

Preemptive policy: This is a P&T approved policy and can be used after the drug is FDA approved until it is superseded by an updated policy



## Clinical Policy: Rebisufligene Etisparvovec (UX111)

Reference Number: CP.PHAR.722

Effective Date: **FDA Approval Date**

Last Review Date: 05.25

Line of Business: Commercial, HIM, Medicaid

[Coding Implications](#)

[Revision Log](#)

See [Important Reminder](#) at the end of this policy for important regulatory and legal information.

### Description

Rebisufligene etisparvovec (Brand Name<sup>®/™</sup>) is adeno-associated virus (AAV) vector-based gene therapy.

### FDA Approved Indication(s) **[Pending]**

UX111 is indicated for the treatment of mucopolysaccharidosis type IIIA (MPS IIIA; Sanfilippo syndrome type A).

### Policy/Criteria

*Provider must submit documentation (such as office chart notes, lab results or other clinical information) supporting that member has met all approval criteria.*

All requests reviewed under this policy **require medical director review**.

It is the policy of health plans affiliated with Centene Corporation<sup>®</sup> that UX111 is **medically necessary** when the following criteria are met:

#### I. Initial Approval Criteria\*

*\*Criteria will mirror the clinical information from the prescribing information once FDA-approved*

##### A. Mucopolysaccharidosis IIIA: Sanfilippo Syndrome Type A (must meet all):

1. Diagnosis of MPS IIIA confirmed by both of the following (a and b):\*
  - a. Genetic confirmation of homozygous or compound heterozygous mutations in the *SGSH* gene;
  - b. Leukocyte assay demonstrating undetectable or significantly reduced SGSH enzyme activity;
2. Prescribed by or in consultation with a specialist with expertise in lysosomal storage diseases (e.g., medical geneticist, pediatric endocrinologist) or neurologist;
3. Age  $\leq$  4 years;\*
4. Member does not have anti-AAV9 antibody titers  $\geq$  1:100;\*
5. Member does not have either of the following via genetic testing (a and b):\*
  - a. Two nonsense or null variants of the *SGSH* gene (*see Appendix D*);
  - b. One or more *S298P* mutation(s) in the *SGSH* gene;
6. Member has not previously received hematopoietic stem cell transplantation;\*
7. Member has not previously received cell or gene therapy;\*
8. Documentation of member's current weight in kg;

9. Dose does not exceed  $3 \times 10^{13}$  vector genomes/kg.\*

**Approval duration: 3 months (one infusion per lifetime)**

**B. Other diagnoses/indications (must meet 1 or 2):**

1. If this drug has recently (within the last 6 months) undergone a label change (e.g., newly approved indication, age expansion, new dosing regimen) that is not yet reflected in this policy, refer to one of the following policies (a or b):
  - a. For drugs on the formulary (commercial, health insurance marketplace) or PDL (Medicaid), the no coverage criteria policy for the relevant line of business: CP.CPA.190 for commercial, HIM.PA.33 for health insurance marketplace, and CP.PMN.255 for Medicaid; or
  - b. For drugs NOT on the formulary (commercial, health insurance marketplace) or PDL (Medicaid), the non-formulary policy for the relevant line of business: CP.CPA.190 for commercial, HIM.PA.103 for health insurance marketplace, and CP.PMN.16 for Medicaid; or
2. If the requested use (e.g., diagnosis, age, dosing regimen) is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized) AND criterion 1 above does not apply, refer to the off-label use policy for the relevant line of business: CP.CPA.09 for commercial, HIM.PA.154 for health insurance marketplace, and CP.PMN.53 for Medicaid.

**II. Continued Therapy\***

*\*Criteria will mirror the clinical information from the prescribing information once FDA-approved*

**A. Mucopolysaccharidosis IIIA: Sanfilippo Syndrome Type A**

1. Re-authorization is not permitted as UX111 is indicated to be dosed one time only.

**Approval duration: Not applicable**

**B. Other diagnoses/indications (must meet 1 or 2):**

1. If this drug has recently (within the last 6 months) undergone a label change (e.g., newly approved indication, age expansion, new dosing regimen) that is not yet reflected in this policy, refer to one of the following policies (a or b):
  - a. For drugs on the formulary (commercial, health insurance marketplace) or PDL (Medicaid), the no coverage criteria policy for the relevant line of business: CP.CPA.190 for commercial, HIM.PA.33 for health insurance marketplace, and CP.PMN.255 for Medicaid; or
  - b. For drugs NOT on the formulary (commercial, health insurance marketplace) or PDL (Medicaid), the non-formulary policy for the relevant line of business: CP.CPA.190 for commercial, HIM.PA.103 for health insurance marketplace, and CP.PMN.16 for Medicaid; or
2. If the requested use (e.g., diagnosis, age, dosing regimen) is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized) AND criterion 1 above does not apply, refer to the off-label use policy for the relevant line of business: CP.CPA.09 for commercial, HIM.PA.154 for health insurance marketplace, and CP.PMN.53 for Medicaid.

**III. Diagnoses/Indications for which coverage is NOT authorized:**

- A. Non-FDA approved indications, which are not addressed in this policy, unless there is sufficient documentation of efficacy and safety according to the off label use policies – CP.CPA.09 for commercial, HIM.PA.154 for health insurance marketplace, and CP.PMN.53 for Medicaid or evidence of coverage documents.

**IV. Appendices/General Information**

*Appendix A: Abbreviation/Acronym Key*

AAV: adeno-associated virus

FDA: Food and Drug Administration

MPS: mucopolysaccharidosis

SGSH: N-sulfolglucosamine sulfohydrolase

*Appendix B: Therapeutic Alternatives*

Not applicable

*Appendix C: Contraindications/Boxed Warnings [Pending]*

- Contraindication(s): **pending**
- Boxed warning(s): **pending**

*Appendix D: Gene Mutation Definitions*

- Nonsense variant: A nonsense mutation occurs in DNA when a sequence change gives rise to a stop codon rather than a codon specifying an amino acid. The presence of the new stop codon results in the production of a shortened protein that is likely non-functional.
- Null variant: Refers to a pathogenic variant that results in either no mRNA, no protein, or a nonfunctional protein

**V. Dosage and Administration [Pending]**

Indication	Dosing Regimen	Maximum Dose
MPS IIIA*	Pending	Pending

**VI. Product Availability [Pending]**

Pending

**VII. References**

1. ClinicalTrials.gov. Phase I/II/III gene transfer clinical trial of ScAAV9.U1a.hSGSH. Available at: <https://clinicaltrials.gov/study/NCT02716246>. Accessed January 27, 2025.
2. Muschol N, Giugliani R, Jones SA, et al. Sanfilippo syndrome: consensus guidelines for clinical care. Orphanet J Rare Dis. 2022;17(1):391.
3. Adam MP, Feldman J, Mirzaa GM, et al. GeneReviews<sup>®</sup> [Internet]. Seattle (WA): University of Washington, Seattle; 1993-2025. Available at: <https://www.ncbi.nlm.nih.gov/books/NBK5191/>. Accessed February 14, 2025.

**Coding Implications [Pending]**

Codes referenced in this clinical policy are for informational purposes only. Inclusion or exclusion of any codes does not guarantee coverage. Providers should reference the most up-to-date sources of professional coding guidance prior to the submission of claims for reimbursement of covered services.

HCPCS Codes	Description
Pending	Pending

Reviews, Revisions, and Approvals	Date	P&T Approval Date
Policy created pre-emptively	03.11.25	05.25

**Important Reminder**

This clinical policy has been developed by appropriately experienced and licensed health care professionals based on a review and consideration of currently available generally accepted standards of medical practice; peer-reviewed medical literature; government agency/program approval status; evidence-based guidelines and positions of leading national health professional organizations; views of physicians practicing in relevant clinical areas affected by this clinical policy; and other available clinical information. The Health Plan makes no representations and accepts no liability with respect to the content of any external information used or relied upon in developing this clinical policy. This clinical policy is consistent with standards of medical practice current at the time that this clinical policy was approved. “Health Plan” means a health plan that has adopted this clinical policy and that is operated or administered, in whole or in part, by Centene Management Company, LLC, or any of such health plan’s affiliates, as applicable.

The purpose of this clinical policy is to provide a guide to medical necessity, which is a component of the guidelines used to assist in making coverage decisions and administering benefits. It does not constitute a contract or guarantee regarding payment or results. Coverage decisions and the administration of benefits are subject to all terms, conditions, exclusions, and limitations of the coverage documents (e.g., evidence of coverage, certificate of coverage, policy, contract of insurance, etc.), as well as to state and federal requirements and applicable Health Plan-level administrative policies and procedures.

This clinical policy is effective as of the date determined by the Health Plan. The date of posting may not be the effective date of this clinical policy. This clinical policy may be subject to applicable legal and regulatory requirements relating to provider notification. If there is a discrepancy between the effective date of this clinical policy and any applicable legal or regulatory requirement, the requirements of law and regulation shall govern. The Health Plan retains the right to change, amend or withdraw this clinical policy, and additional clinical policies may be developed and adopted as needed, at any time.

This clinical policy does not constitute medical advice, medical treatment, or medical care. It is not intended to dictate to providers how to practice medicine. Providers are expected to exercise professional medical judgment in providing the most appropriate care, and are solely responsible

for the medical advice and treatment of members. This clinical policy is not intended to recommend treatment for members. Members should consult with their treating physician in connection with diagnosis and treatment decisions.

Providers referred to in this clinical policy are independent contractors who exercise independent judgment and over whom the Health Plan has no control or right of control. Providers are not agents or employees of the Health Plan.

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**Note:**

**For Medicaid members**, when state Medicaid coverage provisions conflict with the coverage provisions in this clinical policy, state Medicaid coverage provisions take precedence. Please refer to the state Medicaid manual for any coverage provisions pertaining to this clinical policy.

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