

Guidance on reducing medicine and application risks  
**Patients and caregivers**

# Vyjuvek (Beremagen geperpavec)

5×10<sup>9</sup> plaque-forming units/ml suspension and  
gel for gel preparation

Please also read the package insert for Vyjuvek that you received from your doctor, healthcare professional, or pharmacy.

▼ This medicine is subject to additional monitoring. This allows for quick identification of new findings regarding safety. You can help by reporting any side effects (see the last page for information on reporting side effects).



# Contents

<b>Explanation of medical terms</b> .....	<b>4</b>
<b>What is Vyjuvek?</b> .....	<b>5</b>
<b>Please note during treatment</b> .....	<b>6</b>
<b>Use of Vyjuvek in a doctor's office/clinic or at home</b> .....	<b>7</b>
Use of Vyjuvek in a doctor's office/clinic.....	7
Use of Vyjuvek in the home environment by healthcare professionals.....	8
Use of Vyjuvek in the home environment by you or a caregiver.....	8
<b>Use of Vyjuvek</b> .....	<b>9</b>
Video "Applying Vyjuvek" Vyjuvek" and Demo kit.....	9
Step 1: Preparation.....	10
Step 2: Applying Vyjuvek.....	12
Step 3: Dressing the wound.....	13
Cleaning and disposal.....	13
<b>The first dressing change after applying Vyjuvek</b> .....	<b>14</b>
<b>Measures in case of accidental contact with Vyjuvek</b> .....	<b>14</b>
<b>Transport of Vyjuvek</b> .....	<b>15</b>
<b>Storage of Vyjuvek</b> .....	<b>15</b>
<b>Further information</b> .....	<b>16</b>
<b>Reporting side effects</b> .....	<b>17</b>

# Explanation of medical terms used in the guide

Here you will find brief explanations of medical terms used in the guide. (**Terms printed in bold** are also explained elsewhere in this glossary.)

<b>Alpha-1 chain</b>	Term for a specific structure of a <b>protein</b>
<b>Beremagen geperpavec</b>	The active ingredient in Vyjuvek: a genetically modified herpes virus that inserts a specific <b>gene</b> into skin cells so that they can produce <b>type VII collagen</b> .
<b>COL7</b>	Abbreviation for <b>collagen type VII</b>
<b>COL7A1</b>	Name of <b>the gene</b> encoding the <b>alpha-1 chain</b> of <b>collagen type VII protein</b>
<b>DEB</b>	Abbreviation for 'dystrophic epidermolysis bullosa'.
<b>Disinfection</b>	A chemical or physical hygiene measure aimed at killing or inactivating pathogens (bacteria or <b>viruses</b> ) on surfaces.
<b>Gene</b>	A specific section of genetic information that contains the information for building a <b>protein</b> .
<b>Genetically modified organism</b>	An organism whose genetic information has been altered by external intervention. In Vyjuvek, this is done to enable the organism to perform a specific task in the body.
<b>Collagen type VII</b>	Collagens are a specific type of <b>protein</b> that serve as structural elements due to their structure. Collagen type VII is the building block for anchoring structures in the skin that hold the outermost layer of skin and the underlying layer of skin together.
<b>Mutation</b>	A change in genetic information that causes <b>proteins</b> to be constructed differently than they would be without the mutation. Mutations can cause <b>proteins</b> to lose or change their function.
<b>Squamous cell carcinoma</b>	A form of skin cancer that occurs more frequently and is more severe in people with <b>DEB</b> than in others.
<b>Plaque-forming unit</b>	Measure of the amount of <b>virus</b> in a fluid
<b>Protein</b>	Also known as egg white. Proteins are the "building blocks of life" that form important structures and perform functions in all organisms.
<b>Suspension</b>	A liquid containing tiny undissolved particles. In Vyjuvek, these tiny particles are the <b>viruses</b> that carry the <b>COL7A1 gene</b> .
<b>Virus</b>	A virus is a tiny structure made up of proteins and genetic information. In order to reproduce, viruses must invade cells and use their mechanisms. Genetically modified viruses can be used to introduce desired <b>genes</b> into cells.
<b>Virucidal</b>	Destroying <b>viruses</b> (example: virucidal <b>disinfectants</b> ).

# What is Vyjuvek?

Vyjuvek is used to treat wounds in people with a rare genetic disorder called dystrophic epidermolysis bullosa (DEB), which is caused by mutation(s) in the gene for the alpha-1 chain of collagen type VII (*COL7A1*) and mainly affects the skin. Vyjuvek is a gene therapy based on modified viruses. It contains the active ingredient beremagen geperpavec, a genetically modified virus that encodes the human protein COL7.

The modified virus and the genetic material in this medicine **do not** change your own DNA.

Vyjuvek is a gel-like medicine in syringes for application to wounds on the skin. It must be prescribed by a doctor and is mixed by your pharmacy before use.

This guide is intended to provide patients who are prescribed Vyjuvek and/or their caregivers with safety information and instructions for proper use.

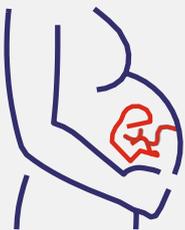


Patients using Vyjuvek are encouraged to participate in a **study to long-term** safety of the medicine.

**If you are interested** in participating, **talk** to your doctor.

# Please note during treatment

- Since this medicine contains genetically modified organisms, people who apply Vyjuvek, dispose of it, or assist in these activities must wear **protective equipment** (e.g., gowns, disposable gloves, masks, and eye protection).
- **Only** treat **wounds** with Vyjuvek that you have discussed with your doctor in your **Treatment plan** has been established.
- Apply Vyjuvek **once a week** to the wound as determined by your doctor.
- Do not use Vyjuvek if you **do not** have **anywounds**.
- 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, apply it as soon as possible and contact your doctor.
- If you have applied **too much Vyjuvek**, tell your doctor.
- Tell your healthcare team if you are taking, have recently taken, or plan to take **any other medications**.
- Do not **stop** treatment without consulting your doctor.
- Treatment with Vyjuvek can be carried out as part of your **routine dressing change**.
- 
- If you are **pregnant or breastfeeding**, or if you suspect you may **be pregnant** or are **planning to become pregnant**, consult your doctor before **using Vyjuvek**.



- If you are **pregnant or breastfeeding**, or if you suspect you are pregnant or intend to become pregnant, consult your doctor before using Vyjuvek.
- 
- If you are **pregnant, you must not come into contact with Vyjuvek** – not even with skin or dressings that have come into contact with Vyjuvek. Do not administer Vyjuvek and do not assist in these activities.



# Use of Vyjuvek in a doctor's office/clinic or at home

After consulting a doctor, it will be determined where and by whom treatment with Vyjuvek will normally be carried out. The application may



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



**at home by healthcare professionals,**



**at home by you or a caregiver** after consultation with a doctor and training.



## Use of Vyjuvek in a doctor's office/clinic

If Vyjuvek is applied in a doctor's office or clinic by healthcare professionals, the appropriate wound dressings will also be applied there. Please note the following:

- The dressing should remain **on the wound for approximately 24 hours.**
- Avoid **touching** the wound during this time.
- How to change **the dressing** for the first time **after application** (including disposal)  
Please refer to the section "The first dressing change after applying Vyjuvek."



## Use of Vyjuvek at home by healthcare professionals

If, after consulting a doctor, it has been determined that healthcare professionals may use Vyjuvek in a home setting, the following steps must be taken together with your doctor **before the first application**:

- **Medical information and advice** for the patient and (if applicable) caregiver on the use of Vyjuvek in the home environment
- **Medical training for persons** who use Vyjuvek or assist in its use
- Preparation of **a treatment plan** by the attending physician (including determination of the **order** in which the wounds are to be treated and **dosage**)
- Handing over and discussing this **guide**



## Application of Vyjuvek at home by you or a caregiver

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

- **Medical information and advice** for the patient and/or caregiver on the use of Vyjuvek at home
- **Medical training for persons** who use Vyjuvek or assist in its use. The application of Vyjuvek must be practiced under medical supervision until it is safely mastered.  
until it is mastered safely. A practice kit that does not contain any active ingredient is available for practical training.
- Preparation of **a treatment plan** (including determination of **the order** in which the wounds are to be treated and **the dosage**)
- Handing out and discussing this **guide**



**Vyjuvek may only be used at home after these steps have been completed!**

# Use of Vyjuvek



As this medicine contains genetically modified organisms, persons applying Vyjuvek or assisting with its application must wear **protective equipment** (e.g., gown, disposable gloves, mask, and eye protection).

## Video "Applying Vyjuvek" and demo kit



A **video on** how to **apply** Vyjuvek is available by scanning the QR code or at <http://ema.krystallabel.com/>.

You can request a **demo kit** with syringes (without active pharmaceutical ingredients) to practice applying Vyjuvek yourself. Order it at:

### Genesis Pharma

Email:

[medinfo@genesishpharmagroup.com](mailto:medinfo@genesishpharmagroup.com)



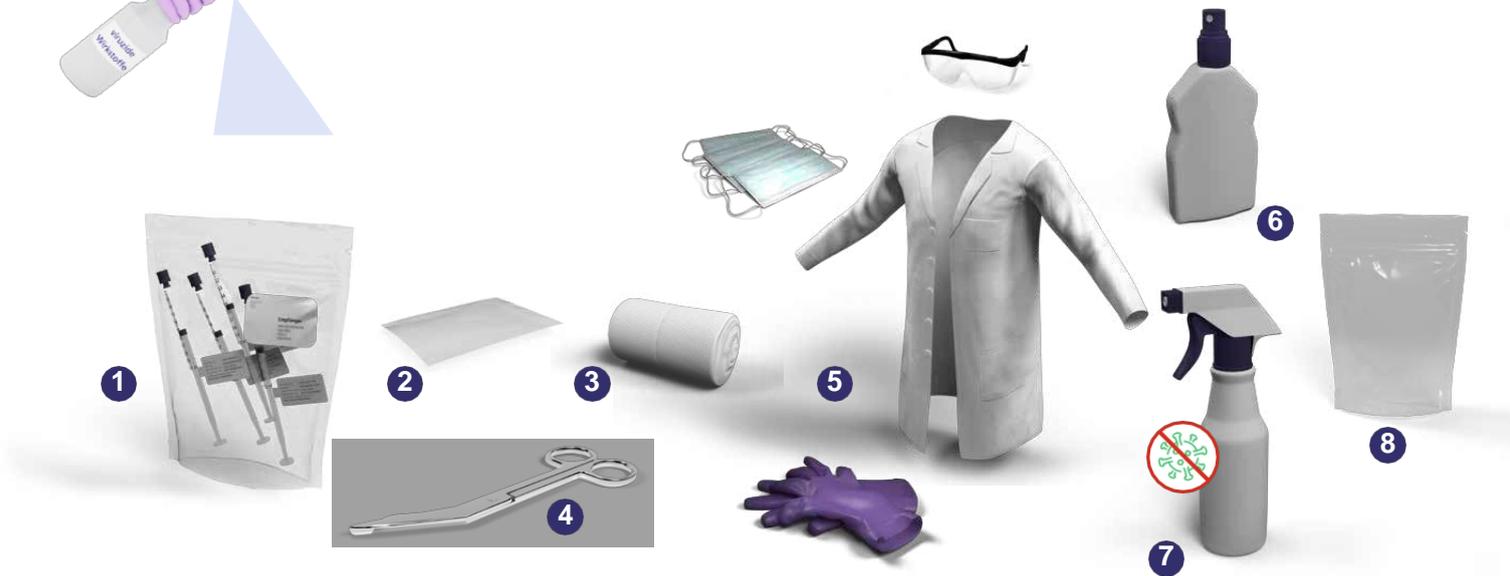
**Practicing with the demo kit does not replace the necessary medical training!**

# Step 1: Preparation

## Step 1.1: Preparing the work area



- Clean the work area with a **virucidal disinfectant** and prepare the necessary materials:



- 1 **Syringes** filled with Vyjuvek
- 2 **Non-absorbent, water-repellent** dressing (which does not absorb Vyjuvek) that is slightly larger than the selected wound
- 3 **Standard wound dressing** slightly larger than the non-absorbent, water-repellent dressing
- 4 **Scissors**
- 5 **Protective equipment** (e.g., gown, disposable gloves, mask, and eye protection)
- 6 **Wound cleanser** (**without** virucidal agents) **Disinfectant**
- 7 (**with** virucidal agents) **Sealable plastic bag** (for disposal)
- 8

Please **note** that you will need **two different cleaning/disinfectant agents**

:



**For the wound: Wound cleanser without virucidal agents**



**For surfaces and materials that have come into contact with Vyjuvek:**

**Disinfectant with virucidal agents**



- Persons applying Vyjuvek or assisting in its application should wear **protective equipment**.



- Check
  - that the number of syringes corresponds to your dosage (1 syringe contains 0.5 ml of Vyjuvek).
  - the **expiry date**. Do not use the medicine after the expiry date.
  - the **color of the medicine**. It may vary from opalescent yellow to colorless. Do not use the medicine if you notice any other discoloration.

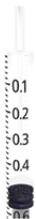
In these cases, inform your doctor or pharmacist.

### Step 1.2: Preparing the selected wound



- Carefully remove dead skin, scabs, and wound fluid, as well as any medicines and ointments from the wound area, and gently clean the wound. Use a **wound cleanser (without** virucidal agents, e.g., sterile saline solution). If you are unsure, ask your doctor.

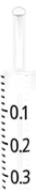
### Step 1.3: Preparing the syringe



- Hold the syringe so that the **tip** is pointing **upwards**.



- **Pull the syringe plunger** back slightly (but not completely out of the syringe).



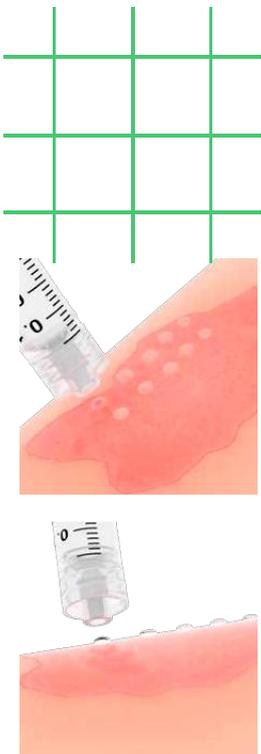
- Then **slowly** push the syringe plunger **back into the syringe** until a small drop of Vyjuvek emerges from the tip of the syringe.

## Step 2: Applying Vyjuvek



### Please note:

- Apply Vyjuvek **only to the wound to be treated**.
- Apply Vyjuvek to a wound until it is **completely closed**. Only then should you treat a new wound (according to the treatment plan/as agreed with your medical team).
- If a wound that has already been treated **reopens**, continue to treat it **first**.
- Do not apply **any other medicines** to the wound **at the same time** as Vyjuvek.



- Imagine **a grid with 1 × 1 cm squares** covering the selected wound (1 cm is approximately the width of the tip of your little finger).
- Now apply **a drop of Vyjuvek to each of these (imaginary) fields**.
- The tip of the syringe must **not touch** the skin!

The **amount** of Vyjuvek you need may vary depending on the size of the wound. It is possible that Vyjuvek will not be sufficient to treat all wounds or that you will have Vyjuvek left over after treating the wounds specified in the treatment plan. In these cases, inform your medical team or pharmacy and dispose of any remaining Vyjuvek (see section "Disposal of Vyjuvek residues and materials").

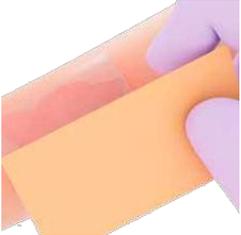
### Maximum dose per weekly application:

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

## Step 3: Dressing the wound



- Cut the **non-absorbent, water-repellent dressing** so that it is **slightly larger than the treated wound** and place it on the wound so that the edges of the dressing are completely **outside the wound area**.

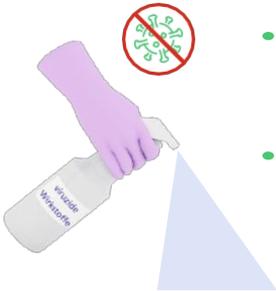


- Then cover **the above dressing completely with your standard dressing** (to prevent the non-absorbent, water-repellent dressing from slipping and transferring Vyjuvek to other areas, and to absorb any wound fluid).



- Leave the dressings on the wound **for about 24 hours** and avoid touching the wound during this time.

## Cleaning and disposal



- **Clean all surfaces** that have come into contact with Vyjuvek with a **virucidal disinfectant**.
- **Disinfect everything that has come into contact with Vyjuvek or could have come into contact with it** (e.g., bandages, used and unused syringes, used materials) with a **virucidal disinfectant**.



- Place the previously disinfected, used materials in a **sealable plastic bag**. Seal the bag and dispose of it in **the household waste**.

Vyjuvek has been tested for **sterility**. However, transmission of infectious agents cannot be completely ruled out. After use, watch for **signs of infection such as fever**. In this case, inform your doctor.  
doctor.



# The first dressing change after using Vyjuvek

- Wear **protective equipment** (e.g., gown, disposable gloves, mask, and eye protection).
- Remove the dressings as usual.
- Disinfect the removed, used dressings with a **virucidal disinfectant**.
- After disinfection, dispose of the used dressings in a separate, sealable plastic bag in the household waste.

After changing the Vyjuvek dressing, you can carry out your usual wound care routine.

## What to do if you accidentally come into contact with Vyjuvek

- Avoid direct contact of Vyjuvek with **skin areas other than the treated wound**.
- **Avoid direct contact** with the treated wound/dressing **for approximately 24 hours** after application.
- The person administering 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 mucous membranes** (e.g., nose, mouth), rinse the affected area **with clean water for at least 5 minutes**.



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



If **surfaces** have accidentally come into contact with Vyjuvek, clean them with a **virucidal disinfectant**.

# Transport of Vyjuvek

- When you collect the Vyjuvek syringes from the pharmacy, transport them in the **container** provided by **the pharmacy**, as it meets the special transport requirements for the medicine (e.g., cooling temperature).
- If the pharmacy delivers the syringes to your home, they will be delivered in a plastic bag in a suitable **insulated container (outer container)**, which should only be **opened** by the **person responsible for administering the medication**.
- Upon receipt, this **person** must ensure that the **outer container** is **intact** and shows no signs of leakage. If this is the case, please inform your pharmacy immediately.

# Storage of Vyjuvek

- Leave the Vyjuvek syringes in the **outer container**.
- Store the syringes in a **clean** place **out of the reach of children**, in accordance with your pharmacy's instructions.
- Use Vyjuvek **within the period specified by your pharmacy**.

# Further information



All training materials on Vyjuvek for patients and caregivers (guide for patients and caregivers, video "Applying Vyjuvek") and the package insert are also available online by scanning the QR code or at <http://ema.krystallabel.com>. Printed copies can be ordered from Genesis Pharma (see contact details below).

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

## Attending physician

Name .....

Address .....

Phone .....

**GENESIS Pharma Bulgaria EOOD**

**Interpred- World Trade Center**

**Dragan Tsankov Blvd., Floor 7, Office 702**

**Sofia 1040**

**Tel: [+359 2 969 3227](tel:+35929693227)**

**Email: [Safety.Bulgaria@genesispharmagroup.com](mailto:Safety.Bulgaria@genesispharmagroup.com)**

REFERENCE: Vyjuvek, current package insert.

# Reported by Side effects

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

If you notice any side effects, talk to your doctor or pharmacist, or report them directly to:

**Bulgarian Drug Agency (BDA)**  
8 Damyan Gruev Str.,  
Sofia 1303, BULGARIA  
phone: +359 2 8903 417  
fax: +359 2 8903 434 e-mail: [bda@bda.bg](mailto:bda@bda.bg)

[www.bda.bg/](http://www.bda.bg/) Form for reporting adverse drug reactions by non-medical persons

or

**GENESIS Pharma Bulgaria EOOD**  
Interpred- World Trade Center  
Dragan Tsankov Blvd., Floor 7, Office 702  
Sofia 1040  
Tel: [+359 2 969 3227](tel:+35929693227)  
Email: [Safety.Bulgaria@genesispharmagroup.com](mailto:Safety.Bulgaria@genesispharmagroup.com)





**Krystal Biotech Germany GmbH**

c/o Mindspace

Friedrichstr. 68

10117 Berlin

Tel.:

**Email:** [info-germany@krystalbio.com](mailto:info-germany@krystalbio.com)

