

Variant: *NM_004360.5(CDH1):c.656del (p.Pro219fs)*

Version: 2.0

CA658683948 [↗](#)

488647 (ClinVar) [↗](#)

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

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

Inheritance Mode: Autosomal dominant inheritance

UUID: 442ddb1-97e7-458e-82c5-7def1bb4661a

Approved on: 2023/8/25

Published on: 2023/8/25

HGVS expressions

NM_004360.5:c.656del

NM_004360.5(CDH1):c.656del (p.Pro219fs)

NC_000016.10:g.68808817del

CM000678.2:g.68808817del

NC_000016.9:g.68842720del

CM000678.1:g.68842720del

NC_000016.8:g.67400221del

NG_008021.1:g.76526del

ENST00000261769.10:c.656del

ENST00000261769.9:c.656del

ENST00000422392.6:c.656del

ENST00000561751.1:c.423del

ENST00000562836.5:n.727del

ENST00000564676.5:n.938del

ENST00000566510.5:c.531+250del

ENST00000566612.5:c.656del

ENST00000567320.1:n.166del

ENST00000611625.4:c.656del

ENST00000612417.4:c.656del

ENST00000621016.4:c.656del

NM_004360.3:c.656del

NM_001317184.1:c.656del

NM_001317185.1:c.-960del

NM_001317186.1:c.-1164del

NM_004360.4:c.656del

NM_001317184.2:c.656del

NM_001317185.2:c.-960del

NM_001317186.2:c.-1164del

Pathogenic

Met criteria codes **4**

PM2_Supporting

PVS1

PM5_Supporting

PS4_Supporting

Not Met criteria codes **22**

Expert Panel

CDH1 VCEP [↗](#)

Criteria Specification Information

PS1

PS2

PS3

BP5

BP7

BP4

BP3

BP1

BP2

PP1

PP2

PP3

PP4

PM6

PM1

PM3

PM4

BA1

BS1

BS4

BS3

BS2

Evidence Links 0

- [Criteria Specification: ClinGen CDH1 Expert Panel Specifications to the ACMG/AMP Variant Interpretation Guidelines Version 3.1](#)
- [Criteria Specification Approval History](#)
- [Criteria Specifications for this VCEP](#)

Evidence submitted by expert panel

CDH1 VCEP

The c.656del p.(Pro219fs) 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://gnomad.broadinstitute.org>). This variant has been reported in at least one family meeting HDGC clinical criteria (PS4_Supporting; SCV000760825.3). Therefore, 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			Absent in population databases.
PVS1			Exon 5 of 16, predicted NMD.
PM5_Supporting			Apply PM5_Supporting to nonsense/frameshift variants that are predicted/proved to undergo NMD.
PS4_Supporting			SCV000760825.3 - one family meets HDGC criteria (2 cases gastric, of which one diffuse).

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			No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
BS1			No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
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.