

Variant: *NM_001110792.2(MECP2):c.1497A>G (p.Ter499Trp)*

Version: 2.0

[CA170257](#) 

[143490 \(ClinVar\)](#) 

Gene: MECP2 ([HGNC:4204](#))

Condition: Rett syndrome ([MONDO:0010726](#))

Inheritance Mode: X-linked inheritance

UID: 2bba9172-4eb1-4098-97c0-63efb3d2ff5b

Approved on: 2025-10-28

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HGVS expressions

NM_001110792.2:c.1497A>G

NM_001110792.2(MECP2):c.1497A>G (p.Ter499Trp)

NC_000023.11:g.154030367T>C

CM000685.2:g.154030367T>C

NC_000023.10:g.153295818T>C

CM000685.1:g.153295818T>C

NC_000023.9:g.152949012T>C

NG_007107.2:g.111761A>G

NG_007107.3:g.111737A>G

ENST00000303391.11:c.1461A>G

ENST00000453960.7:c.1497A>G

ENST00000303391.10:c.1461A>G

ENST00000453960.6:c.1497A>G

ENST00000619732.4:c.1457A>G

ENST00000628176.2:c.*833A>G

NM_001110792.1:c.1497A>G

NM_001316337.1:c.1182A>G

NM_004992.3:c.1461A>G

NM_001316337.2:c.1182A>G

NM_001369391.2:c.1182A>G

NM_001369392.2:c.1182A>G

NM_001369393.2:c.1182A>G

NM_001369394.1:c.1182A>G

NM_001369394.2:c.1182A>G

NM_001386137.1:c.792A>G

NM_001386138.1:c.792A>G

NM_001386139.1:c.792A>G

NM_004992.4:c.1461A>G

Pathogenic

Met criteria codes **4**

PS2

PM2_Supporting

PM4_Strong

PS4_Moderate

Not Met criteria codes **1**

PP4

Expert Panel

[Rett and Angelman-like Disorders VCEP](#) 









Criteria Specification Information

Evidence submitted by expert panel



Rett and Angelman-like Disorders VCEP

The p.Ter487TrpextTer27 variant in MECP2 (NM_004992.4) occurs at the stop codon and causes a change in the length of the protein and an elongated mRNA transcript of MECP2 (PM4_strong). The p.Ter487TrpextTer27 variant in MECP2 has been reported as a de novo occurrence (biological parentage confirmed) in an individual with features of Rett syndrome (internal database - GeneDx) (PS2). The p.Ter487TrpextTer27 variant has been observed in 3 individuals with features of Rett syndrome (PMID 21807996; internal database - LabCorp (formerly Invitae); internal database - GeneDx) (PS4_moderate). The p.Ter487TrpextTer27 variant in MECP2 is absent from gnomAD v4.1.0 (PM2_supporting). In summary, the p.Ter487TrpextTer27 variant in MECP2 is classified as Pathogenic for Rett syndrome based on the ACMG/AMP criteria (PM4_strong, PS2, PS4_moderate, PM2_supporting). (MECP2 Specifications v5.0.0; curation approved on 10/28/2025).

Met criteria codes

PS2			The p.Ter487TrpextTer27 variant in MECP2 (NM_004992.4) has been reported as a de novo occurrence (biological parentage confirmed) in an individual with a neurodevelopmental disorder (internal database - GeneDx) (PS2).
PM2_Supporting			The p.Ter487TrpextTer27 variant in MECP2 is absent from gnomAD (PM2_supporting).
PM4_Strong			The p.Ter487TrpextTer27 variant in MECP2 (NM_004992.4) occurs at the stop codon and causes a change in the length of the protein and an elongated mRNA transcript of MECP2 (PM4_strong).
PS4_Moderate			The p.Ter487TrpextTer27 variant has been observed in 3 individuals with features of Rett syndrome (PMID 21807996; internal database - LabCorp (formerly Invitae); internal database - GeneDx) (PS4_moderate).

Not Met criteria codes

PP4			No code specific comments provided, please refer to the summary above or general recommendations provided in the guideline
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Curation History [↗](#)

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