

Cablivi® 10 mg

powder and solvent for solution for injection caplacizumab

▼ This medicine is subject to additional monitoring. This will allow quick identification of new safety information. You can help by reporting any side effects you may get. See the end of section 4 for how to report side effects.

Read all of this leaflet carefully before you start using this medicine because it contains important information for you.

- Keep this leaflet. You may need to read it again.
- If you have any further questions, ask your doctor or pharmacist.
- This medicine has been prescribed for you only. Do not pass it on to others. It may harm them, even if their signs of illness are the same as yours.
- If you get any side effects, talk to your doctor or pharmacist. This includes any possible side effects not listed in this leaflet. See section 4.

What is in this leaflet

1. What Cablivi is and what it is used for
2. What you need to know before you use Cablivi
3. How to use Cablivi
4. Possible side effects
5. How to store Cablivi
6. Contents of the pack and other information

1. What Cablivi is and what it is used for

Cablivi contains the active substance caplacizumab. It is used to treat an episode of **acquired thrombotic thrombocytopenic purpura** in adults. This is a rare blood clotting disorder in which clots form in small blood vessels. These clots can block blood vessels and damage the brain, heart, kidneys, or other organs. Cablivi prevents the formation of these blood clots by stopping platelets in the blood from clumping together. By doing so, Cablivi reduces the risk of experiencing another episode of aTTP soon after the first.

2. What you need to know before you use Cablivi

Do not use Cablivi

- if you are allergic to caplacizumab or any of the other ingredients in this medicine (listed in section 6).

Warnings and precautions

Tell your doctor if you:

- bleed excessively during treatment. Your doctor may ask you to stop the treatment. The doctor will say when you can start your treatment again.
- are using anticoagulants such as vitamin K antagonists, rivaroxaban, apixaban (which treat blood clots). Your doctor will decide how you should be treated.
- are using anti-platelet agents such as aspirin, or low molecular weight heparin (which prevent blood clots). Your doctor will decide how you should be treated.
- have a bleeding disorder such as haemophilia. Your doctor will decide how you should be treated.
- have severely reduced liver function. Your doctor will decide how you should be treated.
- are going to have an operation or dental treatment. Your doctor will decide if it can be postponed or if you should stop Cablivi before your surgery or dental treatment.

Children and adolescents

Cablivi is not recommended for children under 18 years.

Other medicines and Cablivi

Tell your doctor or pharmacist if you are using, have recently used, or might use any other medicines.

Also tell your doctor if you are using an anticoagulant medicine such as vitamin K antagonists, rivaroxaban, or apixaban which treat blood clots or anti-platelet agents, such as aspirin, or low molecular weight heparin which prevent blood clots.

Pregnancy and breast-feeding

Tell your doctor if you are pregnant or plan to get pregnant. Use of Cablivi is not recommended during pregnancy.

Tell your doctor if you are breastfeeding. Your doctor will advise you whether to discontinue breastfeeding or not use Cablivi, considering the benefit of breastfeeding to the baby and the benefit of Cablivi to you.

Driving and using machines

Cablivi is not expected to influence the ability to drive or use machines.

Cablivi contains sodium

This medicine contains less than 1 mmol sodium (23 mg) per dose, i.e. it is essentially 'sodium-free'.

3. How to use Cablivi

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

Treatment with Cablivi is started by a doctor experienced in blood disorders.

The recommended treatment is

- **first dose :**
 - 1 vial injected into a vein by a healthcare professional
 - the medicine will be given before starting plasma exchange

- **subsequent doses:**

- 1 vial once daily as a subcutaneous injection (under the skin of the belly)
- the subcutaneous injection will be given after each daily plasma exchange
- after the daily plasma exchange finishes, your treatment with Cablivi will continue for at least 30 days with injection of 1 vial once daily.
- your doctor may ask you to continue daily treatment until the underlying signs of your disease are resolved

Your doctor may decide that you or your caregiver may inject Cablivi. In this case, your doctor or healthcare provider will train you or your caregiver on how to use Cablivi.

Instructions for use

The first injection of Cablivi into your vein must be given by a healthcare professional. Instructions for healthcare professionals on how to inject Cablivi into your vein are at the end of the leaflet.

For each injection, use a fresh kit package to prepare the injection solution. Do not try to inject Cablivi until you have been taught how to do so by a healthcare professional. Never use the kit for another injection.

Step 1 – Cleaning

- Wash your hands thoroughly with soap and water.
- Prepare a clean flat surface for placing the kit package.
- Make sure you have a disposal container at hand.

Step 2 – Before use

- Make sure the kit package is complete.
- **Check the expiry date.** Do not use if the expiry date has passed.
- Do not use the kit if the packaging or the items in it are damaged in any way.
- Place all components of the kit on the clean flat surface.
- If the kit was not stored at room temperature, allow the vial and the syringe to reach room temperature (15°C – 25°C) by letting them stand at room temperature for a few minutes. Do not warm them up in any other way.

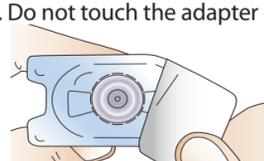
Step 3 – Disinfect the rubber stopper

- Remove the plastic flip-off cap from the vial. Do not use the vial if the green plastic cap is missing.
- Clean the exposed rubber stopper using one of the alcohol pads provided and allow it to dry for a few seconds.
- After cleaning, do not touch the rubber stopper or allow it to touch any surface.



Step 4 – Attaching the adapter

- Take the packed vial adapter and remove the paper cover. Leave the adapter in its opened plastic packaging. Do not touch the adapter itself.

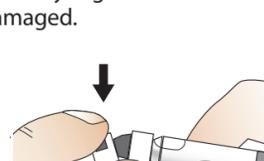


- Place the adapter over the vial, while keeping the adapter in its plastic packaging.
- Press down firmly until the adapter snaps into place, with the adapter spike going through the vial stopper. Leave the adapter attached to the vial, **still in its outer packaging.**



Step 5 – Prepare the syringe

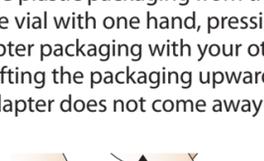
- Holding the syringe in your hand, break off the white cap with your other hand.
- Do not use the syringe if this white cap is missing, loose or damaged.



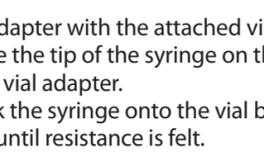
- **Do not touch** the syringe tip or allow it to come into contact with any surfaces.
- Place the syringe on the clean flat surface.

Step 6 – Connect syringe with adapter and vial

- Take the vial with the attached adapter.
- Remove the plastic packaging from the adapter by holding the vial with one hand, pressing the sides of the adapter packaging with your other hand, and then lifting the packaging upwards. Take care that the adapter does not come away from the vial.



- Hold the adapter with the attached vial with one hand. Place the tip of the syringe on the connector part of the vial adapter.
- Gently lock the syringe onto the vial by turning it clockwise until resistance is felt.



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- Keep the vial standing vertically on the surface with the syringe pointing downwards.
- Slowly push the syringe's plunger down until the syringe is empty. Do not remove the syringe from the vial.
- With the syringe still connected to the vial adapter, gently swirl the vial with connected syringe until the powder is dissolved. Avoid foaming. **Do not shake** the vial.

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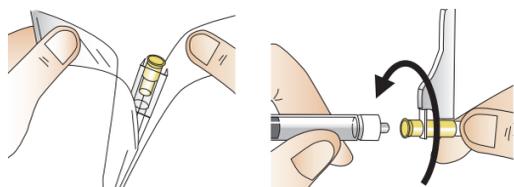
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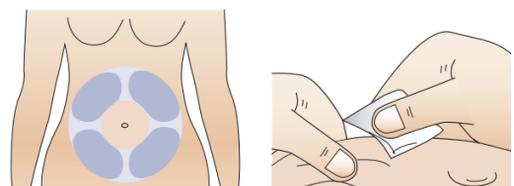
Step 10 – Attach the needle

- Unpack the needle by tearing the paper cover off the needle packaging and removing the needle with protective cap.



- Without removing the needle cap, attach the needle to the syringe by turning clockwise until resistance is felt.
- Pull back the needle safety shield.
- **Check the content of the syringe.** Do not use the medicine if you see any cloudiness, clumps or anything else that looks abnormal. Contact your doctor or nurse if this happens.

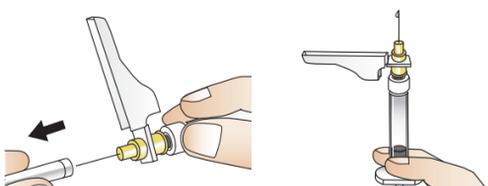
Step 11 – Prepare injection site for injection under the skin



- Select a suitable place ('injection site') on your belly for injection under your skin. Avoid the area around the belly button. Select a different injection site from the one you used on the previous day to help the skin to recover after the injection.
- Use the second alcohol pad to clean the injection site you have chosen.

Step 12 – Administration

- Carefully remove the needle protection cap from the needle and throw it away. Make sure the needle does not touch anything before the injection.
- Hold the syringe at eye level with the needle pointing upwards.
- Remove any air bubbles by tapping the side of the syringe with your finger to make the bubbles rise towards the tip. Then, slowly push the plunger until a small amount of liquid comes out of the needle.
- Gently pinch the cleaned skin between your thumb and forefinger to make a fold.



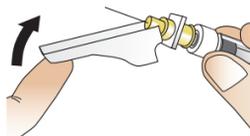
- Hold this skin fold during the entire injection.
- Insert the full length of the needle into the skin fold at an angle as shown in the illustration.
- Press the plunger down as far as it goes.



- Pull out the needle at the same angle you inserted it. Do not rub the injection site.

Step 13 – After administration

- Immediately after the injection, move the needle safety shield over the needle, until it clicks into place.



- Put the syringe with the needle in a disposal container.

If you use more Cablivi than you should

An overdose is unlikely since one vial contains only a single dose. Tell your doctor if you think you have had an overdose.

If you forget to use Cablivi

If you miss a dose you should still take it if it is within 12 hours of the scheduled time. If more than 12 hours have passed since the dose should have been given, do not take the missed dose, but inject the next dose at the usual time.

If you stop using Cablivi

To get the most benefit from your treatment, it is important to use Cablivi as prescribed and for as long as your doctor tells you to use it. Please talk to your doctor before you stop the treatment because stopping it too early can cause your condition to come back.

If you have any further questions on the use of this medicine, ask your doctor or pharmacist.

4. Possible side effects

Like all medicines, this medicine can cause side effects, although not everybody gets them.

Contact your doctor immediately if any of the following serious side effects occur:

Long or excessive bleeding.
Your doctor may decide to keep you under closer observation or change your treatment.

Side effects in a clinical study were reported with the following frequencies:

Very common, may affect more than 1 in 10 people

- bleeding gums
- fever
- tiredness
- headache
- nosebleeds
- hives

Common, may affect up to 1 in 10 people

- bleeding from eye
- vomiting blood
- blood in the stools
- black, tarry stools
- bleeding from the stomach
- bleeding hemorrhoids
- rectal bleeding
- injection site reactions: rash, itching and bleeding
- bleeding in brain as evidenced by severe headache of rapid onset, vomiting, decreased level of consciousness, fever, sometimes seizures and neck stiffness or neck pain
- muscle pain
- stroke
- blood in urine
- excessive bleeding during periods
- vaginal bleeding
- coughing blood
- shortness of breath
- bruise

Reporting of side effects

If you experience any side effects, talk to your doctor or pharmacist. This includes any possible side effect not listed in this leaflet. You can also report side effects directly via the Yellow Card Scheme at www.mhra.gov.uk/yellowcard or search for MHRA Yellow Card in the Google Play or Apple App Store.

By reporting side effects you can help provide more information on the safety of this medicine.

5. How to store Cablivi

Keep this medicine out of the sight and reach of children.

Do not use this medicine after the expiry date stated on the label and carton after EXP. The expiry date refers to the last day of that month.

Store in a refrigerator (2 °C – 8 °C). Do not freeze. Store in the original package in order to protect from light.

Cablivi may be stored at a temperature not above 25 °C for a single period of up to 2 months, but not beyond the expiry date. Do not return Cablivi to refrigerated storage after storage at room temperature. Never expose to temperatures above 30 °C.

Do not use Cablivi if you notice any particulate matter or discolouration prior to administration.

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

6. Contents of the pack and other information

What Cablivi contains

- **powder vial**
 - The active substance is caplacizumab. Each vial contains 10 mg caplacizumab.
 - The other ingredients are sucrose, citric acid anhydrous, trisodium citrate dihydrate and polysorbate 80.

- **pre-filled syringe**

The pre-filled syringe contains 1 mL water for injections.

What Cablivi looks like and contents of the pack

Cablivi is provided as:

- a white powder for solution for injection in a glass vial, and
- water for injections in a pre filled syringe to dissolve the powder.

After dissolving the powder in the solvent, the solution is clear, colourless or slightly yellowish.

Cablivi is available in

- single packs each containing 1 vial with caplacizumab powder, 1 pre-filled syringe with solvent, 1 vial adapter, 1 needle and 2 alcohol swabs
- multipacks each containing 7 single packs
- multidose packs each containing 7 vials with caplacizumab powder, 7 pre-filled syringes with solvent, 7 vial adapters, 7 needles and 14 alcohol swabs.

Not all pack sizes may be marketed.

Marketing Authorisation Holder and Manufacturer

Ablynx NV
Technologiepark 21
9052 Zwijnaarde
Belgium

For any information about this medicine, please contact the local representative of the Marketing Authorisation Holder.

United Kingdom

Sanofi
Tel +44 (0)845 372 7101

This leaflet was last revised in 01/2019.

Detailed information on this medicine is available on the European Medicines Agency web site: <http://www.ema.europa.eu/>

The following information is intended for healthcare professionals only:

The intravenous bolus injection of Cablivi given at the start of the treatment must be administered by a health care professional. Preparing a dose of Cablivi for intravenous injection should be done in the same way as for a subcutaneous injection (see Instructions for Use, step 1 to 9, in section 3).

Cablivi can be intravenously administered by connecting the prepared syringe to standard Luer locks of IV lines or using a suitable needle. The line can be flushed with 0.9% sodium chloride.