Vutrisiran: Hypersensitivity Reactions

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SUMMARY

- In the phase 3 HELIOS-A and HELIOS-B studies, no increased risk of serious or severe hypersensitivity reaction was observed, and no specific risk factors were identified. However, as with all drugs, hypersensitivity reactions are possible. Patients should be treated appropriately for signs of hypersensitivity reactions at the clinical discretion of the healthcare provider.[1]
- A pooled safety analysis including data from 707 patients who received at least one dose of vutrisiran at any time during the HELIOS-A and HELIOS-B studies was conducted to evaluate the safety of vutrisiran in patients with ATTR who received treatment for up to 58 months.[2]
  - Hypersensitivity was not identified as a common adverse event with vutrisiran. Adverse events were consistent with those reported for patients who received vutrisiran during the randomized periods of HELIOS-A and HELIOS-B.[2]
- A cumulative post-marketing review of Alnylam Pharmaceuticals' global safety database did not identify safety concerns involving hypersensitivity reactions related to vutrisiran. Hypersensitivity reactions continue to be closely monitored through routine pharmacovigilance activities.[3]

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VUTRISIRAN FORMULATION AND ADMINISTRATION INFORMATION

Patients who receive vutrisiran do not require premedication to decrease the risk of hypersensitivity reactions.[4] Vutrisiran utilizes GalNAc conjugate technology as the method of drug delivery which allows for subcutaneous injection. GalNAc conjugation facilitates siRNA delivery into the liver via the ASGPR expressed on hepatocytes.[5] Vutrisiran uses second generation ESC, which includes a combination of additional phosphorothioate linkages, as well as 2′-O-methyl nucleotide and 2′-fluoro nucleotide modifications, that provide improved molecular stability and minimized metabolic lability.[6,7]

GLOBAL SAFETY DATABASE

A cumulative post-marketing review of Alnylam Pharmaceuticals' global safety database did not identify safety concerns involving hypersensitivity reactions related to vutrisiran. Hypersensitivity reactions continue to be closely monitored through routine pharmacovigilance activities.[3]

ABBREVIATIONS

ASGPR = asialoglycoprotein receptor; ATTR = transthyretin amyloidosis; ESC = enhanced stabilization chemistry; GalNAc = N-acetylgalactosamine; siRNA = small-interfering ribonucleic acid.

Updated 6 October 2025

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