

## READ THIS FOR SAFE AND EFFECTIVE USE OF YOUR MEDICINE

### PATIENT MEDICATION INFORMATION

#### **PrREVESTIVE®\*** **Teduglutide for injection**

Read this carefully before you start taking **REVESTIVE** 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 **REVESTIVE**.

#### **What is REVESTIVE used for?**

REVESTIVE is used to treat adults and pediatric patients 1 year of age and above with Short Bowel Syndrome. Short Bowel Syndrome is a disorder arising from an inability to absorb food nutrients and fluid across the gastrointestinal tract (gut) due to not enough surface area. It is often caused by surgical removal of all or part of the small intestine.

#### **How does REVESTIVE work?**

REVESTIVE improves the absorption of fluids from your remaining gut by increasing the surface area and function of your remaining gut.

#### **What are the ingredients in REVESTIVE?**

Medicinal ingredients: teduglutide

One vial of powder contains 5 mg of teduglutide. After reconstitution with 0.5 mL of sterile water for injection, a 10 mg/mL sterile solution is obtained.

Non-medicinal ingredients: dibasic sodium phosphate heptahydrate, L-histidine, mannitol, monobasic sodium phosphate monohydrate. The diluent contains sterile water for injection that is provided in a pre-filled syringe.

#### **REVESTIVE comes in the following dosage forms:**

REVESTIVE is a powder and diluent for solution for under the skin (subcutaneous) injection (5 mg powder in a vial and 0.5 mL diluent in a pre-filled syringe). Pack size of 28 each.

The powder is white and the diluent is clear and colorless.

#### **Do not use REVESTIVE if:**

- you are allergic to teduglutide or any of the other ingredients of this medicine;
  - you have or are suspected to have cancer in the gastrointestinal tract, including liver, gallbladder, bile ducts or pancreas;
  - you have had cancer in the gastrointestinal tract, including liver, gallbladder, bile ducts or pancreas, within the last five years.
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**To help avoid side effects and ensure proper use, talk to your healthcare professional before you take REVESTIVE. Talk about any health conditions or problems you may have, including if you:**

- have a history of bowel obstruction. Your doctor will consider this when prescribing this medicine.
- suffer from certain cardiovascular diseases (affecting the heart and/or blood vessels) such as high blood pressure (hypertension) or have a weak heart (cardiac insufficiency). The symptoms include sudden weight gain, swollen ankles and/or shortness of breath. This may require closer monitoring and more frequent changes to the amount of parenteral nutrition/IV fluid that you receive.
- have decreased kidney function. Your doctor may need to give you a lower dose of this medicine.
- have severely decreased liver function. Your doctor will consider this when prescribing this medicine.

### Medical check-ups before and during treatment with REVESTIVE

#### *Colonoscopy Schedule in Adults*

Before you start treatment with this medicine, your doctor will need to perform a colonoscopy (a procedure to see inside your colon and rectum) to check for the presence of polyps (small growths that may be normal or abnormal) and remove them. It is recommended that your doctor performs a follow-up colonoscopy between 1 to 2 years after starting treatment, and then at a minimum of 5 year intervals. If polyps are found either before or during your treatment with REVESTIVE, your doctor will decide whether you should continue using this medicine. REVESTIVE should not be used if a cancer is detected during your colonoscopy.

#### *Colonoscopy Schedule in Children (ages 1 through 17 years)*

Before your child starts treatment with this medicine, your doctor will need to test for any hidden blood in the stool. If there is sign of unexplained blood in the stool, a colonoscopy will be required (a procedure to see inside your child's colon, rectum and the large intestine). It is recommended that your child's doctor tests for hidden blood in the stool every year while your child is receiving REVESTIVE.

Colonoscopy is recommended for all children after 1 year of treatment, every 5 years thereafter while on continuous treatment with REVESTIVE, and if they have new or unexplained blood in the stool.

#### *Laboratory Testing*

Your doctor will take special care and monitor your small bowel function and monitor for signs and symptoms indicating problems with your gallbladder, liver and pancreas by conducting initial laboratory tests (bilirubin, alkaline phosphatase, lipase and amylase) within 6 months before starting treatment with REVESTIVE and every 6 months while you are using REVESTIVE. If there is significant increase in test results, more tests may be needed (for example, imaging of the biliary tract, liver, or pancreas).

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

REVESTIVE may affect how other medicines are absorbed from the gut and therefore how well they work. Your doctor may have to change your dose of other medicines.

**How to take REVESTIVE:**

Always use this medicine exactly as your doctor has told you. Check with your doctor, pharmacist or nurse if you are not sure.

REVESTIVE is injected under the skin (subcutaneously) once daily. The injection can be self-administered or given by another person, for example your doctor, his/her assistant or your home nurse. If you are injecting the medicine yourself, you must receive adequate training by your doctor or nurse. Do not use the same area each day for each injection. Rotate the site of injection between the upper, lower, left and right side of your belly, either upper arm or either thigh.

You will find detailed instructions for the preparation, reconstitution and administration of REVESTIVE at the end of this leaflet.

**Usual dose:**

The recommended daily dose is 0.05 mg per kg body weight. The dose will be given in mL of solution.

Your doctor will choose the dose that is right for you depending on your body weight. Your doctor will tell you which dose to inject. If you are not sure, ask your doctor, pharmacist or nurse.

**Overdose:**

If you think you have taken too much REVESTIVE, contact your healthcare professional, hospital emergency department or regional Poison Control Centre immediately, even if there are no symptoms.
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**Missed Dose:**

If you forget to inject this medicine (or cannot inject it at your usual time), use it as soon as possible on that day. Never use more than one injection in the same day. Do not inject a double dose to make up for a forgotten dose.

Keep using this medicine for as long as your doctor prescribes it for you. Do not stop using this medicine without consulting your doctor, as a sudden stop can cause changes in your fluid balance.

If you have any further questions on the use of this medicine, ask your doctor, pharmacist or nurse.

**What are possible side effects from using REVESTIVE?**

These are not all the possible side effects you may feel when taking REVESTIVE. If you experience any side effects not listed here, contact your healthcare professional.

Very common (may affect more than 1 in 10 people):

- Stomach pain
- Feeling sick (nausea)
- Bloated stomach
- Vomiting
- Respiratory tract infection (any infection of the sinuses, throat, airways or lungs)
- Swelling of hands and/or feet
- Reddening, pain or swelling at the site of the injection
- Swelling of stoma (an artificial opening for waste removal)

Common (may affect up to 1 in 10 people):

- Passing gas (flatulence)
- Decreased appetite
- Problems sleeping
- Rash (hypersensitivity)
- Bleeding of the skin
- Cough

Side effects were similar in adults and children. Other common events seen in children (ages 1 through 17 years) during clinical studies were; fever, infections related to the central line, headache, feeling tired, and runny nose.

Serious side effects and what to do about them			
Symptom / effect	Talk to your healthcare professional in all cases		Stop taking drug and get immediate medical help
	Only if severe	In all cases	
<b>COMMON</b>			
<b>Reduced flow of bile from the gallbladder and/or inflammation of the gallbladder:</b> Yellowing of the skin and the whites in the eyes, itching, dark urine and light-coloured stools or pain in the upper right side or middle of the stomach area		√	
<b>Congestive heart failure:</b> Tiredness, shortness of breath or swelling of ankles or legs, sudden weight gain		√	
<b>Intestinal obstruction (blockage of the bowel):</b> Stomach ache, vomiting and constipation		√	

Serious side effects and what to do about them			
Symptom / effect	Talk to your healthcare professional in all cases		Stop taking drug and get immediate medical help
	Only if severe	In all cases	
<b>Inflammation of the pancreas (pancreatitis):</b> Stomach ache and fever		√	
<b>UNCOMMON</b>			
<b>Fainting:</b> If heart rate and breathing are not normal and you don't awaken fast, seek help as soon as possible		√	

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

<p><b>Reporting Side Effects</b></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"> <li>- Visiting the Web page on Adverse Reaction Reporting (<a href="https://www.canada.ca/en/health-canada/services/drugs-health-products/medeffect-canada/adverse-reaction-reporting.html">https://www.canada.ca/en/health-canada/services/drugs-health-products/medeffect-canada/adverse-reaction-reporting.html</a>) for information on how to report online, by mail or by fax; or</li> <li>- Calling toll-free at 1-866-234-2345.</li> </ul> <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:**

Do not use this medicine after the expiry date which is stated on the carton after EXP. The expiry date refers to the last day of that month.

Store REVESTIVE 5 mg below 25°C. Do not freeze.

Reconstituted solution: Store below 25°C. Do not freeze. The product should be used within 3 hours after reconstitution.

Do not use this medicine if you notice that the solution is cloudy or contains particulate matter.

Do not throw away any medicines via wastewater or household waste. Ask your pharmacist how to throw away medicines you no longer use. These measures will help protect the environment.

Dispose of all needles and syringes in a sharps disposal container.

Keep out of reach and sight of children.

**If you want more information about REVESTIVE:**

- 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 or by calling 1-844-633-6713.

This leaflet was prepared by Shire Pharmaceuticals Ireland Limited.

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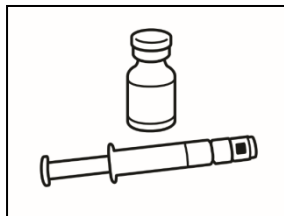
\*REVESTIVE is a registered trade-mark of Shire-NPS Pharmaceuticals, Inc., a Shire plc affiliate.

Last Revised: April 27, 2020

## **Instructions for preparing and injecting REVESTIVE**

### **Important information:**

- Read the Patient Medication Information before using REVESTIVE.
- REVESTIVE is for injection under the skin (subcutaneous injection).
- Do not inject REVESTIVE into a vein (intravenously) or muscle (intramuscularly).
- Keep REVESTIVE out of the sight and reach of children.
- Do not use REVESTIVE after the expiry date which is stated on the carton, the vial and the pre-filled syringe. The expiry date refers to the last day of that month.
- Store below 25°C. Do not freeze.
- The product should be used within 3 hours after reconstitution.
- Do not use REVESTIVE if you notice that the solution is cloudy or contains particulate matter.
- Do not throw away any medicines via wastewater or household waste. Ask your pharmacist how to throw away medicines you no longer use. These measures will help protect the environment.
- Dispose of all needles and syringes in a sharps disposal container.



### Materials in the REVESTIVE pack:

- 28 vials with 5 mg teduglutide as a powder
- 28 pre-filled syringes with diluent

### Materials needed but not included in the pack:

#### *Adults*

- Reconstitution needles (size 22G, length 1½" (0.7 x 40 mm))
- 1 mL injection syringes (with scale intervals of 0.02 mL or smaller)
- Thin injection needles for subcutaneous injection (e.g. size 26G, length 5/8" (0.45 x 16 mm))
- Alcohol swabs
- A puncture-proof container for safe disposal of the used syringes and needles

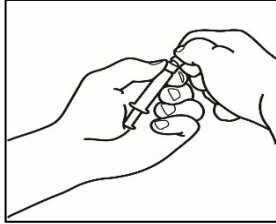
#### *Pediatrics*

- Reconstitution needles (size 22G, length 1½" [0.7 x 40 mm])
- 1 mL injection syringes (with scale intervals of 0.02 mL or smaller)
- Injection needles for subcutaneous injection (e.g. size 30G, length ½" [0.31 x 12.7 mm])
- Alcohol swabs
- A puncture-proof container for safe disposal of the used syringes and needles

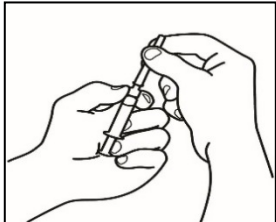
**NOTE:** Before you start, make sure you have a clean work surface and that you have washed your hands before proceeding.

### **1. Assemble the pre-filled syringe**

Once you have all the materials ready, you need to assemble the pre-filled syringe. The following procedure shows how you do this.



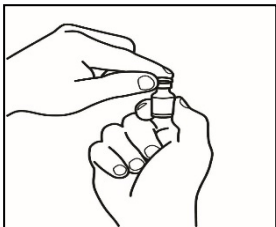
- 1.1 Take the pre filled syringe with diluent and flip off the top part of the white plastic cap so that it is ready for the reconstitution needle to be attached.



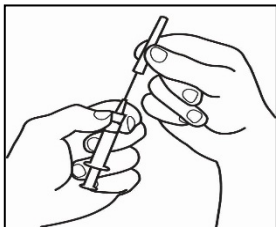
- 1.2 Attach the reconstitution needle (22G, 1½" (0.7 x 40 mm)) to the assembled pre-filled syringe by screwing it on in a clockwise direction.

## 2. Dissolve the powder

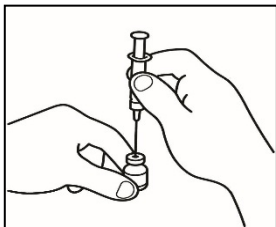
Now you are ready to dissolve the powder with the diluent.



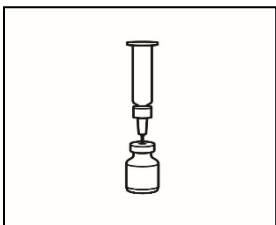
- 2.1 Remove the green flip-off button from the powder vial, wipe the top with an alcohol wipe and allow to dry. Do not touch the top of the vial.



- 2.2 Uncap the reconstitution needle on the assembled pre-filled syringe with diluent without touching the tip of the needle.

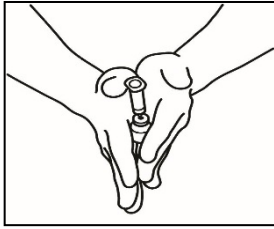


- 2.3 Taking the powder vial, insert the reconstitution needle attached to the assembled pre-filled syringe into the centre of the rubber stopper and gently push the plunger all the way down to inject all the diluent into the vial.



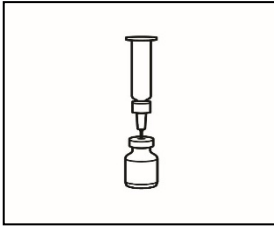
- 2.4 Leave the reconstitution needle and empty syringe in the vial. Let the vial rest for approximately 30 seconds.





2.5 Gently roll the vial between your palms for about 15 seconds. Then gently turn the vial upside-down once with the reconstitution needle and empty syringe still in the vial.

**NOTE:** Do not shake the vial. Shaking the vial may produce foam, which makes it difficult to extract the solution from the vial.



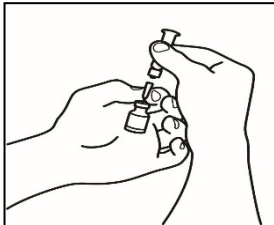
2.6 Let the vial rest for about two minutes.

2.7 Observe the vial for any undissolved powder. If any powder remains, repeat steps 2.5 and 2.6. Do not shake the vial. If there is still some undissolved powder, discard the vial and start the preparation again from the beginning with a new vial.

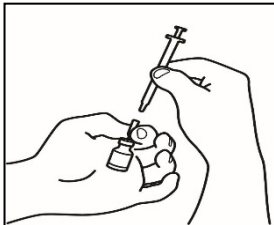
**NOTE:** The final solution should be clear. If the solution is cloudy or contains particulate matter, do not inject it.

**NOTE:** The product should be used within 3 hours after reconstitution.

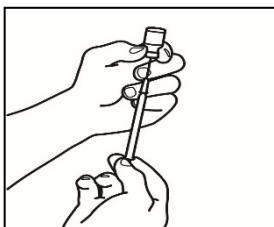
### 3. Prepare the injection syringe



3.1 Remove the reconstitution syringe from the reconstitution needle which is still in the vial and discard the reconstitution syringe.

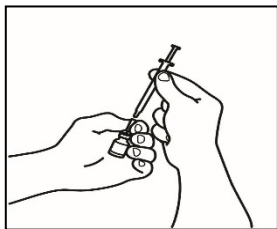


3.2 Take the injection syringe and attach it to the reconstitution needle which is still in the vial.

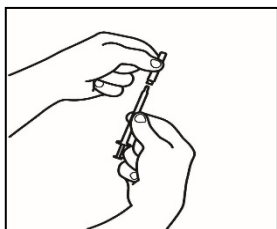


3.3 Turn the vial upside down, slide the tip of the reconstitution needle close to the stopper and allow **ALL** the medicine to fill the syringe by pulling the plunger back gently.

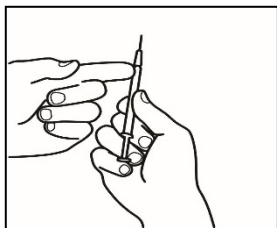
**NOTE:** If your doctor has told you that you need two vials, prepare a second pre-filled syringe with diluent and a second powder vial as shown in the main steps 1 and 2. Withdraw the solution from the second vial into the same injection syringe by repeating step 3.



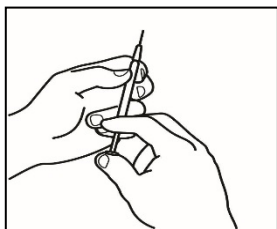
3.4 Remove the injection syringe from reconstitution needle leaving the needle in the vial. Discard the vial and reconstitution needle together into the sharps disposal container.



3.5 Take the injection needle (Adults: 26G, 5/8" [0.45 x 16 mm]) or (Pediatrics: 30G, 1/2" [0.31 x 12.7 mm]), but do not remove the plastic needle cap. Attach the needle to the injection syringe containing the medicine.

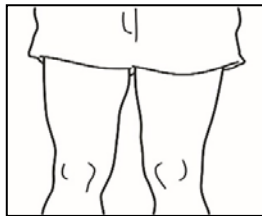
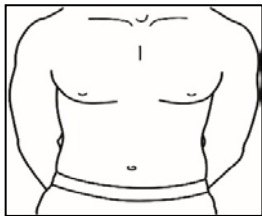


3.6 Check for air bubbles. If air bubbles are present, gently tap the syringe until they rise to the top. Then gently push up the plunger to expel the air.



3.7 Your dose in mL has been calculated by your doctor. Expel any excessive volume from the syringe with the needle cap still on until your dose is reached.

#### 4. Inject the solution

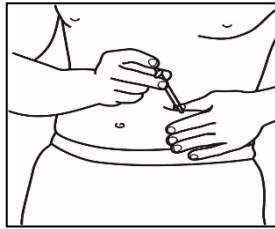


4.1 Find an area on your belly, upper arm, or on your thigh where it is easy for you to give the injection (see the diagram).

**NOTE:** Do not use the same area each day for each injection - rotate sites (use upper, lower, and left and right side of your belly, or either upper arm or thigh) to avoid discomfort. Avoid areas that are inflamed, swollen, scarred or covered by a mole, birthmark or other lesion.



- 4.2 Clean the intended site of injection on your skin with an alcohol swab, using a circular motion, working outwards. Allow the area to air-dry.



- 4.3 Remove the plastic cap from the needle of the prepared injection syringe. Gently grasp the cleaned skin at the injection site with one hand. With the other hand, hold the syringe as you would with a pencil. Bend your wrist back and quickly insert the needle at a 45° angle.

- 4.4 Pull back the plunger slightly. If you see any blood in the syringe, withdraw the needle and replace the needle on the injection syringe with a clean one of the same size. You can still use the medicine that is already in the syringe. Try to inject in another place in the cleaned skin area.
- 4.5 Inject the medicine slowly by pushing steadily on the plunger until all the medicine is injected and the syringe is empty.
- 4.6 Pull the needle straight out of the skin and discard the needle and syringe together into the sharps disposal container. A small amount of bleeding may occur. If necessary, press gently on the injection site with an alcohol swab or 2x2 gauze until any bleeding has stopped.

Dispose all needles and syringes in a sharps disposal container or hard-walled container (for example, a detergent bottle with a lid). This container must be puncture proof (top and sides). If you need a sharps disposal container, please contact your doctor.