

CF150 Antibody (Center)

Affinity Purified Rabbit Polyclonal Antibody (Pab) Catalog # AW5397

Product Information

Application	WB, FC
Primary Accession	<u>Q8N884</u>
Other Accession	<u>NP_612450.2</u>
Reactivity	Human
Host	Rabbit
Clonality	Polyclonal
Calculated MW	58814
Isotype	Rabbit IgG
Antigen Source	HUMAN

Additional Information

Gene ID	115004
Antigen Region	266-295
Other Names	Cyclic GMP-AMP synthase, cGAMP synthase, cGAS, h-cGAS, Mab-21 domain-containing protein 1, MB21D1, C6orf150
Dilution	WB~~1:1000 FC~~1:25
Target/Specificity	This CF150 antibody is generated from rabbits immunized with a KLH conjugated synthetic peptide between 266-295 amino acids from the Central region of human CF150.
Format	Purified polyclonal antibody supplied in PBS with 0.09% (W/V) sodium azide. This antibody is purified through a protein A column, followed by peptide affinity purification.
Storage	Maintain refrigerated at 2-8°C for up to 2 weeks. For long term storage store at -20°C in small aliquots to prevent freeze-thaw cycles.
Precautions	CF150 Antibody (Center) is for research use only and not for use in diagnostic or therapeutic procedures.

Protein Information

Name	CGAS {ECO:0000303 PubMed:23258413, ECO:0000312 HGNC:HGNC:21367}
Function	Nucleotidyltransferase that catalyzes the formation of cyclic GMP-AMP (2',3'-cGAMP) from ATP and GTP and plays a key role in innate immunity

(PubMed:21478870, PubMed:23258413, PubMed:23707061, PubMed:23707065, PubMed:23722159, PubMed:24077100, PubMed:24116191, PubMed:24462292, PubMed:25131990, PubMed:26300263, PubMed:29976794, PubMed:30799039, PubMed:31142647, PubMed:32814054, PubMed:33273464, PubMed:33542149, PubMed:37217469, PubMed:37802025). Catalysis involves both the formation of a 2',5' phosphodiester linkage at the GpA step and the formation of a 3',5' phosphodiester linkage at the ApG step, producing c[G(2',5')pA(3',5')p] (PubMed:<u>28214358</u>, PubMed:<u>28363908</u>). Acts as a key DNA sensor: directly binds double-stranded DNA (dsDNA), inducing the formation of liquid-like droplets in which CGAS is activated, leading to synthesis of 2',3'-cGAMP, a second messenger that binds to and activates STING1, thereby triggering type-I interferon production (PubMed: 28314590, PubMed:28363908, PubMed:29976794, PubMed:32817552, PubMed:33230297, PubMed:33606975, PubMed:35322803, PubMed:<u>35438208</u>, PubMed:<u>35460603</u>, PubMed:<u>35503863</u>). Preferentially recognizes and binds curved long dsDNAs of a minimal length of 40 bp (PubMed: <u>30007416</u>). Acts as a key foreign DNA sensor, the presence of double-stranded DNA (dsDNA) in the cytoplasm being a danger signal that triggers the immune responses (PubMed:<u>28363908</u>). Has antiviral activity by sensing the presence of dsDNA from DNA viruses in the cytoplasm (PubMed:28363908, PubMed:35613581). Also acts as an innate immune sensor of infection by retroviruses, such as HIV-2, by detecting the presence of reverse-transcribed DNA in the cytosol (PubMed:23929945, PubMed:24269171, PubMed:30270045, PubMed:32852081). In contrast, HIV-1 is poorly sensed by CGAS, due to its capsid that cloaks viral DNA from CGAS detection (PubMed:24269171, PubMed:30270045, PubMed:32852081). Detection of retroviral reverse-transcribed DNA in the cytosol may be indirect and be mediated via interaction with PQBP1, which directly binds reverse-transcribed retroviral DNA (PubMed: 26046437). Also detects the presence of DNA from bacteria, such as M.tuberculosis (PubMed: 26048138). 2',3'-cGAMP can be transferred from producing cells to neighboring cells through gap junctions, leading to promote STING1 activation and convey immune response to connecting cells (PubMed:<u>24077100</u>). 2',3'-cGAMP can also be transferred between cells by virtue of packaging within viral particles contributing to IFN- induction in newly infected cells in a cGAS-independent but STING1- dependent manner (PubMed: 26229115). Also senses the presence of neutrophil extracellular traps (NETs) that are translocated to the cytosol following phagocytosis, leading to synthesis of 2',3'-cGAMP (PubMed:<u>33688080</u>). In addition to foreign DNA, can also be activated by endogenous nuclear or mitochondrial DNA (PubMed: 28738408, PubMed:28759889, PubMed:31299200, PubMed:33031745, PubMed:<u>33230297</u>). When self-DNA leaks into the cytosol during cellular stress (such as mitochondrial stress, SARS-CoV-2 infection causing severe COVID-19 disease, DNA damage, mitotic arrest or senescence), or is present in form of cytosolic micronuclei, CGAS is activated leading to a state of sterile inflammation (PubMed:28738408, PubMed:28759889, PubMed:31299200, PubMed:<u>33031745</u>, PubMed:<u>33230297</u>, PubMed:<u>35045565</u>). Acts as a regulator of cellular senescence by binding to cytosolic chromatin fragments that are present in senescent cells, leading to trigger type-I interferon production via STING1 and promote cellular senescence (By similarity). Also involved in the inflammatory response to genome instability and double-stranded DNA breaks: acts by localizing to micronuclei arising from genome instability (PubMed:28738408, PubMed:28759889). Micronuclei, which are frequently found in cancer cells, consist of chromatin surrounded by their own nuclear membrane: following breakdown of the micronuclear envelope, a process associated with chromothripsis, CGAS binds self-DNA exposed to the cytosol, leading to 2',3'-cGAMP synthesis and subsequent activation of STING1 and type-I interferon production (PubMed: 28738408, PubMed:28759889). Activated in response to prolonged mitotic arrest,

	promoting mitotic cell death (PubMed: <u>31299200</u>). In a healthy cell, CGAS is however kept inactive even in cellular events that directly expose it to self-DNA, such as mitosis, when cGAS associates with chromatin directly after nuclear envelope breakdown or remains in the form of postmitotic persistent nuclear cGAS pools bound to chromatin (PubMed: <u>31299200</u> , PubMed: <u>33542149</u>). Nuclear CGAS is inactivated by chromatin via direct interaction with nucleosomes, which block CGAS from DNA binding and thus prevent CGAS-induced autoimmunity (PubMed: <u>31299200</u> , PubMed: <u>32911482</u> , PubMed: <u>32912999</u> , PubMed: <u>33051594</u> , PubMed: <u>33542149</u>). Also acts as a suppressor of DNA repair in response to DNA damage: inhibits homologous recombination repair by interacting with PARP1, the CGAS-PARP1 interaction leading to impede the formation of the PARP1-TIMELESS complex (PubMed: <u>30356214</u> , PubMed: <u>31544964</u>). In addition to DNA, also sense translation stress: in response to translation stress, translocates to the cytosol and associates with collided ribosomes, promoting its activation and triggering type-I interferon production (PubMed: <u>34111399</u>). In contrast to other mammals, human CGAS displays species-specific mechanisms of DNA recognition and produces less 2',3'-cGAMP, allowing a more fine-tuned response to pathogens (PubMed: <u>30007416</u>).
Cellular Location	Nucleus. Chromosome. Cell membrane; Peripheral membrane protein. Cytoplasm, cytosol. Note=Mainly localizes in the nucleus, and at low level in the cytosol (PubMed:31544964, PubMed:31808743). On chromosomes, enriched on centromeric satellite and LINE DNA repeat elements (PubMed:30811988). Exported from the nucleus to the cytosol in a XPO1/CRM1 via the nuclear export signal in response to DNA stimulation (PubMed:33406424). Outside the nucleus, localizes at the cell membrane as a peripheral membrane protein in resting conditions: association to the cell membrane is mediated via binding to phosphatidylinositol 4,5-bisphosphate (PtdIns(4,5)P2) (PubMed:30827685). Localization at the cell membrane is required to limit the recognition of self-DNA (PubMed:30827685). Following detection of double-stranded DNA (dsDNA), released from the cell membrane into the cytosol in order to signal (PubMed:30827685). Upon transfection with dsDNA forms punctate structures that co-localize with DNA and Beclin-1 (BECN1) (PubMed:20048138). Phosphorylation at Tyr-215 promotes cytosolic retention (PubMed:30356214). In response to translation stress, translocates to the cytosol and associates with collided ribosomes (PubMed:34111399).
Tissue Location	Expressed in the monocytic cell line THP1.

Background

The exact function of C6orf150 remains unknown.

References

Rush, J., et al. Nat. Biotechnol. 23(1):94-101(2005) Rush, J., et al. Nat. Biotechnol. 23(1):94-101(2005) Mungall, A.J., et al. Nature 425(6960):805-811(2003)

Images

Overlay histogram showing U-2OS cells stained with AW5397 (green line). The cells were fixed with 2% paraformaldehyde (10 min) and then permeabilized with 90% methanol for 10 min. The cells were then icubated in



2% bovine serum albumin to block non-specific protein-protein interactions followed by the antibody (AW5397, 1:25 dilution) for 60 min at 37°C. The secondary antibody used was Goat-Anti-Rabbit IgG, DyLight® 488 Conjugated Highly Cross-Adsorbed(NA168821) at 1/400 dilution for 40 min at 37°C. Isotype control antibody (blue line) was rabbit IgG (1µg/1x10^6 cells) used under the same conditions. Acquisition of >10, 000 events was performed.

All lanes : Anti-CF150 Antibody (Center) at 1:1000 dilution Lane 1: human cerebellum lysates Lane 2: human spleen lysates Lane 3: HT-29 whole cell lysates Lane 4: MCF-7 whole cell lysates Lane 5: THP-1 whole cell lysates Lysates/proteins at 20 µg per lane. Secondary Goat Anti-Rabbit IgG, (H+L),Peroxidase conjugated at 1/10000 dilution Predicted band size : 59 kDa Blocking/Dilution buffer: 5% NFDM/TBST.

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