

## Vutrisiran: Blood-Brain Barrier

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### SUMMARY

- The PK and ADME properties of vutrisiran were evaluated to support pivotal toxicology studies and clinical development of vutrisiran.<sup>1</sup>
- Vutrisiran did not appear to cross the blood-brain barrier in rats administered radiolabeled medication.<sup>1</sup>

### INDEX

[Pre-Clinical Data](#) – [Abbreviations](#) – [References](#)

### PRE-CLINICAL DATA

The PK and ADME properties of vutrisiran were evaluated in various in vitro and in vivo studies to support pivotal toxicology studies and clinical development of vutrisiran.<sup>1</sup>

#### Vutrisiran Quantitative Whole-body Autoradiography in Rats

Quantitative tissue distribution of total drug-related radioactivity was investigated in male rats following administration of a single 3 mg/kg SC dose of [<sup>3</sup>H]-vutrisiran. [<sup>3</sup>H]-Vutrisiran was radiolabeled on the tenth nucleotide (adenosine) from the 3' end of the antisense strand. [<sup>3</sup>H]-Vutrisiran-derived radioactivity was distributed to limited tissues over time. Vutrisiran did not appear to cross the blood-brain barrier as there was no radioactivity detected in the brain or any part of the central nervous system.<sup>1</sup>

### ABBREVIATIONS

ADME = absorption, distribution, metabolism, and excretion; PK = pharmacokinetics; SC = subcutaneous

*Updated 19 March 2026*

### REFERENCES

1. Alnylam Pharmaceuticals. Data on file. MED-US-TTRSC02-2200053.