



Drug Coverage Policy

Effective Date05/15/2025

Coverage Policy Number.....IP0730

Policy Title.....Alhemo

Hemophilia – Alhemo

- Alhemo® (concizumab-mtci subcutaneous injection – NovoNordisk)

INSTRUCTIONS FOR USE

The following Coverage Policy applies to health benefit plans administered by Cigna Companies. Certain Cigna Companies and/or lines of business only provide utilization review services to clients and do not make coverage determinations. References to standard benefit plan language and coverage determinations do not apply to those clients. Coverage Policies are intended to provide guidance in interpreting certain standard benefit plans administered by Cigna Companies. Please note, the terms of a customer's particular benefit plan document [Group Service Agreement, Evidence of Coverage, Certificate of Coverage, Summary Plan Description (SPD) or similar plan document] may differ significantly from the standard benefit plans upon which these Coverage Policies are based. For example, a customer's benefit plan document may contain a specific exclusion related to a topic addressed in a Coverage Policy. In the event of a conflict, a customer's benefit plan document always supersedes the information in the Coverage Policies. In the absence of a controlling federal or state coverage mandate, benefits are ultimately determined by the terms of the applicable benefit plan document. Coverage determinations in each specific instance require consideration of 1) the terms of the applicable benefit plan document in effect on the date of service; 2) any applicable laws/regulations; 3) any relevant collateral source materials including Coverage Policies and; 4) the specific facts of the particular situation. Each coverage request should be reviewed on its own merits. Medical directors are expected to exercise clinical judgment where appropriate and have discretion in making individual coverage determinations. Where coverage for care or services does not depend on specific circumstances, reimbursement will only be provided if a requested service(s) is submitted in accordance with the relevant criteria outlined in the applicable Coverage Policy, including covered diagnosis and/or procedure code(s). Reimbursement is not allowed for services when billed for conditions or diagnoses that are not covered under this Coverage Policy (see "Coding Information" below). When billing, providers must use the most appropriate codes as of the effective date of the submission. Claims submitted for services that are not accompanied by covered code(s) under the applicable Coverage Policy will be denied as not covered. Coverage Policies relate exclusively to the administration of health benefit plans. Coverage Policies are not recommendations for treatment and should never be used as treatment guidelines. In certain markets, delegated vendor guidelines may be used to support medical necessity and other coverage determinations.

Overview

Alhemo, a tissue factor pathway inhibitor antagonist, is indicated for routine prophylaxis to prevent or reduce the frequency of bleeding episodes in adults and pediatric patients ≥ 12 years

of age with 1) hemophilia A (congenital Factor VIII deficiency) with Factor VIII inhibitors, or 2) hemophilia B (congenital Factor IX deficiency) with Factor IX inhibitors.¹

Alhemo is recommended to be given as a 1 mg/kg loading dose on Day 1 as a subcutaneous (SC) injection once daily (QD).¹ On Day 2, the maintenance dose of 2 mg/kg Alhemo SC QD is recommended with individualization doses based on Alhemo concentrations obtained at 4 weeks following initiation of treatment. After proper training, Alhemo may be self-administered.

Disease Overview

Hemophilia A and B are genetic bleeding disorders caused by a dysfunction or a deficiency of coagulation Factor VIII and Factor IX, respectively.²⁻⁷ Because hemophilia is an X-linked condition, males are primarily impacted. Patients who have these types of hemophilias are not able to properly form clots in blood and may bleed for a longer time than normal following injury or surgery. Patients may also experience spontaneous bleeding in muscles, joints, and organs. Bleeds may be life-threatening. A main morbidity is hemophilic arthropathy, which limits mobility. It is estimated that 33,000 males are living with hemophilia in the US; hemophilia A accounts for around 80% of the cases (approximately 26,400 patients) and hemophilia B comprises 20% of cases (around 6,600 patients). Hemophilias are often classified as mild, moderate, or severe based on reduced Factor VIII or IX levels. Approximately 50% and 30% of patients with hemophilia A and hemophilia B, respectively, have severe disease. The formation of inhibitors (antibodies) to factor products is a challenging complication as it causes Factor VIII and Factor IX therapies to be ineffective, which increases bleeding frequency and severity. Inhibitors develop in around 30% and 10% of patients with severe hemophilia A and hemophilia B, respectively.

Coverage Policy

POLICY STATEMENT

Prior Authorization is required for prescription benefit coverage of Alhemo. All approvals are provided for the duration noted below. Because of the specialized skills required for evaluation and diagnosis of patients treated with Alhemo as well as the monitoring required for adverse events and long-term efficacy, approval requires Alhemo to be prescribed by or in consultation with a hemophilia specialist.

Documentation: Documentation is required where noted in the criteria. Documentation may include, but not limited to, chart notes, laboratory tests, claims records, and/or other information.

Alhemo is considered medically necessary when ONE of the following is met (1 or 2):

FDA-Approved Indications

1. Hemophilia A with Factor VIII Inhibitors. Approve for 1 year if the patient meets ONE of the following (A or B):

A) Initial Therapy. Approve if the patient meets ALL of the following (i, ii, iii, iv and v):

- i.** Patient is ≥ 12 years of age; AND
- ii.** Patient is using Alhemo for routine prophylaxis to prevent or reduce the frequency of bleeding episodes; AND
- iii.** Patient meets BOTH of the following (a and b):
 - a)** Factor VIII inhibitor titer testing has been performed within the past 30 days; AND
 - b)** Patient has a positive test for Factor VIII inhibitors of ≥ 0.6 Bethesda units/mL; AND
- iv.** The medication is prescribed by or in consultation with a hemophilia specialist; AND

- v. Preferred product criteria is met for the product(s) as listed in the below table(s) [Employer Plans]; OR
- B) Patient is Currently Receiving Alhemo.** Approve if the patient meets ALL of the following (i, ii, and iii):
 - i. Patient is using Alhemo for routine prophylaxis to prevent or reduce the frequency of bleeding episodes; AND
 - ii. The medication is prescribed by or in consultation with a hemophilia specialist; AND
 - iii. According to the prescriber, patient experienced a beneficial response to therapy.
Note: Examples of a beneficial response to therapy include a reduction in bleeding events, in the severity of bleeding episodes, in the number of bleeding events that required treatment, and/or in the number of spontaneous bleeds.

2. Hemophilia B with Factor IX Inhibitors. Approve for 1 year if the patient meets ONE of the following (A or B):

- A) Initial Therapy.** Approve if the patient meets ALL of the following (i, ii, iii, and iv):
 - i. Patient is ≥ 12 years of age; AND
 - ii. Patient is using Alhemo for routine prophylaxis to prevent or reduce the frequency of bleeding episodes; AND
 - iii. Patient meets BOTH of the following (a and b):
 - a) Factor IX inhibitor titer testing has been performed within the past 30 days; AND
 - b) Patient has a positive test for Factor IX inhibitors of ≥ 0.6 Bethesda units/mL; AND
 - iv. The medication is prescribed by or in consultation with a hemophilia specialist; OR
- B) Patient is Currently Receiving Alhemo.** Approve if the patient meets ALL of the following (i, ii, and iii):
 - i. Patient is using Alhemo for routine prophylaxis to prevent or reduce the frequency of bleeding episodes; AND
 - ii. The medication is prescribed by or in consultation with a hemophilia specialist; AND
 - iii. According to the prescriber, patient experienced a beneficial response to therapy.
Note: Examples of a beneficial response to therapy include a reduction in bleeding events, in the severity of bleeding episodes, in the number of bleeding events that required treatment, and/or in the number of spontaneous bleeds.

Employer Plans:

Product	Criteria
Alhemo (concizumab-mtci subcutaneous injection)	Patient meets ONE of the following (1 <u>or</u> 2): <ol style="list-style-type: none"> 1. Alhemo is being used for the treatment of Hemophilia A and the patient meets ONE of the following (A, B, <u>or</u> C): <ol style="list-style-type: none"> A. Documentation is provided that the patient has tried and experienced inadequate efficacy or significant intolerance to Hemlibra B. According to the prescriber, there is concern for a drug-drug interaction (e.g., drug interaction with Hemlibra and Feiba) C. The patient has already been started on therapy with Alhemo. 2. Alhemo is being used for the treatment of Hemophilia B

When coverage is available and medically necessary, the dosage, frequency, duration of therapy, and site of care should be reasonable, clinically appropriate, and supported by evidence-based literature and adjusted based upon severity, alternative available treatments, and previous response to therapy.

Receipt of sample product does not satisfy any criteria requirements for coverage.

Alhemo for any other use is considered not medically necessary, including the following (this list may not be all inclusive; criteria will be updated as new published data are available):

1. Concurrent Use with Hemlibra (emicizumab-kxwh subcutaneous injection).

Hemlibra is a bispecific factor IXa- and Factor X-directed antibody indicated for routine prophylaxis to prevent or reduce the frequency of bleeding episodes in adults and pediatric patients ages newborn and older with hemophilia A (congenital factor VIII deficiency) with or without factor VIII inhibitors.⁸ Alhemo has not been studied concurrently with Hemlibra.

2. Concurrent Use with Hympavzi (marstacimab-hncq subcutaneous injection).

Hypmavzi, a tissue factor pathway inhibitor antagonist, is indicated for routine prophylaxis to prevent or reduce the frequency of bleeding episodes in patients ≥ 12 years of age with 1) hemophilia A (congenital Factor VIII deficiency) without Factor VIII inhibitors, and 2) hemophilia B (congenital Factor IX deficiency) without Factor IX inhibitors.⁹ Alhemo and Hympavzi should not be used concomitantly.

3. Concurrent Use of Bypassing Agents for Routine Prophylaxis. Alhemo has not been studied with bypassing agents for routine prophylaxis.

Note: Use of bypassing agents for the treatment of breakthrough bleeding is permitted. Examples of bypassing agents include NovoSeven RT (coagulation Factor VIIa [recombinant] intravenous infusion), Sevenfact (Factor VIIa [recombinant]-jncw intravenous infusion), and FEIBA (anti-inhibitor coagulant complex intravenous infusion).

4. Patient Receiving Immune Tolerance Induction Therapy. The safety and efficacy of concomitant use of Alhemo in patients undergoing immune tolerance induction have not been established.

References

1. Alhemo® subcutaneous injection [prescribing information]. Plainsboro, NJ: Novo Nordisk; December 2024.
2. Mancuso ME, Mahlangu JN, Pipe SW. The changing treatment landscape in haemophilia: from standard half-life clotting factor concentrates to gene editing. *Lancet*. 2021;397:630-640.
3. Franchini M, Mannucci PM. The more recent history of hemophilia treatment. *Semin Thromb Hemost*. 2022;48(8):904-910.
4. Croteau SE. Hemophilia A/B. *Hematol Oncol Clin North Am*. 2022;36(4):797-812.
5. Centers for Disease Control and Prevention. Data and statistics on hemophilia. Available at: <https://www.cdc.gov/hemophilia/data-research/>. Accessed on January 2, 2025.
6. National Bleeding Disorders Foundation. Hemophilia A: An overview of symptoms, genetics, and treatments to help you understand hemophilia A. Available at: <https://www.bleeding.org/bleeding-disorders-a-z/types/hemophilia-a>. Accessed on January 2, 2025.

- 7. National Hemophilia Foundation. Hemophilia B. An overview of symptoms, genetics, and treatments to help you understand hemophilia B. Available at: <https://www.hemophilia.org/bleeding-disorders-a-z/types/hemophilia-b>. Accessed on January 2, 2025.
- 8. Hemlibra® subcutaneous injection [prescribing information]. South San Francisco, CA and Tokyo, Japan: Genentech/Roche and Chugai; January 2024.
- 9. Hympavzi™ subcutaneous injection [prescribing information]. New York, NY: Pfizer; October 2024.

Revision Details

Type of Revision	Summary of Changes	Date
New	New policy	05/15/2025

The policy effective date is in force until updated or retired.

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