PreferredOne®

PRIOR AUTHORIZATION POLICY

POLICY: Hemophilia Factor IX Products Prior Authorization Policy

Extended Half-Life Recombinant Products

- Alprolix® (Coagulation Factor IX [recombinant] Fc fusion protein injection – Bioverativ)
- Idelvion (Coagulation Factor IX [recombinant] albumin fusion protein injection – CSL Behring)
- Rebinyn® (Coagulation Factor IX [recombinant] glycoPEGylated injection – NovoNordisk)

Standard Half-Life Recombinant Products

- BeneFIX® (Coagulation Factor IX [recombinant] injection – Wyeth/Pfizer)
- Ixinity® (Coagulation Factor IX [recombinant] injection – Aptevo BioTherapeutics)
- Rixubis® (Coagulation Factor IX [recombinant] injection – Baxalta)

Plasma-Derived Products

- AlphaNine® SD (Coagulation Factor IX [plasma-derived] injection – Grifols)
- Mononine® (Coagulation Factor IX [plasma-derived] injection – CSL Behring)
- Profilnine® (Factor IX Complex [plasma-derived] injection – Grifols)

REVIEW DATE: 03/03/2021

OVERVIEW

Alprolix, Idelvion, and Rebinyn are extended half-life recombinant Factor IX products; BeneFIX, Ixinity and Rixubis are standard half-life recombinant Factor IX products; and AlphaNine SD, Mononine, and Profilnine plasma-derived Factor IX products.[7-9] All agents are indicated in various clinical scenarios for use in the management of patients with hemophilia B.

Profilnine is used in patients with Factor II and/or X deficiency.[10] Some data are available, albeit limited.

Disease Overview

Hemophilia B is a recessive X-linked bleeding disorder caused by mutations in the factor IX gene that leads to the deficiency or absence of the coagulation factor IX.[11-14] It occurs in 1 out of 30,000 male births and affects about 5,000 people in the US. Hemophilia B predominantly occurs in males; however, approximately 10% of females are carriers and are at risk of usually mild bleeding. The severity of bleeding depends on the degree of the factor IX defect and the phenotypic expression. Factor levels of <1%, 1% to 5%, and > 5% to < 40% are categorized as severe, moderate, and mild hemophilia B, respectively. Patients with mild hemophilia B may only experience abnormal bleeding during surgery, during tooth extractions, or when injured. Patients with moderate hemophilia B generally have prolonged bleeding responses to minor trauma. Severe hemophilia B is marked by spontaneous bleeding such as spontaneous hemarthrosis, soft-tissue hematomas, retroperitoneal bleeding, intracerebral hemorrhage, and delayed bleeding post-surgery. Complications from recurrent bleeding and soft-tissue hematomas include severe arthropathy, and joint contractures, which may lead to pain and disability. The main treatment of hemophilia B is replacement of missing blood coagulation with Factor IX products. Factor IX replacement therapy may be used on-demand when bleeding occurs or given as routine prophylaxis with scheduled infusions. Both plasma-derived and recombinant Factor IX products are available. In general, prophylactic therapy has been associated with a reduction in bleeds and improved outcomes for selected patients (e.g., patients with moderate or severe factor IX deficiency). The goal of therapy is to prevent uncontrolled internal hemorrhage and severe joint damage and to properly manage bleeding episodes. The development of

inhibitors occurs at a lower frequency in patients with severe hemophilia B compared with severe hemophilia A but can occur in up to 5% of patients. Higher doses than that typically used for the uses of standard half-life products can be given if the patient develops an inhibitor.

Guidelines

Guidelines for hemophilia from the National Hemophilia Foundation (2020)[13] and the World Federation of Hemophilia (2020)[14] recognize the important role of Factor IX products in the management of hemophilia B in patients.

POLICY STATEMENT

Prior Authorization is recommended for prescription benefit coverage of the following Factor IX products: Alprolix, Idelvion, Rebinyn, BeneFIX, Ixinity, Rixubis, AlphaNine, Mononine, and Profilnine. All approvals are provided for the duration noted below. Because of the specialized skills required for evaluation and diagnosis of patients treated with recombinant Factor IX products, as well as the monitoring required for adverse events and long-term efficacy, the agent is required to be prescribed by or in consultation with a physician who specializes in the condition being treated.

Automation: None.

RECOMMENDED AUTHORIZATION CRITERIA

I. Coverage of Alprolix, Idelvion, Rebinyn, BeneFIX, Ixinity, and Rixubis is recommended for patients who meet the following criteria:

FDA-Approved Indications

1. **Hemophilia B.** Approve for 1 year if the agent is prescribed by or in consultation with a hemophilia specialist.

II. Coverage of AlphaNine SD, Mononine, and Profilnine is recommended for patients who meet the following criteria:

FDA-Approved Indications

1. **Hemophilia B.** Approve for 1 year if the agent is prescribed by or in consultation with a hemophilia specialist.

III. Coverage of Profilnine is recommended for patients who meet the following criteria:

Other Uses with Supportive Evidence

2. **Factor II Deficiency.** Approve Profilnine for 1 year if the agent is prescribed by or in consultation with a hemophilia specialist.

3. **Factor X Deficiency.** Approve Profilnine for 1 year if the agent is prescribed by or in consultation with a hemophilia specialist.

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CONDITIONS NOT RECOMMENDED FOR APPROVAL

Coverage of the cited Factor IX products is not recommended in the following situations:

1. Coverage is not recommended for circumstances not listed in the Recommended Authorization Criteria. Criteria will be updated as new published data are available.

REFERENCES

1. Alprolix® lyophilized powder for intravenous injection [prescribing information]. Waltham, MA: Bioverativ; October 2020.
2. Idelvion® lyophilized powder for solution for intravenous injection [prescribing information]. Kankakee, IL: CSL Behring; July 2020.
3. Rebinyn® lyophilized powder for solution for intravenous injection [prescribing information]. Plainsboro, NJ: Novo Nordisk; June 2020.
4. BeneFIX® injection for intravenous use [prescribing information]. Philadelphia, PA: Wyeth Pharmaceuticals, Inc. (a subsidiary of Pfizer); June 2020.
5. Ixinity® solution for intravenous injection [prescribing information]. Seattle, WA: Aptevo BioTherapeutics; September 2020.
6. Rixubis® for intravenous injection [prescribing information]. Lexington, MA: Baxalta; June 2020.
7. AlphaNine® SD injection [prescribing information]. Los Angeles, CA: Grifols; June 2018.
8. Mononine® injection [prescribing information]. Kankakee, IL: CSL Behring; December 2018.
9. Profilnine® injection [prescribing information]. Los Angeles, CA: Grifols; June 2018.
10. Menegatti M, Peyvandi F. Treatment of rare factor deficiencies other than hemophilia. *Blood*. 2019;133(5):415-424.
11. Franchini M. Current management of hemophilia B: recommendations, complications and emerging issues. *Expert Rev Hematol.* 2014;7(5):573-581.
12. Peyvandi F, Garagiola I, Young G. The past and future of haemophilia: diagnosis, treatments, and its complications. *Lancet*. 2016;388(10040):187-197.
13. National Hemophilia Foundation. Medical and Scientific Advisory Council (MASAC) recommendations concerning products licensed for the treatment of hemophilia and other bleeding disorders (Revised August 2020). MASAC document #263. Available at: 263_treatment.pdf (hemophilia.org). Accessed on February 22, 2021.
14. Srivastava A, Santagostino E, Dougall A, on behalf of the WFH guidelines for the management of hemophilia panelists and co-authors. Guidelines for the management of hemophilia, 3rd edition. *Haemophilia*. 2020;26(Suppl 6):1-158. Available at: WFH Guidelines for the Management of Hemophilia, 3rd edition (wiley.com). Accessed on February 13, 2021.

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Grievance Specialist
PreferredOne Community Health Plan
PO Box 59052
Minneapolis, MN 55459-0052
Phone: 1.800.940.5049 (TTY: 763.847.4013)
Fax: 763.847.4010
customerservice@preferredone.com

You can file a grievance in person or by mail, fax, or email. If you need help filing a grievance, a Grievance Specialist is available to help you.

You can also file a civil rights complaint with the U.S. Department of Health and Human Services, Office for Civil Rights, electronically through the Office for Civil Rights Complaint Portal, available at https://ocrportal.hhs.gov/ocr/portal/lobby.jsf, or by mail or phone at:

U.S. Department of Health and Human Services
200 Independence Avenue, SW
Room 509F, HHH Building
Washington, D.C. 20201
1-800-368-1019, 800-537-7697 (TDD)

Complaint forms are available at http://www.hhs.gov/ocr/office/file/index.html.

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Fax: 763.847.4010
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U.S. Department of Health and Human Services
200 Independence Avenue, SW
Room 509F, HHH Building
Washington, D.C. 20201
1-800-368-1019, 800-537-7697 (TDD)

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