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_000527.5(LDLR):c.737G>T (p.Gly246Val)

Version: 1.0

CA404079680

440600 (ClinVar)

Gene: LDLR (HGNC:3949)
Condition: hypercholesterolemia, familial (MONDO:0007750)
Inheritance Mode: Semidominant inheritance
UUID: c4b8dd24-6303-4adb-b25c-76981f3203d3
Approved on: 2023/4/1
Published on: 2023/4/1

HGVS expressions

NM_000527.5(LDLR):c.737G>T (p.Gly246Val)
NM_000527.5:c.737G>T
NC_000019.10:g.11106607G>T
CM000681.2:g.11106607G>T
NC_000019.9:g.11217283G>T
CM000681.1:g.11217283G>T
NC_000019.8:g.11078283G>T
NG_009060.1:g.22227G>T
ENST00000252444.10:c.995G>T
ENST00000559340.2:c.737G>T
ENST00000560467.2:c.737G>T
ENST00000558518.6:c.737G>T
ENST00000252444.9:c.991G>T
ENST00000455727.6:c.314-785G>T
ENST00000535915.5:c.614G>T
ENST00000545707.5:c.356G>T
ENST00000557933.5:c.737G>T
ENST00000558013.5:c.737G>T
ENST00000558518.5:c.737G>T
ENST00000558528.1:n.252G>T
ENST00000560467.1:c.337G>T
NM_000527.4:c.737G>T
NM_001195798.1:c.737G>T
NM_001195799.1:c.614G>T
NM_001195800.1:c.314-785G>T
NM_001195803.1:c.356G>T
NM_001195798.2:c.737G>T
NM_001195799.2:c.614G>T
NM_001195800.2:c.314-785G>T
NM_001195803.2:c.356G>T

Uncertain Significance

Met criteria codes 2

PP3 PM2

Not Met criteria codes 20

BA1 BP7 BP4 BP2 BS1 BS4 BS3 BS2 PVS1 PP1 PP4 PM1 PM3 PM5 PM4 PS1 PS2 PS3 PS4 PM6

Evidence Links 0

Expert Panel

Familial Hypercholesterolemia VCEP

Criteria Specification Information

- **Criteria Specification:** *ClinGen Familial Hypercholesterolemia Expert Panel Specifications to the ACMG/AMP Variant Classification Guidelines Version 1.2*
  - **PDF**
- **Criteria Specification Approval History**
- **Criteria Specifications for this VCEP**

Evidence submitted by expert panel

Familial Hypercholesterolemia VCEP

The NM_000527.5(LDLR):c.737G>T (p.Gly246Val) variant is classified as Uncertain significance - insufficient evidence for Familial Hypercholesterolemia by applying evidence codes PM2 and PP3 as defined by the ClinGen Familial Hypercholesterolemia Expert Panel LDLR-specific variant curation guidelines (https://doi.org/10.1016/j.gim.2021.09.012). The supporting evidence is as follows: PM2:This variant is absent from gnomAD (gnomAD version 2.1.1). PP3: REVEL score is 0.87, this is greater than 0.75.

Met criteria codes

PP3	✔	REVEL score is 0.87, this is greater than 0.75.
PM2	✔	This variant is absent from gnomAD (gnomAD version 2.1.1).

Not Met criteria codes

BA1	✖	Not Met.
BP7	✖	Not met.
BP4	✖	Not met.
BP2	✖	Not met.
BS1	✖	Not Met.
BS4	✖	Not met.
BS3	✖	Not met.

BS2		✖	Not met.
PVS1		✖	Not met.
PP1		✖	Not met.
PP4		✖	Not met.
PM1		✖	Not met.
PM3		✖	Not met.
PM5		✖	Not met.
PM4		✖	Not met.
PS1		✖	Not met.
PS2		✖	Not met.
PS3		✖	Not met.
PS4		✖	Not met.
PM6		✖	Not met.

Curation History

owing 1 to 1 of 1 rows

ee Report	Preferred Variant Title	Classification	Condition	Published Date	Version	Criteria Specification	Gene
View	NM_000527.5(LDLR):c.737G>T (p.Gly2...	Uncertain Significance	Hypercholesterolemia, Familial	2023-04-01	1.0	ClinGen Familial Hypercholesterolemia Expert Panel Specifications to the ACMG/AMP Variant Classification Guidelines Version 1.2	LDLR

owing 1 to 1 of 1 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.