

B. PACKAGE LEAFLET

Package leaflet: Information for the patient

Lonsurf 15 mg/6.14 mg film-coated tablets

Lonsurf 20 mg/8.19 mg film-coated tablets
trifluridine/tipiracil

▼ 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 experience. See the end of Section 4 for how to report side effects.

Read all of this leaflet carefully before you 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 or pharmacist.
- This medicine has been prescribed for you 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 experience any side effects, talk to your doctor or pharmacist. This includes any possible side effects not listed in this leaflet. See Section 4.

What is in this leaflet

1. What Lonsurf is and what it is used for
2. What you need to know before you take Lonsurf
3. How to take Lonsurf
4. Possible side effects
5. How to store Lonsurf
6. Contents of the pack and other information

1. What Lonsurf is and what it is used for

Lonsurf is a type of cancer chemotherapy which belongs to the group of medicines called "cytostatic antimetabolite medicines".

Lonsurf contains two different active substances: trifluridine and tipiracil.

- Trifluridine stops the growth of cancer cells.
- Tipiracil stops the trifluridine from being broken down by the body, helping trifluridine to work longer.

Lonsurf is used to treat adults with colon or rectal cancer - sometimes called 'colorectal' cancer.

- It is used when the cancer has spread to other parts of the body.
- It is used when other treatments have not worked - or when other treatments are not suitable for you.

2. What you need to know before you take Lonsurf

Do not take Lonsurf :

- if you are allergic to trifluridine or tipiracil or any of the other ingredients of this medicine (listed in section 6).

Do not take Lonsurf if the above applies to you. If you are not sure, talk to your doctor before taking Lonsurf.

Warnings and precautions

Talk to your doctor or pharmacist before taking Lonsurf if:

- you have kidney problems
- you have liver problems

If you are not sure, talk to your doctor or pharmacist before taking Lonsurf.

Treatment may lead to the following side effects (see section 4):

- a reduced number of certain types of white blood cells (neutropenia) which are important for protecting the body against bacterial or fungal infections. As a consequence of neutropenia, fever (febrile neutropenia) and blood infection (septic shock) may occur
- a reduced number of red blood cells (anaemia)
- a reduced number of platelets in the blood (thrombocytopenia) which are important to stop bleeding and work by clumping and clotting blood vessel injuries
- gastrointestinal problems.

Tests and checks

Your doctor will do blood tests before each cycle of Lonsurf. You start a new cycle every 4 weeks. The tests are needed because Lonsurf can sometimes affect your blood cells.

Children and adolescents

This medicine is not indicated for use in children and adolescents below the age of 18 years. This is because it may not work or be safe.

Other medicines and Lonsurf

Tell your doctor or pharmacist if you are taking, have recently taken or might take any other medicines. This includes medicines obtained without a prescription and herbal medicines. This is because Lonsurf can affect the way some other medicines work. Also some other medicines can affect the way Lonsurf works.

In particular tell your doctor or pharmacist if you are taking medicines used for treatment of HIV, such as zidovudine. This is because zidovudine may not work as well if you are taking Lonsurf. Talk to your doctor about whether to switch to a different HIV medicine.

If the above applies to you (or you are not sure), talk to your doctor or pharmacist before taking Lonsurf.

Pregnancy and breast-feeding

If you are pregnant or breast-feeding, or if you think you might be pregnant or are planning to have a baby, ask your doctor or pharmacist for advice before taking this medicine. Lonsurf may harm your unborn baby.

If you become pregnant, you and your doctor will have to decide if the benefits of Lonsurf are greater than the risk of harm to the baby.

Do not breast-feed if you are taking Lonsurf as it is not known whether Lonsurf passes into the mother's milk.

Contraception

You must not become pregnant while taking this medicine. This is because it may harm your unborn baby.

You and your partner should use effective methods of contraception while taking this medicine. You should also do this for 6 months after you stop taking the medicine. If you or your partner becomes pregnant during this time, you must talk to your doctor or pharmacist straight away.

Driving and using machines

It is not known whether Lonsurf changes your ability to drive or use machines. Do not drive or use any tools or machines if you experience symptoms that affect your ability to concentrate and react.

Lonsurf contains lactose

If you have been told by your doctor that you have an intolerance to some sugars, contact your doctor before taking this medicine.

3. How to take Lonsurf

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

How much to take

- Your doctor will decide the right dose for you - the dose depends on your weight and height.
- Your doctor will tell you how many tablets to take each time.
- You will take a dose 2 times a day.

How to take

You will take Lonsurf for 10 days during the first 2 weeks, and then have 2 weeks off. This 4 week period is called a 'cycle.' The specific dosing schedule is as follows:

- **Week 1**
 - take the dose 2 times a day for 5 days
 - then have 2 days off - no medicine
- **Week 2**
 - take the dose 2 times a day for 5 days
 - then have 2 days off - no medicine
- **Week 3**
 - No medicine
- **Week 4**
 - No medicine

You will then start again with another cycle of 4 weeks following the above pattern.

Lonsurf comes in two strengths. Your doctor may prescribe both strengths for your prescribed dose.

Taking this medicine

- Take this medicine by mouth.
- Swallow the tablets whole with a glass of water.
- Take within 1 hour after your morning and evening meal.
- Wash your hands after handling this medicine.

If you take more Lonsurf than you should

If you take more Lonsurf than you should, talk to a doctor or go to a hospital straight away. Take your pack(s) of medicine with you.

If you forget to take Lonsurf

- If you forget a dose, talk to your doctor or pharmacist.
- 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.

4. Possible side effects

Like all medicines, this medicine can cause side effects, although not everybody gets them. The following side effects may happen with this medicine:

Serious side effects

Tell your doctor immediately if you notice any of the following serious side effects (many of the side effects are shown in laboratory tests - for example those affecting your blood cells):

- Neutropenia (*very common*), febrile neutropenia (*common*) and septic shock (*uncommon*). The signs include chills, fever, sweating or other sign of infection (see section 2).
- Anaemia (*very common*). The signs include feeling short of breath, tiredness or looking pale (see section 2).
- Vomiting (*very common*) and diarrhoea (*very common*), which may lead to a dehydration if severe or persistent.

- Severe gastrointestinal problems: abdominal pain (*common*), ascites (*uncommon*), colitis (*uncommon*), acute pancreatitis (*uncommon*), ileus (*uncommon*) and subileus (*uncommon*). The signs include intense stomach or abdominal pain that can be associated with vomiting, blocked or partly blocked bowel, fever or swelling of the abdomen.
- Thrombocytopenia (*very common*). The signs include unusual bruising or bleeding (see section 2).
- Pulmonary embolism (*uncommon*): blood clots in lungs. The signs include shortness of breath and pain in the chest or in the legs.

Some of these serious side effects may lead to death.

Other side effects

Tell your doctor if you notice any of the following side effects. Many of the side effects are shown in laboratory tests - for example those affecting your blood cells. Your doctor will be looking out for these side effects in your test results.

Very common: may affect more than 1 in 10 people:

- decreased appetite
- feeling very tired (fatigue)
- feeling sick (nausea)
- reduced white blood cells called leucocytes - can increase your risk for infection

Common: may affect up to 1 in 10 people:

- fever
- hair loss
- weight loss
- changes in taste
- constipation
- feeling dizzy, headache
- feeling generally out of sorts (malaise)
- low levels of albumin or total protein in the blood
- increased bilirubin in your blood - can cause yellowing of skin or eyes
- increased number of white blood cells called monocytes
- reduced number of white blood cells called lymphocytes - can increase your risk for infection
- swelling in your hands or legs or feet
- redness, swelling, pain on the palms of your hands and soles of your feet (hand-foot syndrome)
- feeling of numbness or pins and needles in hands or feet
- mouth pain or problems
- swelling of mucous membranes - this could be inside the nose, mouth, throat, eyes, vagina, lungs or gut
- increased liver enzymes
- protein in your urine
- rash, itchy or flaky skin, skin flushing
- cough, feeling short of breath, infection of the sinuses, throat, airway or lungs, chest infections
- problem with sleeping or falling asleep

Uncommon: may affect up to 1 in 100 people

- low or high blood pressure
- blood clots, e.g. in the brain or legs
- blood test results indicating problems with clotting making you bleed more easily
- more noticeable heart-beat, chest pain
- abnormal increase or decrease in heart rate
- changes in your heart trace (ECG - electrocardiogram)
- increased white blood cells
- increased lactate dehydrogenase level in your blood
- low levels of phosphates, sodium, potassium or calcium in your blood
- reduced white blood cells called granulocytes or monocytes - can increase your risk for infection

- high blood sugar (hyperglycaemia), increased salt, urea, creatinine and potassium in your blood
- blood test result indicating inflammation (C-Reactive Protein increased)
- ear pain
- feeling of spinning (vertigo)
- runny or bloody nose, sinus problems
- sore throat, hoarse voice, problems with your voice
- redness, itching of the eye, eye infections, watery eyes
- dry eyes
- vision troubles as blurred vision, double vision, decreased vision, cataracts
- dehydration
- bloating, passing gas, indigestion
- pain or inflammation in upper or lower part of digestive tract
- inflammation, swelling or bleeding in your bowel
- inflammation and infection in your gut
- inflammation or increased acid in your stomach or gullet, reflux
- painful tongue, polyps inside your mouth, mouth ulcers, retching
- bad breath, tooth decay, tooth or gum problems, bleeding gums, gum infections
- dry skin
- swelling or pain in your joints or big toes
- pain or feeling heavy in your arms or legs
- pain, including pain from the cancer
- bone pain, muscle pain, muscle weakness or spasms, pain in tendons, nerves or ligaments
- feeling of being cold
- shingles (pain and vesicular rash on skin over nerve tracts affected by nerve inflammation from herpes zoster virus)
- liver disorder
- inflammation or infection of bile ducts, increase in the diameter of the bile duct
- kidney failure
- infections: bacterial, viral or fungal
- inflammation or infection in your bladder
- changes in urine test, blood in urine
- problems passing water (urine retention), loss of bladder control (incontinence)
- athlete's foot - fungal infection of feet, yeast infections
- accumulation of fluid in the lungs
- changes in the menstrual cycle
- anxiety
- passing out (syncope)
- burning sensation, unpleasant, increased or loss of sense of touch and other non-severe neurological troubles
- raised itchy rash, red skin, blisters, skin sloughing off, hives, acne
- sweating more than normal, sensitivity to light, nail problems

Reporting of side effects

If you experience any side effects, talk to your doctor or pharmacist. This includes any possible side effects not listed in this leaflet. You can also report side effects directly via [the national reporting system listed in Appendix V](#). By reporting side effects you can help provide more information on the safety of this medicine.

5. How to store Lonsurf

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 outer carton or blister tray after "EXP." The expiry date refers to the last day of that month.

This medicine does not require any special storage conditions.

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

6. Contents of the pack and other information

What Lonsurf contains

Lonsurf 15 mg tablet

- The active substances are trifluridine and tipiracil. Each tablet contains 15 mg trifluridine and 6.14 mg tipiracil
- The other ingredients are:
 - Tablet core: lactose monohydrate, starch pregelatinised (Maize) and stearic acid.
 - Film coat: hypromellose, macrogol (8000), titanium dioxide (E171), and magnesium stearate.
 - Printing ink: shellac, iron oxide red (E172), iron oxide yellow (E172), titanium dioxide (E171), indigo carmine aluminium lake (E132), carnauba wax and talc.

Lonsurf 20 mg tablet

- The active substances are trifluridine and tipiracil. Each tablet contains 20 mg trifluridine and 8.19 mg tipiracil.
- The other ingredients are:
 - Tablet core: lactose monohydrate, starch pregelatinised (Maize) and stearic acid.
 - Film coating: hypromellose, macrogol (8000), titanium dioxide (E171), iron oxide red (E172) and magnesium stearate.
 - Printing ink: shellac, iron oxide red (E172), iron oxide yellow (E172), titanium dioxide (E171), indigo carmine aluminium lake (E132), carnauba wax and talc.

What Lonsurf looks like and contents of the pack

- Lonsurf 15 mg is a white, biconvex, round, film-coated tablet, printed with “15” on one side and “102” and “15 mg” on the other side in grey ink.
- Lonsurf 20 mg is a pale red, biconvex, round, film-coated tablet, printed with “20” on one side and “102” and “20 mg” on the other side in grey ink.

Each pack contains 20 film-coated tablets (2 blisters of 10 tablets each) or 40 film-coated tablets (4 blisters of 10 tablets each), or 60 film-coated tablets (6 blisters of 10 tablets each).

Not all pack sizes may be marketed.

Marketing Authorisation Holder

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Other sources of information

Detailed information on this medicine is available on the European Medicines Agency web site:
<http://www.ema.europa.eu>.