflammasome independently of NLRP3 inflammasome and CASP1: the non-canonical inflammasome promotes pyroptosis through GSDMD cleavage

without involving secretion of cytokine IL1B (PubMed:25119034, PubMed:25121752, PubMed:26375003, PubMed:31268602,

PubMed:<u>32109412</u>, PubMed:<u>37993712</u>, PubMed:<u>37993714</u>). In the

non-canonical inflammasome, CASP4 is activated by direct binding to the lipid A moiety of LPS without the need of an upstream sensor (PubMed: 25119034,

PubMed: 25121752, PubMed: 29520027, PubMed: 32510692, PubMed:32581219, PubMed:37993712). LPS-binding promotes CASP4 activation and CASP4-mediated cleavage of GSDMD and IL18, followed by IL18 secretion through the GSDMD pore, pyroptosis of infected cells and their extrusion into the gut lumen (PubMed:25119034, PubMed:25121752, PubMed:37993712, PubMed:37993714). Also indirectly promotes secretion of mature cytokines (IL1A and HMGB1) downstream of GSDMD-mediated pyroptosis via activation of the NLRP3 and NLRP6 inflammasomes (PubMed: 26375003, PubMed: 32109412). Involved in NLRP3-dependent CASP1 activation and IL1B secretion in response to non-canonical activators, such as UVB radiation or cholera enterotoxin (PubMed:22246630, PubMed:23516580, PubMed:24879791, PubMed:25964352, PubMed:26173988, PubMed: 26174085, PubMed: 26508369). Involved in NLRP6 inflammasomedependent activation in response to lipoteichoic acid (LTA), a cell- wall component of Gram-positive bacteria, which leads to CASP1 activation and IL1B secretion (PubMed:33377178). Involved in LPS- induced IL6 secretion; this activity may not require caspase enzymatic activity (PubMed: 26508369). The non-canonical inflammasome is required for innate immunity to cytosolic, but not vacuolar, bacteria (By similarity). Plays a crucial role in the restriction of S.typhimurium replication in colonic epithelial cells during infection (PubMed:25121752, PubMed:25964352). Activation of the non-canonical inflammasome in brain endothelial cells can lead to excessive pyroptosis, leading to blood-brain barrier breakdown (By similarity). Pyroptosis limits bacterial replication, while cytokine secretion promotes the recruitment and activation of immune cells and triggers mucosal inflammation (PubMed:25121752, PubMed:25964352, PubMed:26375003). May also act as an activator of adaptive immunity in dendritic cells, following activation by oxidized phospholipid 1- palmitoyl-2-arachidonoylsn-glycero-3-phosphorylcholine, an oxidized phospholipid (oxPAPC) (By similarity). Involved in cell death induced by endoplasmic reticulum stress and by treatment with cytotoxic APP peptides found in Alzheimer's patient brains (PubMed: 15123740, PubMed: 22246630, PubMed: 23661706). Cleavage of GSDMD is not strictly dependent on the consensus cleavage site but depends on an exosite interface on CASP4 that recognizes and binds the Gasdermin-D, C- terminal (GSDMD-CT) part (PubMed:32109412). Catalyzes cleavage and maturation of IL18; IL18 processing also depends of the exosite interface on CASP4 (PubMed: 15326478, PubMed: 37993712, PubMed:37993714). In contrast, it does not directly process IL1B (PubMed:7743998, PubMed:7797510, PubMed:7797592). During non-canonical inflammasome activation, cuts CGAS and may play a role in the regulation of antiviral innate immune activation (PubMed:28314590).

**Cellular Location** 

Cytoplasm, cytosol. Endoplasmic reticulum membrane; Peripheral membrane protein; Cytoplasmic side. Mitochondrion Inflammasome. Secreted Note=Predominantly localizes to the endoplasmic reticulum (ER) Association with the ER membrane requires TMEM214 (PubMed:15123740) Released in the extracellular milieu by keratinocytes following UVB irradiation (PubMed:22246630).

**Tissue Location** 

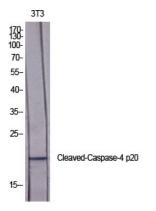
Widely expressed, including in keratinocytes and colonic and small intestinal epithelial cells (at protein level). Not detected in brain.

## **Background**

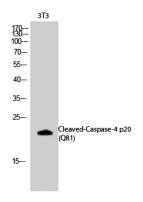
Inflammatory caspase (PubMed: <u>7797510</u>, PubMed: <u>23516580</u>, PubMed: <u>25119034</u>). Essential effector of NLRP3 inflammasome- dependent CASP1 activation and IL1B and IL18 secretion in response to non-canonical activators, such as UVB radiation, cholera enterotoxin subunit B and cytosolic LPS (PubMed: <u>22246630</u>, PubMed: <u>26174085</u>, PubMed: <u>26173988</u>, PubMed: <u>26508369</u>, PubMed: <u>25964352</u>). Independently of NLRP3 inflammasome and CASP1, promotes pyroptosis, through GSDMD cleavage and

activation, and IL1A, IL18 and HMGB1 release in response to non-canonical inflammasome activators (PubMed:24879791, PubMed:25964352). Plays a crucial role in the restriction of Salmonella typhimurium replication in colonic epithelial cells during infection (PubMed:25121752). In later stages of the infection, LPS from cytosolic Salmonella triggers CASP4 activation, which ultimately results in pyroptosis of infected cells and their extrusion into the gut lumen, as well as in IL18 secretion. Pyroptosis limits bacterial replication, while cytokine secretion promotes the recruitment and activation of immune cells and triggers mucosal inflammation. Involved in LPS-induced IL6 secretion; this activity may not require caspase enzymatic activity (PubMed:26508369). Involved in cell death induced by endoplasmic reticulum stress and by treatment with cytotoxic APP peptides found Alzheimer's patient brains (PubMed:15123740, PubMed:22246630, PubMed:23661706). Activated by direct binding to LPS without the need of an upstream sensor (PubMed:25119034). Does not directly process IL1B (PubMed:7743998, PubMed:7797592, PubMed:7797510). During non- canonical inflammasome activation, cuts CGAS and may play a role in the regulation of antiviral innate immune activation (PubMed:28314590).

### **Images**



Western Blot analysis of various cells using Cleaved-Caspase-4 p20 (Q81) Polyclonal Antibody diluted at 1: 2000



Western Blot analysis of 3T3 cells using Cleaved-Caspase-4 p20 (Q81) Polyclonal Antibody diluted at 1: 2000

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