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_000277.2(PAH):c.785T>G (p.Val262Gly)

CA267671

120285 (ClinVar)

Gene: PAH (HGNC:5053)
Condition: phenylketonuria (MONDO:0009861)
Inheritance Mode: Autosomal recessive inheritance
UUID: 9357bfa1-6426-4f30-9c29-28a74e5240df
Approved on: 2018-12-10
Published on: 2018-12-10

HGVS expressions

NM_000277.2:c.785T>G
NM_000277.2(PAH):c.785T>G (p.Val262Gly)
NC_000012.12:g.102852872A>C
CM000674.2:g.102852872A>C
NC_000012.11:g.103246650A>C
CM000674.1:g.103246650A>C
NC_000012.10:g.101770780A>C
NG_008690.1:g.69731T>G
NG_008690.2:g.110539T>G
NM_000277.1:c.785T>G
NM_001354304.1:c.785T>G
NM_000277.3:c.785T>G
ENST00000307000.7:c.770T>G
ENST00000549247.6:n.544T>G
ENST00000553106.5:c.785T>G

Likely Pathogenic

Met criteria codes 4
PM3 PM2 PP3 PP4

Not Met criteria codes 1
PM5

Evidence Links 1

Expert Panel
Phenylketonuria VCEP

Criteria Specification Information

Criteria Specifications for this VCEP

Evidence submitted by expert panel

Phenylketonuria VCEP

The c.785T>G (p.Val262Gly) variant in PAH is observed in 1 patient with classic PKU. BH4 deficiencies were not assessed. It was detected with a known pathogenic variant p.F39L. PMID: 26666653. This variant is absent from ExAC, gnomAD, 1000G, and ESP. It is predicted deleterious in SIFT, Polyphen-2, MutationTaster, and REVEL=0.954. In summary, this variant meets criteria to be classified as likely pathogenic for PAH. PAH-specific ACMG/AMP criteria applied: PM2, PM3, PP4, PP3.

Met criteria codes

PM3	✔	V262G detected with F39L, PMID: 26666653 c. [117C > G]; [785 T > G]/p. [Phe39Leu];[Val262Gly]. F39L, VarID 605, Pathogenic PubMed:26666653
PM2	✔	Absent from ExAC, gnomAD, 1000G, ESP
PP3	✔	Predicted deleterious in SIFT, Polyphen2, MutationTaster. REVEL=0.954
PP4	✔	V262G observed in 1 patient with classic PKU. BH4 deficiencies not assessed. PMID: 26666653. A total of 364 hyperphenylalaninemic patients, including 42 siblings, were investigated. According to their blood Phenylalanine concentration at time of diagnosis, these patients were assigned to one of the three phenotype categories. V262G was observed in 1 patient with classic PKU. PubMed:26666653

Not Met criteria codes

PM5	✖	This is the only variant found in this codon in ClinVar.

Curation History

Showing 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.

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