Special user information card

This card contains important safety information which you should be aware of before and during treatment with Cotellic (Cobimetinib) for skin cancer.

COTELLIC®
cobimetinib

For complete information about the medication, see the patient leaflet as approved by the Ministry of Health prior to use.

Cotellic is an anti-cancer medication that contains the active substance cobimetinib.
The medication is intended for the treatment of a type of skin cancer called metastatic or unresectable melanoma in combination with vemurafenib, in patients with a V600 mutation in a protein called BRAF.

How does the medication work?

The medication is targets a protein called "MEK", which is important for the regulation of cancer cell development. When Cotellic is administered in combination with vemurafenib (a medication that targets mutant BRAF protein), the development of the cancer is further slowed or stopped.

! Special warnings regarding the use of this medication

Before treatment with Cotellic, tell the doctor if you have any:

Bleeding (common side effects which may affect up to 1 in every 10 users).

Cotellic may cause serious bleeding, especially in the brain or stomach. Tell your doctor **immediately** if you have abnormal bleeding or any of the following symptoms: Headaches, dizziness, a feeling of weakness, abdominal pain, blood in the stool or black stools and if you vomit blood.

Eye (vision) disorders (very common side effects which may affect more than 1 in every 10 users).

Cotellic may cause eye disorders, some of which may be the result of a condition called serous retinopathy, caused by the accumulation of fluid under the retina.

Tell your doctor **immediately** if the following symptoms appear: Blurred vision, impaired or distorted vision, partial vision loss or any other change in your vision during the treatment. Your doctor will check your eyes in any case of new or worsening vision disorders during the treatment with Cotellic.

Heart disorders (common side effects which may affect up to 1 in every 10 users).

Cotellic may cause heart failure, manifested by a reduction in the quantity of blood pumped by your heart.

Your doctor will instruct you to perform tests before and during the treatment with Cotellic to examine the ability of your heart to pump blood. Tell your doctor **immediately** if you feel that your heart is pounding, beating rapidly or irregularly; or if you are suffering from dizziness, lightheadedness, shortness of breath, fatigue or swelling in the legs.

Liver disorders (very common side effects which may affect more than 1 in every 10 users).

During the treatment, Cotellic can cause a rise in the blood levels of certain liver enzymes. Your doctor will instruct you to perform blood tests to examine the levels of these enzymes and to monitor your liver function.

Muscle disorders (uncommon side effects which may affect up to 1 in every 100 users).

Cotellic can cause high levels of creatine phosphokinase, an enzyme present mainly in the muscle, heart and brain. This condition may be a sign of muscle damage (rhabdomyolysis). Your attending physician will refer you for blood tests to monitor this condition. Tell your attending physician **immediately** if you have any of the following symptoms: Muscle pain, muscle cramps, weakness, dark or red urine.

Diarrhea (a very common side effect which may affect more than 1 of 10 users).

Tell your doctor immediately if you have diarrhea. Severe diarrhea can cause loss of body fluids (dehydration). Follow your doctor's instructions in order to know what to do to prevent or treat diarrhea.

Reporting side effects
If any side effects appear, contact your doctor, pharmacist or nurse.

Side effects may be reported to the Ministry of Health via the Ministry of Health portal at www.health.gov.il or by entering the link: https://sideeffects.health.gov.il/
You may also report directly to Roche by email at: israel.drugsafety@roche.com

How to use the medication?

Tell your doctor or pharmacist if you are taking or have recently taken any other medications, including non-prescription medications and dietary supplements.

Always take this medication exactly as your doctor has instructed you.

Check with your doctor or pharmacist if you are not sure.

Standard dosage

The dosage and manner of treatment will be determined by the doctor only.
The usual standard dosage is 3 tablets (a total of 60 mg) per day.

- Take the tablets every day for 21 days (this is called the "treatment period").
- After 21 days, do not take Cotellic tablets for 7 days. During the 7 days you are off Cotellic treatment, continue taking vemurafenib as your doctor has instructed you.
- Start your next Cotellic treatment period (21 treatment days) after the 7-day break.
- If you have side effects, your doctor may decide to reduce the dosage, discontinue the treatment temporarily or discontinue the treatment permanently. Always take this medication exactly as your doctor has instructed you.

Taking the medication

- Swallow the tablets whole, with water.
- The medication can be taken with or without food.
- Refrain from taking the medication with grapefruit juice.

If you vomit after taking the medication, do not take another dose of Cotellic on the same day. Continue taking the medication as scheduled the day after.

If you have accidentally taken Cotellic at a higher dosage than that prescribed by your doctor, contact your doctor immediately. Take the medication package and this leaflet with you.

If you forgot to take Cotellic at the scheduled time, follow the instructions below:

- If there are more than 12 hours before the time you should take the next dose, take the forgotten dose as soon as you remember.
- If there are less than 12 hours before the time you should take the next dose, do not take the forgotten dose - skip it. Wait and take the next dose at the regular time.
- If you forgot to take the medication at the scheduled time, do not take a double dose to compensate for the forgotten dose.

Do not stop the treatment with the medication without consulting your doctor, even if your health has improved.

Tests and follow up

- Your doctor will instruct you to perform tests before and during the treatment with Cotellic to examine the ability of your heart to pump blood.
- Your doctor will instruct you to perform blood tests to examine the levels of liver enzymes and to monitor your liver function.
- Your doctor will check your eyes to examine you for new or worsening vision disorders during the treatment with Cotellic.

Roche Pharmaceuticals (Israel) Ltd.
6 HaHarash St.
Tel: 09-9737777
POB 6391, Hod Hasharon 4524079
www.roche.co.il

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