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.91G>A (p.Glu31Lys)

Version: 1.0

CA041752

251013 (ClinVar)

Gene: LDLR (HGNC:3949)
Condition: hypercholesterolemia, familial (MONDO:0007750)
Inheritance Mode: Semidominant inheritance
UUID: 47029d08-db26-42fe-bf8b-bcd8c066753d
Approved on: 2023/3/24
Published on: 2025/4/23

HGVS expressions

NM_000527.5:c.91G>A
NM_000527.5(LDLR):c.91G>A (p.Glu31Lys)
NC_000019.10:g.11100246G>A
CM000681.2:g.11100246G>A
NC_000019.9:g.11210922G>A
CM000681.1:g.11210922G>A
NC_000019.8:g.11071922G>A
NG_009060.1:g.15866G>A
ENST00000252444.10:c.349G>A
ENST00000559340.2:c.91G>A
ENST00000560467.2:c.91G>A
ENST00000558518.6:c.91G>A
ENST00000252444.9:c.345G>A
ENST00000455727.6:c.91G>A
ENST00000535915.5:c.91G>A
ENST00000545707.5:c.91G>A
ENST00000557933.5:c.91G>A
ENST00000557958.1:n.177G>A
ENST00000558013.5:c.91G>A
ENST00000558518.5:c.91G>A
ENST00000560502.5:n.177G>A
NM_000527.4:c.91G>A
NM_001195798.1:c.91G>A
NM_001195799.1:c.91G>A
NM_001195800.1:c.91G>A
NM_001195803.1:c.91G>A
NM_001195798.2:c.91G>A
NM_001195799.2:c.91G>A
NM_001195800.2:c.91G>A
NM_001195803.2:c.91G>A

Uncertain Significance

Met criteria codes 2

PP3 PM2

Not Met criteria codes 23

PP1 PP2 PP4 PM6 PM1 PM3 PM5 PM4 BA1 BS2 PVS1 BS1 BS4 BS3 BP7 BP4 BP3 BP1 BP2 PS1 PS2 PS3 PS4

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.91G>A (p.Glu31Lys) variant is classified as Uncertain significance - insufficient evidence for Familial Hypercholesterolemia by applying ACMG/AMP evidence codes PM2 and PP3 as defined by the ClinGen Familial Hypercholesterolemia Expert Panel LDLR-specific variant curation guidelines (specification version 1.2) on 24 March 2023. The supporting evidence is as follows: PM2: PopMax MAF = 0.00005 (0.005%) in European (Non-Finnish) exomes (gnomAD v2.1.1). PP3: REVEL = 0.777.

Met criteria codes

PP3	✔	REVEL = 0.777, this is greater than 0.75.
PM2	✔	PopMax MAF = 0.00005 (0.005%) in European (Non-Finnish) exomes (gnomAD v 2.1.1).

Not Met criteria codes

PP1	✖	Not met.
PP2	✖	Not applicable
PP4	✖	Clinical criterial not met on patients.
PM6	✖	Not met.
PM1	✖	Not located in exon 4.
PM3	✖	Not met.
PM5	✖	Not met.

PM4		✖	Not an in-frame insertion/deletion.
BA1		✖	Not met.
BS2		✖	Not met.
PVS1		✖	Not a null variant.
BS1		✖	Not met.
BS4		✖	Not met.
BS3		✖	Not met.
BP7		✖	Not a synonymous variant.
BP4		✖	REVEL = 0.777
BP3		✖	Not applicable
BP1		✖	Not applicable
BP2		✖	Not met.
PS1		✖	Not met.
PS2		✖	Not met.
PS3		✖	Not met.
PS4		✖	Variant has been described in several publications (PMID 20145306, 34428338, 35910211), however, clinical criteria not met for identified probands.

Curation History

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ee Report	Preferred Variant Title	Classification	Condition	Published Date	Version	Criteria Specification	Gene
View	NM_000527.5(LDLR):c.91G>A (p.Glu31...	Uncertain Significance	Hypercholesterolemia, Familial	2025-04-23	1.0	ClinGen Familial Hypercholesterolemia Expert Panel Specifications to the ACMG/AMP Variant Classification Guidelines Version 1.2	LDLR

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