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_001754.5(RUNX1):c.1309A>G (p.Thr437Ala)

Version: 1.0

CA410147422

1424427 (ClinVar)

Gene: RUNX1 (HGNC:861)
Condition: hereditary thrombocytopenia and hematologic cancer predisposition syndrome (MONDO:0011071)
Inheritance Mode: Autosomal dominant inheritance
UUID: 6dc6e605-5b43-4d50-b264-78642e75b5f0
Approved on: 2024-07-11
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HGVS expressions

NM_001754.5:c.1309A>G
NM_001754.5(RUNX1):c.1309A>G (p.Thr437Ala)
NC_000021.9:g.34792269T>C
CM000683.2:g.34792269T>C
NC_000021.8:g.36164566T>C
CM000683.1:g.36164566T>C
NC_000021.7:g.35086436T>C
NG_011402.2:g.1197443A>G
ENST00000675419.1:c.1309A>G
ENST00000300305.7:c.1309A>G
ENST00000344691.8:c.1228A>G
ENST00000399240.5:c.1036A>G
ENST00000437180.5:c.1309A>G
ENST00000482318.5:c.*899A>G
NM_001001890.2:c.1228A>G
NM_001754.4:c.1309A>G
NM_001001890.3:c.1228A>G

Uncertain Significance

Met criteria codes 1

BP4

Not Met criteria codes 25

PVS1 PP1 PP2 PP3 PP4 PM1 PM3 PM5 PM4 BA1 PM6 PM2 BS2 BS1 BS4 BS3 PS1 PS2 PS3 PS4 BP3 BP1 BP2 BP5 BP7

Evidence Links 0

Expert Panel

Myeloid Malignancy VCEP

Criteria Specification Information

- **Criteria Specification:** *ClinGen Myeloid Malignancy Expert Panel Specifications to the ACMG/AMP Variant Interpretation Guidelines Version 2*
  - **PDF**
- **Criteria Specification Approval History**
- **Criteria Specifications for this VCEP**

Evidence submitted by expert panel

Myeloid Malignancy VCEP

NM_001754.5(RUNX1): c.1309A>G (p.Thr437Ala) is a missense variant which has a REVEL score < 0.50 (0.154) and a SpliceAI score ≤ 0.20 (0.0) (BP4). In summary, the clinical significance of this variant is uncertain. ACMG/AMP criteria applied, as specified by the Myeloid Malignancy Variant Curation Expert Panel for RUNX1: BP4.

Met criteria codes

BP4	✔	This missense variant has a REVEL score < 0.50 (0.154) and a SpliceAI score ≤ 0.20 (0.0) (BP4).

Not Met criteria codes

PVS1	✖	This variant is not a null variant.
PP1	✖	Segregation data for this variant has not been reported in literature.
PP2	✖	This rule is not applicable for MM-VCEP.
PP3	✖	This missense variant does not have a REVEL score of ≥ 0.88.
PP4	✖	This rule is not applicable for MM-VCEP.
PM1	✖	This variant does not affect any of the following amino acid residues, nor is it located within the RHD: R107, K110, A134, R162, R166, S167, R169, G170, K194, T196, D198, R201, R204 OR within residues 89-204.
PM3	✖	This rule is not applicable for MM-VCEP.
PM5	✖	There has not yet been a different missense change determined to be pathogenic at this amino acid residue.
PM4	✖	This variant is not an in-frame deletion/insertion.
BA1	✖	This variant does not have a MAF ≥ 0.0015 (0.15%) in any general continental population dataset.
PM6	✖	De novo data for this variant has not been reported in literature.
PM2	✖	This variant is present in at least one population database.
BS2	✖	This rule is not applicable for MM-VCEP.
BS1	✖	This variant does not have a MAF between 0.00015 (0.015%) and 0.0015 (0.15%) in any general continental dataset.
BS4	✖	Segregation data for this variant has not been reported in literature.

BS3		✖	In vitro or in vivo functional data has not been reported for this variant in the literature.
PS1		✖	There has not yet been a missense change determined to be pathogenic at this amino acid residue.
PS2		✖	De novo data for this variant has not been reported in literature.
PS3		✖	In vitro or in vivo functional data has not been reported for this variant in the literature.
PS4		✖	Proband data for this variant has not been reported in literature.
BP3		✖	This rule is not applicable for MM-VCEP.
BP1		✖	This rule is not applicable for MM-VCEP.
BP2		✖	This variant has not been observed in trans with a pathogenic variant for a fully penetrant dominant gene/disorder or observed in cis with a pathogenic variant in any inheritance pattern.
BP5		✖	This rule is not applicable for MM-VCEP.
BP7		✖	This variant is not a synonymous or intronic variant.

Curation History

Showing 1 to 1 of 1 rows