

# c.1385\_1393del — WFS1 Molecular Atlas Card

**Variant type:** In-frame indel

**Change:** 3 residue(s) deleted in frame at position 462

**Domain context:** Transmembrane helix 6

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## SCHEMA CATEGORY: I3 — MULTI-RESIDUE IN-FRAME INDEL — LIKELY MAJOR STRUCTURAL DISRUPTION

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3 residues removed in frame around position 462 (Transmembrane helix 6). A change this size usually perturbs local packing and can propagate to the fold. Gene therapy is the primary path unless an AlphaFold prediction of the modified sequence shows a surprisingly intact fold. Predicted structure pending (ColabFold).

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## STRUCTURAL PREDICTION

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- **Reading frame:** preserved (in-frame) — no premature stop, NMD does not apply.
- **Affected domain:** Transmembrane helix 6
- **Predicted modified structure:** \_pending — AlphaFold/ColabFold prediction of the modified sequence and backbone-RMSD vs wild-type backfill here (Wave 2).\_  
[View structure](#)

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## CLINICAL EVIDENCE

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- **Classification:** Likely pathogenic
- **Review status:** criteria provided, single submitter
- **Associated conditions:** Wolfram syndrome 1
- **cDNA change:** c.1385\_1393del
- **ClinVar accession:** VCV000004514
- **Last evaluated:** 1/01/01 00:00

- **Submissions:** 1
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Card generated by `wolfram-atlas-batch` (in-frame indel pipeline) on 2026-06-08T02:41:09.929619Z.

Schema: `reference/card schemaextension.md` (I1–I3). WFS1: UniProt O76024, AlphaFold v6.