YouthCare Health*Choice* Illinois

Clinical Policy: Factor VIII (Human, Recombinant)

Reference Number: IL.PHAR.215 Effective Date: 05.01.16 Last Review Date: 7.2.21 Line of Business: Medicaid

Coding Implications Revision Log

See <u>Important Reminder</u> at the end of this policy for important regulatory and legal information.

Description

The following are factor VIII products requiring prior authorization: human – Hemofil M[®], Koate-DVI[®]; recombinant – Advate[®], Adynovate[®], Afstyla[®], Eloctate[®], Esperoct[®], Helixate FS[®], Jivi[®], Kogenate FS[®], Kogenate FS[®], togenate FS[®], Kogenate FS[®], Kogenate FS[®], NovoEight[®], Nuwiq[®], Obizur[®], Recombinate[®], Xyntha[®], and Xyntha[®] SolofuseTM.

FDA Approved Indication(s)

Factor VIII products are indicated for patients with hemophilia A for the following uses:

- Control and prevention of bleeding episodes:
 - Children and adults: Advate, Adynovate, Afstyla, Eloctate, Esperoct, Helixate FS, Hemofil M, Jivi (in previously treated patients ≥ 12 years of age only), Koate-DVI, Kogenate FS, Kovaltry, Novoeight, Nuwiq, Recombinate, Xyntha
- Perioperative management:
 - Children and adults: Advate, Adynovate, Afstyla, Eloctate, Esperoct, Helixate FS, Hemofil M, Jivi (in previously treated patients ≥ 12 years of age only), Koate-DVI, Kogenate FS, Kovaltry, Novoeight, Nuwiq, Recombinate, Xyntha
- Routine prophylaxis to prevent or reduce the frequency of bleeding episodes:
 - Children and adults: Advate, Adynovate, Afstyla, Eloctate, Esperoct, Helixate FS, Jivi (in previously treated patients ≥ 12 years of age only), Kovaltry, Novoeight, Nuwiq, Xyntha
- Routine prophylaxis to prevent or reduce the frequency of bleeding episodes and to reduce the risk of joint damage in children without pre-existing joint damage:
 Children: Helixate FS, Kogenate FS
- On-demand treatment and control of bleeding episodes in acquired hemophilia A:
 Adults: Obizur

Limitation(s) of use:

- Factor VIII products are not indicated for treatment of von Willebrand disease.
- Obizur is not indicated for the treatment of congenital hemophilia A.
- Safety and efficacy of Obizur have not been established in patients with a baseline anti- porcine factor VIII inhibitor titer of > 20 Bethesda units (BU).
- Jivi is not indicated for use in children < 12 years of age due to a greaterrisk for hypersensitivity reactions.
- Jivi is not indicated for use in previously untreated patients.

YouthCare Health Choice Illinois

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.

It is the policy of health plans affiliated with Centene Corporation[®] that factor VIII products are **medically necessary** when the following criteria are met:

I. Initial Approval Criteria

A. Hemophilia A (must meet all):

- 1. Diagnosis of one of the following (a or b):
 - a. Congenital hemophilia A (factor VIII deficiency) (all products except Obizur);
 - b. Acquired hemophilia A (Obizur only);
- 2. Prescribed by or in consultation with a hematologist;
- 3. Request is for one of the following uses (a, b, or c):
 - a. Control and prevention of bleeding episodes;
 - b. Perioperative management (all products except Obizur);
 - c. Routine prophylaxis to prevent or reduce the frequency of bleeding episodes;
- 4. For all products except Obizur: If factor VIII coagulant activity levels are > 5%, failure of desmopressin acetate, unless contraindicated, clinically significant adverse effects are experienced, or an appropriate formulation of desmopressin acetate is unavailable;
- 5. For Jivi: Member meets both of the following (a and b):
 - a. Age \geq 12 years;
 - b. Has previously been treated for hemophilia A;
- 6. Documentation of member's body weight (in kg);
- 7. Dose does not exceed the FDA-approved maximum recommended dose for the relevant indication.

Approval duration: 3 months (surgical/acute bleeding) or 6 months (prophylaxis)

B. Other diagnoses/indications

1. Refer to the off-label use policy for the relevant line of business if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized): CP.PMN.53 for Medicaid

II. Continued Therapy

- A. Hemophilia A (must meet all):
 - 1. Currently receiving medication via Centene benefit or member has previously met initial approval criteria;
 - 2. Member is responding positively to therapy;
 - 3. Documentation of member's body weight (in kg);
 - 4. If request is for a dose increase, new dose does not exceed the FDA-approved maximum recommended dose for the relevant indication.

Approval duration: 3 months (surgical/acute bleeding) or 6 months (prophylaxis)

YouthCare Health*Choice* Illinois

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

1. Currently receiving medication via Centene benefit and documentation supports positive response to therapy.

Approval duration: Duration of request or 6 months (whichever is less); or

2. Refer to the off-label use policy for the relevant line of business if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized): 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 policy CP.PMN.53 for Medicaid or evidence of coverage documents;
- **B.** Von Willebrand disease.

IV. Appendices/General Information

Appendix A: Abbreviation/AcronymKey BU: Bethesda units FDA: Food and Drug Administration

Appendix B: Therapeutic Alternatives

This table provides a listing of preferred alternative therapy recommended in the approval criteria. The drugs listed here may not be a formulary agent and may require prior authorization

Drug Name	Dosing Regimen	Dose Limit/ Maximum Dose
desmopressin acetate (Stimate [®] nasal spray;	When Factor VIII coagulant activity levels are > 5%	Injection: 0.3 mcg/kg IV every 48 hours
generic injection solution)	Injection: 0.3 mcg/kg IV every 48 hours	Nasal spray: 1 spray intranasally in each
	Nasal spray: < 50 kg: 1 spray intranasally in one nostril only; may repeat based on	nostril
	laboratory response and clinical condition ≥ 50 kg: 1 spray intranasally in each	
	nostril; may repeat based on laboratory response and clinical condition	

Therapeutic alternatives are listed as Brand name[®] (generic) when the drug is available by brand name only and generic (Brand name[®]) when the drug is available by both brand and generic.

Appendix C: Contraindications/Boxed Warnings

- Contraindication(s): life-threatening hypersensitivity reactions, including anaphylaxis, to the product and its constituents* *Including bovine, mouse, or hamster protein for Advate, Adynovate, Afstyla, Esperoct, Helixate FS, Hemofil M, Jivi, Kogenate FS, Kovaltry, Novoeight, Obizur, Recombinate, and Xyntha
- Boxed warning(s): none reported

YouthCare Health Choice All linois

Appendix D: General Information

- Life-threatening bleeding episodes include, but are not limited to, bleeds in the following sites: intracranial, neck/throat, or gastrointestinal.
- Serious bleeding episodes include bleeds in the following site: joints (hemarthrosis).
- Spontaneous bleed is defined as a bleeding episode that occurs without apparent cause and is not the result of trauma.

Drug Name Indication **Dosing Regimen Maximum Dose** Antihemophilic factor Control and Minor episodes: 10-20 50 IU/kg every 6 - recombinant (Advate, prevention of IU/kg IV every 12-24 hours until the Adynovate, Afstyla, bleeding hours bleeding episode is episodes resolved Kovaltry, Novoeight, (Advate: 8-24 hours for Nuwiq, Recombinate, age < 6 years) ReFacto, Xyntha) Moderate episodes: 15-30 IU/kg IV every 12-24 hours (Advate: 8-24 hours for age < 6 years) Major episodes: 30-50 IU/kg IV every 8-24 hours (Advate: 6-12 hours for age < 6 years) Antihemophilic factor Control and Minor and moderate 50 IU/kg every 8 - recombinant prevention of episodes: 20-30 hours until the bleeding IU/kg every 24-48 bleeding episode is (Eloctate) episodes hours (12-24 hours for resolved age < 6 years) Major episodes: 40-50 IU/kg every 12-24 hours (8 to 24 hours for age < 6 years) Antihemophilic factor Control and Minor episodes: 10-20 50 IU/kg single IU/kg IV; repeat dose if dose or 30 – recombinant prevention of (Helixate FS, Kogenate bleeding there is evidence of IU/kg/repeated FS) episodes further bleeding dose Moderate episodes: 15-30 IU/kg IV every 12-24 hours Major episodes: initial 40-50 IU/kg IV, followed by 20-25 IU/kg every 8-24 hours

I. Dosage and Administration



			N V
Drug Name	Indication	Dosing Regimen	Maximum Dose
		(Kogenate FS: every 8- 12 hours)	
Antihemophilic factor – recombinant, glycopegylated (Esperoct)	Control and prevention of bleeding episodes	Minor to moderate episodes: 40-65 IU/kg IV; one dose should be sufficient for minor episodes; additional dose may be administered after 24 hours for moderate episodes.	At least 12 years old: 40 IU/kg < 12 years old: 65 IU/kg
		Major episodes: 50-65 IU/kg IV; additional doses may b eadministered approximately every 24 hours.	
Antihemophilic factor – recombinant (Advate, Adynovate)	Perioperative management	Minor surgery: 30- 50 IU/kg IV as a single dose within 1 hour of the operation and every 12- 24 hours (Adynovate: 24 hours) thereafter as needed to control bleeding	Minor surgery: 50 IU/kg/dose Major surgery: 60 IU/kg/dose
		Major surgery: 40- 60 IU/kg IV as a single dose preoperatively to achieve 100% activity and every 8- 24 hours thereafter to keep factor VIII activity in desired range (Advate: every 6- 24 hours for age < 6 years; Adynovate: every 6-24 hours if age < 12 years)	
Antihemophilic factor – recombinant (Eloctate)	Perioperative management	Minor surgery: 25- 40 IU/kg every 24 hours (12-24 hours age < 6 years) Major surgery: pre- operative dose of 40-60 IU/kg followed by a repeat dose of 40-50 IU/kg after 8-24 hours (6-	Minor surgery: 40 IU/kg/dose Major surgery: 60 IU/kg/dose



			V
Drug Name	Indication	Dosing Regimen	Maximum Dose
		24 hours for age < 6 years) and then every 24 hours to maintain Factor VIII activity within the target range	
Antihemophilic factor – recombinant, glycopegylated (Esperoct)	Perioperative management	Minor and major surgery: 50-65 IU/kg IV; additional doses can be administered after 24 hours if necessary for minor surgeries; additional doses can be administered approximately every 24 hours for the first week and then approximately every 48 hours until wound healing has occurred for major surgeries	At least 12 years old: 50 IU/kg < 12 years old: 65 IU/kg
Antihemophilic factor – recombinant (Helixate FS, Kogenate FS)	Perioperative management	Minor surgery: 15- 30 IU/kg IV every 12-24 hours Major surgery: pre- operative dose of 50 IU/kg IV followed by a repeat dose every 6- 12 hours to maintain Factor VIII activity within the target range	Minor surgery: 30 IU/kg/dose Major surgery: 50 IU/kg/dose
Antihemophilic factor – recombinant (Afstyla, Kovaltry, Novoeight, Nuwiq, Recombinate, Xyntha)	Perioperative management	Minor surgery: 15-30 IU/kg IV every 24 hours (Xyntha: every 12- 24 hours) (Recombinate: 30- 40 IU/kg as a single infusion) Major surgery: 40- 50 IU/kg IV every 8-24 hours (Xyntha: 30-50 IU/kg)	Minor surgery: 30 IU/kg/dose (Recombinate: 40 IU/kg/dose) Major surgery: 50 IU/kg every 8 hours
Antihemophilic factor – recombinant (Xyntha)	Routine prophylaxis	30 IU/kg IV 3 times weekly < 12 years of age: 25 IU/kg every other day.	30 IU/kg/dose



× -			
Drug Name	Indication	Dosing Regimen	Maximum Dose
Antihemophilic factor – recombinant (Advate)	Routine prophylaxis	20-40 IU/kg IV every other day (3 to 4 times weekly)	40 IU/kg every other day
		OR	
		Use every third day dosing regimen targeted to maintain Factor VIII	
		trough levels $\geq 1\%$	
Antihemophilic factor – recombinant (Adynovate)	Routine prophylaxis	≥ 12 years of age: 40-50 IU/kg IV 2 times per week	70 IU/kg/dose
		< 12 years of age: 55 IU/kg IV 2 times per week	
Antihemophilic factor – recombinant (Afstyla)	Routine prophylaxis	 ≥ 12 years of age: 20-50 IU/kg IV 2-3 times per week < 12 years of age: 30-50 IU/kg IV 2-3 times per week 	50 IU/kg/dose
Antihemophilic factor – recombinant (Eloctate)	Routine prophylaxis	times per week 50 IU/kg IV every 4 days	65 IU/kg/dose
		For children < 6 years of age: 50 IU/kg IV twice weekly	
Antihemophilic factor – recombinant, glycopegylated	Routine prophylaxis	At least 12 years old: 50 IU/kg IV every 4 days	At least 12 years old: 50 IU/kg
(Esperoct)		< 12 years old: 65 IU/kg IV twice weekly	< 12 years old: 65 IU/kg
Antihemophilic factor – recombinant (Helixate FS, Kogenate	Routine prophylaxis	Adults: 25 IU/kg IV three times per week	25 IU/kg/dose
FS)		Children: 25 IU/kg every	
Antihemophilic factor – recombinant (Novoeight)	Routine prophylaxis	other day ≥ 12 years of age: 20-50 IU/kg IV 3 times per week OR 20- 40 IU/kg IV every other day	60 IU/kg/dose
		< 12 years of age: 25- 60 IU/kg IV 3	



Drug Name	Indication	Dosing Regimen	Maximum Dose
		times per week OR 25-	
		50 IU every other day	
Antihemophilic factor	Routine	\geq 12 years of age:	50 IU/kg/dose
- recombinant (Nuwiq)	prophylaxis	30-40 IU/kg IV	
	1 1 2	every other day	
		< 12 years of age: 30-	
		50 IU/kg IV	
		every other day or 3	
		times/week	
Antihemophilic factor	Routine	> 12 years of age: 20-40	50 IU/kg every
- recombinant	prophylaxis	IU/kg IV 2-3	other day
(Kovaltry)	propriyrams	times per week	other duy
(Rovani y)		unies per week	
		\leq 12 years of age:	
		25-50 IU/kg twice or	
		three times weekly or	
		every other day	
		according to individual	
		requirements	
Antihamanhilia factor	Treatment of	200 IU/kg every 4-	200 IU every 4
Antihemophilic factor		12 hours	hours
- recombinant, porcine	bleeding	12 nours	nours
sequence (Obizur)	episodes in		
	acquired		
A (1 1 1) C (hemophilia A		100 111/1 0
Antihemophilic factor	Control and	Minor episodes: 10- 20	100 IU/kg every 8
- human (Hemofil M)	prevention of	IU/kg IV every 12-24	hours
	bleeding	hours	
	episodes		
		Moderate episodes: 15-	
		30 IU/kg IV	
		every 12-24 hours	
		Major episodes: 30- 50	
		IU/kg IV every 8-24	
		hours	
Antihemophilic factor	Control and	Minor episodes: 10	25 IU/kg every 8
- human (Koate-DVI)	prevention of	IU/kg IV as a single	hours until the
	bleeding	dose; repeat only if there	bleeding episode i
	episodes	is evidence of further	resolved
		bleeding	
	1		
		Moderate episodes: 15-	
		Moderate episodes: 15- 25 IU/kg IV as a single	
		25 IU/kg IV as a single	
		25 IU/kg IV as a single dose followed by 10-15	
		25 IU/kg IV as a single	



			V.
Drug Name	Indication	Dosing Regimen	Maximum Dose
		Major episodes: 40- 50 IU/kg IV as a single dose followed by 20-25 IU/kg IV every 8-12 hours	
Antihemophilic factor – human (Hemofil M)	Perioperative management	Minor surgery: 30- 40 IU/kg as a single infusion Major surgery: 40- 50 IU/kg every 8-	Minor surgery: 80 IU/kg/dose Major surgery: 100 IU/kg every 8 hours
Antihemophilic factor – human (Koate-DVI)	Perioperative management	24 hours Major surgery: 50 IU/kg pre-operative dose followed by 50 IU/kg every 6-12 hours as needed Minor surgery: less intensive schedules may be adequate	Major surgery: 50 IU/kg every 6 hours
Antihemophilic factor – recombinant, PEGylated-aucl (Jivi)	Control and prevention of bleeding episodes	Minor episodes: 10- 20 IU/kg every 24- 48 hours Moderate episodes: 15- 30 IU/kg every 24-48 hours Major episodes: 30- 50 IU/kg every 8-24 hours	50 IU/kg every 8 hours
	Perioperative management	Minor surgery: 15- 30 IU/kg every 24 hours Major surgery: 40- 50 IU/kg every 12- 24 hours	Minor surgery: 30 IU/kg/dose Major surgery: 50 IU/kg/dose
	Routine prophylaxis	30-40 IU/kg twice weekly; may be adjusted to 45-60 IU/kg every 5 days with further individual adjustment to less or more frequent dosing	60 IU/kg/dose; frequency varies based on bleeding episodes

I. Product Availability



Drug NameAvailabilityAntihemophilic factor - recombinant (Advate)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000, 4,000 IUAntihemophilic factor - recombinant (Adynovate)Vial: 250, 500, 750, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor - recombinant (Afstyla)Vial: 250, 500, 750, 1,000, 1,500, 2,000, 3,000 4,000, recombinant (Eloctate)Antihemophilic factor - recombinant (Eloctate)Vial: 250, 500, 750, 1,000, 1,500, 2,000, 3,000 4,000, recombinant (glycopegylated- exei (Esperot)Antihemophilic factor - recombinant (Helixate FS, Kogenate FS, Kovaltry)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor - recombinant (Noveight)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor - recombinant (Noveight)Vial: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor - recombinant (Nawiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000 IUAntihemophilic factor - recombinant (Nawiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000 IUAntihemophilic factor - recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor - recombinant (Xyntha Solofuse)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor - recombinant (Refacto, Xyntha)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor - recombinant (Koate - V)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor - recombinant (Koate - V)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor - recombinant (Koate - V)Vial: 250, 500, 1,000 IUAntihemophilic factor - recombinant (Koate - V)Vial: 250, 500, 1,000 I		N. N
recombinant (Advate)Vial: 250, 500, 750, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor – recombinant (Adynovate)Vial: 250, 500, 750, 1,000, 1,500, 2,000, 2,500, 3,000 IUAntihemophilic factor – recombinant (Eloctate)Vial: 250, 500, 750, 1,000, 1,500, 2,000, 3,000 4,000, 5,000, 6,000 IUAntihemophilic factor – recombinant (Eloctate)Vial: 250, 500, 750, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor – recombinant, glycopegylated- exei (Esperoct)Vial: 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor – recombinant (Helixate FS, Kogenate FS, Kovaltry)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor – recombinant (Novoeight)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor – recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Recombinat)Vial: 250, 500, 1,000, 2,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor – recombinant (Xyntha Solofuse)Prefiled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Cobizur)Vial: 500 IUAntihemophilic factor – recombinant (Cobizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – recombinant (Cobizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – recombinant (Cobizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – recombinant (Kater-DVI)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – recombinant, PEGylated-Vial: 500, 1,000, 2,	Drug Name	Availability
Antihemophilic factor – recombinant (Adynovate)Vial: 250, 500, 750, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor – recombinant (Afstyla)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 4,000, recombinant (Eloctate)Antihemophilic factor – recombinant, glycopegylated- exei (Esperoct)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 4,000, recombinant (Helixate FS, Kogenate FS, Kovaltry)Antihemophilic factor – recombinant (Melixate FS, Kogenate FS, Kovaltry)Vial: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor – recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor – recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Xyntha Solofuse)Vial: 250, 500, 1,000, 2,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Cbizur)Vial: 500 IUAntihemophilic factor – recombinant (Cbizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – recombinant (Cbizur)Vial: 250, 500, 1,000 IUAntihemophilic factor – recombinant (Cbizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – recombinant (Cbizur)Vial: 250, 500, 1,000 IUAntihemophilic factor – recombinant (PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU	Antihemophilic factor –	Vial: 250, 500, 1,000, 1,500, 2,000, 3,000, 4,000 IU
recombinant (Adynovate)Antihemophilic factor – recombinant (Afstyla)Vial: 250, 500, 1,000, 1,500, 2,000, 2,500, 3,000 IUAntihemophilic factor – recombinant (Eloctate)Vial: 250, 500, 750, 1,000, 1,500, 2,000, 3,000 4,000, recombinant (Eloctate)Antihemophilic factor – recombinant (glycopegylated- exei (Esperoct)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor – recombinant (Helixate FS, Kogenate FS, Kovaltry)Vial: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Novoeight)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor – recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor – recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor – recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 1,240, 1241-1800, 1801-2400 IUAntihemophilic factor – recombinant (Kyntha)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Qbizur)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – recombinant (Cbizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – necombinant (Kate-DVI)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – recombinant (Kate-DVI)Vial: 500, 1,000, 2,000, 3,000 IU	recombinant (Advate)	
Antihemophilic factor - recombinant (Afstyla)Vial: 250, 500, 1,000, 1,500, 2,000, 2,500, 3,000 IUAntihemophilic factor - recombinant (Eloctate)Vial: 250, 500, 750, 1,000, 1,500, 2,000, 3,000 4,000, 5,000, 6,000 IUAntihemophilic factor - recombinant, glycopegylated- exei (Esperoct)Vial: 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor - recombinant (Helixate FS, Kogenate FS, Kovaltry)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor - recombinant (Novoeight)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor - recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor - recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor - recombinant (Recombinate)Vial: 220-400, 401-800, 801-1240, 1241-1800, 1801-2400 IUAntihemophilic factor - recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor - recombinant (Xyntha Solofuse)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor - recombinant (Obizur)Vial: 500 IUAntihemophilic factor - recombinant (Obizur)Vial: 500 IUAntihemophilic factor - recombinant (Obizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor - recombinant (Obizur)Vial: 250, 500, 1,000 IUAntihemophilic factor - recombinant (Obizur)Vial: 250, 500, 1,000 IUAntihemophilic factor - recombinant (Koate-DVI)Vial: 500, 1,000, 2,000, 3,000 IUAntihemophilic factor - recombinant, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU <td>Antihemophilic factor –</td> <td>Vial: 250, 500, 750, 1,000, 1,500, 2,000, 3,000 IU</td>	Antihemophilic factor –	Vial: 250, 500, 750, 1,000, 1,500, 2,000, 3,000 IU
recombinant (Afstyla)Antihemophilic factor - recombinant (Eloctate)Vial: 250, 500, 750, 1,000, 1,500, 2,000, 3,000 4,000, 5,000, 6,000 IUAntihemophilic factor - recombinant, glycopegylated- exei (Esperoct)Vial: 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor - recombinant (Helixate FS, Kogenate FS, Kovaltry)Vial: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor - recombinant (Novoeight)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor - recombinant (Novoeight)Vial: 250, 500, 1,000, 2,000, 2,000, 3,000 IUAntihemophilic factor - recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor - recombinant (Refacto, Xyntha)Vial: 250, 500, 1,000, 2,000, 2,000, 3,000 IUAntihemophilic factor - recombinant (Xyntha Solofuse)Vial: 250, 500, 1,000, 2,000, 2,000, 3,000 IUAntihemophilic factor - recombinant (Obizur)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor - recombinant (Obizur)Vial: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor - recombinant (Obizur)Vial: 500 IUAntihemophilic factor - recombinant (Obizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor - recombinant (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor - recombinant, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU	recombinant (Adynovate)	
Antihemophilic factor – recombinant (Eloctate)Vial: 250, 500, 750, 1,000, 1,500, 2,000, 3,000 4,000, 5,000, 6,000 IUAntihemophilic factor – recombinant, glycopegylated- exei (Esperoct)Vial: 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor – recombinant (Helixate FS, Kogenate FS, Kovaltry)Vial: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Novoeight)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor – recombinant (Novoeight)Vial: 250, 500, 1,000, 2,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor – recombinant (Recombinate)Vial: 220-400, 401-800, 801-1240, 1241-1800, 1801-2400 IUAntihemophilic factor – recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor – recombinant (Xyntha Solofuse)Vial: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Cotizur)Vial: 500 IUAntihemophilic factor – recombinant (Cotizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – recombinant (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor – recombinant, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU	Antihemophilic factor –	Vial: 250, 500, 1,000, 1,500, 2,000, 2,500, 3,000 IU
recombinant (Eloctate)5,000, 6,000 IUAntihemophilic factor - recombinant, glycopegylated- exei (Esperoct)Vial: 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor - recombinant (Helixate FS, Kogenate FS, Kovaltry)Vial: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor - recombinant (Novoeight)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor - recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor - recombinant (Nuwiq)Vial: 220-400, 401-800, 801-1240, 1241-1800, 1801-2400 IUAntihemophilic factor - recombinant (Recombinate)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor - recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor - recombinant (Quruta)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor - recombinant (Obizur)Vial: 500 IUAntihemophilic factor - recombinant (Obizur)Vial: 500, 1,000, 1,700 IUAntihemophilic factor - recombinant (Cbizur)Vial: 250, 500, 1,000 IUAntihemophilic factor - recombinant (Cbizur)Vial: 250, 500, 1,000 IUAntihemophilic factor - recombinant (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor - recombinant (Koate-DVI)Vial: 500, 1,000, 2,000, 3,000 IU	recombinant (Afstyla)	
Antihemophilic factor – recombinant, glycopegylated- exei (Esperoct)Vial: 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor – recombinant (Helixate FS, Kogenate FS, Kovaltry)Vial: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Novoeight)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor – recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor – recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor – recombinant (Recombinate)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor – recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor – recombinant (Xyntha Solofuse)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – recombinant (Obizur)Vial: 500, 1,000, 1,700 IUAntihemophilic factor – recombinant (Koate-DVI)Vial: 250, 500, 1,000 IU	Antihemophilic factor –	Vial: 250, 500, 750, 1,000, 1,500, 2,000, 3,000 4,000,
recombinant, glycopegylated- exei (Esperoct) Vial: 250, 500, 1,000, 2,000, 3,000 IU recombinant (Helixate FS, Kogenate FS, Kovaltry) Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IU recombinant (Novoeight) Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IU recombinant (Nuwiq) Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IU recombinant (Nuwiq) Vial: 220-400, 401-800, 801-1240, 1241-1800, 1801-2400 IU Recombinant (Recombinate) IU Antihemophilic factor – recombinant (ReFacto, Xyntha) Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IU recombinant (Xyntha Solofuse) Vial: 500 IU Antihemophilic factor – recombinant (Dbizur) Vial: 250, 500, 1,000, 1,700 IU Antihemophilic factor – recombinant (Dbizur) Vial: 250, 500, 1,000, 1,700 IU Antihemophilic factor – recombinant (Dbizur) Vial: 250, 500, 1,000, 1,700 IU Antihemophilic factor – Nantihemophilic factor – Vial: 250, 500, 1,000, 1,700 IU Antihemophilic factor – Nial: 250, 500, 1,000 IU Vial: 250, 500, 1,000 IU Vial: 250, 500, 1,000 IU Vial: 250, 500, 1,000 IU Vial: 250, 500, 1,000 IU	recombinant (Eloctate)	5,000, 6,000 IU
recombinant, glycopegylated- exei (Esperoct) Vial: 250, 500, 1,000, 2,000, 3,000 IU recombinant (Helixate FS, Kogenate FS, Kovaltry) Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IU recombinant (Novoeight) Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IU recombinant (Nuwiq) Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IU recombinant (Nuwiq) Vial: 220-400, 401-800, 801-1240, 1241-1800, 1801-2400 IU Recombinant (Recombinate) IU Antihemophilic factor – recombinant (ReFacto, Xyntha) Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IU recombinant (Xyntha Solofuse) Vial: 500 IU Antihemophilic factor – recombinant (Dbizur) Vial: 250, 500, 1,000, 1,700 IU Antihemophilic factor – recombinant (Dbizur) Vial: 250, 500, 1,000, 1,700 IU Antihemophilic factor – recombinant (Dbizur) Vial: 250, 500, 1,000, 1,700 IU Antihemophilic factor – Nantihemophilic factor – Vial: 250, 500, 1,000, 1,700 IU Antihemophilic factor – Nial: 250, 500, 1,000 IU Vial: 250, 500, 1,000 IU Vial: 250, 500, 1,000 IU Vial: 250, 500, 1,000 IU Vial: 250, 500, 1,000 IU	Antihemophilic factor –	Vial: 500, 1,000, 1,500, 2,000, 3,000 IU
Antihemophilic factor – recombinant (Helixate FS, Kogenate FS, Kovaltry)Vial: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Novoeight)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor – recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor – recombinant (Recombinate)Vial: 220-400, 401-800, 801-1240, 1241-1800, 1801-2400 IUAntihemophilic factor – recombinant (Recombinate)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor – recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor – recombinant (Xyntha Solofuse)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – recombinant (Obizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – recombinant (Obizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – recombinant (Koate-DVI)Vial: 500, 1,000, 2,000, 3,000 IU	-	
Antihemophilic factor – recombinant (Helixate FS, Kogenate FS, Kovaltry)Vial: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Novoeight)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor – recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor – recombinant (Recombinate)Vial: 220-400, 401-800, 801-1240, 1241-1800, 1801-2400 IUAntihemophilic factor – recombinant (Recombinate)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor – recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor – recombinant (Xyntha Solofuse)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – recombinant (Obizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – recombinant (Obizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – recombinant (Koate-DVI)Vial: 500, 1,000, 2,000, 3,000 IU		
recombinant (Helixate FS, Kogenate FS, Kovaltry) Antihemophilic factor – recombinant (Novoeight) Antihemophilic factor – recombinant (Nuwiq) Antihemophilic factor – recombinant (Nuwiq) Antihemophilic factor – recombinant (Refacto, Xyntha) Antihemophilic factor – recombinant (ReFacto, Xyntha) Antihemophilic factor – recombinant (Xyntha Solofuse) Antihemophilic factor – recombinant (Obizur) Antihemophilic factor – recombinant (Obizur) Antihemophilic factor – recombinant (Obizur) Antihemophilic factor – tecombinant (Obizur) Antihemophilic factor – Nathemophilic factor – tecombinant (Obizur) Antihemophilic factor – Nathemophilic factor – Nial: 250, 500, 1,000, 2,000, 3,000 IU Vial: 250, 500, 1,000, 1,700 IU human (Hemofil M) Antihemophilic factor – Nial: 250, 500, 1,000, 1,700 IU human (Koate-DVI) Antihemophilic factor – Vial: 500, 1,000, 2,000, 3,000 IU		Vial: 250, 500, 1,000, 2,000, 3,000 IU
Kogenate FS, Kovaltry)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor – recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor – recombinant (Recombinate)Vial: 220-400, 401-800, 801-1240, 1241-1800, 1801-2400 IUAntihemophilic factor – recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor – recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor – recombinant (Xyntha Solofuse)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – recombinant (Koate-DVI)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – recombinant (Koate-DVI)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – recombinant (Koate-DVI)Vial: 500, 1,000, 2,000, 3,000 IU	1	
Antihemophilic factor – recombinant (Novoeight)Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IUAntihemophilic factor – recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor – recombinant (Recombinate)Vial: 220-400, 401-800, 801-1240, 1241-1800, 1801-2400 IUAntihemophilic factor – recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor – recombinant (Xyntha Solofuse)Vial: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – human (Hemofil M)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000, 2,000, 3,000 IU		
recombinant (Novoeight)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor – recombinant (Recombinate)Vial: 220-400, 401-800, 801-1240, 1241-1800, 1801-2400 IUAntihemophilic factor – recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor – recombinant (Xyntha Solofuse)Vial: 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – recombinant (Koate-DVI)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – recombinant, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU		Vial: 250, 500, 1,000, 1,500, 2,000, 3,000 IU
Antihemophilic factor – recombinant (Nuwiq)Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IUAntihemophilic factor – recombinant (Recombinate)Vial: 220-400, 401-800, 801-1240, 1241-1800, 1801-2400 IUAntihemophilic factor – recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor – recombinant (Xyntha Solofuse)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – recombinant (Obizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – recombinant (Koate-DVI)Vial: 250, 500, 1,000 IU	recombinant (Novoeight)	
recombinant (Nuwiq)Vial: 220-400, 401-800, 801-1240, 1241-1800, 1801-2400 IUAntihemophilic factor – recombinant (Recombinate)Vial: 220-400, 401-800, 801-1240, 1241-1800, 1801-2400 IUAntihemophilic factor – recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor – recombinant (Xyntha Solofuse)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – recombinant (Obizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – human (Hemofil M)Vial: 250, 500, 1,000 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor – recombinant, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU		Vial: 250, 500, 1,000, 2,000, 2,500, 3,000, 4,000 IU
recombinant (Recombinate)IUAntihemophilic factor – recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor – recombinant (Xyntha Solofuse)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – recombinant (Obizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – human (Hemofil M)Vial: 250, 500, 1,000 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor – human, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU	recombinant (Nuwiq)	
recombinant (Recombinate)IUAntihemophilic factor – recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor – recombinant (Xyntha Solofuse)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – recombinant (Obizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – human (Hemofil M)Vial: 250, 500, 1,000 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor – human, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU	· · · · · · · · · · · · · · · · · · ·	Vial: 220-400, 401-800, 801-1240, 1241-1800, 1801-2400
Antihemophilic factor – recombinant (ReFacto, Xyntha)Vial: 250, 500, 1,000, 2,000 IUAntihemophilic factor – recombinant (Xyntha Solofuse)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – human (Hemofil M)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor – human (Roate-DVI)Vial: 250, 500, 1,000 IU	-	
recombinant (ReFacto, Xyntha)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Xyntha Solofuse)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – human (Hemofil M)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor – recombinant, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU	(Recombinate)	
recombinant (ReFacto, Xyntha)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Xyntha Solofuse)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – human (Hemofil M)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor – recombinant, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU	Antihemophilic factor –	Vial: 250, 500, 1,000, 2,000 IU
Antihemophilic factor – recombinant (Xyntha Solofuse)Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – human (Hemofil M)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000 IU	recombinant (ReFacto,	
recombinant (Xyntha Solofuse) Antihemophilic factor – recombinant (Obizur) Antihemophilic factor – human (Hemofil M) Antihemophilic factor – human (Koate-DVI) Antihemophilic factor – Yial: 250, 500, 1,000, 1,700 IU Vial: 250, 500, 1,000 IU Vial: 250, 500, 1,000 IU Vial: 500, 1,000, 2,000, 3,000 IU	Xyntha)	
Solofuse)Vial: 500 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – human (Hemofil M)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor – recombinant, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU	Antihemophilic factor –	Prefilled syringe: 250, 500, 1,000, 2,000, 3,000 IU
Solofuse)Vial: 500 IUAntihemophilic factor – recombinant (Obizur)Vial: 500 IUAntihemophilic factor – human (Hemofil M)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor – recombinant, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU	recombinant (Xyntha	
recombinant (Obizur)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – human (Hemofil M)Vial: 250, 500, 1,000 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor – recombinant, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU	Solofuse)	
Antihemophilic factor – human (Hemofil M)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor – recombinant, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU	Antihemophilic factor –	Vial: 500 IU
Antihemophilic factor – human (Hemofil M)Vial: 250, 500, 1,000, 1,700 IUAntihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor – recombinant, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU	recombinant (Obizur)	
Antihemophilic factor – human (Koate-DVI)Vial: 250, 500, 1,000 IUAntihemophilic factor – recombinant, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU		Vial: 250, 500, 1,000, 1,700 IU
human (Koate-DVI)Vial: 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU	human (Hemofil M)	
human (Koate-DVI)Vial: 500, 1,000, 2,000, 3,000 IUAntihemophilic factor – recombinant, PEGylated-Vial: 500, 1,000, 2,000, 3,000 IU	Antihemophilic factor –	Vial: 250, 500, 1,000 IU
recombinant, PEGylated-	-	
	Antihemophilic factor –	Vial: 500, 1,000, 2,000, 3,000 IU
	recombinant, PEGylated-	
	aucl (Jivi)	

V. References

- 1. Advate Prescribing Information. Westlake Village, CA: Baxalta US, Inc.; December 2018. Available at: <u>www.advate.com</u>. Accessed December 1, 2020.
- 2. Adynovate Prescribing Information. Westlake Village, CA: Baxalta US, Inc.; May 2018. Available at: <u>www.adynovate.com</u>. Accessed December 1, 2020.
- 3. Afstyla Prescribing Information. Kankakee, IL: CSL Behring LLC; April 2020. Available at: <u>http://labeling.cslbehring.com/PI/US/Afstyla/EN/Afstyla-Prescribing-Information.pdf</u>. Accessed December 1, 2020.
- 4. Eloctate Prescribing Information. Cambridge, MA: Biogen, Inc.; September 2019. Available at: <u>www.eloctate.com</u>. Accessed December 1, 2020.
- 5. Esperoct Prescribing Information. Plainsboro, NJ: Novo Nordisk Inc.; October 2019.

YouthCare Health*Choice* Illinois

Available at: https://www.novo-pi.com/esperoct.pdf. Accessed December 1, 2020.

- 6. Helixate FS Prescribing Information. Whippany, NJ: Bayer HealthCare LLC; May 2016. Available at: http://www.helixate.com/. Accessed December 1, 2020.
- Hemofil M Prescribing Information. Westlake Village, CA: Baxter Healthcare Corporation; June 2018. Available at: <u>http://www.shirecontent.com/PI/PDFs/HEMOFILM_USA_ENG.pdf</u>. December 1, 2020.
- 8. Jivi Prescribing Information. Whippany, NJ: Bayer HealthCare LLC; August 2018. Available at: <u>www.jivi.com</u>. Accessed December 1, 2020.
- 9. Koate-DVI Prescribing Information. Research Triangle Park, NC: Grifols Therapeutics, Inc.; June 2018. Available at: <u>www.koate-dviusa.com</u>. Accessed December 1, 2020.
- 10. Kogenate FS. Whippany, NJ: Bayer HealthCare LLC; December 2019. Available at: <u>www.kogenatefs.com</u>. Accessed December 1, 2020.
- 11. Kovaltry Prescribing Information. Whippany, NJ: Bayer HealthCare LLC; March 2016. Available at: <u>www.kovaltry-us.com</u>. Accessed December 1, 2020.
- 12. Novoeight Prescribing Information. Plainsboro, NJ: Novo Nordisk, Inc.; July 2020. Available at: <u>http://www.novoeight.com/</u>. Accessed December 1, 2020.
- 13. Nuwiq Prescribing Information. Hoboken, NJ: Octapharma; July 2017. Available at: <u>www.nuwiq.com</u>. Accessed December 1, 2020.
- 14. Obizur Prescribing Information. Westlake Village, CA: Baxalta US, Inc.; July 2020. Available at: <u>www.obizur.com</u>. Accessed December 1, 2020.
- 15. Recombinate Prescribing Information. Westlake Village, CA: Baxalta US Inc.; June 2018. Available at: <u>www.recombinate.com</u>. Accessed December 1, 2020.
- 16. Xyntha Prescribing Information. Philadelphia, PA: Wyeth Pharmaceuticals, Inc.; August 2020. Available at: <u>www.xyntha.com</u>. Accessed December 1, 2020.
- 17. Xyntha Solofuse Prescribing Information. Philadelphia, PA: Wyeth Pharmaceuticals, Inc.; August 2020. Available at: <u>www.xyntha.com</u>. Accessed December 1, 2020.
- 18. Srivastava A, Brewer AK, Mauser-Bunschoten EP, et al. Guidelines for the management of hemophilia. *Haemophilia*. Jan 2013; 19(1): e1-47.
- Medical and Scientific Advisory Council (MASAC) of the National Hemophilia Foundation (NHF): Database of treatment guidelines. Available at <u>https://www.hemophilia.org/Researchers-Healthcare-Providers/Medical-and-Scientific-Advisory-Council-MASAC/MASAC-Recommendations</u>. Accessed December 1, 2020.

Coding Implications

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	Description
Codes	
J7207	Injection, factor VIII (antihemophilic factor, recombinant) PEGylated, 1 IU
J7209	Injection, factor VIII (antihemophilic factor, recombinant) (Nuwiq), 1 IU
J7182	Injection, factor VIII, (antihemophilic factor, recombinant), (NovoEight), per IU
J7185	Injection, factor VIII (antihemophilic factor, recombinant) (Xyntha), per IU
J7188	Injection, factor VIII (antihemophilic factor, recombinant) (Obizur), per IU
J7190	Factor VIII (antihemophilic factor, human) per IU
J7191	Factor VIII (antihemophilic factor, porcine) per IU
J7192	Factor VIII (antihemophilic factor, recombinant) per IU, not otherwise specified



Reviews, Revisions, and Approvals	Date	P&T Approval Date
Policy split from CP.PHAR.12.Blood Factors and converted to new template. Added Kovaltry; removed requests for documentation; added 12 and older per PI indications if Adynovate. Removed preferencing for Helixate before Kogenate and Refacto. Under initial criteria, removed requirement for "severe hemophilia" and "history of 2 or more joint bleeds for prophylaxis indication." Non-prophylactic approval duration changed to 3 months initially with one 3-month re- auth. Removed denial based on inhibitor titer of \geq 5 BU/mL. Reviewed by specialist.	04.01.16	05.16
Product updates: Afstyla added (new drug); Adynovate updated to include perioperative management and use in children; Koate added	04.01.17	05.17



	63	
Reviews, Revisions, and Approvals	Date	P&T Approval Date
- Koate-DVI being phased out; Kogenate is available via three		
different PIs as Kogenate FS, Kogenate FS with Vial Adapter and		
Kogenate FS with Bio-Set; Obizur added (new drug for acquired		
hemophilia); ReFacto – removed "short term" use from criteria;		
Xyntha Solofuse added (same indications as Xyntha).		
Required trial of desmopressin is edited to avoid necessity of testing		
for coagulation factors. Safety information removed.		
Removed age >18 age restriction for Obizur per specialist		
recommendation.		
Wording for uses of all blood factor products made consistent across		
all policies. Per specialist review, for congenital hemophilia A,		
opened indications for routine prophylaxis up to all drugs listed in the		
policy, except Obizur. Approval periods across all blood factor		
policies made consistent.		
Efficacy statement added to renewal criteria. Hemophilias are		
specified as "congenital" versus "acquired" across blood factor		
policies where indicated.		
Reviewed by specialist- hematologist/internal medicine.		
Changed to new Centene Medicaid template	10.01.17	
1Q18 annual review:	11.27.17	02.18
-No significant changes.		
-References reviewed and updated.		
1Q 2019 annual review: added HIM-Medical Benefit; added Jivi;	10.29.18	02.19
removed Monoclate-P since it is no longer available on market;		
removed requirement for failure of Advate for Xyntha requests as it is		
not clinically necessary nor contractually driven; allowed use of		
Kovaltry for routine prophylaxis per FDA indication; moved criterion		
that member does not have VWD to section III Diagnoses/Indications		
Not Covered; references reviewed and updated.		
No significant changes: Esperoct added to the policy; referenced	03.13.19	
reviewed and updated.		
Removed reference to Centene preferred product strategy for	12.11.19	
migration to HFS PDL.		
Removed references to HIM, Medicaid policy only.		
3Q 2021 Annual Review: added diagnosis Routine prophylaxis	7.2.21	
to prevent or reduce the frequency of bleeding episodes;		
Removed Refacto; Added Documentation of member's body		
weight (in kg);		
Updated Appendix D general information; Undated table dosage and		
adminstration;Updated table product availability;Updated Coding		
Implications; Reviewed and updated references		

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

YouthCare Health Choice Illinois

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.

YouthCare HealthChoice/Illinois

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.

This clinical policy is the property of the Health Plan. Unauthorized copying, use, and distribution of this clinical policy or any information contained herein are strictly prohibited. Providers, members and their representatives are bound to the terms and conditions expressed herein through the terms of their contracts. Where no such contract exists, providers, members and their representatives agree to be bound by such terms and conditions by providing services to members and/or submitting claims for payment for such services.

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.

©2016 Centene Corporation. All rights reserved. All materials are exclusively owned by Centene Corporation and are protected by United States copyright law and international copyright law. No part of this publication may be reproduced, copied, modified, distributed, displayed, stored in a retrieval system, transmitted in any form or by any means, or otherwise published without the prior written permission of Centene Corporation. You may not alter or remove any trademark, copyright or other notice contained herein. Centene[®] and Centene Corporation[®] are registered trademarks exclusively owned by Centene Corporation.