

Variant: *NM_004360.5(CDH1):c.1408del (p.Thr470fs)*

Version: 2.0

CA645369638 [↗](#)

428633 (ClinVar) [↗](#)

Gene: CDH1 ([HGNC:999](#))

Condition: CDH1-related diffuse gastric and lobular breast cancer ([MONDO:0100488](#))

Inheritance Mode: Autosomal dominant inheritance

UUID: e338dcbc-38e5-47e2-bb62-ac1e7907225c

Approved on: 2023/8/25

Published on: 2023/8/25

HGVS expressions

NM_004360.5(CDH1):c.1408del (p.Thr470fs)

NM_004360.5:c.1408del

NC_000016.10:g.68815602del

CM000678.2:g.68815602del

NC_000016.9:g.68849505del

CM000678.1:g.68849505del

NC_000016.8:g.67407006del

NG_008021.1:g.83311del

ENST00000261769.10:c.1408del

ENST00000261769.9:c.1408del

ENST00000422392.6:c.1225del

ENST00000562836.5:n.1479del

ENST00000566510.5:c.*74del

ENST00000566612.5:c.1408del

ENST00000611625.4:c.1471del

ENST00000612417.4:c.1408del

ENST00000621016.4:c.1408del

NM_004360.3:c.1408del

NM_001317184.1:c.1225del

NM_001317185.1:c.-141del

NM_001317186.1:c.-412del

NM_004360.4:c.1408del

NM_001317184.2:c.1225del

NM_001317185.2:c.-141del

NM_001317186.2:c.-412del

Pathogenic

Met criteria codes 4

PM2_Supporting

PVS1

PM5_Supporting

PS4_Supporting

Not Met criteria codes 22

PS1

PS2

PS3

BP5

BP7

BP4

BP3

BP1

BP2

PP1

PP2

PP3

PP4

PM6

PM1

PM3

PM4

BA1

BS1

BS4

BS3

BS2

Expert Panel

CDH1 VCEP [↗](#)

Criteria Specification Information

[↗](#) **Criteria Specification:** ClinGen CDH1 Expert Panel Specifications to the ACMG/AMP Variant Interpretation Guidelines Version 3.1

[↗](#) **Criteria Specification Approval History**

Evidence submitted by expert panel

CDH1 VCEP

The c.1408del (p.Thr470fs) variant is predicted to result in a premature stop codon that leads to a truncated or absent protein (PVS1, PM5_Supporting). The variant is absent in the gnomAD cohort (PM2_Supporting; <http://https://gnomad.broadinstitute.org/>). The variant has been reported in a family meeting clinical criteria for HDGC (PS4_Supporting; SCV000580711.3). In summary, this variant meets criteria to be classified as Pathogenic based on the ACMG/AMP criteria applied as specified by the CDH1 Variant Curation Expert Panel (Variant Interpretation Guidelines Version 3.1): PVS1, PM2_Supporting, PS4_Supporting, PM5_Supporting.

Met criteria codes

PM2_Supporting			Variant absent from gnomAD
PVS1			Variant predicted to result in NMD with stop codon occurring at amino acid 480
PM5_Supporting			Apply PM5_Supporting to nonsense/frameshift variants that are predicted/proved to undergo NMD.
PS4_Supporting			Two gastric cancer cases in a family with one confirmed DGC (SCV000580711.3).

Not Met criteria codes

PS1			No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
PS2			No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
PS3			No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
BP5			No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
BP7			No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
BP4			No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
BP3			

	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline		
BP1		✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
BP2		✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
PP1		✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
PP2		✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
PP3		✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
PP4		✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
PM6		✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
PM1		✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
PM3		✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
PM4		✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
BA1		✖	Variant absent from gnomAD
BS1		✖	Variant absent from gnomAD
BS4		✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
BS3		✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
BS2		✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline

Curation History [↗](#)

⛶

☰ ▾

⬇️ ▾

Showing 1 to 2 of 2 rows

--

The information on this website is not intended for direct diagnostic use or medical decision-making without review by a genetics professional. Individuals should not change their health behavior solely on the basis of information contained on this website. If you have questions about the information contained on this website, please see a health care professional.