

Variant: NM_000277.2(PAH):c.826A>G (p.Met276Val)

CA229794 [↗](#)

102856 (ClinVar) [↗](#)

Gene: PAH ([HGNC:5053](#))

Condition: phenylketonuria ([MONDO:0009861](#))

Inheritance Mode: Autosomal recessive inheritance

UUID: f085841b-65ba-4950-9a05-678a254062ce

Approved on: 2018-12-09

Published on: 2018-12-10

HGVS expressions

NM_000277.2(PAH):c.826A>G (p.Met276Val)

NM_000277.2:c.826A>G

NC_000012.12:g.102852831T>C

CM000674.2:g.102852831T>C

NC_000012.11:g.103246609T>C

CM000674.1:g.103246609T>C

NC_000012.10:g.101770739T>C

NG_008690.1:g.69772A>G

NG_008690.2:g.110580A>G

NM_000277.1:c.826A>G

NM_001354304.1:c.826A>G

NM_000277.3:c.826A>G

ENST00000307000.7:c.811A>G

ENST00000549247.6:n.585A>G

ENST00000553106.5:c.826A>G

Likely Pathogenic

Met criteria codes **3**

PM2

PP4_Moderate

PM3_Strong

Not Met criteria codes **1**

PP3

Evidence Links **2**

Expert Panel

[Phenylketonuria VCEP](#) [↗](#)

Criteria Specification Information **!**

[↗](#) **Criteria Specifications for this VCEP**

Evidence submitted by expert panel

Phenylketonuria VCEP

The c.826A>G (p.Met276Val) variant in PAH is absent from population databases. It has been identified in trans with pathogenic variants in two patients (A300S and R261Q; PMID: 16198137), and identified in an additional patient in which a defect in BH4 metabolism was excluded as a cause of elevated phenylalanine (PMID: 9634518). In summary, this variant meets criteria to be classified as likely pathogenic for PAH. PAH-specific ACMG/AMP criteria applied: PP4_Moderate, PM2, PM3_strong.

Met criteria codes

PM2		Absent from population databases	
PP4_Moderate			BH4 defect excluded. Single patient (Guldborg, 1998) PMID: 9634518
			BH4 defect excluded in all patients PubMed:9634518
PM3_Strong			Two patients, A300S and R261Q (both pathogenic)(PMID: 16198137). Both with MHP
			Two patients, both in trans with pathogenic variants (A300S and R261Q) PubMed:16198137

Not Met criteria codes

PP3			SIFT = Damaging Provean = Neutral REVEL = 0.786
-----	--	--	---

Curation History		
		<div><div></div><div>▼</div><div>▼</div></div>
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.