PATIENT INFORMATION LEAFLET

SCHEDULING STATUS

S4

RAMOLEP 20 mg/ml solution for injection in pre-filled syringe

RAMOLEP 40 mg/ml solution for injection in pre-filled syringe

Glatiramer acetate

Contains sugar (mannitol 40 mg/ml).

Read all of this leaflet carefully before you start using RAMOLEP

- Keep this leaflet. You may need to read it again.
- If you have any further questions, please ask your doctor, pharmacist, nurse or other health care provider.
- RAMOLEP has been prescribed for you personally and you should not share your medicine with other people. It may harm them, even if their symptoms are the same as yours.

What is in this leaflet

1. What RAMOLEP is and what it is used for

RAMOLEP is a medicine used for the treatment of relapsing forms of multiple sclerosis (MS). It modifies the way in which your body's immune system works and

it is classed as an immunomodulating medicine. The symptoms of MS are thought to be caused by a defect in the body's immune system. This produces patches of inflammation in the brain and spinal cord.

RAMOLEP is used to reduce the number of times you suffer attacks of MS (relapses). It has not been demonstrated to help if you have any form of MS which does not relapses or hardly ever relapses. RAMOLEP may not have any effect on the length of time an MS attack lasts, or how badly or suffer during an attack.

2. What you need to know before you use RAMOLEP

Do not use RAMOLEP

- If you are hypersensitive (allergic) to glatiramer or any of the ingredients of RAMOLEP.
- If you are pregnant (see Pregnancy and breastfeeding).

Warnings and precautions

Special care should be taken with RAMOLEP:

- If you have kidney or heart problems, as you may need to have regular tests and check-ups.

Children and adolescents

RAMOLEP should not be used in children or adolescents under the age of 18 years.

Other medicines and RAMOLEP

Always tell your health care provider if you are taking any other medicine. (This includes all complementary or traditional medicines.)

An increased incidence of injection site reactions has been seen in RAMOLEP patients receiving concurrent administration of corticosteroids.

If you are using phenytoin, carbamazepine or any other medicines for fits or seizures, inform your doctor or healthcare provider before you start using RAMOLEP, as it can have an effect on the effectiveness of the protein-bound medicines.

Pregnancy and breastfeeding

If you are pregnant or breastfeeding, think you may be pregnant or are planning to have a baby, please consult your doctor, pharmacist or other health care provider for advice before using this medicine.

You should not use RAMOLEP if you are pregnant.

Additional contraception should be considered when using RAMOLEP.

Driving and using machines

You may feel confused, your vision may be impaired or you may feel dizzy after being given RAMOLEP. If this happens, do not drive or use any tools or machines. It is not always possible to predict to what extent RAMOLEP may interfere with your daily activities. You should not engage in the above activities until you are aware of the measure to which RAMOLEP affects you.

3. How to use RAMOLEP

Do not share medicines prescribed for you with any other person.

Always use RAMOLEP exactly as your doctor has told you. Check with your doctor if you are not sure.

The usual dose in adults is 20 mg of glatiramer acetate (one pre-filled syringe), administered as a subcutaneous injection once daily, or 40 mg of glatiramer

acetate (one pre-filled syringe), administered as a subcutaneous injection (under the skin) three times per week, at least 48 hours apart.

It is very important to inject RAMOLEP properly:

- Only administer RAMOLEP under the skin (see Instructions for use)
- Only use the dose as prescribed by your doctor
- Never use the same syringe more than once. Any unused medicine or waste must be discarded
- Do not mix or co-administer the content of RAMOLEP with any other medicine
- Do not use RAMOLEP if the solution contains particles. Use a new pre-filled syringe.

Your doctor or nurse will assist you when you administer RAMOLEP for the first time.

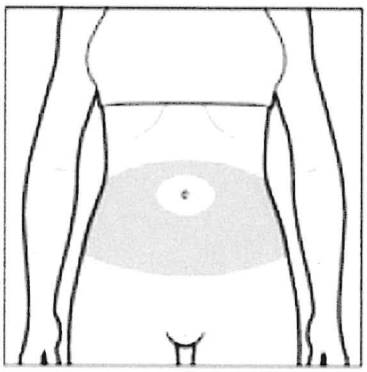
They will monitor you for half an hour afterwards, to make sure that there are no problems.

Instructions for use

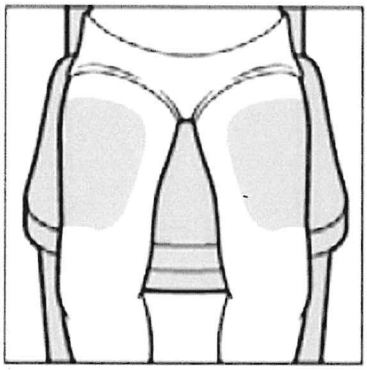
Read the following instructions carefully before you use RAMOLEP

1. Make sure you have the following:
    - Disposable container to place the used syringe and needle
    - One tray of RAMOLEP pre-filled syringe
    - If RAMOLEP is stored in the fridge, take the tray out at least 20 minutes before you will inject it, so that it warms to room temperature.
    - Only opens one tray containing RAMOLEP pre-filled syringe at the time you will inject yourself.
2. Wash your hands thoroughly with soap and water.
3. Choose the injection site. There are seven possible areas on your body for the injection:

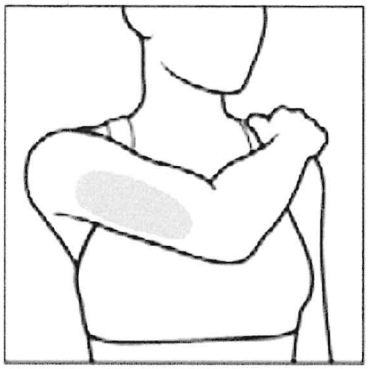
- Area 1: Stomach area (abdomen) around the belly button. Avoid 5 cm around the belly button

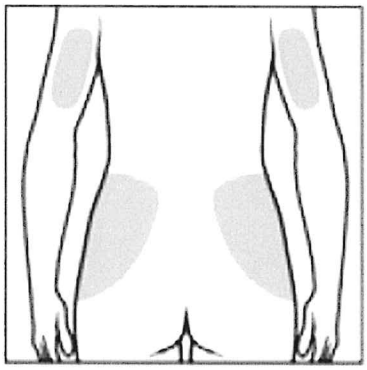


- Area 2 and 3: Thighs (above your knees)



- Area 4, 5, 6 and 7: back of the upper arms and upper hips (below your waist).





Choose a different site for the injection every day. Rotate the injection sites.

Do not use the same site each time.

Please note:

- Do not use any area that is painful or discoursed or where you feel firm knots or lumps.
- You should consider having a planned schedule for rotating injection sites and making a note of it in a diary.
- There are some sites that may be difficult to self-inject (like the back of your arm). If you want to use these sites, you may require assistance.

4. Inject RAMOLEP
   - Remove the syringe from the tray by peeling back the paper foil lid.
   - Remove the shield from the needle. Do not remove the shield with our mouth or teeth.
   - With the thumb and forefinger of the free hand, gently pinch up the skin as shown in Figure 1.
   - Push the needle into the skin as shown in Figure 2.
   - Inject the medicine by steadily pushing the plunger all the way down until the syringe is empty.
   - Pull the syringe and needle straight out.
   - Discard the syringe and needle in a safe disposable container.

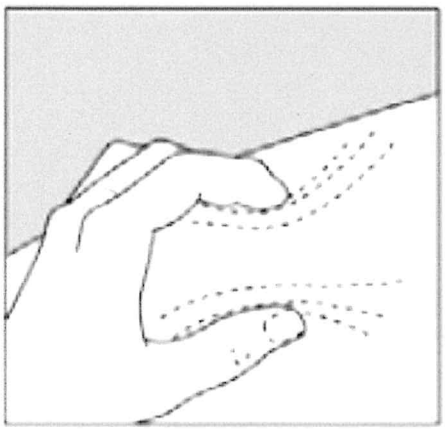


Figure 1

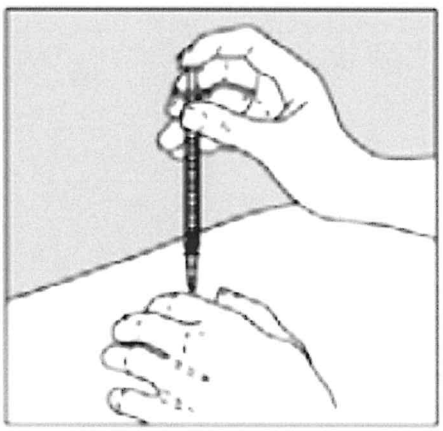


Figure 2

Your doctor will tell you how long your treatment with RAMOLEP will last. If you have the impression that the effect of RAMOLEP is too strong or too weak, tell your doctor or pharmacist.

If you used more RAMOLEP than you should

In the event of overdosage, consult your doctor or pharmacist. If neither is available, contact the nearest hospital or poison centre.

If you forget to use RAMOLEP

Do not use a double dose to make up for forgotten individual doses. Inject RAMOLEP as soon as you remember and use the next dose 24 hours later.

4. Possible side effects

RAMOLEP can have side effects.

Not all side effects reported for RAMOLEP are included in this leaflet. Should your general health worsen or if you experience any untoward effects while using RAMOLEP, consult your health care provider for advice.

If any of the following happens, stop using RAMOLEP and tell your doctor immediately or go to the casualty department at your nearest hospital:

- Swelling of the face, lips, tongue and throat or other parts of the body
- Wheezing, nasal congestion or trouble breathing
- Rash, itching or eczema
- Fainting.

These are all very serious side effects. If you have them, you may have had a serious reaction to RAMOLEP. You may need urgent medical attention or hospitalisation.

Tell your doctor immediately or go to the casualty department at your nearest hospital if you notice any of the following:

Reactions immediately after the injection. It is uncommon, but some people may get one or more of the following symptoms within minutes after injecting RAMOLEP. These symptoms usually disappear within half an hour. However, if they last longer than 30 minutes tell your doctor or go to the casualty department at your nearest hospital

- Flushing or reddening of the chest or face (vasodilatation)
- Chest pain
- Shortness of breath (dyspnoea)
- Pounding and rapid heartbeat (palpitation or tachycardia).

These are all serious side effects. You may need urgent medical attention.

Tell your doctor if you notice any of the following:

Frequent side effects

- Inflammation of the respiratory track, flu, bronchitis, inflammation of the stomach (gastroenteritis), cold sores (herpes simplex), inflammation of the ears (otitis media), tooth abscess, vaginal candidiasis
- Abnormal mass or growth in a part of your body
- Lymph node swelling
- Allergic reaction
- Loss of appetite, weight gain
- Nervousness

- Headache, altered taste, increased tightness of muscle tone, migraine, speech disorder, fainting, tremor
- Dizziness, light-headedness
- Double vision, eye disorder
- Ear disorder
- Nausea, disorder of the anus rectum, constipation, tooth decay, indigestion, difficulty in swallowing, lack of control over defecation, vomiting
- Abnormal liver function test
- Rash, bruising, excessive sweating, itching, skin disorder, nettle rash
- Joint pain, back pain, neck pain
- Urge to empty bladder, frequent urination, inability to empty bladder appropriately
- Lack or energy, chest pain, injection site reactions, pain, chills, face swelling, wasting of tissue at injection site, swelling due to build-up of fluid of hands or feet, fever.

Less frequent side effects

- Abscess, inflammation of the skin and soft tissue underneath, boils, shingles, inflammation of kidney
- Skin cancer
- Increased white blood cell count, reduced white blood cell count, spleen enlargement, low blood platelet count, change in form of white blood cells
- Enlarged thyroid (Goitre), overactive thyroid
- Alcohol intolerance, gout, increased blood fat levels, increased blood sodium levels, decreased serum iron (ferritin)
- Abnormal dreams, confusion, extreme excitement (euphoric) mood, hallucination, aggression, abnormal elevated mood, personality disorder,

suicide attempt

- Hand numbness and pain (carpal tunnel syndrome), mental disorder, convulsions, problems with handwriting and reading, muscle disorders, problems with movement, muscle spasm, nerve inflammation, abnormal nerve-muscle link leading to abnormal muscle function, involuntary rapid movement of the eyeballs, loss of ability to move, unconscious state (stupor), visual blind spots
- Cataract, eye lesion in the cornea, dry eye, eye bleeding, droopy upper eyelid, pupil widening, wasting of the optic nerve leading to visual problems
- Heart beat changes, e.g. extra, slower or faster heart beat
- Varicose vein
- Periodic stops in breathing, nose bleeding, abnormally fast or deep breathing (hyperventilation), tight feeling in the throat, lung disorder, inability to breathe due to throat tightness (choking sensation)
- Bowel inflammation, polyps in the colon, intestine inflammation, burping, ulcer in the throat, inflammation of the gums, rectal bleeding, enlarged salivary glands
- Tingling or itching in the mouth
- Gallstones, liver enlargement
- Swelling of the skin and soft tissues, skin contact rash, painful red skin lumps, skin lumps
- Swelling, inflammation and pain of joints, flank pain, decrease in the mass of muscles
- Blood in the urine, kidney stones, urinary tract disorder, urine abnormality
- Abortion
- Inflammation of the breast, difficulties getting an erection, drop of pelvic organ e.g. bladder from normal place (pelvic prolapse), sustained or prolonged erections, disorders of prostate, abnormal PAP smear test, testicle disorder,

vaginal bleeding, vaginal disorder

- Cyst, hangover, low body temperature (hypothermia), non-specific inflammation, death of tissue at the injection site, mucous membrane disorders
- Post vaccination syndrome.

If you notice any side effects not mentioned in this leaflet, please inform your doctor or pharmacist.

Reporting of side effects

If you get side effects, talk to your doctor or pharmacist. You can also report side effects to SAHPRA via the "6.04 Adverse Drug Reaction Reporting Form", found online under SAHPRA's publications:

https://www.sahpra.org.za/Publications/Index/8.

By reporting side effects, you can help provide more information on the safety of RAMOLEP.

5. How to store RAMOLEP

Store all medicines out of reach of children.

- Store between 2 °C and 8 °C (in a refrigerator)
- RAMOLEP pre-filled syringes can be kept for up to one month outside the refrigerator between 15 °C and 25 °C. You can do this only once. After one month, any RAMOLEP that have not been used and are still in their original packaging must be returned to the refrigerator.
- Do not freeze
- Store the pre-filled syringe in the original package (tray)
- Keep the tray in the outer carton in order to protect it from light
- Do not use after the expiry date stated on the tray and outer carton

- Do not use RAMOLEP if you notice any particles or discolouration of the solution.

Return all unused medicine to your pharmacist.

Do not dispose of unused medicine in drains or sewerage systems (e.g. toilets).

6. Contents of the pack and other information

What RAMOLEP contains

- The active substance is glatiramer acetate. Each pre-filled syringe (1 ml) of solution for injection contains 20 mg glatiramer acetate, equivalent to 18 mg of glatiramer or 40 mg glatiramer acetate, equivalent to 36 mg of glatiramer
- The other ingredients are mannitol and water for injection.

What RAMOLEP looks like and contents of the pack

RAMOLEP solution for injection is a clear colourless to slightly yellow/brownish solution free from visible particles.

The container closure system consists of a single use glass syringe barrel with an integrated needle. A rubber stopper (bromobutyl, type 1) is fitted in the barrel for closure and acts as a piston during injection. A driving rod is screwed in the rubber stopper. The needle is covered with a needle shield.

The pre-filled syringe is packed in a PVC tray that is closed with a paper lidding foil. Each tray holds 1 syringe. Trays holding the pre-filled syringes are packed in a carton box.

The volume of solution in the syringe is 1,0 ml.

RAMOLEP 20 mg/ml:

7 pre-filled syringes

28 pre-filled syringes

30 pre-filled syringes

90 (3x30) pre-filled syringes

RAMOLEP 40 mg/ml

3 pre-filled syringes

12 pre-filled syringes

36 (3x12) pre-filled syringes

Not all pack sizes may be marketed.

Holder of Certificate of Registration

Kahma Biotech (Pty) Ltd

106, 16th Road

Midrand

This leaflet was last revised in

Registration numbers

RAMOLEP 20 mg/ml: 54/26/0697

RAMOLEP 40 mg/ml: 54/26/0698

Access to the corresponding Professional Information

To be confirmed