

PATIENT MEDICATION INFORMATION

READ THIS FOR SAFE AND EFFECTIVE USE OF YOUR MEDICINE

BREZTRI™ AEROSPHERE®

budesonide / glycopyrronium / formoterol fumarate dihydrate pressurized inhalation suspension

Read this carefully before you start taking **BREZTRI™ AEROSPHERE®** and each time you get a refill. This leaflet is a summary and will not tell you everything about this drug. Talk to your healthcare professional about your medical condition and treatment and ask if there is any new information about **BREZTRI AEROSPHERE**.

What is BREZTRI AEROSPHERE used for?

- BREZTRI AEROSPHERE is used in adults for the long-term treatment of a lung disease called chronic obstructive pulmonary disease (COPD), including chronic bronchitis and emphysema.
- BREZTRI AEROSPHERE is used to make breathing easier and help prevent worsening of COPD symptoms (“flare ups”). You may experience “flare-ups” during which your symptoms become worse. If you have a history of experiencing “flare-ups” BREZTRI AEROSPHERE can help prevent them from developing in the first place.

BREZTRI AEROSPHERE is not meant to treat asthma, or to provide immediate relief of an asthma attack. Use a fast acting ‘rescue’ inhaler for any sudden asthma attacks. If you do not have a rescue inhaler, ask your doctor to prescribe one for you.

How does BREZTRI AEROSPHERE work?

BREZTRI AEROSPHERE contains 3 medicinal ingredients:

- budesonide is an inhaled corticosteroid (ICS).
- glycopyrronium is a long-acting muscarinic antagonist (LAMA).
- formoterol fumarate dihydrate is a long-acting beta₂ agonist (LABA).

Budesonide belongs to a group of medicines called corticosteroids. It reduces inflammation in the airways of your lungs, which can ease breathing problems, and help prevent “flare-ups” of your COPD. Glycopyrronium and formoterol fumarate dihydrate belong to a group of medicines called bronchodilators. These two medicines work together to help open and relax the muscles of the airways. This allows more air to get in and out of the lungs. This makes it easier for people with COPD to breathe and helps prevent shortness of breath and wheezing. BREZTRI AEROSPHERE is not a cure for COPD, but it can help to control it. Therefore, it is important to continue to take it regularly, even if you feel fine.

What are the ingredients in BREZTRI AEROSPHERE?

Medicinal ingredients: budesonide, glycopyrronium (as bromide), and formoterol fumarate dihydrate

Non-medicinal ingredients: hydrofluoroalkane (HFA-134a), porous particles (comprised of: 1,2-distearoyl-sn-glycero-3-phosphocholine (DSPC) and calcium chloride).

BREZTRI AEROSPHERE comes in the following dosage forms:

Pressurized inhalation suspension in a metered dose inhaler (for oral inhalation): 182 mcg

budesonide / 8.2 mcg glycopyrronium (as bromide) / 5.8 mcg formoterol fumarate dihydrate per actuation.

Do not use BREZTRI AEROSPHERE:

- if you are allergic to budesonide, glycopyrronium, formoterol fumarate dihydrate or any of the other ingredients of BREZTRI AEROSPHERE;
- to treat sudden symptoms of COPD, such as shortness of breath or wheezing. Always have a rescue inhaler with you to treat sudden symptoms (“flare ups”). If you do not have a rescue inhaler, ask your healthcare professional to prescribe one for you;
- to treat asthma.

To help avoid side effects and ensure proper use, talk to your healthcare professional before you take BREZTRI AEROSPHERE. Talk about any health conditions or problems you may have, including if you:

- are a smoker. It is important to quit smoking if you use BREZTRI AEROSPHERE, this will help decrease the symptoms of COPD and may increase your lifespan;
- are using a rescue inhaler on a regular basis;
- have heart problems, such as:
 - heart disease
 - rapid or irregular heart beat or any problems with how your heart beats
 - a condition called “QT prolongation”
- have high blood pressure;
- have eye problems such as increased pressure in the eye, glaucoma, cataracts, blurry vision or other changes in vision;
- have prostate or bladder problems, or any problems passing urine;
- used to take 20 mg or more of prednisone per day;
- have or are at risk for decreased bone mineral content;
- have ever had seizures;
- have problems with your thyroid gland;
- have diabetes;
- have ever had to stop taking another medication for your breathing problems because you were allergic to it or it caused problems;
- have been taking other corticosteroids by mouth or by inhalation;
- have any allergies to food or drugs;
- have or have a history of lung infections such as pneumonia;
- have low levels of potassium in your blood;
- have kidney or liver problems;
- have or have ever had tuberculosis;
- have chronic or untreated infections:
 - bacterial infection
 - viral infection
 - fungal infection (yeast infection or thrush)
 - parasitic infection
 - herpes simplex infection of the eye
- are pregnant or trying to become pregnant. Your healthcare professional will consider the benefit to you and the risk to your unborn baby;
- are breastfeeding or planning to breastfeed.

Other warnings you should know about:

Do not stop taking BREZTRI AEROSPHERE without talking to your doctor first. If you stop

treatment on your own, your symptoms may worsen. Talk to your doctor **right away** if:

- there is a change in your symptoms such as more coughing, wheezing, chest tightness or breathlessness;
- you find that you need to use your rescue inhaler more often than usual.

These could be warning signs that your condition may be getting worse.

Paradoxical bronchospasm: if you feel tightness of the chest, coughing, wheezing or breathlessness right after using BREZTRI AEROSPHERE, you may have a serious condition called “paradoxical bronchospasm” (an unexpected closing of your airways). Stop using BREZTRI AEROSPHERE and seek medical help right away.

Monitoring: During treatment with BREZTRI AEROSPHERE, your doctor may monitor:

- your bones and eyes;
- the effects of corticosteroid therapy on your body;
- your blood sugar levels, especially if you have diabetes;
- your blood potassium levels, especially if you already have low levels of potassium.

Eye problems: BREZTRI AEROSPHERE can cause eye disorders such as:

- cataracts: clouding of the lens in the eye, blurry vision, eye pain;
- glaucoma: an increased pressure in your eyes, eye pain, halos around lights or coloured images, red eyes. Untreated, it may lead to permanent vision loss.

If you have any changes in your vision, tell your healthcare professional **right away**. You may need regular eye exams.

Chicken pox and measles: You should avoid exposure to chicken pox and measles, and tell your doctor if you are exposed. This is important if you are taking any cortisone-type medicine and your immune system is not working well (if you have difficulty in fighting an infection).

Eosinophilic granulomatosis with polyangiitis (EGPA): Some people experience a flu-like illness called EGPA when taking an inhaled corticosteroid. Tell your healthcare professional if you have:

- a rash,
- pins and needles or numbness of your arms or legs,
- a severe sinus infection and
- worsening lung or breathing problems.

Driving and using machines: BREZTRI AEROSPHERE can cause headaches, or nausea which may affect your ability to drive and use machines. Use caution when driving and using machines until you know how BREZTRI AEROSPHERE affects you.

Tell your healthcare professional about all the medicines you take, including any drugs, vitamins, minerals, natural supplements or alternative medicines.

The following may interact with BREZTRI AEROSPHERE:

- medicines called beta-blockers (such as atenolol or propranolol), to treat high blood pressure or other heart problems, or to treat glaucoma (such as timolol);
- ketoconazole or itraconazole, to treat fungal infections;
- ritonavir, nelfinavir or cobicistat, to treat HIV infection;
- oral corticosteroids such as prednisone;

- medicines that lower the amount of potassium in your blood, such as some diuretics (“water pills”);
- other long-acting medicines similar to this medicine that are used to treat breathing problems, such as medicines containing tiotropium, ipratropium, aclidinium, umeclidinium or salmeterol, arformoterol, vilanterol, olodaterol or indacaterol;
- medicines which are used to treat heart rhythm problems, such as amiodarone;
- medicines used in the treatment of depression (antidepressants or monoamine oxidase inhibitors);
- medicines called “anticholinergics”.

Ask your healthcare professional if you are not sure. The use of BREZTRI AEROSPHERE with these medicines is not recommended.

How to take BREZTRI AEROSPHERE:

- Read the Instructions for Use at the end of this Patient Medication Information for instructions on how to use BREZTRI AEROSPHERE. If you have any questions, talk to your healthcare professional.
- This medicine has been prescribed for you personally and you should not pass it on to others. It may harm them, even if their symptoms are the same as yours.
- Take BREZTRI AEROSPHERE exactly as your healthcare professional tells you to.
- Take every day as prescribed, even if you have no symptoms of COPD.
- After you take your dose, always rinse your mouth with water and spit it out, do not swallow.

Usual dose:

The recommended dose is 2 oral inhalations (puffs) taken twice daily in the morning and evening.

Overdose:

Signs of an overdosage may include:

- blurred vision;
- dry mouth;
- nausea;
- muscle spasm;
- tremor;
- headache;
- increased heart rate; and
- increased blood pressure.

If you think you, or a person you are caring for, have taken too much BREZTRI AEROSPHERE, contact a healthcare professional, hospital emergency department or regional poison control centre immediately, even if there are no symptoms.

Missed Dose:

Do not take a double dose to make up for a forgotten dose. If you miss a dose, take it as soon as you remember. However, if it is nearly time for your next dose, skip the missed dose and go back to your regular dosing schedule. Do not take more than 2 puffs twice a day.

What are possible side effects from using BREZTRI AEROSPHERE?

These are not all the possible side effects you may have when taking BREZTRI AEROSPHERE. If you experience any side effects not listed here, tell your healthcare professional.

- feeling anxious;
- trouble sleeping;
- headache;
- feeling sick (nausea);
- cough and/or a hoarse voice;
- muscle spasms;
- shaking/tremors;
- dry mouth;
- mild irritation in the throat;
- bruising of the skin;
- feeling restless, nervous or agitated;
- feeling dizzy;
- changes in behaviour.

Serious side effects and what to do about them			
Symptom / effect	Talk to your healthcare professional		Stop taking drug and get immediate medical help
	Only if severe	In all cases	
COMMON			
Hyperglycemia (high blood sugar): increased thirst, frequent urination, headache, blurred vision and fatigue		✓	
Palpitation (fast-beating, fluttering or pounding heart): skipping beats, beating too fast, pounding, fluttering rapidly		✓	
Pneumonia (infection in the lungs): chest pain when you breath or cough, confusion, cough which may produce phlegm, fatigue, fever, sweating and shaking, chills, shortness of breath		✓	
Thrush (yeast infection of the mouth or throat): thick white patches in the mouth, tongue or on the throat, sore throat		✓	
Urinary tract infection (infection in urinary system including kidneys, ureters, bladder and urethra): pain or burning sensation while urinating, frequent urination,		✓	

Serious side effects and what to do about them			
Symptom / effect	Talk to your healthcare professional		Stop taking drug and get immediate medical help
	Only if severe	In all cases	
blood in urine, pain in the pelvis, strong smelling urine, cloudy urine			
UNCOMMON			
Allergic Reaction: difficulty swallowing or breathing, wheezing, feeling weak or light headed, feeling sick to your stomach and throwing up, hives or rash, swelling of the face, lips, tongue or throat			✓
Angina (not enough oxygen to the heart muscle): discomfort in the shoulder, arm, back, throat, jaw or teeth; pain or pressure in the chest		✓	
Arrhythmia (abnormal heart rhythms): rapid, slow or irregular heartbeat		✓	
Bronchospasm (when there is a sudden narrowing of the airway): difficulty breathing with wheezing or coughing			✓
Depression (sad mood that won't go away): difficulty sleeping or sleeping too much, changes in appetite or weight, feelings of worthlessness, guilt, regret, helplessness or hopelessness, withdrawal from social situations, family, gatherings and activities with friends, reduced sex drive and thoughts of death or suicide. If you have a history of depression, your depression may become worse		✓	
Tachycardia (abnormally fast heartbeat): dizziness, light headedness, shortness of breath, racing heart		✓	
Urinary retention (inability to pass urine or to empty the bladder): painful urination, pain or swelling in the lower abdomen		✓	

Serious side effects and what to do about them			
Symptom / effect	Talk to your healthcare professional		Stop taking drug and get immediate medical help
	Only if severe	In all cases	
VERY RARE			
Decreased Adrenal Function: tiredness, weakness, nausea and vomiting, low blood pressure	✓		
UNKNOWN			
Cataracts: clouding of the lens in the eye, blurry vision, dim vision and/or eye pain		✓	
Glaucoma: increased pressure in your eyes, eye and head pain, swelling or redness in or around the eye, changes in vision, hazy or blurred vision, sudden sight loss		✓	

If you have a troublesome symptom or side effect that is not listed here or becomes bad enough to interfere with your daily activities, tell your healthcare professional.

<p>Reporting Side Effects</p> <p>You can report any suspected side effects associated with the use of health products to Health Canada by:</p> <ul style="list-style-type: none"> • Visiting the Web page on Adverse Reaction Reporting (https://www.canada.ca/en/health-canada/services/drugs-health-products/medeffect-canada/adverse-reaction-reporting.html) for information on how to report online, by mail or by fax; or • Calling toll-free at 1-866-234-2345. <p><i>NOTE: Contact your health professional if you need information about how to manage your side effects. The Canada Vigilance Program does not provide medical advice.</i></p>
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Storage:

Store between 15 to 30°C. Store in a dry place.

Warning: The canister should not be broken, punctured or burnt, even when it seems empty. Do not use or store near heat or open flames. Do not expose to temperatures higher than 50°C and protect from freezing temperatures. If exposed to freezing temperatures, the inhaler should be equilibrated for 1 hour at room temperature prior to taking a dose.

Use by date: Do not use BREZTRI AEROSPHERE after the expiry date on the label/carton. The expiry date refers to the last day of the month.

After opening the pouch, the inhaler must be used within:

- 6 weeks for an inhaler that contains 56 actuations;
- 3 months for an inhaler than contains 120 actuations.

Dispose of your inhaler after the last dose, following local guidelines. Ask your pharmacist how to dispose of medicines no longer required. These measures will help protect the environment.

Keep out of reach and sight of children.

If you want more information about BREZTRI AEROSPHERE:

- Talk to your healthcare professional.
- Find the full product monograph that is prepared for healthcare professionals and includes this Patient Medication Information by visiting the Health Canada website (<https://www.canada.ca/en/health-canada/services/drugs-health-products/drug-products/drug-product-database.html>); the manufacturer's website www.astrazeneca.ca, or by calling 1-800-668-6000.
- This Patient Medication Information is current at the time of printing. The most up-to-date version can be found at www.astrazeneca.ca.

This leaflet was prepared by AstraZeneca Canada Inc. Mississauga, Ontario L4Y 1M4

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Last Revised: September 30, 2021



INSTRUCTIONS FOR USE

BREZTRI™ AEROSPHERE®

budesonide / glycopyrronium / formoterol fumarate dihydrate pressurized inhalation suspension

Read this Instructions for Use before you start using BREZTRI AEROSPHERE and each time you get a new pack. There may be new information. This information does not take the place of talking to your healthcare professional about your medical condition or treatment.

Important Information:

- **For oral inhalation use only.**
- Use BREZTRI AEROSPHERE exactly as your doctor tells you to.
- If you have any questions about the use of your inhaler, ask your doctor or pharmacist.
- **Clean your inhaler once each week.** It is very important to keep the plastic actuator clean so that medicine will not build-up and block the spray through the mouthpiece. See Steps 1 through 8 under “**How to clean your BREZTRI AEROSPHERE inhaler**”.

Parts of your BREZTRI AEROSPHERE inhaler (see Figure 1):

- BREZTRI AEROSPHERE comes as a canister that fits into an actuator with a dose indicator.
 - **Do not** use the BREZTRI AEROSPHERE actuator with a canister of medicine from any other inhaler.
 - **Do not** use the BREZTRI AEROSPHERE canister with an actuator from any other inhaler.

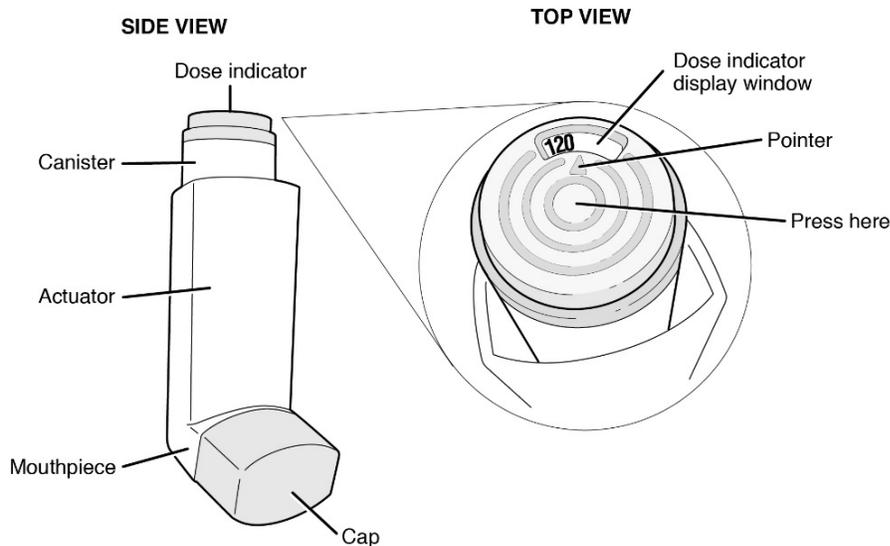


Figure 1

- BREZTRI AEROSPHERE comes with a dose indicator located on the top of the canister (see Figure 1). The dose indicator display window will show you how many puffs of medicine you have left. A puff of medicine is released each time you press the centre of the dose indicator.

Before you use BREZTRI AEROSPHERE for the first time make sure that the pointer on the dose indicator is pointing to the right of the “120” inhalation mark in the dose indicator display window (**see Figure 1**). (Note: The pointer will point to the right of the “60” inhalation mark if you have a 14-day inhaler, 56 inhalation canister.)

- The pointer will be pointing to 120 after 10 puffs are delivered from BREZTRI AEROSPHERE. This means that there are 120 puffs of medicine left in the canister (**see Figure 2a**).
- The pointer will be pointing between 100 and 120 after you take 10 more puffs. This means that there are 110 puffs of medicine left in the canister (**see Figure 2b**).
- The pointer will be pointing to 100 after you take 10 more puffs. This means that there are 100 puffs of medicine left in the canister (**see Figure 2c**).



Figure 2a
120 puffs



Figure 2b
110 puffs

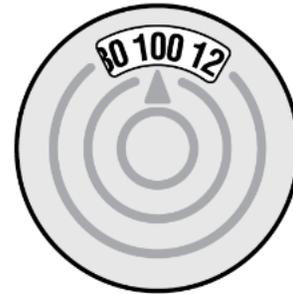


Figure 2c
100 puffs

- The dose indicator display window will continue to move after every 10 puffs. The number in the dose indicator display window will continue to change after every 20 puffs.

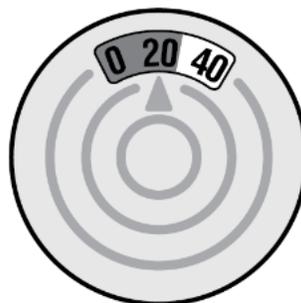


Figure 2d

- The colour in the dose indicator display window will change to red, as shown in the shaded area, when there are only 20 puffs of medicine left in your inhaler (**see Figure 2d**).
- The dose indicator for the 14-day inhaler, 56 inhalation canister, moves after every 10 puffs; with markings for 60, 40, 20 and 0 puffs. The colour in the 14-day inhaler, 56 inhalation canister dose indicator display window will change to red when there are only 20 puffs of medicine left in your inhaler.

Preparing your BREZTRI AEROSPHERE inhaler for use:

- Your BREZTRI AEROSPHERE inhaler comes in a foil pouch that contains a drying packet (desiccant).
 - Take the BREZTRI AEROSPHERE inhaler out of the foil pouch.

- Throw away the pouch and the drying packet. Do not eat or breathe in the contents of the drying packet.
- BREZTRI AEROSPHERE should be at room temperature before you use it.
- Store BREZTRI AEROSPHERE in a dry place.

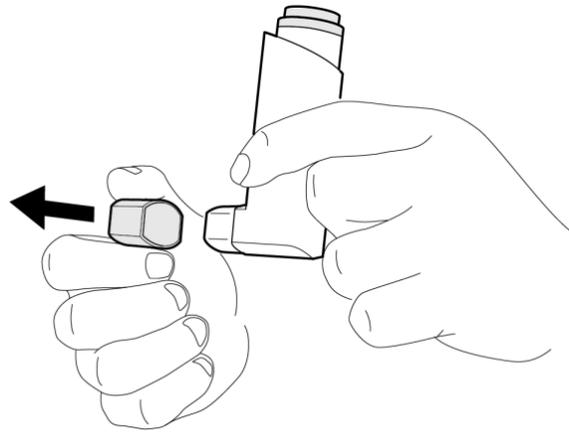


Figure 3

Priming your BREZTRI AEROSPHERE inhaler:

Before you use BREZTRI AEROSPHERE for the first time, you must prime the inhaler.

- Remove the cap from the mouthpiece (see **Figure 3**). Check inside the mouthpiece for objects before use.
- Hold the inhaler in the upright position away from your face and shake the inhaler well (see **Figure 4**).

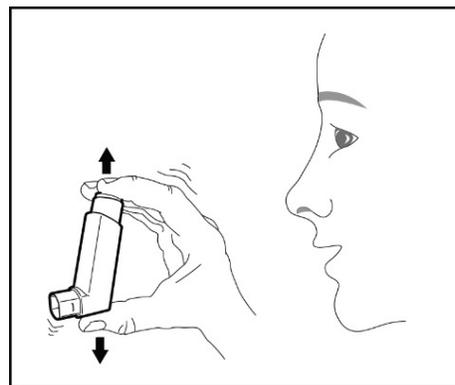


Figure 4

- Press down firmly on the centre of the dose indicator until the canister stops moving in the actuator, to release a puff of medicine from the mouthpiece (see **Figure 5**). You may hear a soft click from the dose indicator as it counts down during use.

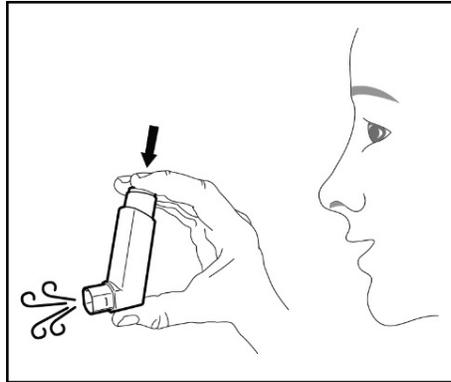


Figure 5

- **Repeat the priming steps 3 more times (see Figure 4 and Figure 5).** Shake the inhaler well before each priming puff.
- After priming 4 times, the dose indicator should be pointing to the right of “120” and your inhaler is now ready to use.

Using your BREZTRI AEROSPHERE inhaler:

Step 1: Remove the cap from the mouthpiece (see Figure 6).

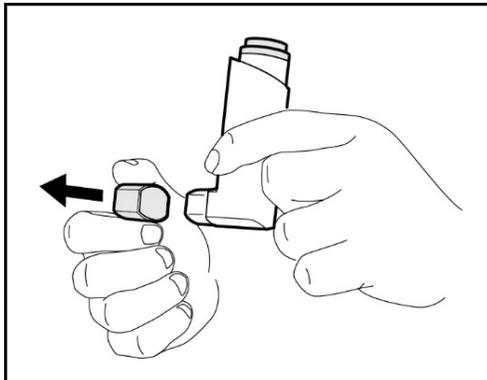


Figure 6

Step 2: Shake the inhaler well before each use (see Figure 7).

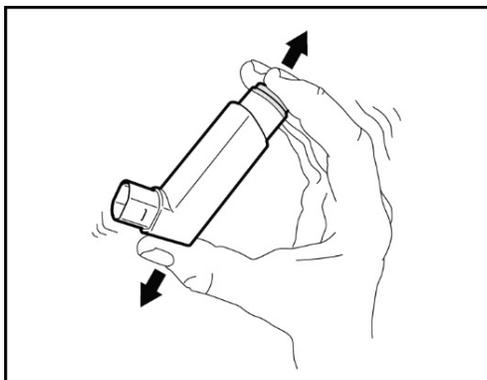


Figure 7

Step 3: Hold the inhaler with the mouthpiece pointing towards you and breathe out as fully as you comfortably can through your mouth (see **Figure 8**).

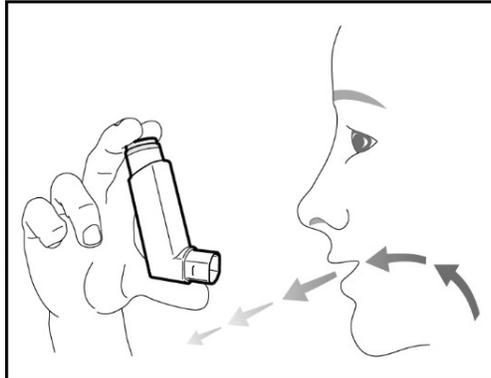


Figure 8

Step 4: Close your lips around the mouthpiece and tilt your head back, keeping your tongue below the mouthpiece (see **Figure 9**).

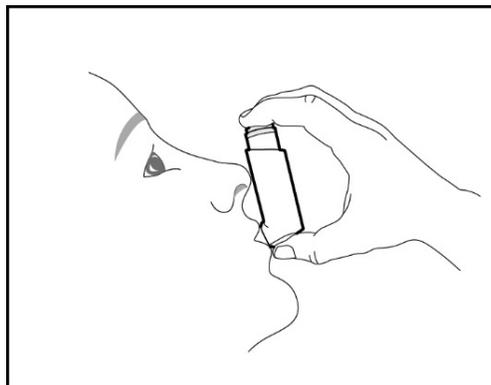


Figure 9

Step 5: While breathing in deeply and slowly, press down on the centre of the dose indicator until the canister stops moving in the actuator and a puff of medicine has been released (see **Figure 10**). Then stop pressing the dose indicator.

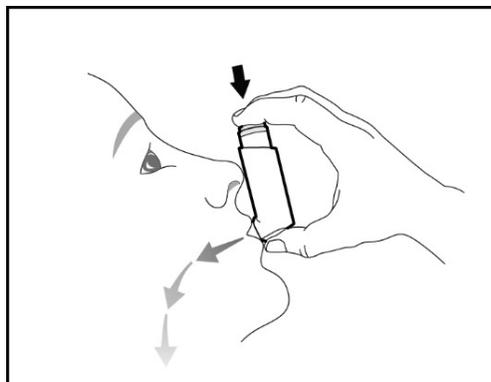


Figure 10

Step 6: When you have finished breathing in, remove the mouthpiece from your mouth.

Hold your breath as long as you comfortably can, up to 10 seconds (see **Figure 11**).

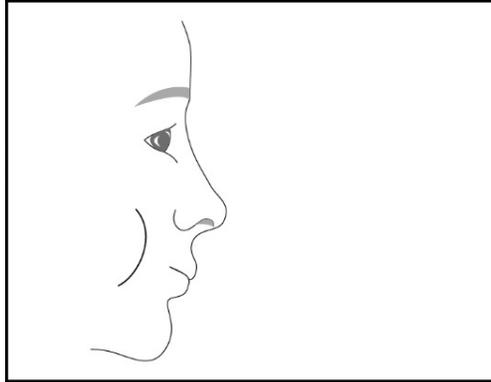


Figure 11

Step 7: Breathe out gently (see **Figure 12**). Repeat **Steps 2 through 7** to take your second puff of BREZTRI AEROSPHERE.

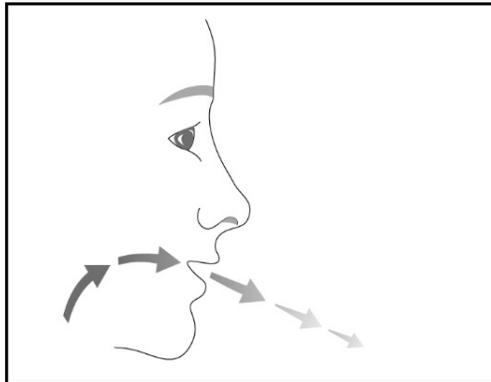


Figure 12

Step 8: Replace the cap over the mouthpiece right away after use (see **Figure 13**).

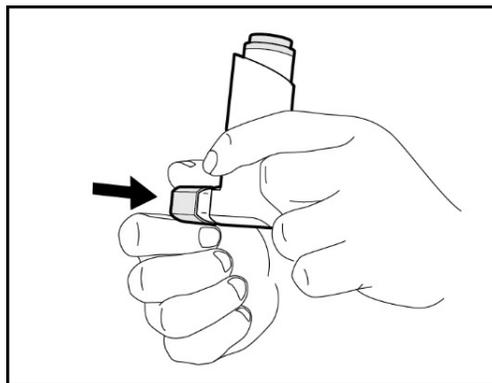


Figure 13

Step 9: Rinse your mouth with water to remove any excess medicine. Do not swallow.

How to clean your BREZTRI AEROSPHERE inhaler:

Clean the inhaler once each week. It is very important to keep your inhaler clean so that medicine will not build-up and block the spray through the mouthpiece (see **Figure 14**).

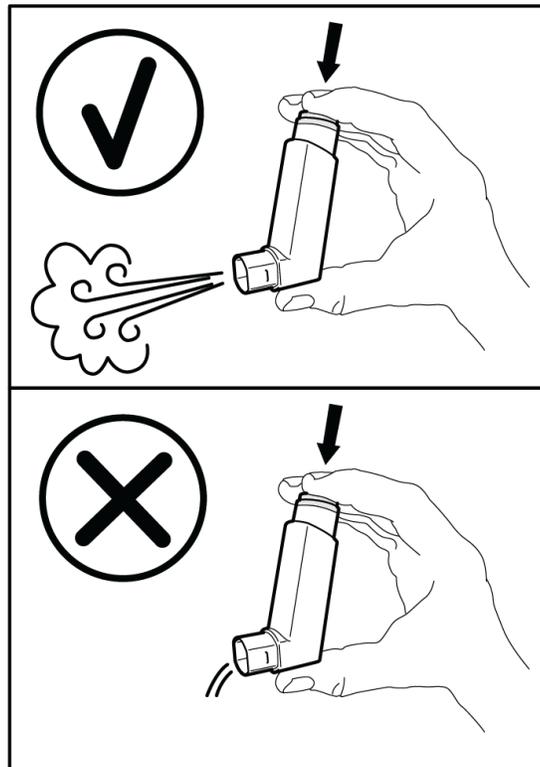


Figure 14

Step 1: Take the canister out of the actuator (see **Figure 15**). Do not clean the canister or let it get wet.

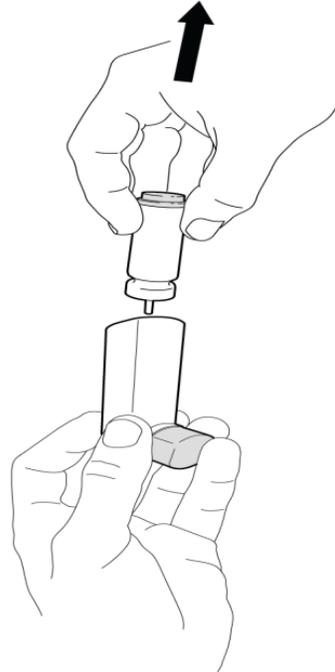


Figure 15

Step 2: Take the cap off the mouthpiece.

Step 3: Hold the actuator under the tap and run warm water through it for about 30 seconds. Turn the actuator upside down and run warm water through it again for about 30 seconds (see **Figure 16**).

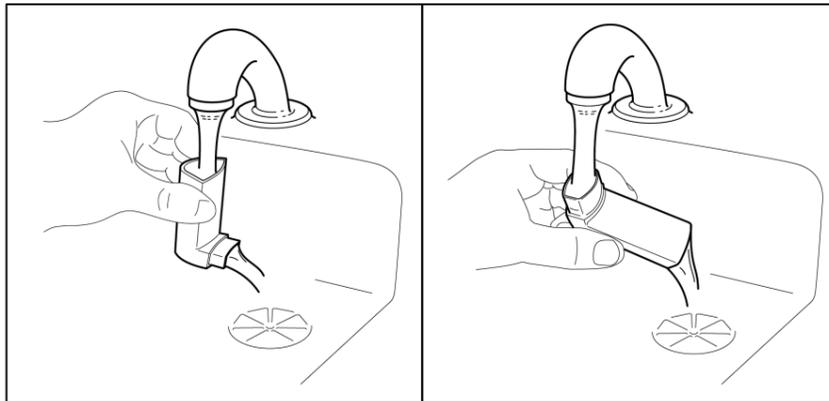


Figure 16

Step 4: Shake off as much water from the actuator as you can.

Step 5: Look into the actuator and the mouthpiece to make sure any medicine build-up has been completely washed away. If there is any build-up, repeat Steps 3 through 5 in the section “**How to clean your BREZTRI AEROSPHERE inhaler**”.

Step 6: Let the actuator air-dry overnight (see **Figure 17**). **Do not** put the canister back into the actuator if it is still wet.

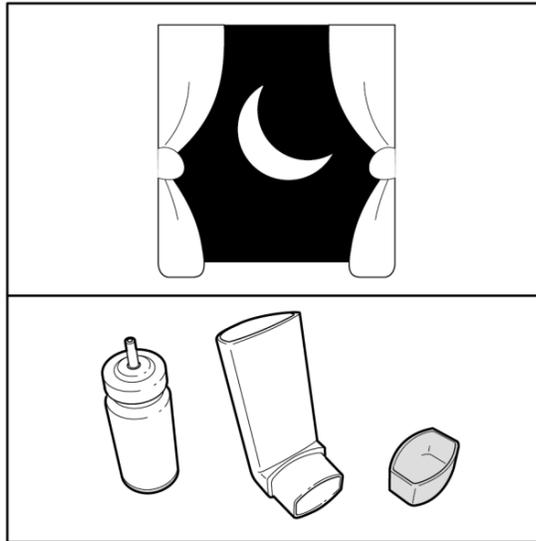


Figure 17

Step 7: When the actuator is dry, gently press the canister down in the actuator (see **Figure 18**). Do not press down too hard on the canister. This could cause a puff of medicine to be released.

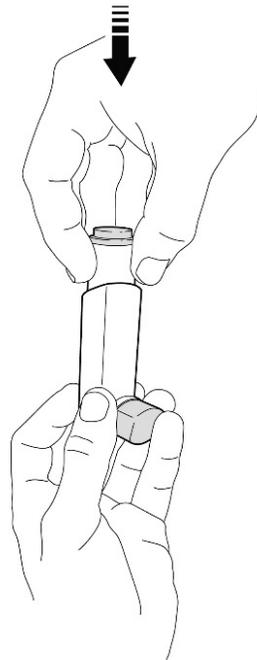


Figure 18

Step 8: Re-prime your **BREZTRI AEROSPHERE** inhaler after each cleaning. To re-prime the inhaler, shake the inhaler well and press down on the centre of the dose indicator 2 times to release a total of 2 puffs into the air away from your face. Your inhaler is now ready to use.

If you do not use your BREZTRI AEROSPHERE for more than 7 days, or your BREZTRI

AEROSPHERE is dropped, you will need to re-prime it before use.

To re-prime the inhaler, shake the inhaler well and press down on the centre of the dose indicator 2 times to release a total of 2 puffs into the air away from your face. Your inhaler is now ready to use.

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Last Revised: September 30, 2021

