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.1587dup (p.Ala530fs)

Version: 2.0

CA16620251

422539 (ClinVar)

Gene: CDH1 (HGNC:999)
Condition: CDH1-related diffuse gastric and lobular breast cancer (MONDO:0100488)
Inheritance Mode: Autosomal dominant inheritance
UUID: 55e01934-c703-4d75-ae7e-ad1eb3df27e3
Approved on: 2023/8/25
Published on: 2023/8/25

HGVS expressions

NM_004360.5(CDH1):c.1587dup (p.Ala530fs)
NM_004360.5:c.1587dup
NC_000016.10:g.68819301dup
CM000678.2:g.68819301dup
NC_000016.9:g.68853204dup
CM000678.1:g.68853204dup
NC_000016.8:g.67410705dup
NG_008021.1:g.87010dup
ENST00000261769.10:c.1587dup
ENST00000261769.9:c.1587dup
ENST00000422392.6:c.1404dup
ENST00000562836.5:n.1658dup
ENST00000566510.5:c.*253dup
ENST00000566612.5:c.1566-2700dup
ENST00000611625.4:c.1650dup
ENST00000612417.4:c.1587dup
ENST00000621016.4:c.1587dup
NM_004360.3:c.1587dup
NM_001317184.1:c.1404dup
NM_001317185.1:c.39dup
NM_001317186.1:c.-254-2700dup
NM_004360.4:c.1587dup
NM_001317184.2:c.1404dup
NM_001317185.2:c.39dup
NM_001317186.2:c.-254-2700dup

Pathogenic

Met criteria codes 4

PS4_Supporting PVS1 PM2_Supporting PM5_Supporting

Not Met criteria codes 22

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

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 Links 0

Criteria Specifications for this VCEP

Evidence submitted by expert panel

CDH1 VCEP

The c.1587dup (p.Ala530fs) 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/). This variant has been identified in a family meeting HDGC clinical criteria (PS4_Supporting; SCV000580697.3). 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

PS4_Supporting	✔	Variant present in female with history of signet cell adenocarcinoma diagnosed @ 20s (SCV000580697.3).
PVS1	✔	Variant creates a premature stop codon at position 7 (amino acid 536) of the new reading frame
PM2_Supporting	✔	Variant not present in gnomAD
PM5_Supporting	✔	Apply PM5_Supporting to nonsense/frameshift variants that are predicted/proved to undergo NMD.

Not Met criteria codes

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
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	ℹ	✖	Variant not present in gnomAD
PM6	ℹ	✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
BS1	ℹ	✖	Variant not present in gnomAD
BS4	ℹ	✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
BS3	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
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
BP5	ℹ	✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
BP7	ℹ	✖	No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
BP4	ℹ	✖	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

Curation History

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