

PATIENT PACKAGE INSERT IN ACCORDANCE WITH
THE PHARMACISTS' REGULATIONS (PREPARATIONS) - 1986
The dispensing of this medicine requires a doctor's prescription
Read the package insert carefully in its entirety
before using this medicine

TOBI® Solution for Inhalation Solution for inhalation

COMPOSITION:
Each ampoule of 5 ml contains:
Tobramycin 300 mg

Inactive ingredients:
Sodium chloride, water for injections, sodium hydroxide, sulfuric acid.

THERAPEUTIC GROUP: Aminoglycoside antibiotics.
Each ampoule (5 ml) contains 11.25 mg sodium

THERAPEUTIC ACTIVITY:

For the management of cystic fibrosis patients with *Pseudomonas aeruginosa*.

Tobi Solution for Inhalation contains antibiotic called tobramycin.

Tobi Solution for Inhalation is a solution to be inhaled with a specific nebulizer and a suitable compressor; the solution is contained in a ready-to-use ampoule.

When **Tobi Solution for Inhalation** is inhaled, the antibiotic can get directly to the lungs to fight against the bacteria causing the infection, and to improve your breathing.

Pseudomonas aeruginosa is a very common bacterium that infects the lungs of nearly everyone with cystic fibrosis at some time during their lives. Some people do not get this infection until later on in their lives, while others get it very young. It is one of the most damaging bacteria for people with cystic fibrosis. If the infection is not properly fought, it continues to damage the lungs causing further breathing problems.

WHEN SHOULD THE PREPARATION NOT BE USED?

Do not use this medicine if you are breastfeeding.
Do not use this medicine if you are allergic (hypersensitive) to tobramycin, to any of its ingredients or to any other aminoglycoside antibiotic.
If you think you may be allergic, ask the doctor for advice.

Do not take this medicine without consulting a doctor before starting treatment:

Inform the doctor if you have or if you have ever had hearing problems (including noises in the ears and dizziness); kidney problems; unusual difficulty in breathing with wheezing or coughing, chest tightness; muscle weakness that lasts or becomes worse in time, a symptom mostly related to conditions such as myasthenia or Parkinson's disease.

If you are pregnant. It is not known whether inhaling this medicine during pregnancy cause side effects. If you want to become pregnant or are pregnant, you should talk with the doctor about the possibility that this medicine will cause any harm to you or to the unborn child.

When given by injection, tobramycin and other aminoglycoside antibiotics can cause harm to the unborn child, such as deafness.

HOW WILL THIS MEDICINE AFFECT YOUR EVERYDAY LIFE?

Tobi Solution for Inhalation should not affect the ability to drive and use machines.

WARNINGS:

Inhaling medicines can cause chest tightness and wheezing and this can happen immediately after inhalation with **Tobi Solution for Inhalation**.

If you are taking tobramycin or another aminoglycoside antibiotic by injection, it can sometimes cause hearing loss, dizziness and kidney damage, and can harm an unborn child.

In patients 65 years and older the doctor may perform additional tests to decide if **Tobi Solution for Inhalation** is right for them.

If you are sensitive to any type of food or medicine, inform your doctor before commencing treatment with this medicine.

DRUG INTERACTIONS:

If you are taking another drug concomitantly, including non-prescription drugs and nutritional supplements, or if you have just finished treatment with another medicine, inform the attending doctor, in order to prevent hazards or lack of efficacy arising from drug interactions. This is especially important for medicines from the following groups: furosemide or ethacrynic acid, a diuretic; urea or mannitol; other medicines which may harm the nervous system, kidneys or hearing.

Tobi Solution for Inhalation should not be diluted or mixed with other medications in the nebulizer.

SIDE EFFECTS:

In addition to the desired effect of the medicine, adverse reactions may occur during the course of taking this medicine.

Very common side effects (may affect more than 1 in 10 patients): runny or stuffy nose, sneezing; voice alteration (hoarseness); discoloration of the substance you cough up; decreased results for the tests of lung function. If any of these affects you severely, **tell the doctor**.

Common side effects (may affect between 1 and 10 in every 100 patients): generally feeling unwell; muscle pain; voice alteration (hoarseness) with sore throat and difficulty swallowing (laryngitis).
Other side effects: itching; itchy rash; rash; loss of voice; disturbed sense of taste; sore throat.

If any of these affects you severely, **tell the doctor**.

Side effects that require special attention:

if you have any of the following side effects, stop taking **Tobi Solution for Inhalation** and **tell the doctor straight away**: unusual difficulty in breathing with wheezing or coughing and chest tightness; allergic reactions including hives and itching.

If you have any of the following side effects, **tell the doctor straight away**: worsening of your lung disease (very common); decrease in hearing (ringing in the ears is a potential warning sign of hearing loss), noises (such as hissing) in the ears.

In the event that you experience side effects not mentioned in this leaflet, or if there is a change in your general health, consult your doctor immediately.

DOSAGE:

Use **Tobi Solution for Inhalation** exactly according to the doctor's instructions. Check with the doctor if you are not sure.

The dose is the same for all patients aged 6 years old and older: the content of two ampoules per day (one in the morning and one in the evening) to be inhaled with the PARI LC PLUS nebulizer and a suitable compressor.

Taking **Tobi Solution for Inhalation** at the same time each day will help you remember when to take the medicine.

- The content of one ampoule in the morning to be inhaled with the PARI LC PLUS nebulizer.
- The content of one ampoule in the evening to be inhaled with the PARI LC PLUS nebulizer.

You must stay as close as possible to a 12-hour time difference between the dose of the morning and the dose of the evening.

After taking the medicine for 28 days, you then have a 28-day break, where you don't inhale any **Tobi Solution for Inhalation**, before starting another course. It is important that you keep using the product twice each day during the 28 days on treatment and that you keep to the 28-day on and 28-day off cycle.

If you forget to take **Tobi Solution for Inhalation** and there are at least 6 hours to the next dose, take the dose as soon as you can. Otherwise, wait for the next dose. Do not double the dose to make up for the missed dose.

Continue taking **Tobi Solution for Inhalation** as the doctor instructed. If you have questions about how long to take **Tobi Solution for Inhalation**, talk to the doctor or pharmacist.

If you are taking several different inhaled treatments and performing therapies for cystic fibrosis, **Tobi Solution for Inhalation** should be taken after all other treatments are done. The order of medicines should be checked with the doctor.

This medicine is not usually intended for administration to infants and children under 6 years of age.

DIRECTIONS FOR USE:

See the instructions for the use of **Tobi Solution for Inhalation**.

Follow the doctor's instructions carefully.
Do not exceed the recommended dose.

HOW CAN YOU CONTRIBUTE TO THE SUCCESS OF THE TREATMENT?

Complete the full course of treatment as instructed by the doctor.
Even if there is an improvement in your health, do not discontinue use of this medicine, without consulting your doctor.

AVOID POISONING!

This medicine, and all other medicines, must be stored in a safe place out of the reach and sight of children and/or infants, in order to avoid poisoning.

If you inhaled an overdose, inform the doctor as soon as possible.
If **Tobi Solution for Inhalation** is swallowed, do not worry but tell the doctor as soon as possible.

Do not induce vomiting unless explicitly instructed to do so by a doctor!
This medicine has been prescribed for the treatment of your ailment; in another patient it may cause harm. **Do not give this medicine to your relatives, neighbors or acquaintances.**

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

STORAGE:

Do not use **Tobi Solution for Inhalation** if the solution is cloudy or if there are particles in it.

Store at 2- 8°C (in a refrigerator). Do not freeze.
The foil pouches can be kept (opened or unopened) at room temperature (up to 25°C) for up to 28 days. Do not use **Tobi Solution for Inhalation** ampoules which have been stored at room temperature for more than 28 days.

Store the ampoules in the original pack to protect from light and moisture.

Tobi Solution for Inhalation is normally a slightly yellow color but this can vary and sometimes it can be a darker yellow. This does not change the way **Tobi Solution for Inhalation** works provided that the storage instructions have been followed.

Never store an opened ampoule. Once opened, the ampoule should be used immediately.

Even if kept in their original container and stored as recommended, medicines may be kept for a limited period only. Please note the expiry date of the medicine!

In case of doubt, consult the pharmacist who dispensed the medicine to you. Do not store different medications in the same package.

LICENSE NUMBER: 133 76 29825 00

MANUFACTURER: Catalent Pharma Solutions, Woodstock, Illinois, USA for Novartis Pharma AG, Basel, Switzerland.

REGISTRATION HOLDER: Novartis Israel Ltd., 36 Shacham St., Petach-Tikva.

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Instructions for the use of Tobi Solution for Inhalation

Please read carefully and follow these instructions

The equipment needed for inhaling Tobi Solution for Inhalation should be used with a clean and dry PARI LC PLUS reusable nebulizer and a suitable compressor. The doctor or physiotherapist can advise you on the proper use of Tobi Solution for Inhalation and the equipment needed. You may need different nebulizers for the other inhaled medicines for cystic fibrosis.

Preparing Tobi Solution for Inhalation for Inhalation

- Wash your hands thoroughly with soap and water and fully dry hands.
- Each Tobi Solution for Inhalation foil pouch contains four ampoules. Cut or tear open the pouch. Remove one Tobi Solution for Inhalation ampoule by gently pulling apart from the rest of the ampoules attached at the bottom tabs (Figure 1).
- Put the other ampoules back in the foil pouch and keep it in the refrigerator.
- Lay out all the nebulizer pieces: nebulizer top (a), nebulizer bottom (b), inspiratory valve cap (c), mouthpiece with valve (d), and tubing (e) on a clean dry paper or cloth towel (Figure 2).
- Make sure you have the suitable compressor and tubing to connect the nebulizer and compressor.
- Be careful to follow the appropriate instructions for use for your type of nebulizer, you must read the leaflet provided with the nebulizer by the manufacturer. Check that the nebulizer and compressor are working properly according to the manufacturer's instructions before you start to take the medicine.

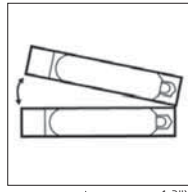


Figure 1 ציור 1 רשמ 1

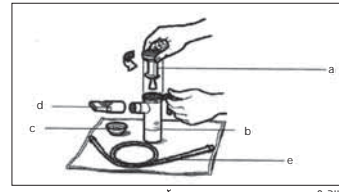


Figure 2 ציור 2 רשמ 2

Use of Tobi Solution for Inhalation with PARI LC PLUS nebulizer

For more detailed instructions on the use and care of the nebulizer, please read the leaflet provided with the PARI LC PLUS.

- Remove the nebulizer top from the nebulizer bottom by twisting the top anticlockwise and lifting it. Place the nebulizer top on the towel and stand the nebulizer bottom upright on the towel.
- Connect one end of the tubing to the compressor air outlet. Make sure that the tubing is well fitted. Plug in the compressor into the electrical outlet.
- Open the Tobi Solution for Inhalation ampoule by holding the bottom tab with one hand and twisting off the top of the ampoule with the other hand (Figure 3).
- Squeeze all the contents of the ampoule into the nebulizer bottom (Figure 4).
- Replace the nebulizer top (a), put the mouthpiece (d) and the inspiratory valve cap (c) in place on the nebulizer and then connect the compressor as indicated in the PARI LC PLUS nebulizer leaflet (Figure 5).
- Turn on the compressor. Check that there is a steady mist coming from the mouthpiece. If there is no mist, check all tubing connections and that the compressor is working properly.
- Sit or stand in an upright position so that you can breathe normally.
- Place the mouthpiece between the teeth and on top of the tongue. Breathe normally, but only through the mouth (you may use a nose clip if your doctor agrees). Try not to block the airflow with the tongue (Figure 6).
- Continue until all of the Tobi Solution for Inhalation is gone and there is no longer any mist being produced. It should take about 15 minutes to take all the treatment. You may hear splattering sound when the nebulizer bottom is empty.
- Remember to clean and disinfect the nebulizer after treatment according to the manufacturer's instructions. Never use a dirty or a clogged nebulizer. You should not share your nebulizer with other people.

If you are interrupted or need to cough or rest during treatment, turn off the compressor to save the medicine. Turn the compressor on again when you are ready to restart treatment. Leave out this dose if the next dose is due in less than 6 hours.

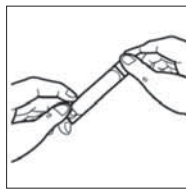


Figure 3 ציור 3 רשמ 3



Figure 4 ציור 4 רשמ 4

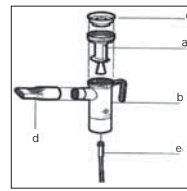


Figure 5 ציור 5 רשמ 5

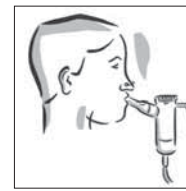


Figure 6 ציור 6 רשמ 6

إرشادات استعمال توبي محلول للإستنشاق

يجب قراءة التعليمات بإمعان والعمل وفقاً.

التجهيزات المطلوبة لإستنشاق توبي محلول للإستنشاق يجب استعمال توبي محلول للإستنشاق بواسطة نيبوليزر PARI LC PLUS متعدد الاستعمال بحيث يكون نظيفاً وجافاً مع ضغط هواء مناسب. يمكن للطبيب أو المعالج الفيزيائي أن يقدم لك المشورة حول الإستعمال الصحيح بتوبي محلول للإستنشاق كذلك التجهيزات المطلوبة. من الجائز أن تحتاج /ي أجهزة نيبوليزر أخرى لأدوية التثيف الخويصلي الأخرى المتناولة عن طريق الإستنشاق.

تحضير توبي محلول للإستنشاق

- يجب غسل اليدين بشكل جيد بالصابون والماء، وتجفيفهما تماماً.
- كل كيس الألمنيوم من توبي محلول للإستنشاق يحتوي على 4 أمبولات. يجب قص أو تمزيق الكيس.
- يجب إخراج الأمبولة واحدة من توبي محلول للإستنشاق عن طريق فصلها بملف من باقي الأمبولات المتصقة في قسمها السفلي (الرسم رقم 1).
- يجب إعادة الأمبولات المتبقية إلى داخل كيس الألمنيوم وحفظها في البراد.
- يجب تجهيز كافة أقسام النيبوليزر: نيبوليزر علوي (a)، نيبوليزر سفلي (b)، غطاء صمام الإستنشاق (c)، فوهة مع صمام (d)، وأنبوية (e) على منديل من الورق أو القماش النظيف والجاف (رسم 2).
- يجب التأكد من أن لديك ضغط هواء مناسب، وأنبوية من أجل وصلها للنيبوليزر والضاغط.
- يجب الإنتباه إلى متابعة تعليمات الإستعمال المناسبة لنوع النيبوليزر خاصتكم، عليك قراءة النشرة المرفقة مع النيبوليزر من قبل المنتج.
- يجب التأكد بأن النيبوليزر والضاغط الهوائي يعملان كالمطلوب حسب تعليمات المنتج قبل البدء بإستعمال الدواء.

في حال كنت بحاجة إلى مزيد من المعلومات، يرجى قراءة النشرة المرفقة مع النيبوليزر.

הוראות שימוש בטובי תמיסה לשיפה

יש לקרוא בעיון ולפעול בהתאם להוראות.

הציוד הדרוש לשיפה טובי תמיסה לשיפה עם נבולייזר PARI LC PLUS רב פעמי שהיה נקי ויבש ולמרות מתיאם. הרופא או הפיזיותרפיסט יכולים ליעץ לך לגבי השימוש הנכון בטובי תמיסה לשיפה וציוד הדרוש. ייתכן כי תצטרף/י נבולייזר אחרים לתרופות האחרות לאיטסיק פיברוזיס הגלוחות בשיפה.

הכנת טובי תמיסה לשיפה

- יש לרחוץ את הידיים בסבון עם מים ולבוב אותן לחלוטין.
- כל שקית אלומיניום של טובי תמיסה לשיפה מכילה ארבע אמפולות. יש לתפור או לקרוע את השקית. יש להוציא אמפולה אחת של טובי תמיסה לשיפה על-ידי הפרדה עדינה משאר האמפולות הצמודות בחלקו התחתון (ציור 1).
- יש לשים את האמפולות הנותרות בחזרה בתוך שקית האלומיניום ולשמור במקרר.
- יש להכין את כל חלקי הנבולייזר: נבולייזר עליון (a), נבולייזר תחתון (b), מכסה שסתום השאיפה (c), פיה עם שסתום (d), וצינורית (e) על מגבת נקי או בד נקי ויבש (ציור 2).
- יש לוודא כי יש לך מרחס מתיאם, וצינורית שתתחבר לנבולייזר ולמרחס.
- שים/י לב כי היקר עוקבת/ת אחר הוראות השימוש המתאימות של סוג הנבולייזר שלך. עליך לקרוא את העלון המסופק עם הנבולייזר על-ידי היצרן. יש לבדוק כי הנבולייזר והמרחס עובדים כדורש בהתאם להוראות היצרן לפני שהיך מתחיל/ת לקחת את התרופה.

במידת הצורך, ניתן להתייעץ עם הרופא או הפיזיותרפיסט.

שימוש בטובי תמיסה לשיפה עם נבולייזר PARI LC PLUS

להוראות נוספות מפורטות לגבי השימוש והטיפול בנבולייזר, אנא קרא/י את העלון המצורף ל- PARI LC PLUS.

- יש להוציא את הנבולייזר העליון מהנבולייזר התחתון על-ידי סיבוב של החלק העליון בנגד כיוון השעון והרמתו. יש לשים את הנבולייזר העליון על המגבת ולהעמיד את הנבולייזר התחתון במצב אנכי על המגבת.
- יש לחבר קצה אחד של הצינורית לפתח האוויר של המרחס. יש לוודא שהצינורית מותאמת בצורה טובה. יש לחבר את המרחס לשקע החשמל.
- יש לפתוח את אמפולת טובי תמיסה לשיפה על-ידי אחיזה על תחתית האמפולה ביד אחת וסיבוב החלק העליון של האמפולה ביד השנייה (ציור 3).
- יש לסחוט את כל תכולת האמפולה לתוך הנבולייזר התחתון (ציור 4).
- יש להחזיר את החלק העליון של הנבולייזר (a), לשים את הפיה (d) ואת מכסה שסתום השאיפה (c) במקום על הנבולייזר ואז לחבר את המרחס כפי שצויין בעלון של הנבולייזר PARI LC PLUS (ציור 5).
- יש להדליק את המרחס. יש לבדוק שעלום אדים מופיעה באופן קבוע, אם אין אדים, יש לבדוק את כל חיבורי הצינורית והמרחס פעול כהלכה.
- יש לשבת או לעמוד במצב מאונך כך שתוכלו/י לנשום כרגיל.
- יש לשים את הפיה בין השיניים ומעל הלשון. יש לנשום כרגיל אבל רק דרך הפה ואפשר להשתמש באטם לאף אם הרופא שלך מסכים. יש לנסות שלא לנחום את מעבר האוויר עם הלשון (ציור 6).
- יש להמשיך עד שלח טובי תמיסה לשיפה נגמרת ולא נוצרים יותר אדים. לקחת כל הטיפול אמורה לקחת כ- 15 דקות. ייתכן כי תשמע/י קולות חזקה כאשר נבולייזר התחתון ריק.
- יש לזכור לקנות ולחסא את הנבולייזר לאחר השימוש בהתאם להוראות היצרן. לעולם אין להשתמש בנבולייזר מלוכלך או חסום. אין לחלוק את הנבולייזר שלך עם אנשים אחרים.

אם הלה הפרעה או אם עליך להשתעל או לנוח במהלך הטיפול, יש לכבות את המרחס כדי לשמור את התרופה. יש להדליק שוב את המרחס כאשר היקר מוכן/ה להתחיל שוב בטיפול. יש לוודא על המנה חוה אם ישנאו פחות מ- 6 שעות ללקחת המנה הבאה.