

Title: Hemophilia Agents

Policy #: Rx.01.149

Application of pharmacy policy is determined by benefits and contracts. Benefits may vary based on product line, group, or contract. Some medications may be subject to precertification, age, quantity, or formulary restrictions (ie limits on non-preferred drugs). Individual member benefits must be verified.

This pharmacy policy document describes the status of pharmaceutical information and/or technology at the time the document was developed. Since that time, new information relating to drug efficacy, interactions, contraindications, dosage, administration routes, safety, or FDA approval may have changed. This Pharmacy Policy will be regularly updated as scientific and medical literature becomes available. This information may include new FDA-approved indications, withdrawals, or other FDA alerts. This type of information is relevant not only when considering whether this policy should be updated, but also when applying it to current requests for coverage.

Members are advised to use participating pharmacies in order to receive the highest level of benefits.

Intent:

The intent of this policy is to communicate the medical necessity criteria for **Advate®**, **Adynovate®**, **Afstyla®**, **Alphanate®**, **Alphanine SD®**, **Alprolix®**, **Bebulin®**, **Benefix®**, **Coagadex®**, **Corifact®**, **Eloctate®**, **Esperoct®**, **Feiba NF®**, **Helixate FS®**, **Hemlibra®**, **Hemofil M®**, **Humate P®**, **Idelvion®**, **Ixinity®**, **Jivi®**, **Koate DVI®**, **Kogenate FS®**, **Kovaltry®**, **Monoclate P®**, **Mononine®**, **Novoeight®**, **Novoseven RT®**, **Nuwiq®**, **Obizur®**, **Profilnine®**, **Rebinyn®**, **Recombinate®**, **Riastap®**, **Rixubis®**, **Sevenfact®**, **Tretten®**, **Vonvendi®**, **Wilate®**, and **Xyntha®/Xyntha® Solofuse™** as provided under the member's prescription drug benefit.

Description:

Hemophilia is a bleeding disorder caused by clotting factor deficiencies. Hemophilia is a rare X-linked, congenitally acquired disease that leads to a deficiency in coagulation factor, VIII (hemophilia A) or IX (hemophilia B). Hemophilia A or B occurs in approximately 1 out of 10,000 births. Hemophilia A is more common than hemophilia B, representing approximately 80-85% of hemophilia cases. The disorder is further characterized by the percentage of serum clotting factor activity as compared to the normal range:

- **Severe hemophilia:** clotting factor levels are less than 1% of normal
- **Moderate hemophilia:** clotting factor levels are 1-5% of normal
- **Mild hemophilia:** clotting factors are 5-40% of normal

Treatment of hemophilia generally involves an infusion of the deficient factors to allow for more normalized clotting response and decreased incidence of uncontrolled bleeding events.

Treatment regimens are individualized based on disease severity and may include prophylactic infusions of clotting factors, on demand infusions of clotting factors, or a combination of both. Perioperative infusions are also part of the treatment regimen.

While Hemophilia A and hemophilia B are the most common types of hemophilia, 1 in every 500,000 to 2 million people have coagulation disorders caused by deficiencies in clotting factors other than factor VIII or IX. The products listed below include those with indications to treat these rare forms of coagulation disorders.

Riastap®	Fibrinogen concentrate, human					X			X		
Rixubis®	Factor IX, recombinant		X						X	X	X
Sevenfact®	Factor VIIa, recombinant	X	X								
Tretten®	Factor XIII A subunit, recombinant							X (A subunit)			X
Vonvendi®	Von Willebrand Factor Recombinant				X				X		
Wilate®	von Willebrand/ factor VIII complex, human				X				X		
Xyntha® [Solufuse]	Recombinant factor VIII	X							X	X	X

Policy:

Advate®, Adynovate®, Afstyla®, Alphanate®, Alphanine SD®, Alprolix®, Bebulin®, Benefix®, Coagadex®, Corifact®, Eloctate®, Esperoct®, Feiba NF®, Helixate FS®, Hemlibra®, Hemofil M®, Humate P®, Idelvion®, Ixinity®, Jivi®, Koate DVI®, Kogenate FS®, Kovaltry®, Monoclate P®, Mononine®, Novoeight®, Novoseven RT®, Nuwiq®, Obizur®, Profilnine®, Rebinyn®, Recombinate®, Riastap®, Rixubis®, Sevefact®, Tretten®, Vonvendi®, Wilate®, Xyntha®/Xyntha® Solofuse are approved when there is a diagnosis of an FDA approved indication.

Approval duration: 2 years

Black Box Warning as shown in the drug Prescribing Information:

Feiba® NF

Thrombotic/Thromboembolic events: Thrombotic and thromboembolic events have been reported during postmarketing surveillance following infusion of anti-inhibitor coagulant complex, particularly following the administration of high doses and/or in patients with thrombotic risk factors. Monitor patients receiving anti-inhibitor coagulant complex for signs and symptoms of thromboembolic events.

Hemlibra®

Cases of thrombotic microangiopathy and thrombotic events were reported when on average a cumulative amount of >100 U/kg/24 hours of activated prothrombin complex concentrate was administered for 24 hours or more to patients receiving HEMLIBRA prophylaxis. Monitor for the development of thrombotic microangiopathy and thrombotic events if aPCC is administered. Discontinue aPCC and suspend dosing of HEMLIBRA if symptoms occur.

Novoseven®, Sevenfact®

Serious arterial and venous thrombotic events following administration of Novoseven and Sevenfact RT have been reported. Discuss the risks and explain the signs and symptoms of thrombotic and thromboembolic events to patients who

will receive Novoseven and Sevenfact RT. Monitor patients for signs and symptoms of activation of the coagulation system and for thrombosis.

Guidelines:

Refer to the specific manufacturer's prescribing information for administration and dosage details and any applicable Black Box warnings.

BENEFIT APPLICATION

Subject to the terms and conditions of the applicable benefit contract, the applicable drug(s) identified in this policy is (are) covered under the prescription drug benefits of the Company's products when the medical necessity criteria listed in this pharmacy policy are met. Any services that are experimental/investigational or cosmetic are benefit contract exclusions for all products of the Company.

References:

Advate® (antihemophilic factor, human recombinant) [package insert]. Westlake Village, CA. Baxter Healthcare Corporation; December 2018. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=80fa03d2-cd4c-4155-9b57-1396c4fa42da&type=display#section-9>. Accessed July 22, 2021.

Adynovate® (antihemophilic factor, recombinant, PEGylated) [package insert]. Westlake Village, CA. Baxalta US Inc. May 2018. Available from: https://www.shirecontent.com/PI/PDFs/ADYNOVATE_USA_ENG.pdf. Accessed July 22, 2021.

Afstyla® (antihemophilic factor, recombinant, single chain) [package insert]. Kankakee, IL. CSL Behring, LLC. April 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=51f19873-a63f-4229-9477-5da4ecf31cde&type=display>. Accessed July 22, 2021.

Alphanate® (antihemophilic factor/ von willebrand factor complex, human) [package insert]. Los Angeles, CA. Grifols Biologicals Inc. June 2018. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=5a7aea94-654c-4113-9014-c3137de0a931&type=display>. Accessed July 22, 2021.

Alphanine SD® (coagulation factor IX, human) [package insert]. Los Angeles, CA. Grifols Biologicals Inc. June 2018. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=3e99052d-4442-4283-8915-c9a796c77008&type=display>. Accessed July 22, 2021.

Alprolix® (coagulation factor IX, recombinant, Fc fusion protein) [package insert]. Cambridge, MA. Biogen Idec Inc. October 2020. Available from: <http://products.sanofi.us/Alprolix/alprolix.html>. Accessed July 22, 2021.

Bebulin® (factors IX complex, human) [package insert]. Westlake Village, CA. Baxter Healthcare Corporation. September 2018. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=b375a2fb-7bd4-486c-87f6-72237713d528&type=display>. Accessed July 22, 2021.

BeneFIX® (coagulation factor IX, recombinant) [package insert]. Philadelphia, PA. Wyeth Pharmaceuticals Inc. June 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=85faa5bc-cee5-4ef1-8d80-bdcb7eba1e4&type=display>. Accessed July 22, 2021.

Bolton-Maggs PHB. Rare coagulation disorders. Available from: <http://www1.wfh.org/publication/files/pdf-1188.pdf>. Accessed July 22, 2021.

Coagadex® (coagulation factor X, human) [package insert]. Durham, NC. Bio Products Laboratory USA, Inc. September 2018. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=ad20e6ff-98bc-47e6-aa00-7fb441edd2b0&type=display>. Accessed July 22, 2021.

Corifact® (coagulation factor XIII, human) [package insert]. Kankakee, IL. CSL Behring LLC. December 2019. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=96f1b658-01b0-43a7-9aba-693271389a78&type=display>. Accessed July 22, 2021.

Eloctate® (antihemophilic factor, recombinant, Fc fusion protein) [package insert]. Cambridge, MA. December 2020. Biogen, Inc. Available from: <http://products.sanofi.us/Eloctate/Eloctate.pdf>. Accessed July 22, 2021.

Esperoct® (antihemophilic factor, recombinant, glycopegylated-exei) [prescribing information]. Bagsvaerd, Denmark: Novo Nordisk Inc.; October 2019. Available from: <https://www.novo-pi.com/esperoct.pdf>. Accessed July 22, 2021.

Feiba® (antiinhibitor coagulant complex) [package insert]. Westlake Village, CA. Baxter Healthcare Corporation. February 2020. Available from: http://www.baxalta.com/assets/documents/feiba_us_pi.pdf. Accessed July 22, 2021.

Feiba NF® (antiinhibitor coagulant complex) [package insert]. Westlake Village, CA. Baxter Healthcare Corporation. February 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=752604e5-4ea2-44f4-83ed-1569373f6412&type=display>. Accessed July 22, 2021.

Helixate FS® (antihemophilic factor recombinant) [package insert]. Whippany, NJ. Bayer HealthCare LLC. July 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=4e4bc43f-892a-4ca4-ad43-5383c276c61e&type=display>. Accessed July 22, 2021.

Hemlibra® (antihemophilic factor, humanized) [package insert]. South San Francisco, CA. Genentech, Inc. June 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/drugInfo.cfm?setid=2483adba-fab6-4d1b-96c5-c195577ed071>. Accessed July 22, 2021.

Hemofil M® (antihemophilic factor, human) [package insert]. Westlake Village, CA. Baxter Healthcare Corporation. June 2018. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=b8953ff7-3bba-4a0b-a486-f26fb81f05d9&type=display>. Accessed July 22, 2021.

Humate-P® (antihemophilic factor/ von Willebrand factor complex, human) Kankakee, IL. CSL Behring LLC. June 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=c5f4b63b-8d8b-45b3-b530-2f1c32a1b852&type=display>. Accessed July 22, 2021.

Idelvion® (coagulation factor IX, recombinant, albumin fusion protein) [package insert]. Kankakee, IL. CSL Behring. July 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=f74301bf-95e2-44ab-a8b6-98aa07b04683&type=display>. Accessed July 22, 2021..

Ixinity® (coagulation factor IX, recombinant) [package insert]. Baltimore, MD. Cangene Corporation. September 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=1cae46d1-f463-488a-9e8b-3431ef1de1e4&type=display>. Accessed July 22, 2021.

Jivi® [antihemophilic factor (recombinant), PEGylated-aucl] [prescribing information]. Whippany, NJ. Bayer Healthcare LLC. August 2018. Available from: https://labeling.bayerhealthcare.com/html/products/pi/Jivi_PI.pdf. Accessed July 22, 2021.

Koate-DVI® (antihemophilic factor, human) [package insert]. Research Triangle Park, NC. Grifols Therapeutics, Inc. June 2018. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=2f77d685-e382-b2e7-2414-0d3a3448688a&type=display>. Accessed July 22, 2021.

Kogenate FS® (antihemophilic factor, recombinant) [package insert] Whippany, NJ. Bayer HealthCare LLC. December 2019. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=01c6292d-b670-415b-9d8e-7df92c0adc0f&type=display>. Accessed July 22, 2021.

Kovaltry® (antihemophilic factor recombinant) [package insert]. Whippany, NJ. Bayer HealthCare LLC. January 2021. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=89ba5048-469c-4f08-a173-81b0c6f3004d&type=display>. Accessed July 22, 2021.

Monoclate-P® (antihemophilic factor, human) [package insert]. Kankakee, IL. CSL Behring LLC. September 2018. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=d9c17023-cbff-4d95-8f44-ef842d59d938&type=display>. Accessed July 22, 2021.

Mononine® (coagulation factor IX, human) [package insert]. Kankakee, IL. CSL Behring LLC. February 2019. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=16f35cdb-7a3b-481b-a25b-0a065aea2c9b&type=display>. Accessed July 22, 2021.

Novoeight® (antihemophilic factor, recombinant) [package insert]. Plainsboro, NJ. Novo Nordisk, Inc, July 2020. Available from: <https://www.novo-pi.com/novoeight.pdf>. Accessed July 22, 2021.

Novoseven RT® (coagulation factor VIIa, recombinant) [package insert]. Plainsboro, NJ. Novo Nordisk, Inc, July 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=ea65c32f-8518-4383-b621-17056c0eebc0&type=display>. Accessed July 22, 2021.

Nuwiq® (antihemophilic factor, recombinant) [package insert]. Hoboken, NJ. Octapharma USA, Inc. September 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=d6cb51f8-3acd-412c-93ec-41a8017dff3e&type=display>. Accessed July 22, 2021.

Obizur® (antihemophilic factor, recombinant, porcine sequence) [package insert]. Westlake Village, CA. Baxter Healthcare Corporation. July 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=a69ccbb3-7648-4978-8c6d-7ca6b95a9b01&type=display>. Accessed July 22, 2021.

Profilnine SD® (factor IX complex) [package insert]. Los Angeles, CA. Grisfols Biologicals Inc. June 2018. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=79821042-86a5-4a64-b621-76de0a0bbab2&type=display>. Accessed July 22, 2021.

Rebiny® (factor IX, recombinant) [package insert]. Plainsboro, NJ. Novo Nordisk Inc. June 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/drugInfo.cfm?setid=0ea37235-35fd-410d-b8c4-40ba15fe1294>. Accessed July 22, 2021.

Recombinate® (antihemophilic factor, recombinant) [package insert]. Westlake Village, CA. Baxter Healthcare Corporation. June 2019. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=2a569ec5-08d7-44ab-b649-384a496e173c&type=display>. Accessed July 22, 2021.

Riastap® (fibrinogen injection) [package insert]. Kankakee, IL. CSL Behring LLC. July 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=903dc8d0-39da-462c-9dac-004e0c7a26cc&type=display>. Accessed July 22, 2021.

Rixubis® (coagulation factor IX, recombinant) [package insert]. Westlake Village, CA. Baxter Healthcare Corporation. June 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=f5070a92-96b8-476a-a2dc-18b22d95e5e0&type=display>. Accessed July 22, 2021.

Sevenfact® (coagulation factor VIIa (recombinant)-jncw) [package insert]. Louisville, KY. HEMA Biologics. April 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/drugInfo.cfm?setid=b23ad666-8ce0-4e0b-9eff-c6983ac5f7e6>. Accessed July 22, 2021.

Srivastava A, et al. Guidelines for the management of hemophilia. Available from: <https://www1.wfh.org/publication/files/pdf-1472.pdf>. Accessed July 22, 2021.

Tretten® (coagulation factor XIII a-subunit, recombinant) [package insert]. Plainsboro, NJ. Novo Nordisk, Inc, June 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=8664978e-1814-4930-aca5-97a24455f6df&type=display>. Accessed July 22, 2021.

Vonvendi® (von Willebrand factor, recombinant) [package insert]. Westlake Village, CA. Baxalta US, Inc. February 2019. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=53b6d198-6175-4a26-8606-dc0d2a0f12d6&type=display>. Accessed July 22, 2021.

Wilate® (Factor VII/VWF) [package insert]. Hoboken, NJ. Octapharma USA, Inc. September 2019. Available from: https://www.wilateusa.com/wp-content/uploads/2019/10/20190923_pil_18x_USA_26.pdf. Accessed July 22, 2021.

Xyntha® (antihemophilic factor, recombinant) [package insert]. Philadelphia, PA. Wyeth Pharmaceuticals Inc. August 2020. Available from: <https://dailymed.nlm.nih.gov/dailymed/fda/fdaDrugXsl.cfm?setid=52e97c7f-8240-4cd6-afd2-c452ab2f6e32&type=display>. Accessed July 22, 2021.

Applicable Drugs:

Inclusion of a drug in this table does not imply coverage. Eligibility, benefits, limitations, exclusions, precertification/referral requirements, provider contracts, and Company policies apply.

Drug Names

Advate®

Adynovate®

Afstyla®

Alphanate®

Alphanine SD®

Alprolix®

Bebulin®

BeneFIX®

Coagadex®

Corifact®

Eloctate®

Esperoct®

Feiba NF®

Helixate FS®

Hemlibra®

Hemofil M®

Humate-P®

Idelvion®

Ixinity®

Jivi®

Koate-DVI®

Kogenate FS®

Kovaltry®

Monoclate-P®

Mononine®

Novoeight®

Novoseven RT®

Nuwiq®

Obizur®

Profilnine SD®

Rebinyn®

Recombinate®

Riastap®

Rixubis®

Sevenfact®

Tretten®

Vonvendi®

Wilate®

Xyntha®

Cross References:

Rx.01.33 Off-Label Use

Policy Version Number:	20.00
P&T Approval Date:	June 10, 2021
Policy Effective Date:	October 01, 2021
Next Required Review Date:	June 10, 2022

The Policy Bulletins on this web site were developed to assist the Company in administering the provisions of the respective benefit programs, and do not constitute a contract. If you have coverage through the Company, please refer to your specific benefit program for the terms, conditions, limitations and exclusions of your coverage. Company does not provide health care services, medical advice or treatment, or guarantee the outcome or results of any medical services/treatments. The facility and professional providers are responsible for providing medical advice and treatment. Facility and professional providers are independent contractors and are not employees or agents of the Company. If you have a specific medical condition, please consult with your doctor. The Company reserves the right at any time to change or update its Policy Bulletins.