

# R228= — WFS1 Molecular Atlas Card

**Variant type:** Synonymous (silent)

**Codon:** position 228 (Arginine, R) — amino acid unchanged

**Domain context:** N-terminal cytoplasmic (intrinsically disordered)

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## SCHEMA CATEGORY: SILENT — SILENT — NO AMINO-ACID CHANGE

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No amino-acid change (R228 is unchanged): the codon is altered but the protein sequence is identical to wild-type. No structural, stability or AlphaMissense effect applies. Synonymous variants are typically benign unless they affect splicing or regulatory elements; this one is not adjacent to an exon boundary.

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## CLINICAL EVIDENCE

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- **Classification:** Benign
- **Review status:** criteria provided, multiple submitters, no conflicts
- **Associated conditions:** WFS1-Related Spectrum Disorders; Wolfram syndrome 1; Autosomal dominant nonsyndromic hearing loss 6
- **cDNA change:** c.684C>G
- **ClinVar accession:** VCV000045460
- **Last evaluated:** 2026/02/04 00:00
- **Submissions:** 1

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*Card generated by wolfram-atlas-batch (synonymous pipeline) on 2026-06-08T02:51:38.093954Z.*

*WFS1: UniProt O76024, AlphaFold v6. Synonymous variants carry no protein-structural effect.*