

ALEVIATE® HUMAN COAGULATION FACTOR VIII/VWF COMPLEX, POWDER FOR INJECTION

Human coagulation factor VIII (FVIII) / human von Willebrand factor (VWF) complex (250 IU FVIII/600 IU VWF, 500 IU FVIII/1200 IU VWF)

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What Aleviate® is used for

Aleviate® is for individuals with reduced levels or abnormality of either von Willebrand factor (VWF) or factor VIII (FVIII) in the bloodstream resulting in abnormal blood clotting, which may result in excessive bleeding.

Aleviate® is used to prevent or treat bleeding, including during surgery, due to lack of VWF in individuals with von Willebrand disease, when treatment with another medicine, desmopressin, is not effective or cannot be given.

Aleviate® is also used to prevent or treat bleeding, including during surgery, due to lack of FVIII in individuals with Haemophilia A.

How Aleviate® works

Aleviate® contains FVIII and VWF in a purified and concentrated form. Both FVIII and VWF are blood proteins that are essential for normal blood clotting.

Ask your Haemophilia Treatment Centre if you have any questions about why Aleviate® has been prescribed for you.

Before you use Aleviate®

- When you must not use it

Tell your doctor if you:

- have allergies to any medicines
- have allergies to any of the ingredients in this medicine (listed in Product Description)

- Before you start to use it

Tell your doctor if you:

- have any other medical conditions
- are pregnant or breast feeding
- become pregnant during your treatment

If you want further information, consult your doctor or Haemophilia Treatment Centre.

- have Von Willebrand disease

If you have a known risk of developing blood clots, you must be monitored for early signs of thrombosis (blood clotting). Your doctor should give you treatment to prevent thrombosis.

About blood products

Aleviate® is manufactured from human plasma (the liquid component of blood). When medicines are made from human blood or plasma, certain measures are put in place by the manufacturer to prevent viruses/infections being passed on to individuals. Despite these safety measures, the possibility of passing on infection cannot be totally excluded.

Vaccines are available against some viruses and your doctor can help you decide if vaccines are a suitable option for you.

Please discuss the risks and benefits of Aleviate® with your doctor.

- Taking other medicines

Tell your doctor if you are taking any other medicines, including any that you buy without a prescription from a pharmacy, supermarket or health food shop.

How to use Aleviate®

- How much to use

Your doctor will determine the exact dose for your condition.

Aleviate® will usually be given in a hospital. Should your doctor decide that treatment at

home is appropriate, your Haemophilia Treatment Centre will provide detailed instructions on how to use Aleviate®.

- When to use it

Use as directed by your doctor.

- How long to use it

Continue using Aleviate® for the duration that the doctor recommends.

- If you forget to use it

Consult your doctor or pharmacist on what you should do if you forget to use it.

Do not take a double dose to make up for the missed dose.

- If you use too much (overdose)

Cases of overdose have been observed. No severe adverse reactions were associated with these cases. The risk of blood clots cannot be excluded in cases of major overdose, especially in individuals with VWD. If you have any questions, consult your doctor.

Contact your doctor immediately or go to the Emergency Department of your nearest hospital, if you think you or anyone else may have used too much of this medicine. Do this even if there are no signs of discomfort or poisoning. You may need urgent medical attention.

While you are using Aleviate®

- Things you must do

Use your medicine exactly as your doctor has told you.

Tell all the doctors, dentists and pharmacists treating you that you are using Aleviate®.

Tell your doctor immediately if you become pregnant while using this medication.

Take Aleviate® only with a doctor's prescription.

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The following procedures are given as a guide only:

Preparing Aleviate[®] for administration

You will need one 5 mL Water for Injections vial for each 250 IU and 500 IU (100 IU/mL) vial of Aleviate[®].

1. Ensure you have all the required equipment to administer Aleviate[®].
2. Allow the vials of Aleviate[®] and Water for Injections to reach room temperature prior to use, which may take up to one hour. Do not warm the Water for Injections in hot water.
3. Remove jewellery, watches, rings, etc
4. Wash hands with soap and water, dry with a clean towel.
5. Select an appropriate work area with good lighting and a surface which can be cleaned (such as a kitchen table).
6. Using a clean cloth or paper towel, clean the preparation area with methylated spirits.
7. Gather the equipment to be used.

Equipment

- One carton of Aleviate[®] containing:
 - one vial of Aleviate[®]
 - one vial of Water for Injections
 - one Mix2Vial[™] filter transfer set.
 Check the expiry date of each item. Do not use if expired.
- two alcohol wipes
- sharps container
- waste container for discarding biological material
- plastic syringe(s)
- adhesive tape
- cotton balls
- intravenous injection set
- gloves.

Instructions for Aleviate[®] reconstitution

Follow these steps to prepare the injection.

1. Wash hands with soap and water, dry with a clean towel.

2. Ensure Aleviate[®] and Water for Injections are at room temperature.

3. Remove flip top caps from Aleviate[®] and Water for Injections vials.

4. Wipe the rubber stoppers of both the Aleviate[®] and Water for Injections vials with the alcohol wipes and allow to dry for two minutes. Do not leave the alcohol wipes resting on the stoppers. Do not touch the rubber stoppers with your fingers.

5. Open the lid of the Mix2Vial[™] packaging. If the seal of the lid is not intact or you have any other concerns about the integrity of the Mix2Vial[™], do not use it but return it to your Haemophilia Treatment Centre or the Manufacturer.

Place the Water for Injections vial on a level surface and hold the vial firmly. Take the Mix2Vial[™] together with the outer package, invert it and push the blue plastic cannula of the Mix2Vial[™] firmly through the rubber stopper of the Water for Injections vial.

See Figure 1, (5 mL Water for Injections vial is provided for 250 IU vial and 10 mL Water for Injections vial is provided for 500 IU (50 IU/mL) vial).

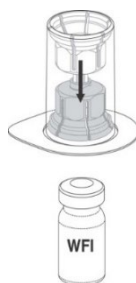


Figure 1



Figure 2



Figure 3



Figure 4

WFI = Water for Injections

6. While holding onto the vial of Water for Injections, carefully remove the outer package from the Mix2Vial[™], being careful to leave the Mix2Vial[™] firmly attached to the vial of Water for Injections, see Figure 2. Make sure that you only remove the outer package and not the Mix2Vial[™].
7. With the Aleviate[®] vial held firmly on a level surface, invert the Water for Injections with the Mix2Vial[™] attached and push the transparent plastic cannula end of the Mix2Vial[™] firmly through the stopper of the Aleviate[®] vial, see Figure 3.

The water will be drawn into the vial by the vacuum contained within the Aleviate[®] vial.

If water is not drawn into the vial, it means that there is no vacuum in the vial and the seal may be faulty. Do not use the product but return it to your Haemophilia Treatment Centre or the Manufacturer.

Note: The Mix2Vial[™] is intended to filter the contents of a single vial of Aleviate[®] only. If multiple vials of Aleviate[®] are to be given, a separate Mix2Vial[™] must be used for each vial.

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8. With the Water for Injections and Aleviate® vials still attached to the Mix2Vial™, gently swirl (do not shake) the Aleviate® vial until all of the product is dissolved. Ensure the contents of the vial are completely dissolved.

The solution should be colourless to slightly yellow with a clear or slightly opalescent appearance (meaning it might sparkle when held up to the light). The solution must not contain any obvious particles.

If the solution is cloudy, contains flakes or particles, or a clot or gel forms, do not use the product but return it to your Haemophilia Treatment Centre or Manufacturer.

9. Once the contents of the Aleviate® vial are completely dissolved, firmly hold both the transparent and blue parts of the Mix2Vial™. Unscrew the Mix2Vial™ into two separate pieces, see Figure 4. Discard the empty Water for Injections vial with the blue part of the Mix2Vial™ still attached into an appropriate waste container.
10. While the Aleviate® vial is upright, attach a plastic disposable syringe to the transparent part of the Mix2Vial™. Invert the system and draw the Aleviate® into the syringe by pulling the plunger back slowly.
11. Once the Aleviate® has been transferred into the syringe, firmly hold the barrel of the syringe (keeping the syringe plunger facing down) and detach the Mix2Vial™ from the syringe. Do not use the Mix2Vial™ for injection.

Note: One large syringe may be used to withdraw Aleviate® from multiple vials.

12. Discard the empty vial of Aleviate® with the transparent part of the Mix2Vial™ attached, into an appropriate waste container.

Use Aleviate® as soon as you can after preparation. (Make sure it is used for one person on one occasion only). Any unused portion remaining in the vial must be discarded appropriately.

Injection administration procedure guidelines

The injection is best given by following each of the steps outlined in turn.

1. Apply tourniquet. Select injection site.
2. Wash hands with soap and water, dry with a clean towel.
3. Cleanse the skin area with an alcohol wipe, allow to dry.
4. Put on gloves.
5. Insert intravenous needle into vein.
6. Secure needle with adhesive tape.
7. Attach the syringe containing Aleviate® to the intravenous needle.
8. Gently pull back the plunger until the tubing is filled with blood. Take care not to get any blood in the syringe containing the product.
9. Release the tourniquet.
10. Administer the Aleviate® solution slowly (usually within 5-6 minutes, or as tolerated). The infusion rate should not be more than 6 mL per minute.
11. Carefully remove adhesive tape.
12. Carefully remove the intravenous needle with syringe attached and place directly into the sharps container.
13. Apply pressure to the injection site using a cotton ball for one to two minutes. Apply dressing if necessary.
14. Discard all used sharps into the sharps container, and dispose of the other used equipment appropriately.
15. Wash hands with soap and water, dry with a clean towel.
16. It is recommended that treatment details and the lot number from the

Aleviate® vial label are recorded every time you use Aleviate®.

- Things you must not do

Do not stop using the medicine unless advised by your doctor.

Do not use any new medicines without consulting your doctor or pharmacist.

Do not give Aleviate® to anyone else, even if they have the same symptoms or condition as you.

The solution must not be stored for longer than 8 hours and the infusion should be completed as soon as practicable, as Aleviate® does not contain an antimicrobial preservative.

Do not refrigerate Aleviate® once it has been prepared.

- Things to be careful of

Should any of the symptoms listed under 'Side effects' develop, stop the infusion immediately and contact your Haemophilia Treatment Centre.

Side effects

You may experience unwanted effects while using Aleviate® although severe reactions are rare.

Stop using this product immediately and contact your doctor if any of the following side effects occur:

- headache
- pain (such as back pain, joint pains and bone pains)
- dizziness
- anxiety
- reddening of the face or neck
- chest pain
- sweating
- feeling sick or vomiting
- strange taste in the mouth
- irritation of the vein used for infusion (which may include swelling, redness or tenderness)

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- shortness of breath or difficulty breathing.

Contact your doctor immediately if you experience any of these symptoms at any time:

- fever
- loss of appetite
- extreme tiredness
- abdominal pain
- jaundice (yellow skin and eyes)
- dark urine
- joint pains
- skin rashes.

Inhibitors

Treatment with factor VIII products such as Aleviate® may sometimes lead to the formation of antibodies (inhibitors) which neutralise factor VIII and reduce the effectiveness of the treatment. Your doctor will monitor you for development of these inhibitors and if they suspect that an inhibitor is present, the level of inhibitor will be measured using the appropriate laboratory tests. In some individuals, inhibitors can be successfully overcome with larger doses of factor VIII, but for some individuals it may be necessary to switch to a different treatment.

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

Storage and Disposal of Aleviate®

- Storage

Store at 2°C to 8°C (Refrigerate. Do not freeze). Protect from light. Do not use after the expiry date.

- Disposal

Do not throw away any medicines via waste water or household waste.

Ask your doctor or healthcare professional how to throw away medicines you no longer use. These measures will help protect the environment.

Product Description

- What it looks like

Aleviate® is a white or pale yellow powder contained in a glass vial. Aleviate® is registered as two different presentations. Each pack size contains:

1. 250 IU vial of Aleviate® (with a FVIII concentration of 50 IU/mL and a VWF concentration of 120 IU/mL), 5 mL vial of Water for Injections, and a special filter transfer set called a Mix2Vial™.
2. 500 IU vial of Aleviate® (with a FVIII concentration of 50 IU/mL and a VWF concentration of 120 IU/mL), 10 mL vial of Water for Injections and a special filter transfer set called a Mix2Vial™.

- Ingredients

- Active ingredient(s)

Aleviate® contains FVIII and VWF as the active ingredients.

- Inactive ingredients

- sucrose
- sodium citrate dihydrate
- sodium chloride
- trometamol
- calcium chloride dihydrate
- human albumin
- human plasma proteins.

This product is packaged in latex free materials.

- MAL number:

MAL20040991AZ

Manufacturer

Aleviate® is manufactured in Australia by:

CSL Behring (Australia) Pty Ltd
189–209 Camp Road
Broadmeadows VIC 3047
Australia

Product Registration Holder

DKSH Malaysia Sdn Bhd
B-11-01, The Ascent Paradigm,
No.1, Jalan SS7/26A Kelana Jaya 47301
Petaling Jaya Selangor.

Date of revision

12/04/2022

Serial Number

NPRA(R4/1)080321/00007

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