Package leaflet: Information for the patient

Ezmekly 1 mg dispersible tablets
mirdametinib

▼This medicine is subject to additional monitoring. This will allow quick identification of new safety information. You can help by reporting any side effects you may get. See the end of section 4 for how to report side effects.

Read all of this leaflet carefully before you (or your child) start taking this medicine because it contains important information for you.
- Keep this leaflet. You may need to read it again.
- If you have any further questions, ask your doctor, pharmacist or nurse.
- This medicine has been prescribed for you (or your child) only. Do not pass it on to others. It may harm them, even if their signs of illness are the same as yours.
- If you get any side effects, talk to your doctor, pharmacist or nurse. This includes any possible side effects not listed in this leaflet. See section 4.

What is in this leaflet

1. What Ezmekly is and what it is used for

Ezmekly contains the active substance mirdametinib, and is a mitogen-activated protein kinase (MEK) inhibitor.

Ezmekly is used to treat plexiform neurofibromas in adults, adolescents, and children 2 years of age and older with a genetic condition called neurofibromatosis type 1 (NF1).

People with NF1 have a mutation (change) in the gene that encodes for the protein neurofibromin. This leads to the loss of this protein, which allows the tumours to grow on the nerves in the body. Ezmekly works by blocking certain proteins involved in the growth of these tumour cells, which is expected to shrink them.

If you have any questions about how Ezmekly works or why this medicine has been prescribed for you, ask your doctor.

2. What you (or your child) need to know before you take Ezmekly

Do not take Ezmekly
If you (or your child) are allergic to mirdametinib or any of the other ingredients of this medicine (listed in section 6).

Warnings and precautions
Talk to your doctor, pharmacist or nurse before taking Ezmekly, or if the following happen to you (or your child) while taking Ezmekly. Your doctor may decide to lower the dose, stop treatment temporarily or permanently:

Eye problems
Ezmekly can cause eye problems that can lead to blindness (see section 4 'Possible side effects'). A doctor (eye specialist) will have to check your vision (the vision of your child) before and regularly during treatment. Tell your doctor straight away if you get blurred vision, or any other changes to your vision during treatment.

Heart problems
Ezmekly can lower the amount of blood pumped by your heart (see section 4 'Possible side-effects'). In clinical studies of Ezmekly at the recommended dose, this did not cause any symptoms, however it can make you feel tired. Tell your doctor if you feel tired. Your doctor will check how well your heart works before and during your treatment with Ezmekly.

Skin rash
Skin rashes are very common with Ezmekly and can also become severe. Tell your doctor if you get a skin rash which may be bumpy or acne-like (see section 4 'Possible side effects'). Your doctor may provide guidance for managing your skin rash, including treatment (e.g. cream).

Pregnancy
Ezmekly is not recommended during pregnancy. Tell your doctor straight away if you or your partner becomes pregnant while taking this medicine. See 'Pregnancy, breast-feeding and fertility' section below.

Children younger than 2 years of age
Do not give Ezmekly to children younger than 2 years old. It has not been studied in this age group.

Other medicines and Ezmekly
Tell your doctor or pharmacist if you (or your child) are taking, have recently taken or might take any other medicines, including herbal medicines while receiving Ezmekly.

Pregnancy, breast-feeding and fertility
Ezmekly is not recommended during pregnancy. It may cause harm to your unborn baby.

If you are pregnant, think you may be pregnant or are planning to have a baby, ask your doctor for advice before taking this medicine. Your doctor may ask you to take a pregnancy test before starting treatment.

You or your partner should not become pregnant while taking this medicine. If you are able to become pregnant, you must use effective contraception (birth control). See 'Contraception - information for women and men' below.
If you or your partner become pregnant during treatment, tell your doctor straight away.

Contraception – information for women and men
If you are a woman who could become pregnant, you should use effective contraception while you are taking this medicine and for 6 months after the last dose. If you are a man whose partner could become pregnant, you should use effective contraception while you are taking this medicine and for 3 months after the last dose. It is not known if Ezmekly reduces how well hormonal contraceptives work. Please tell your doctor if you or your partner are taking a hormonal contraceptive, as your doctor may recommend using a second form of birth control (e.g. condoms).

Breast-feeding
It is not known if Ezmekly passes into breast milk. A risk to the breast-fed child cannot be excluded. Do not breast-feed if you are taking Ezmekly and for one week after the last dose of Ezmekly.

Fertility (ability to have children)
Ezmekly may reduce fertility in men and women. You should discuss this with your doctor before starting treatment.

Driving and using machines
Ezmekly can cause side effects, including fatigue (feeling tired or weak) and blurred vision (see section 4 'Possible side-effects'), that may affect your ability to drive or use machines. Do not drive or use machines if you feel tired or if you have problems with your vision (such as blurred vision).

Ezmekly contains sodium
Each dispersible tablet contains less than 23 mg sodium (main component of cooking/table salt), that is to say it is essentially 'sodium-free'.

3. How to take Ezmekly

Always take this medicine exactly as your doctor or pharmacist told you. Check with your doctor or pharmacist if you are not sure.

How much to take
Your doctor will work out the correct dose for you (or your child) based on your height and weight. The maximum recommended dose is 8 mg a day (4 mg every 12 hours). The doctor will tell you how many tablets of Ezmekly to take.

How to take
- Take Ezmekly by mouth twice a day, about 12 hours apart. You should take this medicine every day for 21 days, and then stop taking it for 7 days. Each 28-day period is called a treatment cycle.
- Take Ezmekly with or without food.
- You can continue taking this medicine in treatment cycles until your disease gets worse or side effects become unacceptable. If you get side effects, your doctor may decide to lower your dose, stop treatment temporarily or permanently (see section 2 'Warnings and precautions').

The dispersible tablets can be either:
1) Swallowed whole with drinking water. Do not split tablets.

OR

2) Dispersed in drinking water. Follow these instructions for dispersing the tablet:
   a) A dosing cup (and a 10 mL syringe if necessary), provided by your doctor or pharmacist can be used to prepare and administer the medicine.
   b) Disperse the prescribed number of dispersible tablets in a small amount (about 5-10 mL) of drinking water in the dosing cup.
   c) Swirl gently so that no lumps remain. Take care not to spill any medicine. The medicine will be white and cloudy.
   d) Administer the prepared medicine by mouth. Alternatively, the medicine can be drawn up from the dosing cup into a syringe and administered.
   e) After swallowing the suspension, there may be some medicine (residue) still inside the dosing cup or syringe that is difficult to see.
   f) Rinse the container with an additional small amount (about 5-10 mL) of drinking water and administer.
   g) Only water should be used to disperse the dispersible tablet.

OR

3) Administered via feeding tube
   a) Always talk to your doctor or pharmacist before administering Ezmkely suspension via a feeding tube. Your doctor, nurse or pharmacist should show you how to administer Ezmekly suspension via a feeding tube.
   b) Ezmekly suspension can be administered via a nasogastric (NG) or gastric (G) feeding tube of size French 8 or greater.
   c) Only use feeding tubes made of polyvinylchloride (PVC) or polyurethane (PUR).
   d) You may need an ENFIT adapter to connect the syringe to the feeding tube.

e) Follow steps above to disperse the tablet appropriately.
f) After dispersing the tablet in 5-10 mL of water, the oral suspension should be drawn up from the dosing cup into a syringe.
g) Slowly push the suspension into the feeding tube while holding the syringe in a horizontal position.
h) Use another 5-10 mL water to rinse the cup and draw the rinsate into the syringe and slowly push that through the feeding tube.
i) Flush the feeding tube according to the manufacturer's instructions immediately before and after administering Ezmekly suspension.

If you (or your child) are sick after taking Ezmekly
If you are sick (vomit) at any time after taking Ezmekly, do not take an extra dose. Take the next dose at the normal time.

If you (or your child) take more Ezmekly than you should
If you have taken more Ezmekly than you should, contact your doctor or pharmacist immediately.

If you (or your child) forget to take Ezmekly
If you forget to take a dose of Ezmekly, skip the missed dose and then take the next dose at the normal time. Do not take a double dose to make up for a forgotten dose.

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

If you (or your child) stop taking Ezmekly
Do not stop taking Ezmekly unless your doctor tells you as it may worsen your condition.

4. Possible side effects

Like all medicines, this medicine can cause side effects, although not everybody gets them. Contact your doctor if you experience any side effects. Your doctor may pause treatment with Ezmekly until your symptoms improve and/or they may reduce the dose you receive (see Section 3 'How to take').

Serious side effects
Eye problems (common: may affect up to 1 in 10 people)

Ezmekly can cause eye problems in adults and children. Some of these eye problems may lead to a blockage of the vein draining to the eye (retinal vein occlusion) or detachment of different layers of the eye (retinal pigment epithelial detachment). Tell your doctor straight away if you get blurred vision or any other changes to your sight during treatment. Your doctor may ask you to stop taking this medicine or send you to a specialist, if you develop symptoms that include:
- blurred vision
- other changes to your vision (such as reduced vision, coloured dots in your vision)

Tell your doctor straight away if you notice any of the serious side effects above.

Other side effects
Talk to your doctor if you get any of the following side effects.

<u>Adults</u>
Very common (may affect more than 1 in 10 people):
- skin rash with a flat discoloured area or raised acne-like bumps (dermatitis acneiform)
- diarrhoea
- increased blood levels of creatine phosphokinase, an enzyme found mainly in heart, brain, and skeletal muscle

- feeling sick (nausea)
- being sick (vomiting)
- pain in the muscles or bone
- feeling tired, weak or lacking energy (fatigue)
- stomach (abdominal) pain
- constipation
- rash
- headache
- increased levels of liver enzymes, as shown in blood tests
- swelling of the hands or feet (oedema peripheral)
- a decrease in the amount of blood that the heart is pumping (decreased left ventricular ejection fraction)
- dry skin
- itching
- hair loss or thinning (alopecia)

Common (may affect up to 1 in 10 people):
- decreased blood levels of neutrophils and leukocytes (types of white blood cells that help fight infections)
- mouth sores (stomatitis)
- dry mouth
- infection around the nails (paronychia)
- dry or itchy skin rash (eczema)
- changes in hair color
- changes in hair texture

Paediatrics
Very common (may affect more than 1 in 10 people):
- increased blood levels of creatine phosphokinase, an enzyme found mainly in heart, brain, and skeletal muscle
- diarrhoea
- skin rash with a flat discoloured area or raised acne-like bumps (dermatitis acneiform)
- pain in muscles or bone
- stomach (abdominal) pain
- feeling sick (nausea)
- being sick (vomiting)
- a decrease in the amount of blood that the heart is pumping (decreased ejection fraction)
- headache
- rash
- infection around the nails (paronychia)
- decreased blood levels of neutrophils and leukocytes (types of white blood cells that help fight infections)
- mouth sores (stomatitis)
- increased levels of liver enzymes as shown in blood tests
- dry skin
- hair loss or thinning (alopecia)
- dry or itchy skin rash (eczema)
- itching
- changes in hair color
- feeling tired, weak or lacking energy (fatigue)
- constipation

Common (may affect up to 1 in 10 people):
- changes in hair texture
- swelling of the hands or feet (oedema peripheral)

Reporting of side effects
If you get any side effects, talk to your doctor, pharmacist or nurse. This includes any possible side effects not listed in this leaflet. You can also report side effects directly via the Yellow Card Scheme Website: www.mhra.gov.uk/yellowcard or search MHRA Yellow Card in the Google Play or Apple App Store.

5. How to store Ezmekly

Keep this medicine out of the sight and reach of children.

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

Store below 30°C. Store in the original package to protect from light. After dispersing the tablet (s) in water, suspension is stable up to 6 hours.

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

6. Contents of the pack and other information

What Ezmekly contains
The active substance is mirdametinib.

Each dispersible tablet contains 1 mg mirdametinib.

The other ingredients are:

Microcrystalline cellulose (E460)
Croscarmellose sodium (E468) (see section 2 'Ezmekly contains sodium')
Sucralose (E955)
Magnesium stearate (E572)

Grape flavour:
Dried glucose liquid
Natural flavour
Modified corn starch (E1422)
Triacetin (E1518)

What Ezmekly looks like and contents of the pack

Ezmekly 1 mg dispersible tablets
Oval, white to off-white dispersible tablets 6 × 9 mm, debossed with 'S' on one side.

Ezmekly is packaged in plastic bottles, secured with child-resistant closures and aluminium foil induction seals. The bottles include cotton coil.

Ezmekly 1 mg dispersible tablets are provided in a carton box containing one bottle of 42 or 84 dispersible tablets.

Not all pack sizes may be marketed.

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This leaflet was last revised in 08/2026

This medicine has been given ‘conditional approval’. This means that there is more evidence to come about this medicine. The Agency will review new information on this medicine at least every year and this leaflet will be updated as necessary.