

PATIENT PACKAGE INSERT IN ACCORDANCE WITH THE PHARMACISTS' REGULATIONS (PREPARATIONS) – 1986

The medicine is dispensed with a doctor's prescription only

ARIKAYCE® LIPOSOMAL 590 mg, Suspension for inhalation for use with a nebuliser

Active ingredient and its quantity per dosage unit

Each vial contains:
590 mg liposomal amikacin (as sulfate)

The mean delivered dose per vial is approximately 312 mg of amikacin. Inactive and allergenic ingredients in the preparation: see section 6 "Further Information".

Read the leaflet carefully in its entirety before using the medicine. This leaflet contains concise information about the medicine. If you have further questions, refer to the doctor or pharmacist. This medicine has been prescribed for the treatment of your ailment. Do not pass it on to others. It may harm them, even if it seems to you that their ailment is similar.

In addition to the leaflet, there is a Patient Safety Information Card for **Arikayce Liposomal 590 mg** preparation. This card contains important safety information that you must know and adhere to before starting and during the course of treatment with **Arikayce Liposomal 590 mg**. Read the Patient Safety Information Card and the patient leaflet before starting to use the preparation. Keep the card for further reading, if necessary.

1. WHAT IS THE MEDICINE INTENDED FOR?

For treatment of an infectious lung inflammation, nontuberculous mycobacteria (NTM), caused by *Mycobacterium avium* Complex (MAC) in adults without cystic fibrosis, with limited treatment options.

Therapeutic group: systemic aminoglycoside antibiotic.

Arikayce Liposomal 590 mg is an **antibiotic** that contains the active ingredient amikacin, from the group of antibiotics called aminoglycosides, which stop the growth of certain bacteria that cause infection.

Arikayce Liposomal 590 mg is delivered by inhalation.

2. BEFORE USING THE MEDICINE

Do not use the medicine if:

- You are sensitive (allergic) to the active ingredient or to other aminoglycosides, to soya, or to any of the additional ingredients contained in the medicine (see the list of inactive ingredients in section 6 "Further Information").
- You are taking other aminoglycosides (oral or injection).
- You have very poor kidney function.

Special warnings regarding use of the medicine

Before treatment with Arikayce Liposomal 590 mg, tell the doctor if:

- You use a bronchodilator for breathing problems, as you will be asked to use that first, before using Arikayce Liposomal 590 mg;
- You have **kidney problems**; you may need to have a kidney test before starting treatment;
- You have **hearing difficulties, ringing or buzzing in the ears** (tinnitus) or **balance problems** including spinning sensation, lack of coordinated muscle movements or dizziness. You may have to have a hearing test before starting or during treatment if you have any hearing problems;
- You suffer from **other lung diseases**;
- You have a disease that causes muscle weakness and fatigue, such as **myasthenia gravis**.

Tell your doctor immediately if whilst using Arikayce Liposomal 590 mg, any of the following occur to you:

- Loss of consciousness, skin rash, fever, worsening or new breathing problems;
- Worsening of kidney problems;
- Ear problems, such as ringing in the ears or loss of hearing.

See section 4 "Side effects".

Children and adolescents

This medicine is not intended for children and adolescents under the age of 18. There is no information regarding the safety and effectiveness of this medicine in children and adolescents under the age of 18.

Drug interactions

If you are taking, or have recently taken, other medicines, including non-prescription medicines and nutritional supplements, tell the doctor or pharmacist. Especially if you are taking:

- Diuretics such as ethacrynic acid, furosemide or mannitol.
- Other medicines that may affect the kidneys, hearing, balance or reduce muscle strength.

Pregnancy, breastfeeding and fertility

If you are pregnant or breastfeeding, think you are pregnant or are planning to become pregnant, avoid using Arikayce Liposomal 590 mg. Consult with a doctor or pharmacist before using this medicine.

If you become pregnant while using Arikayce Liposomal 590 mg, inform the doctor, and he will advise whether to stop using Arikayce Liposomal 590 mg.

It is not known if amikacin passes into breast milk in humans. If you are breastfeeding, your doctor will advise you whether to stop breastfeeding or stop treatment with this medicine.

Driving and operating machinery

Arikayce Liposomal 590 mg may cause dizziness and other vestibular disturbances, such as lightheadedness (vertigo) and balance disturbances. Driving or operating machinery is not recommended during the course of treatment with Arikayce Liposomal 590 mg. If you have questions, talk to your doctor.

3. HOW SHOULD YOU USE THE MEDICINE?

Always use the medicine according to the doctor's instructions.

Check with the doctor or pharmacist if you are uncertain about the dosage and treatment regimen of the medicine.

The dosage and treatment regimen will be determined by the doctor only. The usual dosage is generally:

One vial of Arikayce Liposomal 590 mg inhaled in your mouth once a day, via the Lamira Nebuliser system.

After 6 months of treatment, the doctor will advise whether to continue or to stop treatment.

Do not exceed the recommended dose.

Duration of treatment

The maximum duration of treatment is 18 months.

Using Arikayce Liposomal 590 mg

If you use a bronchodilator, use that first, before using Arikayce Liposomal 590 mg.

Each vial is for **single use only**.

- Use Arikayce Liposomal 590 mg **only** with the Lamira Nebuliser Handset and Lamira Aerosol Head connected to a Lamira Control Unit. See section 7 "Instructions for Use" – how to use the medicine together with the Lamira Nebuliser System.
- **Do not use** Arikayce Liposomal 590 mg with any other type of handset or aerosol head.
- **Do not put** other medicines in the Lamira Nebuliser Handset.
- **Do not drink** the liquid in the vial.
- **Read the instructions for use**, which are provided at the end of this leaflet.

How and when do you replace the Lamira Nebuliser Handset?

One Lamira Nebuliser Handset should be used for one 28-day treatment course. The aerosol head should be replaced weekly. There are 4 aerosol heads provided in each Arikayce Liposomal 590 mg carton package.

Please read the manufacturer's instructions for use for information regarding cleaning and storage.

If you accidentally take a higher dosage

Consult a doctor immediately if you are concerned that you may have taken too much of this medicine.

If you took an overdose or if a child has accidentally swallowed the medicine, refer immediately to a doctor or proceed to an hospital emergency room and bring the package of the medicine with you.

If you forget to take the medicine

If you forget to take the medicine, take it as soon as possible on the day of the missed dose. Do not take more than one dose on the same day to compensate for a forgotten dose.

Adhere to the treatment regimen as recommended by the doctor.

If you stop using the medicine

Consult a doctor if you decide to stop using Arikayce Liposomal 590 mg for any reason.

Do not take medicines in the dark! Check the label and the dose each time you take medicine. Wear glasses if you need them.

If you have further questions regarding use of the medicine, consult the doctor or pharmacist.

4. SIDE EFFECTS

As with any medicine, use of Arikayce Liposomal 590 mg may cause side effects in some users. Do not be alarmed when reading the list of side effects. You may not suffer from any of them.

Refer to a doctor immediately if:

- You experience hypersensitivity or severe allergic reactions when taking Arikayce Liposomal 590 mg (e.g., with low blood pressure, loss of consciousness, severe skin rash or severe wheezing and breathlessness). The frequency of these side effects is not known.
- You experience worsening of your usual lung problems or new breathing problems (e.g., breathlessness or wheezing). This may be a sign of severe inflammation in the lungs that requires treatment and may mean that you will have to stop taking Arikayce Liposomal 590 mg. These severe side effects are common to very common.

Other side effects

Very common side effects (effects that occur in more than 1 in 10 users)

- Difficulty in speaking
- Difficulty in breathing
- Cough
- Coughing up blood

Common side effects (effects that occur in 1-10 users in 100)

- Infection causing worsening of your lung condition
- Increase in mucus coughed up from lungs
- Chesty cough
- Wheezing
- Throat irritation
- Sore throat
- Loss of voice
- A fungal infection in the mouth
- Pain in the mouth
- Change in sense of taste
- Lung inflammation
- Headache
- Dizziness
- Feeling unsteady
- Diarrhea
- Nausea
- Vomiting
- Dry mouth
- Decrease in appetite
- Itching of the skin
- Deafness
- Ringing in your ears
- Kidney problems including impaired kidney function
- Joint pain
- Muscle pain
- Rash
- Tiredness
- Discomfort in chest
- Fever
- Loss of weight

Uncommon side effects (effects that occur in 1-10 in 1,000 users)

- Anxiety

If a side effect occurs, if one of the side effects worsens, or if you suffer from a side effect not mentioned in the leaflet, consult with the doctor.

Reporting side effects

Side effects can be reported to the Ministry of Health by clicking on the link "Report Side Effects of Drug Treatment" found on the Ministry of Health homepage (www.health.gov.il) that directs you to the online form for reporting side effects, or by entering the link: <https://sideeffects.health.gov.il>

5. HOW SHOULD THE MEDICINE BE STORED?

- Avoid poisoning! This medicine and any other medicine should be kept in a safe place out of the reach and sight of children and/or infants in order to avoid poisoning. Do not induce vomiting unless explicitly instructed to do so by the doctor.
- Do not use the medicine after the expiry date (exp. date) that appears on the package and on the label of the vial. The expiry date refers to the last day of that month.

Storage conditions:

Store in a refrigerator (2°C-8°C).

Do not freeze. Dispose of any vial that has frozen.

If the dose you want to use is in the refrigerator, take the vial out of the refrigerator and allow it to reach room temperature before using it.

Alternatively, Arikayce Liposomal 590 mg can be stored at room temperature, below 25°C, but only for up to 4 weeks. When storing at room temperature, any unused preparation should be disposed of after 4 weeks.

This medicine is a milky white liquid in a clear vial. Do not use the medicine if you notice a change in color or small particles floating in the vial.

Do not dispose of medicines via the wastewater or household waste bin. Ask the pharmacist how to dispose of medicines you no longer need. These measures will help protect the environment.

6. FURTHER INFORMATION

In addition to the active ingredient, the medicine also contains:

dipalmitoylphosphatidylcholine (DPPC), cholesterol high purity (HP), sodium chloride, sodium hydroxide and water for injection.

What the medicine looks like and the contents of the package:

Arikayce Liposomal 590 mg is a milky white to off-white suspension for inhalation, provided in a glass 10 ml vial with a rubber stopper and metal seal, with a removable cover.

The carton package contains 28 vials which are sufficient for 28 days; one vial per day.

One carton package of Arikayce Liposomal 590 mg contains 4 inner carton packages, with 7 vials in each and one aerosol head. In addition, the package contains one Lamira Nebuliser Handset.

Registration holder and address:

Truemed Ltd., 10 Beni Gaon St., Poleg Industrial Park, P.O. Box 8105, Netanya.

Manufacturer and address:

Insmed Netherlands BV, Utrecht, Netherlands.

This leaflet was checked and approved by the Ministry of Health in May 2022.

Registration number of the medicine in the National Drug Registry of the Ministry of Health:
169-59-36844

7. INSTRUCTIONS FOR USE

Arikayce Liposomal 590 mg is intended for oral inhalation use with the Lamira Nebuliser System.

Before you use your Lamira Nebuliser System, be sure you read and understand the detailed information in the full Instructions for Use that come with the Lamira Nebuliser System. These will provide more complete information about how to assemble, prepare, use, clean, and disinfect the nebuliser.

Wash your hands with soap and water and dry them well.

Assemble the handset including the connection to the controller as illustrated in the full Instructions for Use.

Preparing the medicine for use:

1. It is recommended to take the vial out of the refrigerator at least 45 minutes before use to allow it to reach room temperature. Do not use other medicines in the Lamira Nebuliser Handset.
2. Shake the Arikayce Liposomal 590 mg vial vigorously, until the medicine looks the same throughout and is well mixed.
3. Lift the orange cap from vial and put it aside (Figure 1).

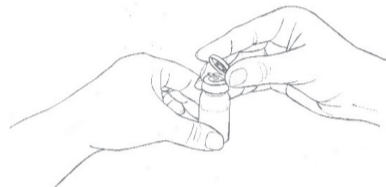


Figure 1

4. Grip the metal ring on top of the vial and pull it down gently until one side breaks away from the vial (Figure 2).



Figure 2

5. Pull the metal band from around the vial top in a circular motion until it comes off completely from the vial (Figure 3).

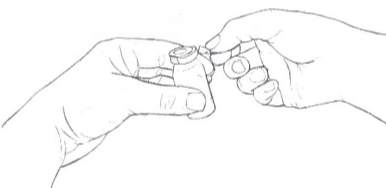


Figure 3

6. Put aside the metal ring after it is detached. Carefully remove the rubber stopper (Figure 4).



Figure 4

7. Pour the contents of the Arikayce Liposomal 590 mg vial into the medication reservoir of the Lamira Nebuliser Handset (Figure 5).



Figure 5

8. Close the medication reservoir (Figure 6).



Figure 6

9. Sit in a relaxed, upright position. This makes inhaling easier and helps the medicine get into your lungs.
10. Insert the mouthpiece and take slow, deep breaths. Then, breathe normally in and out through the mouthpiece until the treatment is complete. The treatment should take about 14 minutes but could take up to 20 minutes. Be sure to hold the handset level throughout the treatment (Figure 7).

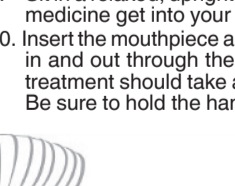


Figure 7