

IVOSIDENIB for biliary tract cancers

If your doctor recommended the administration of ivosidenib to treat your cancer, here is some important information about this medication and some aspects related to it.

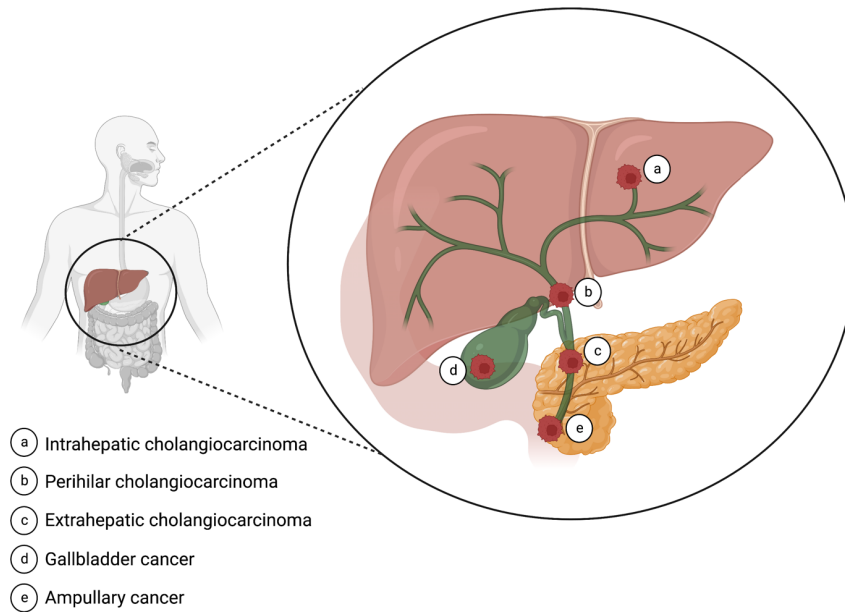
Biliary tract cancer

Biliary tract cancers develop from cells (called cholangiocytes) that normally make up the lining of the biliary system (depicted in green in the picture below). The biliary ducts are small tubes that connect the liver and gallbladder to the small bowel and normally carry the bile, which is a fluid that helps to digest food.

Biliary tract cancers include cholangiocarcinoma, gallbladder cancer and ampullary cancer:

- **Cholangiocarcinoma** - arises from the bile ducts. This cancer is further classified into three different types, depending on which part of the bile duct the cancer develops:
 - *intrahepatic cholangiocarcinoma* - the cancer develops in the ducts inside the liver
 - *perihilar cholangiocarcinoma* - it develops in the ducts just outside the liver
 - *distal/extrahepatic cholangiocarcinoma* - it develops in the ducts further away from the liver
- **Gallbladder cancer** - arises from the cells of the gallbladder
- **Ampullary cancer** - arises from the join between the bile ducts and the small bowel, called ampulla of Vater

You have been given this leaflet because you have been diagnosed with a biliary tract cancer. Your oncologist can discuss with you which biliary tract cancer you have and how extended it is.



What is IVOSIDENIB?

Ivosidenib is a targeted drug treatment for biliary tract cancers, and it is used when the cancer has a change in a gene called isocitrate dehydrogenase-1 (IDH-1), which leads to the differentiation of normal cells into malignant cells. As an inhibitor of IDH-1, ivosidenib blocks cancer cell from growing and proliferating.

How is IVOSIDENIB administered?

You will take ivosidenib orally as tablets. Tablets need to be swallow with a glass of water and can be taken with or without food (you should avoid high-fat meals). You should take the tablets at the same time, every day. You should not crush or chew the tablets. If you miss a dose, you should talk to your doctor or nurse.

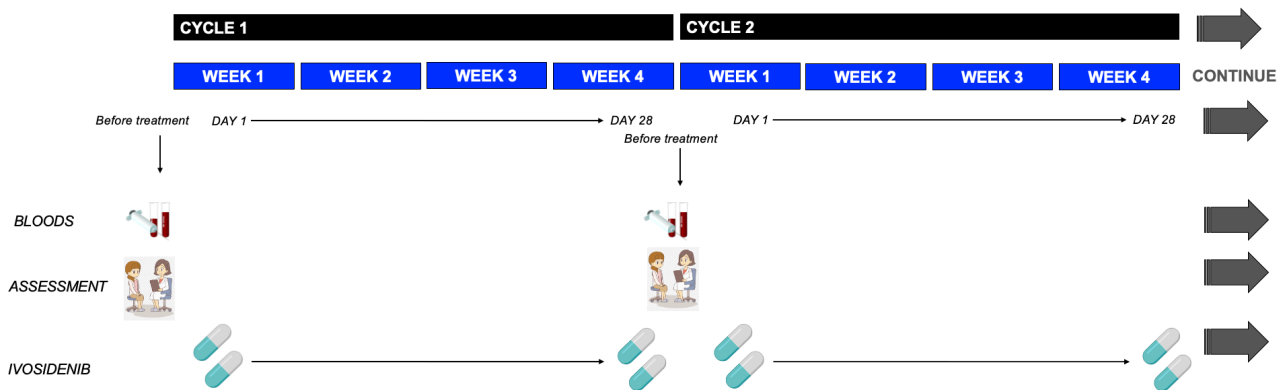
Your doctor will tell you the exact dosage of ivosidenib you will receive. Before you start this treatment, your doctor will collect information on your general health and underlying conditions.

You will see a cardiologist before starting with the treatment, in order to assess your cardiovascular function. An electrocardiography (ECG) will be perform to evaluate the electrical activity of your heart.

Schedule of IVOSIDENIB treatment

You will receive ivosidenib in cycles of treatment. The standard schedule of ivosidenib treatment follows a four-week cycle. You should take ivosidenib once a day for 28 days. Before each cycle, you will have a blood test and your oncological team will check whether you are fit enough for the treatment. It is important that you report any symptom(s) and problem(s) since your last session so that the dose can be adjusted for you. The doctor may ask you to have the bloods taken at your GP practice the day before your treatment appointment.

IVOSIDENIB once daily in continuous 28-day cycles



Duration of IVOSIDENIB treatment

The doctor will discuss the duration of your treatment with you. Each ivosidenib cycle will last for 4 weeks. If you are tolerating the treatment well, you will receive ivosidenib for at least 3 months before you will repeat scan to understand if the treatment is working. If the first 3 months scan shows your cancer is stable or shrinking, you may continue the treatment. You will continue taking ivosidenib for as long as the treatment is helping you. You will repeat a scan every 3 months.

What are the side effects of the IVOSIDENIB treatment?

There are known side effects with this treatment, but you may not experience any of them. This does not mean that the treatment is not working. There is no relationship between the presence or severity of side effects and the efficacy of the treatment.

You may get some of the side effects, but you are unlikely to get them all. Importantly, side effects are often predictable in terms of their onset, duration, and severity; they are almost always reversible and should completely go away after treatment.

However, their frequency and severity may vary from person to person.

Fortunately, there are many medications available that can be taken during treatment to control and minimise the impact of any side effects you may have.

Some of the most common side effects related to treatment (IVOSIDENIB)

Low number of red blood cells: red blood cells normally carry oxygen to every part of your body. If the number of red blood cells is too low (this is called *anaemia*) you can feel tired and breathless. If the *anaemia* is severe, you will need a blood transfusion.

Fatigue: a very common side effect, which may increase during the treatment course.

Nausea: sometimes accompanied by vomiting but is usually well controlled by anti-sickness drugs (injections and tablets). It is important to take anti-sickness medicines even if you don't feel sick, because it is easier to prevent sickness rather than treat it once it has started. Contact your doctor or nurse straight away if you've been feeling sick more than once in a day.

Diarrhoea: you need to get in touch with your team if you have 4 or more loose poos (stools) in 24 hours. Your doctor will give you anti-diarrhoea medicines. Remember to drink plenty of water to replace fluids lost. You should eat low-fibre food and avoid raw fruits, fruit juice, cereals and vegetables. It can help to also avoid alcohol, caffeine, dairy products and high-fat-foods.

Poor appetite: don't worry if you don't eat much for a day or two. Also, your taste can change.

Build-up of fluid: You may put on weight and stomach may swell. The swelling will get better after the end of treatment.

Skin changes: your skin may be drier or develop a rash.

Changes in liver function test: you will have blood tests before each cycle in order to check your liver function.

Some serious side effects related to treatment (IVOSIDENIB)

QTc prolongation: this is a change in the electrical activity of your heart. QTc prolongation can cause irregular heartbeats. Your doctor and nurse will check the electrical activity of your heart before and during treatment with ivosidenib. If you feel dizzy, lightheaded or faint, you should contact immediately your hospital.

Guillain-Barré Syndrome: this syndrome has been reported in people treated with ivosidenib. Your doctor will monitor your nervous system functions and will permanently stop the treatment if you develop this syndrome. Tell your doctor or nurse if you have any of these symptoms:

- weakness or tingling feeling in your legs, arms, or upper body
- numbness and pain on one side or both sides of your body
- any changes in the ability to see, touch, hear, taste
- burning or prickling sensation
- difficult breathing

Medication to help control side effects

Make sure to let your doctor know of any side effects; there are useful medications to control the symptoms.

Should I continue to take all my usual medications?

Yes, you have to keep taking all your usual medications. Please report to your oncology team all the medications you are taking, so that they can advise. Cancer drugs can have interaction with some other medicines and herbal products. You should not eat grapefruit or drink its juice because it can react with the drug.

Can I have the flu and COVID vaccination?

Yes, it is advised you have the flu and COVID vaccination before you start your treatment. If you have already started your treatment, please ask your doctor who can advise on the best timing to have the vaccination.

Tips during treatment

- Drink plenty of fluids (at least 2 litres per day)
- Maintain good nutrition. Eating small frequent meals may help to reduce nausea. You should avoid fatty or fried foods. You can take anti-sickness drugs if you need.
- Avoid sun exposure. Wear SPF 15 (or higher) sunblock and protective clothing.
- Get plenty of rest.
- Keep at home the medications you may need to help control the symptoms.
- Before starting treatment, tell your doctor about any medication you are taking. Sometimes side effects may be related to your medicines.
- If your symptoms are severe or do not improve after 24 hours, do not hesitate to contact the hospital.
- Do not receive any kind of vaccination without your doctor's approval while taking treatment.
- If you are a woman of childbearing age:
 - inform your doctor if you are pregnant or may be pregnant prior to starting this treatment
 - avoid getting pregnant during treatment
 - do not breastfeed during treatment

When to contact the hospital?

If your symptoms are severe or do not improve after 24 hours, do not hesitate to contact the hospital.

Where can I get more information?

If you want to get more information in this field you can visit **ESMO** website for *Biliary tract cancer: Guide for Patients* and on the **AMMF** *The Cholangiocarcinoma Charity* website.

You can find the related link below:

<https://www.esmo.org/for-patients/patient-guides/biliary-tract-cancer>

<https://ammf.org.uk/cholangiocarcinoma/>

You can also visit the Precision-BTC-Network COST action website:

<https://precision-btc.eu/info/patientmaterials>