

UTILIZATION MANAGEMENT MEDICAL POLICY

POLICY: Gaucher Disease – Enzyme Replacement Therapy – Vpriv Utilization Management Medical Policy

- Vpriv® (velaglucerase intravenous infusion – Shire Human Genetic Therapies)

REVIEW DATE: 04/10/2024; selected revision 07/17/2024

OVERVIEW

Vpriv, an analogue of β -glucocerebrosidase, is indicated for long-term enzyme replacement therapy for **Type 1 Gaucher disease**.¹

The efficacy and safety of Vpriv have not been established in pediatric patients younger than 4 years of age.¹

Disease Overview

Gaucher disease is a rare autosomal recessive, inherited, lysosomal storage disorder caused by a deficiency of the lysosomal enzyme β -glucocerebrosidase.²⁻⁴ Glucocerebrosidase is responsible for the breakdown of glucosylceramide (GluCer) into glucose and ceramide. A deficiency of this enzyme is characterized by an excessive accumulation of GluCer in the visceral organs such as the liver, spleen, and bone marrow. GluCer remains stored within lysosomes causing enlarged lipid-laden macrophages called “Gaucher cells”.

Gaucher disease is classified into three phenotypes (Types 1 through 3).²⁻⁵ Type 1 is a non-neuronopathic variant with asymptomatic or symptomatic clinical manifestations of splenomegaly, hepatomegaly, anemia, thrombocytopenia, skeletal complications, and occasional lung involvement. Type 2 is an acute neuronopathic form characterized by an early onset (3 to 6 months of age) of rapidly progressive neurological disease with visceral manifestations; death generally occurs by the time patients reach 1 to 2 years of age. Type 3 is referred to as a chronic neuronopathic form and characterized by a later onset. Patients present with neurological, hematological, and visceral symptoms. Type 1 is most prevalent in the Western world, accounting for an estimated 94% of patients with Gaucher disease.^{2,6} Types 2 and 3 represent < 1% and 5%, respectively, in Europe, North America, and Israel.^{2,5} The diagnosis of Gaucher disease is established by demonstrating deficient β -glucocerebrosidase activity in leukocytes or fibroblasts, or mutations in the glucocerebrosidase gene.^{7,8}

Guidelines

Treatment guidelines for Type 1 Gaucher disease (non-neuronopathic form) recommend initiating enzyme replacement therapy (ERT) in patients with significant and/or progressive disease.^{9,10} Additionally, ERT should be initiated immediately in all patients with Type 3 Gaucher disease (chronic neuronopathic form).¹¹ Guidelines note that there is no evidence that ERT has reversed, stabilized, or slowed the progression of neurological involvement. However, ERT ameliorates systemic involvement (skeletal deterioration, visceromegaly, hematological abnormalities) in non-neuronopathic as well as chronic neuronopathic disease, ultimately enhancing the quality of life. Additionally, it is noted that higher doses may be needed to control visceral symptoms associated with chronic neuronopathic disease.

POLICY STATEMENT

Prior Authorization is recommended for medical benefit coverage of Vpriv. Approval is recommended for those who meet the **Criteria** and **Dosing** for the listed indications. Extended approvals are allowed if the

patient continues to meet the Criteria and Dosing. Requests for doses outside of the established dosing documented in this policy will be considered on a case-by-case basis by a clinician (i.e., Medical Director or Pharmacist). All approvals are provided for the duration noted below. Because of the specialized skills required for evaluation and diagnosis of patients treated with Vpriv as well as the monitoring required for adverse events and long-term efficacy, approval requires Vpriv to be prescribed by or in consultation with a physician who specializes in the condition being treated.

Automation: None.

RECOMMENDED AUTHORIZATION CRITERIA

Coverage of Vpriv is recommended in those who meet one of the following criteria:

FDA-Approved Indication

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- 1. Gaucher Disease – Type 1.** Approve for 1 year if the patient meets ALL of the following (A, B, and C):

Note: Type 1 Gaucher disease is also known as non-neuronopathic Gaucher disease.

A) Patient is ≥ 4 years of age; AND

B) The diagnosis is established by ONE of the following (i or ii):

i. Demonstration of deficient β -glucocerebrosidase activity in leukocytes or fibroblasts; OR

ii. Molecular genetic testing documenting biallelic pathogenic variants in the glucocerebrosidase (*GBA*) gene; AND

C) Vpriv is prescribed by or in consultation with a geneticist, endocrinologist, a metabolic disorder sub-specialist, or a physician who specializes in the treatment of lysosomal storage disorders.

Dosing. Each individual dose must not exceed 60 U/kg administered intravenously no more frequently than once every 2 weeks.

Other Uses with Supportive Evidence

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- 2. Gaucher Disease – Type 3.** Approve for 1 year if the patient meets ALL of the following (A, B, C, and D):

Note: Type 3 Gaucher disease is also known as chronic neuronopathic Gaucher disease.

A) Patient is ≥ 4 years of age; AND

B) The diagnosis is established by ONE of the following (i or ii):

i. Demonstration of deficient β -glucocerebrosidase activity in leukocytes or fibroblasts; OR

ii. Molecular genetic testing documenting biallelic pathogenic variants in the glucocerebrosidase (*GBA*) gene; AND

C) The patient meets BOTH of the following (i and ii):

i. Medication is not being used for the management of neurological manifestations; AND

Note: Examples of neurological manifestations may include abnormal ocular movement, auditory impairment, cognitive impairment, and seizures.

ii. Medication is being used for the management of impaired growth, hematologic, or visceral symptoms; AND

Note: Examples of visceral symptoms include splenomegaly and hepatomegaly. Examples of hematologic symptoms include anemia and thrombocytopenia.

- D)** Vpriv is prescribed by or in consultation with a geneticist, endocrinologist, a metabolic disorder sub-specialist, or a physician who specializes in the treatment of lysosomal storage disorders.

Dosing. Each individual dose must not exceed 120 U/kg administered intravenously no more frequently than once every 2 weeks.

CONDITIONS NOT RECOMMENDED FOR APPROVAL

Coverage of Vpriv is not recommended in the following situations:

- 1. Concomitant Use with Other Approved Therapies for Gaucher Disease.** Concomitant use with other treatments approved for Gaucher disease has not been evaluated. Of note, examples of medications approved for Gaucher disease include Cerdelga (eliglustat capsules), Elelyso (taliglucerase alfa intravenous infusion), Cerezyme (imiglucerase intravenous infusion), and Zavesca (miglustat capsules).
- 2.** Coverage is not recommended for circumstances not listed in the Recommended Authorization Criteria. Criteria will be updated as new published data are available.

REFERENCES

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8. Baris HN, Cohen IJ, Mistry PK. Gaucher disease: The metabolic defect, pathophysiology, phenotypes and natural history. *Pediatr Endocrinol Rev.* 2014;12:72-81.
9. Kishnani PS, Al-Hertani W, Balwani M, et al. Screening, patient identification, evaluation, and treatment in patients with Gaucher disease: Results from a Delphi consensus. *Mol Genet Metab.* 2022 Feb;135(2):154-162.
10. Kaplan P, Baris H, De Meirleir L, et al. Revised recommendations for the management of Gaucher disease in children. *Eur J Pediatr.* 2013 Apr;172(4):447-58.
11. Vellodi A, Tytki-Szymanska A, Davies EH, et al. Management of neuronopathic Gaucher disease: revised recommendations. *J Inherit Metab Dis.* 2009 Oct;32(5):660-664.

HISTORY

Type of Revision	Summary of Changes	Review Date
Annual Revision	No criteria changes.	04/05/2023
Annual Revision	No criteria changes.	04/10/2024
Selected Revision	Gaucher Disease – Type 1: Added qualifier “Type 1” to the condition name and Note to indicate Type 1 disease is also referred to as non-neuronopathic disease. Added age ≥ 4 years as a condition of approval. For diagnosis established by genetic testing, genetic testing demonstrating a mutation in the glucocerebrosidase (<i>GBA</i>) gene was further specified to state a genetic test documenting biallelic pathogenic variants in the <i>GBA</i> gene. Gaucher Disease – Type 3: This new condition of approval was added under other uses with supportive evidence. Concomitant use with other approved therapies for Gaucher disease was added under conditions not recommended for approval.	07/17/2024