(Preparations) - 1986

The medicine is dispensed with a doctor's prescription only

Mekinist® 0.5 mg, film-coated tablets

Each film-coated tablet contains trametinib (as dimethyl sulfoxide) 0.5 mg

Mekinist® 2 mg, film-coated tablets

Each film-coated tablet contains trametinib (as dimethyl sulfoxide) 2 mg

Inactive ingredients and allergens: See section 6 under 'Additional information' and section 2 under 'Important information about some of this medicine's inaredients'

Read the entire leaflet carefully before using the medicine. This leaflet contains concise information about the medicine. If you have any further questions, refer to your doctor or pharmacist.

This medicine has been prescribed for you. Do not pass it on to others. It may harm them, even if it seems to you that their medical condition is similar to yours

1. What is the medicine intended for?

Mekinist is used as monotherapy or in combination with an additional medicine containing dabrafenib for the treatment of adults with a type of skin cancer called melanoma that has spread to other parts of the body or cannot be removed by

Mekinist in combination with dabrafenib is also used for the adjuvant treatment of stage III melanoma in adults, following complete removal of the lesion by surgery

Mekinist in combination with dabrafenib is also used for the treatment of adults with an advanced stage of lung cancer called non-small cell lung cancer (NSCLC).

Mekinist in combination with dabrafenib is also intended for the treatment of a type of thyroid cancer called anaplastic thyroid cancer (ATC) that has a mutation in the BRAF gene (BRAF V600E mutation), which has spread to other parts of the body or is in one place and in advanced state, and after no satisfactory results have been obtained from local treatment options.

Therapeutic group:

protein kinase inhibitor, antineoplastic agent.

The three types of cancer have a particular change (mutation) in a gene called BRAF at the V600 position.

This mutation in the gene may have caused the cancer to develop. The medicine targets proteins made from this mutated gene and slows down or stops the development of the cancer.

2. Before using the medicine

Mekinist should only be used to treat melanomas, non-small cell lung cancer (NSCLC) and anaplastic thyroid cancer (ATC) with the BRAF mutation Therefore, before starting treatment, your doctor will test for this mutation

If your doctor decides that you will receive a combined treatment of Mekinist and dabrafenib, read the dabrafenib leaflet carefully as well as this leaflet. If you have any further questions on the use of this medicine, refer to your doctor. Do not use this medicine if:

You are sensitive (allergic) to trametinib or to any of the other ingredients in this medicine (listed in section 6 of this leaflet). Check with your doctor if you think this applies to you

Special warnings about using this medicine Before using Mekinist, tell your doctor if:

you have any liver problems. Your doctor may perform blood tests to monitor

your liver function while you are taking this medicine you have or have ever had kidney problems.

 you have or have ever had breathing or lung problems.
 you have heart problems such as heart failure (which can cause shortness of breath, difficulty in breathing when lying down, swelling of the feet or legs) or problems with the way your heart beats. Your doctor should check your heart function before and during treatment.

 you have eye problems, including blockage of the vein draining the eye (retinal vein occlusion) or swelling in the eye which may be caused by fluid blockage (chorioretinopathy).

Before treatment with Mekinist in combination with dabrafenib, tell your

 you have had a different type of cancer other than melanoma, non-small cell lung cancer (NSCLC) or anaplastic thyroid cancer (ATC), as you may be at greater risk of developing non skin cancers when taking Mekinist.

Check with your doctor if you think any of these may apply to you. Conditions you need to look out for

Some people taking Mekinist develop other conditions which can be serious. You need to know about important symptoms to look out for.

 ${\it Bleeding}$ Taking Mekinist or the combination of Mekinist and dabrafenib can cause serious bleeding including in the brain, the digestive system (such as stomach, rectum or intestine), lungs, and other organs, and can lead to death. Symptoms may include:

héadaches, dizziness, or feeling weak

blood in the stools or black stools

blood in the urine

 stomach pain coughing/vomiting up blood

Tell your doctor as soon as possible if you get any of these symptoms

Fever Taking Mekinist or the combination of Mekinist and dabrafenib may cause fever although it is more likely if you are taking the combination treatment (see also section 4). In some cases, people with fever may develop low blood pressure, dizziness or other symptoms.

Tell your doctor immediately if you get a temperature above 38°C or if you feel a fever coming on while you are taking this medicine.

Heart disorder

Taking Mekinist or the combination of Mekinist with dabrafenib can cause heart problems or make existing heart problems worse (see also "Heart conditions" in section 4).

Tell your doctor if you have a heart disorder. Your doctor will run tests to check that your heart is working properly before and during treatment with this medicine. Tell your doctor immediately if you feel like your heart is pounding, racing, or beating irregularly, or if you experience dizziness, tiredness, light headedness, shortness of breath or swelling in the legs. If necessary, your doctor may decide to interrupt your treatment or to stop it altogether.

Changes in your skin which may indicate new skin cancer

Your doctor will check your skin before you start taking this medicine and regularly while vou are táking it. Tell your doctor immediately if you notice any changes to your skin while

taking this medicine or after treatment (see also section 4).

Eve problems

You should have your eyes examined by your doctor while you are taking this medicine.

Tell your doctor immediately if you get eye redness and irritation, blurred vision, eye pain or other vision changes during your treatment (see also section 4). iniet can cause ove problems including blindness Mekinist is recommended if you have ever had blockage of the vein draining the eye (retinal vein occlusion). Tell your doctor immediately if you get the following symptoms of eye problems: blurred vision, loss of vision or other vision changes, coloured dots in your vision or halos (seeing blurred outlines around objects) during your nt. If necessary, your doctor may decide to interrupt your treatme to stop it altogether.

Liver problems Mekinist or Mekinist in combination with dabrafenib can cause problems with

Mekhist of Mekhist III Combination with daplatering can cause problems who your liver which may develop into serious conditions such as hepatitis and liver failure, which may be fatal. Your doctor will monitor you periodically. Signs that your liver may not be working properly may include loss of appetite

- nausea
- vomitina stomach nain
- yellowing of your skin or the whites of your eyes (jaundice) dark-coloured urine
- Tell your doctor as soon as possible if you get any of these symptoms Breathing or lung problems

Tell your doctor if you have any breathing or lung problems, including difficulty breathing often accompanied by a dry cough, shortness of breath and fatigue. Your doctor may conduct tests to check your lung function before you start taking your medicine. Muscle pain Mekinist can result in the breakdown of muscle (destruction of muscle cells -

rhabdomyolysis).

Tell your doctor as soon as possible if you get any of these symptoms:

→ Read the information "Possible serious side effects" in section 4 of

- dark urine due to kidney damage
- If necessary, your doctor may decide to interrupt your treatment or to stop it

Hole in the stomach or intestine (perforation)
Taking Mekinist or the combination of Mekinist and dabrafenib may increase the

risk of developing holes in the gut wall.

Tell your doctor as soon as possible if you have severe abdominal pain.

Serious skin reactions Serious skin reactions have been reported in people taking Mekinist in

Tell your doctor immediately if you notice any changes to your skin (see

section 4 for symptoms to be aware of). Inflammatory disease mainly affecting the skin, lung, eyes and lymph nodes

An inflammatory disease mainly affecting the skin, lung, eyes and lymph nodes (sarcoidosis). Common symptoms of sarcoidosis may include coughing, shortness of breath, swollen lymph nodes, visual disturbances, fever, fatigue pain and swelling in the joints and tender bumps on your skin. Tell your doctor if you get any of these symptoms.

Children and adolescents

Mekinist is not intended for children and adolescents since the effects of Mekinist

Drug interactions

in people younger than 18 years old are not known. efore starting treatment, tell your doctor or pharmacist if you are taking,

have recently taken or might take any other medicines. This includes nonprescription medications and dietary supplements. Keep a list of the medicines you take, so you can show it to your doctor or

pharmacist when you get a new medicine.

Using this medicine and food It is important to take Mekinist on an empty stomach, because food affects the way the medicine is absorbed into your body (see section 3).

Pregnancy, breastfeeding and fertility Mekinist is not recommended for use during pregnancy.

- If you are pregnant, think you may be pregnant or are planning to become pregnant, consult your doctor before taking this medicine. Mekinist can harm the unborn baby.
- If you are a woman who could become pregnant, you must use reliable birth control while you are taking Mekinist and for at least 16 weeks after you stop
- Birth control using hormones (such as pills, injections or patches) may not. work properly if you are taking Mekinist in combination with dabrafenib. You need to use another effective method of birth control so you do not become pregnant while you are taking this combination of medicines. Ask your doctor or pharmacist for advice.
- If you become pregnant while you are taking Mekinist, tell your doctor immediately.

Mekinist is not recommended while breastfeeding

It is not known whether the ingredients of Mekinist can pass into breast milk. If you are breastfeeding, or planning to breastfeed, you must tell your doctor. It is recommended that you do not breastfeed while you are taking Mekinist. You and your doctor will decide whether you will take Mekinist or breastfeed. Fertility - in both men and women

Mekinist may impair fertility in both men and womer

Taking Mekinist with dabrafenib: Dabrafenib may permanently reduce male fertility. In addition, men who are taking dabrafenib may have a reduced sperm count, and their sperm count may not return to normal levels after they stop taking this medicine.

Prior to starting treatment with dabrafenib, talk to your doctor about options to improve your chances to have children in the future If you have any further questions on the effect of this medicine on fertility, ask

Driving and using machines

Mekinist can have side effects that may affect your ability to drive and use machines. Avoid driving or using machines if you feel tired or weak, if you have problems with your vision or if your energy levels are low. Descriptions of these effects can be found in other sections (see sections 2 and

4). Read all the information in this leaflet for guidance. Consult your doctor or pharmacist if you are unsure about anything. Your disease symptoms and treatment situation may also affect your ability to drive

Important information about some of this medicine's ingredients This medicine contains less than 1 mmol sodium (23 mg) per tablet, that is to

say essentially 'sodium-free'

3. How to use the medicine? Always use this medicine according to your doctor's instructions

with your doctor or pharmacist if you are not sure about the dosage and treatment regimen of Mekinist.

The dosage and treatment regimen will be determined only by your doctor The usual dosage of Mekinist either used alone or in combination with dabrafenib is one 2 mg tablet once a day.

The usual dosage of dabrafenib, when used in combination with Mekinist, is 150 mg twice dailv.

Your doctor may decide to lower the dose if you get side effects.

Do not exceed the recommended dose, since this may increase the risk of

Swallow the tablet whole, with a full glass of water. Do not chew, crush or split the tablets, since they are film-coated and there is no information about their being chewed, crushed or split. Take Mekinist once a day, on an empty stomach (at least one hour before a meal or two hours after a meal). This means that:

after taking Mekinist, you must wait at least one hour before eating, or

after eating, you must wait at least two hours before taking Mekinist.

Take Mekinist at about the same time each day. If you take a higher dose of Mekinist

your next dose at the regular time.

If you take too many tablets of Mekinist, refer to your doctor or pharmacist. If possible, show them the Mekinist package and this leaflet. In case of vomiting after taking the medicine, do not take another dose. Take

If you forget to take the medicine less than 12 hours have passed since your regular dosing time, take the medicine as soon as you remember. If more than 12 hours have passed since your regular dosing time, skip this dose

and take your next dose at the usual time. Then carry on taking your tablet at

regular times as usual Do not take a double dose to make up for a forgotten dose.

Adhere to the treatment as recommended by your doctor.

If you stop taking this medicine Take Mekinist for as long as your doctor recommends. Do not stop unless your doctor advises you to.

How should you take Mekinist in combination with dabrafenib Take Mekinist in combination with dabrafenib exactly as your doctor tells vou. Do not change your dosage or stop taking Mekinist or dabrafenib unless your doctor tells you to.

- Take Mekinist once daily and dabrafenib- twice daily. It may be better for you to get into the habit of taking both medicines at the same times each day. Mekinist should be taken with either the morning dose of dabrafenib **or** the evening dose of dabrafenib. The dabrafenib doses should be taken about 12 hours apart. Take Mekinist and dabrafenib on an empty stomach, at least one hour before
- or two hours after a meal. Take them whole with a full glass of water.

 Do not chew, crush or split the tablets, since they are film-coated and there is
- no information about their being chewed, crushed or split. If you forget to take a dose of Mekinist or dabrafenib

- Mekinist: If there are more than 12 hours left until your next scheduled dose of Mekinist (which is taken once daily) take the medicine as soon as you remember o Dabrafenib: If there are more than 6 hours left until your next scheduled dose
- of dabrafenib (which is taken twice daily) take the medicine as soon as you remember. In any other case, continue taking the next dose at the regular time. Do not compensate for forgotten doses.

If you accidentally took a higher dose

If you take too much Mekinist or dabrafenib, immediately refer to your doctor, take the medicines with you, including the package and leaflets from each of them. If a child has accidentally swallowed some medicine, immediately refer to a doctor or go to a hospital emergency room and bring the medicine package with vou. If you get side effects, your doctor may decide that you should take a lower dosage of Mekinist and dabrafenib. Take the doses of Mekinist and dabrafenib

exactly as your doctor tells you. Do not take medicines in the dark! Check the label and dose every time you take medicine. Wear glasses if you need them.

If you have any further questions about using this medicine, consult your doctor or pharmacist.

4. Side effects Like with any medicine, using Mekinist may cause side effects in some users. Do not be alarmed when reading the list of side effects. You may not suffer from

Possible serious side effects

Heart conditions

Mekinist can affect how well your heart pumps blood. It is more likely to affect people who have an existing heart problem. You will be checked for any heart em while you are taking Mekinist

Signs and symptoms of heart problems include:

- tiredness
- feeling light-headed
- shortness of breath
- swelling in the legs

High blood pressure

Mekinist can cause new or worsening high blood pressure (hypertension). Your doctor or nurse will check your blood pressure during treatment with Mekinist. Call your doctor or nurse right away if you develop high blood pressure, your blood pressure worsens or you have a severe headache, dizziness or light headedness.

Bleeding problems Mekinist can cause serious bleeding problems, especially in your brain or stomach. Call your doctor and get medical help right away if you have any

- ınusual signs of bleeding, includi
- headache, dizziness or weakness
- coughing up blood or coughing up blood clots
 vomit containing blood or that looks like "coffee grounds"
- red or black stools that look like tar

Eye (vision) problems Mekinist can cause eye problems. Mekinist is not recommended if you have ever had blockage of the vein draining the eye (retinal vein occlusion). Your doctor may advise an eye examination before you take Mekinist and while you Your doctor may ask you to stop taking Mekinist or refer you to a

- specialist, if you develop signs and symptoms in your vision that include: loss of vision eye redness and irritation
- coloured dots in your vision
- halo (seeing blurred outlines around objects)
- blurred vision
- Changes in your skin
- Serious skin reactions have been reported in people taking Mekinist in combination with dabrafenib (frequency not known). If you notice one of the following: · reddish patches on the skin that are circular or target-shaped, with central blisters. Skin peeling. Ulcers of the mouth, throat, nose, genitals and eyes.
- These serious skin rashes can be preceded by fever and flu-like symptoms (Stevens-Johnson syndrome). widespread rash, fever and enlarged lymph nodes (DRESS-syndrome or drug
- hypersensitivity syndrome). → Stop using the medicine and seek medical attention immediately

Up to 3 in 100 people taking Mekinist in combination with dabrafenib may develop a different type of skin cancer called cutaneous squamous cell carcinoma (cuSCC). Others may develop a type of skin cancer called basal cell carcinoma (BCC). Usually, these skin changes remain local and can be removed with surgery, and treatment with Mekinist and dabrafenib can be continued without interrupt Some people taking Mekinist in combination with dabrafenib may also notice that new melanomas have appeared. These melanomas are usually removed

Your doctor will check your skin before you start taking dabrafenib, then check it again every month while you are taking dabrafenib and for 6 months after you stop taking it. This is to look for any new skin cancers.

by surgery and treatment with Mekinist and dabrafenib can be continued without

Your doctor will also check your head, neck, mouth and lymph glands, and you will have scans of your chest and stomach area (called CT scans) regularly. You may also have blood tests. These checks are to detect if any other cancer, including cutaneous squamous cell carcinoma, develops inside your body Pelvic examinations (for women) and anal examinations are also recommended before and at the end of your treatment.

Mekinist as monotherapy or in combination with dabrafenib can cause rash or acne-like rash. Follow your doctor's instructions for what to do to help prevent rash. Tell your doctor as soon as possible if you get any of these symptoms for the first time or if they get worse Refer to your doctor immediately if you get a severe skin rash with any of

Tell your doctor as soon as possible if you get any skin rash, or if you have a rash that gets worse. Muscle pain Mekinist can result in the breakdown of muscle (destruction of muscle cells -

the following symptoms: blisters on your skin, blisters or sores in your mouth, peeling of your skin, fever, redness or swelling of your face or your feet.

rhabdomyolysis). Tell your doctor if you have any new or worsening symptoms, includina dark urine due to kidney damage

Breathing or Lung problems Mekinist can cause inflammation of the lung (pneumonitis or interstitial lung disease). Tell your doctor if you have any new or worsening symptoms of lung or

- breathing problems, including: shortness of breath cough
- Possible side effects in patients taking Mekinist alone Side effects that you may experience when you take Mekinist alone are

fatique

- as follows: Very common side effects (may appear in more than one in 10 users):
- High blood pressure
- Bleeding at various sites in the body, which may be mild or serious Cough
- Shortness of breath Diarrhoea
- Constipation Stomach ache
- Drv mouth Skin rash, acne-like rash, redness of the face, dry or itching skin (see also 'Changes in your skin" at the beginning of section 4)
- Lack of energy or feeling weak or tired Swelling of the hands or feet (oedema peripheral)

Unusual hair loss or thinning

- Fever Very common side effects that may show up in your blood tests Abnormal blood test results related to the liv
- Common side effects (may appear in 1-10 in 100 users):

 Inflammation of hair follicles in the skin Nail disorders such as nail bed changes, nail pain, infection and swelling of
- Infection of the skin (cellulitis) Skin rash with pus-filled blisters (see also "Changes in your skin" at the beginning of section 4)

Heart rate that is lower than the normal range and/or a decrease in heart rate

- Allergic reaction (hypersensitivity)
- Dehydration (low levels of water or fluids) Blurred vision
- Swelling around the eyes
 Eyesight problems (see also "Eye (vision) problems" at the beginning of section 4) Changes in how the heart pumps blood (left ventricular dysfunction) (see also "Heart conditions" at the beginning of section 4)
- Localised tissue swelling Inflammation of the lung (pneumonitis or interstitial lung disease) Sore mouth or mouth ulcers, inflammation of mucous membranes Redness, chapping or cracking of the skin

Red and painful hands and feet

- Swelling of the face Inflammation of the mucosa Feeling weak
- Decreased red blood cells (anaemia), abnormal results of creatine phosphokinase values (an enzyme found mainly in heart, brain, and skeletal muscle). Uncommon side effects (may appear in 1-10 in 1,000 users):

Common side effects that may show up in your blood tests

- Swelling in the eve caused by fluid leakage (chorioretinopathy) (see also "Eve Swelling in the eye caused by induced learning of section 4)

 Swelling of nerves at the beginning of section 4)

 Swelling of nerves at the back of the eye (optic nerve head oedema papilloedema) (see also "Eye (vision) problems" at the beginning of section 4) Separation of the light-sensitive membrane in the back of the eye (the
- retina) from its supporting layers (retinal detachment) (see also "Eve (vision) problems" at the beginning of section 4).

 Blockage of the vein draining the eye (retinal vein occlusion) (see also "Eye (vision)) (vision) problems at the beginning of section 4)
 Heart pumping less efficiently, causing shortness of breath, extreme tiredness
- and swelling in ankles and legs (heart failure)
 A hole (perforation) in the stomach or intestines
- Inflammation of the intestines (colitis)

- Breakdown of muscle which can cause muscle pain and kidney damage (destruction of the muscle cells - rhabdomyolysis) If you experience any side effects, refer to your doctor or pharmacist. This includes any possible side effects not listed in this leaflet.

- feeling like your heart is pounding, racing or beating irregularly

- Tell your doctor as soon as possible if you get any of these symptoms, either for the first time or if they get worse.

- Nasal and throat inflammation
- Headache
- Cough Stomach ache
- Diarrhoea
- Nausea, vomitingRash, dry skin, itching, skin reddening
- Feeling weak, lack of energy Chills
- Very common side effects that may show up in your blood tests:
- follicles in the skin, nail disorders such as nail bed changes, nail pain, infection
- tumour which is usually not harmful), wart-like growths, increased sensitivity of the skin to sun (see also "Changes in your skin" at the beginning of section 4)
 Dehydration (low levels of water or fluids)
- Shortness of breath Dry mouth • Sore mouth or mouth ulcers, inflammation of mucous membranes
- Increased sweating, night sweats Unusual hair loss or thinning

Inflammation of the fatty layer under the skin (panniculitis)

Red and painful hands and feet

Inflammation of the pancreas

- Inflammation of the mucosa Swelling of the face
- Low levels of sodium (hyponatraemia) or phosphate (hypophosphataemia) in Increase in blood sugar level Increase in creatine phosphokinase, an enzyme found mainly in heart, brain,
- Skin tags
- Allergic reactions (hypersensitivity)
 Eye changes including swelling in the eye caused by fluid leakage
- Heart rate that is lower than the normal range and/or a decrease in heart rate Inflammation of the lung (pneumonitis)
- Inflammation of the intestines (colitis) Kidney failure
- (sarcoidosis) Rare side effects (may appear in 1-10 in 10.000 users): A hole (perforation) in the stomach or intestines
- Inflammation of the heart muscle (myocarditis) which can result in breathlessness, fever, palpitations and chest pain
 Inflamed, flaky skin (exfoliative dermatitis)
- If a side effect appears, if any side effect gets worse, or if you suffer from a side effect not mentioned in this leaflet, consult your doctor. You can report side effects to the Ministry of Health by clicking the link 'Report Side Effects of Drug Treatment' found on the Ministry of Health home page (www.health.gov.il) which directs you to the online form for reporting side effects, or by using the link: https://sideeffects.health.gov.il

5. How to store the medicine? Prevent poisoning! To prevent poisoning, this medicine, and all other medicines should be kept in a closed place, out of the reach and sight of children and/or infants. Do not induce vomiting unless explicitly instructed to do so by the doctor

Do not use the medicine after the expiry date (exp. date) which appears on the

Storage conditions: Store in a refrigerator (2°C to 8°C).
Store in the original package in order to protect from light and moisture. Keep the bottle tightly closed with the desiccant inside (small cylinder-shaped

neasures will help protect the environment

In addition to the active ingredient, the medicine also contains:

package. The expiry date refers to the last day of that month

Magnesium stearate (E470b) Sodium lauryl sulfate Colloidal silicon dioxide (E551) Film coating of 0.5 mg tablet

Film coating of 2 mg tablet

Hypromellose (E464)

Mekinist 2 mg:

The Mekinist 0.5 mg film-coated tablets are yellow, modified oval, biconvex, with "GS" debossed on one side and "TFC" on the opposing side.

The Mekinist 2 mg film-coated tablets are pink, round, biconvex, with "GS" debossed on one side and "HMJ" on the opposing side.

The bottles also include a silica gel desiccant in a small cylinder-shaped container. The desiccant must be kept inside the bottle and must not be swallowed.

Registration number of the medicine in the Ministry of Health's National Drug Registry: Mekinist 0.5 mg: 154 95 34309

154 96 34322 DOR-Mek-PIL-0422-07 MEK APL MAR22 V3

- together with Mekinist that appear in the list below

Side effects when Mekinist and dabrafenib are taken together

(increase or decrease).

- Tell your doctor as soon as possible if you get any of these symptoms, either for the first time or if they get worse.

When you take Mekinist and dabrafenib together, you may experience any of the side effects given in the lists above, although their frequency may change

You may also suffer from additional side effects due to taking dabrafenib

- Please read the dabrafenib Package Leaflet for details of the side effects

- you may get when taking this medicine.

- The side effects that you may notice when you take Mekinist in combination
- with dabrafenib:

- Very common side effects (may appear in more than one in 10 users):
- Decreased appetite
- High blood pressure Bleeding, at various sites in the body, which may be mild or serious (haemorrhage)
- Constipation
- Joint pain, muscle pain or pain in the hands or feet
- · Swelling of the hands or feet (oedema peripheral)
- Fever
- Abnormal blood test results related to the liver Common side effects (may appear in 1-10 in 100 users): Infection of the urinary system · Skin effects including infection of the skin (cellulitis), inflammation of hair
- and swelling of the cuticles, skin rash with pus-filled blisters, cutaneous squamous cell carcinoma (a type of skin cancer), papilloma (a type of skin
- Blurred vision, eyesight problems, inflammation of the eye (uveitis)
 Heart pumping less efficiently Low blood pressure Localised tissue swelling
- Acne-like problems • Thickening of the outer layer of the skin (hyperkeratosis), patches of thick scaly, or crusty skin (actinic keratosis), chapping or cracking of the skin
- Common side effects that may show up in your blood tests: Low levels of white blood cells Decrease in number of red blood cells (anaemia), blood platelets (cells that help blood to clot), and a type of white blood cells (leukopenia)
- and skeletal muscles • Increase in enzymes produced by the liver Uncommon side effects (may appear in 1-10 in 1,000 users):
 • Appearance of new skin cancer (melanoma)
- (chorioretinopathy), separation of the light-sensitive membrane in the back of the eye (the retina) from its supporting layers (retinal detachment) and swelling around the eyes
- Inflammation of the kidneys Inflammatory disease mainly affecting the skin, lung, eyes and lymph nodes
- Side effects of unknown frequency (the frequency of these effects has not yet been established):
- Reporting side effects
- After first opening of the bottle: It should be kept below 30°C and used within 30 days.

 Do not throw away medicines via wastewater or household waste. Ask the

pharmacist how to throw away medicines which are no longer in use. These

6. Additional information

- Tablet core Mannitol (E Microcrystalline cellulose (E460) Hypromellose (E464) Croscarmellose sodium (E468)
- Titanium dioxide (E171) Polyethylene glycol Iron oxide yellow (E172)
- Iron oxide red (È172) What the medicine looks like and contents of the pack:
- The film coated tablets are supplied in opaque white plastic bottles with plastic closures. One bottle contains 30 tablets
- POB 7126 Tel Aviv
 - Revised in March 2022 according to MOH guidelines.

- Hypromellose (E464) Titanium dioxide (E171) Polyethylene glycol Polysorbate 80 (E433)
- Registration holder and importer's name and address: Novartis Israel Ltd..