

PACKAGE LEAFLET: INFORMATION FOR THE USER
Saizen 8 mg click.easy powder and solvent for solution for injection
somatropin

Read all of this leaflet carefully before you start using this medicine:

- Keep this leaflet. You may need to read it again.
- If you have any further questions, ask your doctor or pharmacist.
- This medicine has been prescribed for you . Do not pass it on to others. It may harm them, even if their symptoms are the same as yours.
- If any of the side effects gets serious, or if you notice any side effects not listed in this leaflet, please tell your doctor or pharmacist.

In this leaflet:

1. What Saizen is and what it is used for
2. Before your use Saizen
3. How to use Saizen
4. Possible side effects
5. How to store Saizen
6. Further information

1. WHAT SAIZEN 8 MG CLICK.EASY IS AND WHAT IT IS USED FOR

Saizen is a growth hormone. Saizen's main action is to increase growth in children and adolescents and to treat adults with growth hormone deficiency.

The growth hormone (somatropin) contained in Saizen is almost the same as human's natural growth hormone except that it is made outside the body by a process called "recombinant DNA technology" (genetic engineering).

Saizen is used:

In children and adolescents:

- in the treatment of children with short stature who have failed to grow because their body produces no growth hormone or insufficient levels of growth hormone
- in the treatment of girls who have failed to grow due to gonadal dysgenesis (also referred to as Turner Syndrome), confirmed by a test on the chromosomes
- in the treatment of pre-pubertal children who have failed to grow due to chronic renal failure, a condition in which kidneys are damaged
- in the treatment of growth problems in children who were born small and who have not reached normal height by the age of 4 years or later.

In adults:

- in the treatment of adults with a marked lack of growth hormone deficiency (growth hormone deficiency).

This therapy is given to adults who have a serious growth hormone deficiency that has been medically diagnosed by a test.

The doctor or pharmacist will be able to explain why this medicine has been given to you or to your child.

2. BEFORE YOU USE SAIZEN 8 MG CLICK.EASY

Do not use Saizen

- If you (or your child) are allergic (hypersensitive) to somatropin or any of the other ingredients of Saizen or the liquid used to make up the injection.
- If you have been told that your child's bones have stopped growing and that he/she has therefore reached his/her final height.
- If you (or your child) have been or are currently treated for any tumour or a brain tumour in particular.
 - Any evidence of active malignant tumours.
- If you or your child have an acute critical illness, suffering complications following open heart surgery, abdominal surgery, multiple accidental trauma, acute respiratory failure or similar conditions.

Treatment with Saizen has to be stopped in children with chronic kidney disease at time of kidney transplantation.

Take special care with Saizen

Saizen therapy should be carried out under regular guidance of a doctor who is experienced in the diagnosis and management of patients with growth hormone deficiency.

Shortly after Saizen is given, you (or your child) may feel shaky or light-headed due to low blood sugar levels. These feelings will quickly disappear. Your (or your child's) blood sugar levels may then rise above normal 2-4 hours after administration. Since treatment with growth hormone can alter how your body handles sugar, your (or your child's) sugar levels will be tested regularly by a doctor. Somatropin may cause you (or your child's) blood sugar to increase.

If you (or your child) are diabetic or a member of your family has diabetes, your doctor will monitor closely your blood sugar level and may adjust the treatment for diabetes while you are being treated with Saizen.

Please be aware that you may need regular eye tests after having this medicine.

Saizen may affect how your thyroid works. Your doctor may test your blood for levels of thyroid hormones and prescribe another hormone if you (or your child) are found to have developed a lack of thyroid hormone.

Saizen may cause fluid retention in adult patients. This may appear as swelling and joint or muscle pain. If you feel these symptoms tell your doctor who may decide to adjust your dose of Saizen.

If in the past you (or your child) have had a condition affecting the brain, e.g. a tumour, the doctor will examine you (or your child) regularly to check that this has not come back again. Please note that there is no evidence to suggest that the use of Saizen increases the chances of such conditions coming back.

Some patients may develop swelling of the brain, while taking Saizen. If you (or your child) suffer from a bad or recurrent headache, have problems with your eyesight, feel sick (nausea) and/or being sick (vomiting), contact your doctor immediately. In this case it may be necessary to stop the growth hormone treatment, although treatment may be re-started at a later date. If the symptoms of brain swelling come back, treatment with Saizen should be discontinued.

When the medicine is injected into the same place over a long period of time, it can cause damage to this area. It is therefore important to keep changing the injection site. Your doctor or pharmacist can speak to you about which parts of the body should be used (see HOW TO USE SAIZEN).

Some children with growth hormone deficiency have developed leukaemia (increased number of white blood cells), whether or not they have received treatment with growth hormone. However there is no evidence that leukaemia incidence is increased in growth hormone recipients without predisposing factors.. No cause and effect relationship with growth hormone treatment has been proven.

Hip problems may occur more commonly in children with hormone or kidney problems. If your child has chronic renal failure, which can occur when kidneys are damaged, he or she should be examined periodically for evidence of bone disease. It is uncertain whether the bone disease in children with hormone or kidney problems is affected by growth hormone therapy. X-rays of the hip should be obtained prior to initiating therapy. If your child develops a limp or complains of hip or knee pain while being treated with Saizen, tell your doctor.

Treatment with Saizen will be stopped in children with chronic kidney failure at the time of kidney transplant.

Saizen is not indicated for the long-term treatment of paediatric patients who have growth failure due to genetically confirmed Prader-Willi Syndrome, unless they also have a diagnosis of growth hormone deficiency. There have been reports of sleep apnoea and sudden death after initiating therapy with growth hormone in paediatric patients with Prader-Willi Syndrome who had one or more of the following risk factors: severe obesity, history of upper airway obstruction or sleep apnoea, or unidentified respiratory infection.

Growth hormone should generally not be used by people who are seriously ill.

If you do not respond to the therapy with Saizen, you may have developed antibodies to growth hormone. Your doctor will conduct appropriate tests to determine this.

If you are over 60 years old or if you are taking Saizen for a long time, you should be examined more frequently by your doctor. As there is less experience both in the treatment of older people and in prolonged treatment with Saizen, special care is required.

Using other medicines

It is usually safe to use other medicines. However, if you (or your child) are using corticosteroids, it is important to tell the doctor or pharmacist. These medicines could interact with Saizen and therefore your doctor may need to adjust the dose of these medicines or your Saizen dose. Corticosteroids are used to treat several illnesses including asthma, allergies, kidney rejection and rheumatoid arthritis. Please tell your doctor or pharmacist if you are taking or have recently taken any other medicines, including medicines obtained without a prescription.

Pregnancy and breast feeding

Ask your doctor or pharmacist for advice before taking any medicine.

There is currently insufficient evidence from human studies on the safety of growth hormone treatment during pregnancy and breast feeding. Saizen should be discontinued, if pregnancy occurs.

If you become pregnant tell your doctor immediately. Consult your doctor before taking Saizen if you are breast feeding.

Driving and using machines

No studies on the effects of Saizen on the ability to drive and use machines have been performed. Somatropin-containing products have no influence on the ability to drive and use machines.

3. HOW TO USE SAIZEN 8 MG CLICK.EASY

Always use Saizen exactly as your doctor has told you. You should check with your doctor or pharmacist if you are not sure.

Dosage

It is recommended that Saizen be administered at bedtime.

The dose and frequency of administration of Saizen will be decided by your doctor and will depend on your (or your child's) body weight or body surface area.

Children and adolescents:

- Short stature due to a lack of, or insufficient levels of natural growth hormone:
0.7-1.0 mg/m² body surface area each day or 0.025-0.035 mg/kg body weight each day by subcutaneous administration (under the skin).
- Growth failure in girls due to gonadal dysgenesis (Turner Syndrome):
1.4 mg/m² body surface area per day or 0.045-0.050 mg/kg body weight each day by subcutaneous administration (under the skin).
If your daughter is treated for Turner Syndrome and she receives non-androgenic anabolic steroids as well, growth response may be enhanced. Ask your doctor or pharmacist if you are not sure about these drugs.
- Growth failure in pre-pubertal children due to chronic renal failure, a condition in which kidneys are damaged:
- 1.4 mg/m² body surface area, approximately equal to 0.045-0.050 mg/kg body weight each day by subcutaneous administration (under the skin).
- Growth problems in children who were born small:
1 mg/m² body surface area, approximately equal to 0.035 mg/kg body weight each day by subcutaneous administration (under the skin).

Adults:

- Growth Hormone Deficiency in adults:
At the start, low doses of 0.15-0.30 mg each day are recommended by subcutaneous injection (under the skin).. The dose will be adjusted in stages by your doctor. The recommended final dose of growth hormone rarely exceeds 1.0 mg/day. In general the lowest efficacious dose that works for you should be administered. If you are older or overweight, a lower dose may be necessary.

Method and route of administration

The dose and frequency of administration of Saizen will be decided by your doctor and will depend on your (or your child's) size or body weight. In general Saizen 8 mg click.easy should be administered each day by subcutaneous injection (under the skin).

The product (powder in vials) must be reconstituted with the enclosed bacteriostatic solvent using the click.easy reconstitution device.

The reconstituted solution is intended for subcutaneous administration (under the skin) and should be clear with no particles. If the solution contains particles, it must not be injected. The reconstituted solution contains 8 mg of Saizen (5.83 mg/ml).

Important information

Patients should be thoroughly instructed in the reconstitution procedure.

For young children, the reconstitution process should be supervised by an adult.

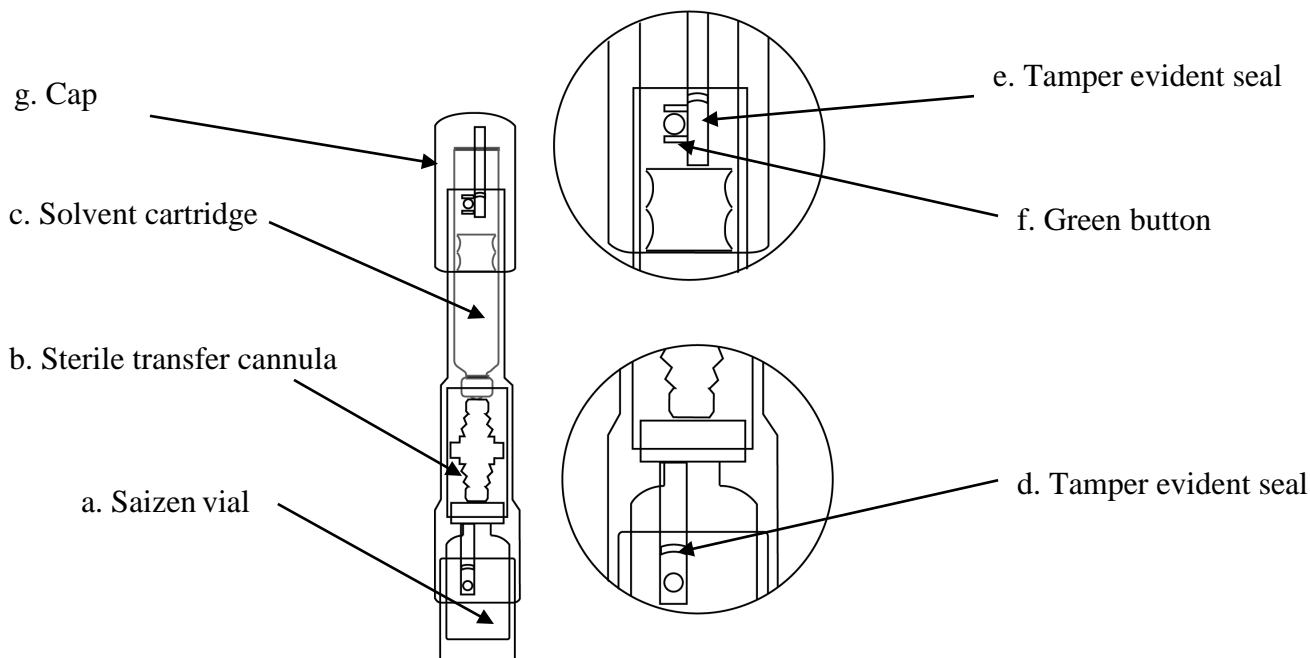
For administration of Saizen 8 mg click.easy, please read the following instructions carefully.

When the medicine is injected into the same place every time for a long time, it can cause damage. It is important to keep changing the place where you have your injection. Your doctor or pharmacist can speak to you about which part of the body you should use. Do not use any areas in which you feel

lumps, firm knots, depressions, or pain; talk to your doctor or pharmacist about anything you find. Clean the skin at the injection site with soap and water.

A separate leaflet in this box gives you detailed instructions in pictures about how to use this reconstitution kit.

Please consult your doctor, or pharmacist, if you have any questions concerning the reconstitution process.



How to prepare your solution of Saizen 8 mg click.easy

- Make sure the click.easy is complete by checking that the Saizen vial (a), the sterile transfer cannula (b) and the solvent cartridge (c) are present.
 - Check that the tamper evident seals on the click.easy housing (d) and on the cap (e) are not broken. If either of the tamper evident seals are broken return it to your pharmacist or doctor.
 - Place all elements needed for the preparation of the solution on a clean surface.
 - Wash your hands with soap and water.
1. Place the click.easy reconstitution device vertically on a flat surface with the vial at the bottom and the cap (g) facing upward.
 2. Push the cap down until it will go no further. (Note: The tamper evident seal on the click.easy housing is now broken).
 3. Turn slightly the cap clockwise until the green button (f) is positioned in the vertical opening.
 4. Continue pushing the cap down very slowly until it will go no further to transfer the solvent from the cartridge into the vial (Note: The tamper evident seal on the cap is now broken). It is important to push slowly to avoid foam formation in the vial. Check that all the solvent has been transferred into the vial.
 5. Dissolve the powder with the solvent by gently swirling the click.easy (Please note that you should not shake the solution strongly or you will cause the solution to foam). Avoid vigorous shaking to prevent creation of foam. Let the solution stand until the powder is completely dissolved. If the solution contains particles, it must not be injected. Push the cap back down until it will go no further and keep it in position.
 6. Turn the click.easy so that the vial is at the top and pull the cap slowly downwards until the solution is completely drawn back into the cartridge. Check that no more than one or two drops of solution remain in the vial.

If there are more than one or two drops of solution remaining in the vial, slowly push the cap up until some of the solution is back in the vial and gently tap the click.easy. Then draw the solution slowly again back into the cartridge.

7. Remove any excess air that has been drawn into the cartridge by pushing slowly the cap up until no air is visible in the cartridge.
(Note: Avoid pulling the cap down too fast, as this will draw air into the cartridge).
8. Keeping the click.easy in this position (vial on the top) unscrew the cap and remove it. Still keeping the same position (vial on the top) remove the cartridge containing the reconstituted solution for injection from the click.easy.
9. Carefully peel off the outer label using the tab provided. Write the reconstitution date on the transparent inner label on the cartridge. Discard the click.easy safely in accordance with your local requirements.

The cartridge containing the reconstituted solution of Saizen is now ready to be used for administration with your one.click auto-injector, cool.click needle-free auto-injectors or easypod auto-injector.

The reconstituted solution should be clear with no particles. If the solution contains particles, it must not be injected.

How to perform your daily self-administration of Saizen

For instructions on how to load the cartridge into the one.click auto-injector, how to use the cool.click needle-free auto-injectors or easypod auto-injector and inject the reconstituted solution of Saizen, please carefully read the corresponding instruction manual provided with each auto-injector. Intended users of easypod are primarily children starting from the age of 7 up to adults. Use of the devices by children should always be made under adult's supervision.

Frequency of administration

It is recommended that Saizen be administered at bedtime.

Duration of treatment

Your child should stop using this treatment when he or she reaches a satisfactory adult height or his or her bones cannot grow any longer, as assessed by his or her doctor. Treatment with Saizen will be stopped in children with chronic kidney failure at the time of kidney transplant.

If you use more Saizen than you should

If you inject too much Saizen, you should tell your doctor as it may be necessary to change slightly the dose to make up for this. Injecting too much can lead to changes in blood sugar levels which could mean that you (or your child) will feel shaky and light-headed. If this happens contact your doctor as soon as possible.

If you forget to use Saizen

If you miss a dose, tell your doctor as it may be necessary to slightly change the dose to make up for this.

If you stop taking Saizen

Do not stop taking Saizen without speaking with your doctor.

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

4. POSSIBLE SIDE EFFECTS

Like all medicines, Saizen can cause side effects, although not everybody gets them.

Tell your doctor immediately if you experience strong and recurrent headaches associated with feeling sick (nausea), vomiting or sight disturbances. These are symptoms of a side effect called benign intracranial hypertension and it is uncommon.

Side effects may occur with certain frequencies, which are defined as follows:

- common: affects 1 to 10 users in 100
- uncommon: affects 1 to 10 users in 1,000
- rare: affects 1 to 10 users in 10,000
- very rare: affects less than 1 user in 10,000
- not known: frequency cannot be estimated from the available data.

Common side effects:

- Redness and itching at the site of injection. If this appears to be particularly troublesome, you should discuss it with your doctor.
- Oedema (swelling), muscle pain, joint pain, and joint disorders in adult users. These side-effects appear usually early in the treatment and are short-lived.

Uncommon side effects:

- Carpal tunnel syndrome that is characterized by persistent stinging, burning sensation, pain and/or numbness in the hand. (Isolated) headache.
- Oedema (swelling), muscle pain, joint pain, and joint disorders in children. These side-effects appear usually early in the treatment and are short-lived.

Very rare side effects:

- Slipped capital femoral epiphysis (a hip problem that starts if the growing end of the thigh bone slips from the ball of the hip joint) and avascular necrosis of the femoral head. If your child shows an unexplained limp and hip or knee pain, please contact your doctor or pharmacist.
- Growth hormone treatment may reduce the levels of thyroid hormone. This can be tested by your doctor and if necessary your doctor will prescribe the adequate treatment.

You (or your child) can experience high blood sugar further to growth hormone treatment.

Leukemia has been reported in a small number of growth hormone deficiency patients, some of whom have been treated with somatropin. However, there is no evidence that leukaemia incidence is increased in growth hormone recipients without predisposing factors.

Very rarely a patient could develop antibodies (type of protein that helps protect the body) to somatropin. These are usually not associated with any side effects and do not usually interfere with growth.

If any of the side effects gets serious, or if you notice any side effects not mentioned in this leaflet, please inform your doctor or pharmacist.

5. HOW TO STORE SAIZEN

Keep out of the reach and sight of children.

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

Do not store above 25° C. Do not freeze. Store in the original package in order to protect from light.. Saizen 8 mg click.easy reconstituted solution should be stored in a refrigerator (2°C-8°C) in the cartridge and should be used within 28 days.

When using the easypod and one.click auto-injectors, the cartridge is kept in the device and the device must be stored in the refrigerator. When using the cool.click needle-free auto-injectors, only the cartridge of reconstituted Saizen should be stored in a refrigerator (2°C-8°C).

Medicines should not be disposed of via wastewater or household waste. Ask your pharmacist how to dispose of medicines no longer required. These measures will help to protect the environment.

6. FURTHER INFORMATION

What Saizen contains

The active substance is somatropin (Recombinant Human Growth Hormone) 8 mg.
The other ingredients are sucrose, phosphoric acid and sodium hydroxide (for pH-adjustment).
The solvent is metacresol (as preservative 0.3% w/v) solution in water for injection.

What Saizen looks like and contents of the pack

Saizen 8 mg click.easy is a powder for solution for injection. It is available in the following pack sizes:

- 1 vial of Saizen 8 mg product and 1 cartridge of bacteriostatic solvent (water for injection and metacresol) pre-assembled in 1 reconstitution device (click.easy).
- 5 vials of Saizen 8 mg product and 5 cartridges of bacteriostatic solvent (water for injection and metacresol) pre-assembled in 5 reconstitution devices (click.easy).

Not all pack sizes may be marketed.

THIS IS A MEDICINE

- A medicine is a product which affects your health and its consumption, contrary to instructions, is dangerous for you.
- Closely follow your doctor's prescription, the method of use and the instructions of the pharmacist who sold the product.
- Your doctor and the pharmacist are experts in medicine, its benefits and risks.
- Do not interrupt the period of treatment prescribed without your doctor's permission.
- Do not repeat the same prescription without consulting your doctor.

Keep medicines out of reach of children.

Council of Arab Health Ministers

Union Arab Pharmacists

Marketing Authorisation Holder

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