

Guide for reducing the risks of medicinal products and their application
Patients and Caregivers

Vyjuvek (beremagene geperpavec)

5×10⁹ plaque-forming units/mL suspension
and gel for gel

Please also refer to the Vyjuvek instructions for use
that you have received from your healthcare provider or pharmacy.

▼ This medicinal product is subject to additional monitoring. This will allow quick identification of new safety information. You can help by reporting any adverse events that occur (for information on how to report adverse events, please see last page).

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Medical terms explained

These are brief explanations of medical terms used in the guide.

(**bolded terms** are also explained elsewhere in this glossary)

Alpha-1 chain	Name of a specific form of a protein
Beremagene geperpavec	The active ingredient in Vyjuvek; a genetically modified herpes virus that transports a specific gene into skin cells to enable them to produce type VII collagen .
COL7	Abbreviation for type VII collagen
COL7A1	The name of the gene that encodes the alpha-1 chain of the type VII collagen protein .
DEB	Acronym of "dystrophic epidermolysis bullosa".
Disinfect	A chemical or physical hygiene measure to kill or inactivate pathogens (bacteria or viruses) on surfaces.
Gene	A specific part of your genetic material that contains the information for the structure of a protein .
Genetically modified organism	An organism whose genetic information has been altered by an external process. In Vyjuvek, this was done to allow it to fulfil a specific role in the body.
Type VII collagen	Collagens are a specific type of structural protein . Type VII collagen is the building block for anchoring structures within the skin that bind together the outer and the underlying layer of the skin.
Mutation	A change in genetic information that causes a change in the structure of a protein . Mutations can cause proteins to lose or change their function.
Squamous cell carcinoma	A type of skin cancer that occurs more frequently and is more severe in people with DEB .
Plaque-forming unit	A unit of measurement of the amount of viruses in a liquid.
Protein	A substance that plays a crucial role in biological processes. Proteins are the "building blocks of life" that form important structures in all organisms and play a role in various functions.
Suspension	A liquid containing tiny, undissolved particles. In Vyjuvek, these tiny particles are viruses transporting the COL7A1 gene .
Virus	A virus is a tiny structure made up of proteins and genetic information. In order to multiply, viruses have to enter cells and use these cells' mechanisms. Specially modified viruses can be used to insert specific genes into the cells.
Virucidal	Virus -destroying (e.g. virucidal disinfectant).

What is Vyjuvek?

Vyjuvek is used to treat wounds in people with a rare genetic condition called dystrophic epidermolysis bullosa (DEB) caused by mutation(s) in the alpha 1 chain of type VII collagen gene (*COL7A1*) that primarily affects the skin. Vyjuvek is a gene therapy based on genetically modified viruses. It contains beremagene geperpevac as an active ingredient, a genetically modified virus that encodes the human protein COL7.

Neither the modified virus nor the genetic material in this medicinal product will change your DNA.

Vyjuvek is a gel-like medicine made to be applied to skin wounds via syringes. It has to be prescribed by a doctor and is mixed and prepared for use by your pharmacy.

This guide was created to provide safety-relevant information and instructions on the correct use of Vyjuvek to patients who have been prescribed this medication and/or to their caregivers.

Please note the following when applying Vyjuvek

- Because this medicinal product contains genetically modified organisms, anyone applying or disposing of Vyjuvek (or assisting with these tasks) must wear **protective equipment** (e.g. gown, disposable gloves, mask and eye protection).
- **Only treat those wounds** with Vyjuvek that you and your doctor have added to your **treatment plan**.
- Apply Vyjuvek to the wound you and your doctor have chosen for treatment **once a week**.
- Do not use Vyjuvek if you **do not have any wounds**.
- **Do not** apply Vyjuvek to wounds with a confirmed or suspected diagnosis of **squamous cell carcinoma**, a form of skin cancer.
- If you **forget** a treatment with Vyjuvek, make up for it as soon as possible and contact your doctor.
- Notify your doctor if you have applied **too much Vyjuvek**.
- Let your medical team know if you take, have recently taken or plan to take **other medications**.
- **Do not stop or interrupt treatment** without consulting with your doctor.
- Treatment with Vyjuvek can be done as part of your **routine dressing change**.
- Consult with your doctor on how to proceed when **documenting treated wounds**.



- If you are **pregnant or breastfeeding**, or if you may be pregnant or are planning a pregnancy, please consult with your doctor before applying Vyjuvek.
- If you are **pregnant**, you must **not come into contact with Vyjuvek**, or with skin or dressings that have come into contact with Vyjuvek. Do not apply Vyjuvek and do not assist in its application.

Application of Vyjuvek in an office/clinic setting or in a home environment

Consult with your doctor to determine where and by whom your Vyjuvek treatment will generally be carried out. Vyjuvek can be applied



in a **doctor's office or clinic** by healthcare professionals,



in a **home environment** by medical professionals,



in a **home environment** by you or a caregiver after consultation with and training by your healthcare provider.



Application of Vyjuvek in an office/clinic setting

If Vyjuvek is applied in a doctor's office or clinic by healthcare professionals, these also dress the wounds accordingly. After the treatment, please note the following:

- The dressing should remain on the treated wound for **approx. 24 hours**.
- **Refrain** from **touching** the wound during this time.
- For information on how to **perform the first dressing change after application** (including the disposal of the wound dressing), see "The first dressing change after applying Vyjuvek".



Application of Vyjuvek in a home environment by medical professionals

If, after consultation with a doctor, it has been determined that medical personnel may apply Vyjuvek in a home environment, the following steps must be taken together with your doctor **before the first application**:

- **Information and counseling** of the patient and/or caregiver on the use of Vyjuvek in a home environment by a healthcare provider
- **Training** of persons who apply or assist in applying Vyjuvek **by a healthcare provider**
- Development of a **treatment plan** by the treating physician (including determining the **order** of wounds to be treated and **dosage**)
- Provision and discussion of this **guide**



Application of Vyjuvek in a home environment by you or a caregiver

If, after consultation with a doctor, it has been determined that you or a caregiver may use Vyjuvek in a home environment, the following steps must be taken together with your doctor **before the first application**:

- **Information and counseling** of the patient and/or caregiver on the use of Vyjuvek in a home environment by a healthcare provider
- **Training** of persons who apply or assist in applying Vyjuvek **by a healthcare provider**. The application of Vyjuvek must be practiced under the supervision of a healthcare provider until the patient/caregiver has fully mastered the application process. A practice kit without any active ingredients is available.
- Development of a **treatment plan** (including determining the **order** of wounds to be treated and **dosage**)
- Provision and discussion of this **guide**



Vyjuvek may only be used in a home environment after all these conditions have been met!

Application of Vyjuvek



Because this medicinal product contains genetically modified organisms, anyone applying or disposing of Vyjuvek (or assisting with these tasks) must wear **protective equipment** (e.g. gown, disposable gloves, mask and eye protection).

Please **note** that you will need to use **two different cleaning/disinfecting products**:



For the wound:
a wound cleanser
WITHOUT virucidal ingredients



For surfaces and materials
that have been in contact with Vyjuvek:
a disinfectant **WITH** virucidal ingredients



Vyjuvek administration" video and demo kit

The Vyjuvek **administration video** is available by flashing the QR code or on the website: <http://ema.krystallabel.com>.

You can request a **demonstration kit** with syringes (without active ingredient) to practice administering Vyjuvek. Order it from the following address
medinfo.estonia@swixxbiopharma.com



Practice with the kit does not replace the medical training required before using Vyjuvek at home.

Step 1: Preparation

Step 1.1: Preparation of the work area



- Clean the work area with a **virucidal disinfectant** and set up the materials you will need:

- 1 syringes filled with Vyjuvek
- 2 a **non-absorbent, water-repellent** dressing (that does not absorb Vyjuvek) that is slightly larger than the selected wound
- 3 a **standard wound dressing** that is slightly larger than the non-absorbent, water-repellent dressing
- 4 a pair of scissors
- 5 **protective equipment** (e.g. gown, disposable gloves, mask and eye protection)
- 6 **wound cleanser** (**without** virucidal ingredients)
- 7 **disinfectant** (**with** virucidal ingredients)
- 8 a sealable **plastic bag** (for disposal)



- Anyone applying or assisting with the application of Vyjuvek should now don their **protective equipment**.



- Verify
 - that the number of syringes corresponds to your dosage (1 syringe contains 0.5 mL of Vyjuvek).
 - the **expiry date**. Do not use the medicinal product after the expiry date has passed.
 - the **color of the medicinal product**. It may range from opalescent yellow to colorless. Do not use the medicinal product if you notice any discoloration outside this range.

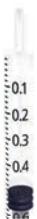
If this happens, notify your doctor and/or your pharmacy.

Step 1.2: Preparing the selected wound



- **Carefully** remove dead skin, scabs and wound fluid as well as any medicinal products and ointments from the wound area and gently **cleanse** the wound. Use a **wound cleanser WITHOUT** virucidal ingredients, e.g. sterile saline solution. If you are unsure, ask your doctor.

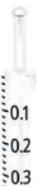
Step 1.3: Preparation of the syringe



- Hold the syringe with **its tip pointing upwards**.



- **Pull the plunger a little**, but not completely, out of the syringe.



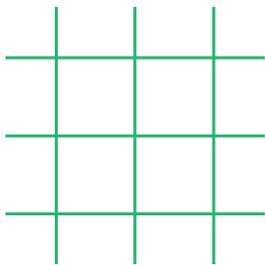
- **Now slowly push** the plunger **back into the syringe** until a small drop of Vyjuvek exits the syringe tip.

Step 2: Applying Vyjuvek



Note:

- Apply Vyjuvek **only to the wound to be treated**.
- Keep applying Vyjuvek to the same wound until it is **completely closed**. Only then start treating a new wound (according to the treatment plan/as discussed with your medical team).
- If a previously treated wound **reopens**, continue by treating **this wound first**.
- **Do not apply any other medicines** to the wound at the same time as Vyjuvek.



- Imagine a **grid with fields sized 1×1 cm** covering the selected wound (1 cm is roughly the width of the tip of your little finger).



- Then apply **one drop of Vyjuvek to each of these (imaginary) fields**.



- Make sure that the tip of the syringe **does not touch the skin!**

The **amount** of Vyjuvek required may **vary** depending on the size of the wound. It is possible that one dose of Vyjuvek is not sufficient to treat all wounds or that you still have Vyjuvek left after treating the wounds specified in your treatment plan. If this happens, contact your medical team or your pharmacy and dispose of any remaining Vyjuvek (see “Cleaning and disposal”).

Maximum dose per weekly application:

- Adults and children aged 3 years and over: 4 syringes filled with 0.5 mL Vyjuvek each
- Children under 3 years of age: 2 syringes filled with 0.5 mL Vyjuvek each

Step 3: Bandaging the wound



- Cut the **non-absorbent, water-repellent dressing** to a size **slightly larger than the treated wound** and place it on the wound in such a way that the edges of the dressing **fully extend beyond the wound surface area on all sides**.



- Afterwards, **completely cover** the **above wound dressing with your standard bandage** (in order to prevent the non-absorbent, water-repellent dressing from moving and Vyjuvek coming into contact with other areas of the skin as well as to absorb wound fluid).



- Leave the dressings on the wound for **about 24 hours** and do not touch the wound during this time if possible.

Cleaning and disposal



- **Clean any surfaces** that have come into contact with Vyjuvek using a **virucidal disinfectant**.
- **Disinfect everything that came or could have come into contact with Vyjuvek** (e.g. dressings, used and unused syringes, used materials) **with a virucidal disinfectant**.



- Afterward, place the previously disinfected, used materials in an **sealable plastic bag**. Seal it and dispose of it with **household waste**.

Vyjuvek is tested for **sterility**. Nevertheless, a transmission of infectious agents cannot be completely ruled out. After application, monitor the patient for **signs of infection** (fever etc.). If this happens, notify your doctor.



The first dressing change after applying Vyjuvek

- Wear **protective equipment** (e.g. gown, disposable gloves, mask and eye protection).
- **Remove** the dressings/bandages as usual.
- **Disinfect** the removed, used dressings using a **virucidal disinfectant**.
- **Dispose** of used wound dressings in a separate, sealable plastic bag with household waste after disinfecting.

After changing the Vyjuvek wound dressing, you can continue with your regular wound care regimen.

Measures to take after accidental contact with Vyjuvek

- Avoid direct contact of Vyjuvek with **areas other than the treated wound**.
- **Avoid direct contact** with the treated wound/the dressing for **approx. 24 hours** after application.
- The person performing the treatment or disposing of syringes and dressings should always wear **protective equipment** (e.g. gown, disposable gloves, mask and eye protection).



If Vyjuvek accidentally gets **into your eyes or on your mucous membranes** (e.g. nose, mouth), thoroughly rinse the affected area(s) **with clean water for at least 5 minutes**.



If your **intact skin** comes into contact with Vyjuvek, **wash** the affected area(s) thoroughly with **soap and water and/or a virucidal disinfectant**.



If **surfaces** unintentionally come into contact with Vyjuvek, clean them using a **virucidal disinfectant**.

Transportation of Vyjuvek

- When you collect the Vyjuvek syringes from the pharmacy, transport them in the **container provided by the pharmacy** for this purpose, as it meets special transportation requirements for the medicinal product (e.g. refrigeration temperature).
- If the pharmacy ships the syringes to your home, these will be delivered in a plastic bag in a suitable **insulated container (outer container)** that should only be opened **by the person responsible for applying the medicinal product**.
- Upon receipt, this **person** must also ensure the **integrity of the outer container** and that there are no signs of leaking. If any damage or leakage has occurred, please immediately notify your pharmacy.

Storage of Vyjuvek

- Leave the syringes with Vyjuvek in their **outer container**.
- Store the syringes in a **clean place** and **keep them away from children**, as instructed by your pharmacy.
- Use Vyjuvek **before the use-by date provided by the pharmacy**.

Reporting adverse events

By reporting adverse events, you can help to ensure that more information on the safety of Vyjuvek becomes available.

If you notice any side effects, contact your doctor or pharmacist.

REFERENCE: Vyjuvek, current patient information leaflet

Additional information



All Vyjuvek training documents for patients (guide for patients treated with Vyjuvek and their caregivers, Vyjuvek administration video) and all information on the drug, are also available online by flashing the QR code or at <http://ema.krystalabel.com>.

If you have any questions about your health or your treatment with Vyjuvek, please contact your doctor.