The ClinGen Evidence Repository is an **FDA-recognized human genetic variant database** containing expert-curated assertions regarding variants' pathogenicity and supporting evidence summaries. [Disclaimer]

Variant: NM_004360.5(CDH1):c.2062_2063TG[1] (p.Cys688_Glu689delinsTer)

Version: 2.0

CA163542

140781 (ClinVar)

Gene: CDH1
Condition: CDH1-related diffuse gastric and lobular breast cancer (MONDO:0100488)
Inheritance Mode: Autosomal dominant inheritance
UUID: 3f59f180-03c4-40bf-8632-9084331cc8d5
Approved on: 2023/8/25
Published on: 2023/8/25

HGVS expressions

NM_004360.5:c.2062_2063TG[1]
NM_004360.5(CDH1):c.2062_2063TG[1] (p.Cys688_Glu689delinsTer)
NC_000016.10:g.68823526_68823527del
CM000678.2:g.68823526_68823527del
NC_000016.9:g.68857429_68857430del
CM000678.1:g.68857429_68857430del
NC_000016.8:g.67414930_67414931del
NG_008021.1:g.91235_91236del
ENST00000261769.10:c.2064_2065del
ENST00000261769.9:c.2064_2065del
ENST00000422392.6:c.1881_1882del
ENST00000562118.1:n.282_283del
ENST00000562836.5:n.2135_2136del
ENST00000566510.5:c.*730_*731del
ENST00000566612.5:c.*304_*305del
ENST00000611625.4:c.2127_2128del
ENST00000612417.4:c.1830+1407_1830+1408del
ENST00000621016.4:c.1865+1372_1865+1373del
NM_004360.3:c.2064_2065del
NM_001317184.1:c.1881_1882del
NM_001317185.1:c.516_517del
NM_001317186.1:c.99_100del
NM_004360.4:c.2064_2065del
NM_004360.5:c.2064_2065del
NM_001317184.2:c.1881_1882del
NM_001317185.2:c.516_517del
NM_001317186.2:c.99_100del

Pathogenic

Met criteria codes 4

PS4 PM5_Supporting PM2_Supporting PVS1

Not Met criteria codes 22

Expert Panel

CDH1 VCEP

Criteria Specification Information

BP5 BP7 BP4 BP3 BP1 BP2
PS1 PS2 PS3 PP1 PP2 PP3
PP4 PM1 PM3 PM4 PM6
BA1 BS2 BS1 BS4 BS3

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.2064_2065delTG (p.Cys688Terfs) 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 five families meeting HDGC clinical criteria (PS4; PMID: 15235021, 17545690, 21424370, 26072394, 29307626). 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: PVS1, PM2_Supporting, PS4, PM5_Supporting.

Met criteria codes

PS4	✔	5 families that meet HDGC clinical criteria
PM5_Supporting	✔	Apply PM5_Supporting to nonsense/frameshift variants that are predicted/proved to undergo NMD.
PM2_Supporting	✔	Absent in gnomAD v2.1.1 in a region of sufficient coverage
PVS1	✔	PTC created in exon 13 of 16, expected to undergo NMD

Not Met criteria codes

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	ℹ	✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
PS1		✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
PS2	ℹ	✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
PS3	ℹ	✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
PP1	ℹ	✖	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	ℹ	✖	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
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	ℹ	✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
PM6	ℹ	✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
BA1	ℹ	✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
BS2	ℹ	✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
BS1	ℹ	✖	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

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.

ClinGen Terms of Use.
¤ Powered by BCM's Genboree.