

ADYNOVATE 250, 500, 750, 1000, 1500, 2000, 3000IU**Antihemophilic Factor (Recombinant), PEGylated****What is in this leaflet**

1. What *ADYNOVATE* is used for
2. How *ADYNOVATE* works
3. Before you use *ADYNOVATE*
4. How to use *ADYNOVATE*
5. While you are using it
6. Side effects
7. Storage and Disposal of *ADYNOVATE*
8. Product Description
9. Manufacturer and Product Registration Holder
10. Date of revision

What *ADYNOVATE* is used for

ADYNOVATE is an injectable medicine that is used to help treat and control bleeding in children and adults with hemophilia A (congenital Factor VIII deficiency). Your healthcare provider may give you *ADYNOVATE* when you have surgery. *ADYNOVATE* can reduce the number of bleeding episodes when used regularly (prophylaxis).

ADYNOVATE is not used to treat von Willebrand disease.

How *ADYNOVATE* works

ADYNOVATE, a PEGylated form of recombinant antihemophilic factor (*ADVATE*), temporarily replaces the missing coagulation factor VIII needed for effective hemostasis in congenital hemophilia A patients. *ADYNOVATE* exhibits an extended terminal half-life through pegylation of the parent molecule, *ADVATE*, which reduces binding to the physiological factor VIII clearance receptor (LRP1).

Before you use *ADYNOVATE*When you must not use it

You should not use *ADYNOVATE* if you:

- are allergic to any ingredients in *ADYNOVATE* or *ADVATE*. See the end of this leaflet for a complete list of ingredients in *ADYNOVATE*.
- are allergic to mice or hamster protein.

If you are unsure about this, ask your doctor.

Before you start to use it

You should tell your healthcare provider if you:

- Have or have had any medical problems.
- Have any allergies, including allergies to mice or hamsters.
- Are breastfeeding. It is not known if *ADYNOVATE* passes into your milk and if it can harm your baby.
- Are pregnant or planning to become pregnant. It is not known if *ADYNOVATE* may harm your unborn baby.
- Have been told that you have inhibitors to factor VIII (because *ADYNOVATE* may not work for you).

Taking other medicines

You should tell your healthcare provider if you are taking any medicines, including prescription and non-prescription medicines, such as over-the-counter medicines, supplements or herbal remedies.

How to use *ADYNOVATE*

ADYNOVATE is given directly into the bloodstream.

You may infuse *ADYNOVATE* at a hemophilia treatment center, at your healthcare provider's office or in your home. You should be trained on how to do infusions by your healthcare provider or hemophilia treatment center. Many people with hemophilia A learn to infuse their *ADYNOVATE* by themselves or with the help of a family member.

You may have to have blood tests done after getting *ADYNOVATE* to be sure that your blood level of factor VIII is high enough to clot your blood.

Talk to your healthcare provider before travelling. Plan to bring enough *ADYNOVATE* for your treatment during this time.

Reconstituted product (after mixing dry product with wet diluent) must be used

within 3 hours and cannot be stored or refrigerated. Discard any *ADYNOVATE* left in the vial at the end of your infusion as directed by your healthcare professional.

Always follow the specific instructions given by your healthcare provider. The steps listed below are general guidelines for using *ADYNOVATE*. If you are unsure of the procedures, please call your healthcare provider before using.

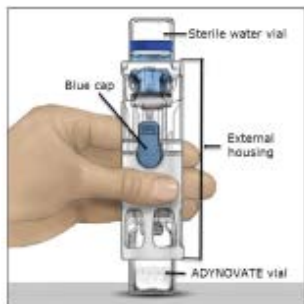
Call your healthcare provider right away if your bleeding does not stop after taking *ADYNOVATE*.

Step-by-step instructions for reconstituting *ADYNOVATE*:

1. Prepare a clean flat surface and gather all the materials you will need for the infusion.
 - Check the expiration date and let the *ADYNOVATE* warm up to room temperature.
 - Wash your hands and put on clean exam gloves. If infusing yourself at home, the use of gloves is optional.
2. Open the *ADYNOVATE* package by peeling away the lid. Remove the *ADYNOVATE* from the package and visually inspect the contents of the product and diluent vial. The *ADYNOVATE* powder should be white to off-white in color and the diluent should not contain particles. Do not use if discoloration or particles are seen.
3. Place on a flat surface with the diluent vial on top. The diluent vial has a blue stripe.

ADYNOVATE 250, 500, 750, 1000, 1500, 2000, 3000IU

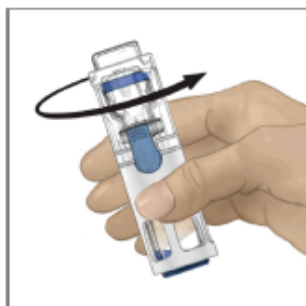
Antihemophilic Factor (Recombinant), PEGylated



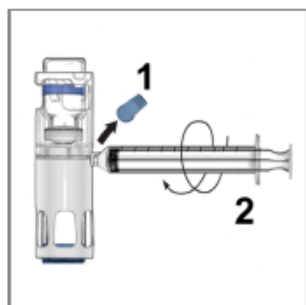
4. With one hand holding the *ADYNOVATE* housing, press down firmly on the diluent vial with the other hand until the system is fully collapsed and the diluent flows down into the *ADYNOVATE* vial. Both vials will move into the housing when pressed. If you don't see the diluent transfer to the product vial, press the vials again to assure they are completely inserted. Do not remove the blue cap until instructed in a later step.



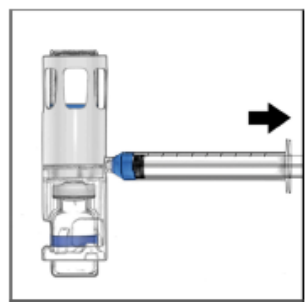
5. Swirl the *ADYNOVATE* gently and continuously until the *ADYNOVATE* is completely dissolved. Do not shake. Do not refrigerate after reconstitution. Inspect the *ADYNOVATE* solution for particulate matter and discoloration prior to administration. The solution should be clear and colorless in appearance. If not, do not use the solution and notify your healthcare provider immediately.



6. Take off the blue cap from the housing and connect the syringe. Be careful to not inject air into the *ADYNOVATE*.



7. Turn over the *ADYNOVATE* so that vial containing the *ADYNOVATE* solution is on top. Draw the *ADYNOVATE* solution into the syringe by pulling back the plunger slowly. If the solution does not draw into the syringe, be sure that both vials are pressed firmly together. The contents of more than one vial may be drawn into a single, appropriately sized syringe if you are using more than one vial of *ADYNOVATE*.



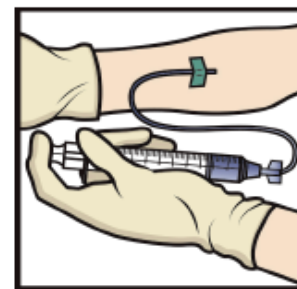
8. Disconnect the syringe from the system. Attach the infusion needle to the syringe using a winged (butterfly) infusion set, if available. Point the needle up

and remove any air bubbles by gently tapping the syringe with your finger and slowly and carefully pushing air out of the syringe and needle.

9. Apply a tourniquet and get the injection site ready by wiping the skin well with an alcohol swab (or other suitable solution suggested by your healthcare provider or hemophilia center).



10. Insert the needle into the vein and remove the tourniquet. Slowly infuse the *ADYNOVATE*. Do not infuse any faster than 10 mL per minute.



11. Take the needle out of the vein and use sterile gauze to put pressure on the infusion site for several minutes.
12. Remove the peel-off label from blister lid and place it in your logbook. Clean any spilled blood with a freshly prepared mixture of 1 part bleach and 9 parts water, soap and water, or any household disinfecting solution.
13. Do not recap the needle. Place needle, syringe and *ADYNOVATE* system in a hard-walled sharps container for proper disposal. Do not dispose

ADYNOVATE 250, 500, 750, 1000, 1500, 2000, 3000IU**Antihemophilic Factor (Recombinant), PEGylated**

of these supplies in ordinary household trash.

Important: Contact your healthcare provider or local hemophilia treatment center if you experience any problems.

How much to use

Your healthcare provider will tell you how much ADYNOVATE to use based on your individual weight, level of physical activity, the severity of your hemophilia A, and where you are bleeding.

When to use it

Always use this medicine exactly as your doctor has told you. Check with your doctor if you are not sure.

How long to use it

Do not stop using ADYNOVATE without consulting your doctor. If you have any further questions on the use of this medicine, ask your doctor.

If you forget to use it

Do not inject a double dose to make up for a forgotten dose. Proceed with the next injection as scheduled and continue as advised by your doctor.

If you use too much (overdose)

Always take ADYNOVATE exactly as your doctor has told you. You should check with your doctor if you are not sure. If you inject more ADYNOVATE than recommended, tell your doctor as soon as possible.

While you are using ADYNOVATEThings you must do

You must carefully follow your healthcare provider's instructions regarding the dose and schedule for infusing ADYNOVATE so that your treatment will work best for you.

Things you must not do

Do not attempt to do an infusion to yourself unless you have been taught

how by your healthcare provider or hemophilia center.

Medicines are sometimes prescribed for purposes other than those listed here. Do not use ADYNOVATE for a condition for which it is not prescribed. Do not share ADYNOVATE with other people, even if they have the same symptoms that you have.

Things to be careful of

Your body may form inhibitors to Factor VIII. An inhibitor is part of the body's normal defense system. If you form inhibitors, it may stop ADYNOVATE from working properly. Consult with your healthcare provider to make sure you are carefully monitored with blood tests for the development of inhibitors to Factor VIII.

Side effects

You can have an allergic reaction to ADYNOVATE.

Call your healthcare provider right away and stop treatment if you get a rash or hives, itching, tightness of the throat, chest pain or tightness, difficulty breathing, lightheadedness, dizziness, nausea or fainting.

The common side effects of ADYNOVATE are headache, diarrhea, rash, nausea, dizziness and hives. Tell your healthcare provider about any side effects that bother you or do not go away.

These are not all the possible side effects with ADYNOVATE. You can ask your healthcare provider for information that is written for healthcare professionals.

You may report any side effects or adverse drug reactions directly to the National Centre for Adverse Drug Reaction Monitoring by calling Tel: 03-78835490, or visiting the website npra.gov.my [Consumers → Reporting Side Effects to Medicines (ConSERF) or Vaccines (AEFI)].

Storage and Disposal of ADYNOVATEStorage

- Do not freeze.
- Store at refrigerated temperature 2°C to 8°C (36°F to 46°F).
- May store at room temperature not to exceed 30°C (86°F) for up to 3 months.
- Write the date on the carton when ADYNOVATE is removed from refrigerator.
- After storage at room temperature, do not return product back to the refrigerator.
- Do not use beyond the expiration date printed on the carton or vial.
- Store ADYNOVATE in the original box and protect from extreme exposure to light.

Disposal

- Do not throw away any medicines via waste water or household waste. Ask your pharmacist how to throw away medicines you no longer use. These measures will help protect the environment.

Product DescriptionWhat it looks like

ADYNOVATE with 2 mL or 5 mL Sterile Water for Injection in a BAXJECT III system comes in seven different dosage strengths: 250IU, 500IU, 750IU, 1000IU, 1500IU, 2000IU, and 3000IU. The actual strength will be imprinted on the label and on the box.

Always check the actual dosage strength printed on the label to make sure you are using the strength prescribed by your healthcare provider. Always check the expiration date printed on the box. Do not use the product after the expiration date printed on the box.

Ingredients

Active ingredient
Antihemophilic factor (recombinant), PEGylated

Inactive ingredients

ADYNOVATE 250, 500, 750, 1000, 1500, 2000, 3000IU

Antihemophilic Factor (Recombinant), PEGylated

Tris (Hydroxymethyl) Aminomethane,
Calcium Chloride, Mannitol, Sodium
Chloride, Trehalose Dihydrate,
Glutathione, Histidine, Polysorbate 80

MAL number(s):

MAL21056010AZ (250IU)
MAL21056011AZ (500IU)
MAL21056012AZ (750IU)
MAL21056013AZ (1000IU)
MAL21056014AZ (1500IU)
MAL21056015AZ (2000IU)
MAL21056016AZ (3000IU)

Manufacturer

Baxalta US Inc., 1700 Rancho Conejo
Boulevard,
Thousand Oaks, CA 91320, US

Product Registration Holder

Takeda Malaysia Sdn Bhd (868989-K),
Unit TB-L 13-1, Level 13, Tower B,
Plaza 33, No.1, Jalan Kemajuan,
Seksyen 13, 46200 Petaling Jaya,
Malaysia

Date of revision

06/10/2021

Serial Number

NPRA (R3/01) 14122021/0156