



DASATINIB NEAPOLIS®

DASATINIB 20 mg-50mg-70mg-140mg

Film-coated tablet - DASATINIB

Read 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, pharmacist or nurse.
- 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 get any side effects, talk to your doctor, pharmacist or nurse. This applies to any side effect that is not mentioned in this leaflet. See section 4.

What is in this leaflet:

1. What is DASATINIB NEAPOLIS and in which case is it used ?
2. What you need to know before you take DASATINIB NEAPOLIS ?
3. How to take DASATINIB NEAPOLIS ?
4. Possible side effects ?
5. How to store DASATINIB NEAPOLIS ?
6. Content of the packaging and other information

1. What is DASATINIB NEAPOLIS and in which case is it used ?

DASATINIB NEAPOLIS contains the active substance dasatinib. This treatment is used to treat chronic myeloid leukemia (CML) in adults, adolescents and children at least 1 year old. Leukemia is a cancer affecting the white blood cells. These cells usually help the body to fight against infections. People with CML have an uncontrolled proliferation of white blood cells called granulocytes. DASATINIB NEAPOLIS helps fight against the proliferation of these leukemia cells.

DASATINIB NEAPOLIS is also used to treat the acute lymphoblastic leukemia (ALL) with chromosome Philadelphia (Ph +) in adult patients, adolescents and children aged at least one year, and CML in lymphoid blast phase, in adult patients for whom previous treatments are not beneficial. In people with ALL, white cells called lymphocytes were growing too fast and too long alive. DASATINIB NEAPOLIS helps fight against the proliferation of these leukemia cells.

If you have any questions about how DASATINIB NEAPOLIS works or why it has been prescribed for you, ask your doctor.

2. What you need to know before you take DASATINIB NEAPOLIS ?

Never take DASATINIB NEAPOLIS

- if you are **allergic** to dasatinib or to one of the other components contained in this medicine listed in section 6.

If you think you are allergic, ask for your doctor advice.

Warnings and precautions

Talk to your doctor or pharmacist before taking DASATINIB NEAPOLIS

- if you are taking **blood-thinning medicines** or medicines to prevent blood clots (see “Other medicines and DASATINIB NEAPOLIS”)
- if you have liver problems, heart problems, or if you have ever had any.
- if you have **breathing difficulty, chest pain or cough** while taking DASATINIB NEAPOLIS : this may be a sign of fluid in the lungs or chest (may be more common in patients 65 years of age and older), or a sign of changes in the blood vessels that bring blood to the lungs.
- if you have ever had or may now have hepatitis B. In deed, DASATINIB NEAPOLIS could reactivate your hepatitis B which can be fatal in some cases. Patients **will** be closely monitored by their doctor for any sign of infection before starting treatment.
- if you have bruising (bruising on the skin), bleeding, fever, fatigue and confusion while taking DASATINIB NEAPOLIS, contact your doctor. This can be a sign of damage to the blood vessels called thrombotic microangiopathy (TMA).

Your doctor **will** monitor you regularly to check that treatment with DASATINIB NEAPOLIS is working as expected. You **will** also have regular blood tests throughout the duration of treatment with DASATINIB NEAPOLIS.

Children and adolescents

Do not give this medication to children under one year of age. Experience with its use is limited in this age group. Bone growth and development will be closely monitored in children taking DASATINIB NEAPOLIS.

Other medicines and DASATINIB NEAPOLIS

Tell your doctor or pharmacist if you are taking, have recently taken or might take any other medicines. DASATINIB NEAPOLIS is mainly eliminated through the liver. Some medicines may interfere with the effect of DASATINIB NEAPOLIS when taken together.

These medicines should not be administered with DASATINIB NEAPOLIS :

- ketoconazole, itraconazole - these are **antifungal medicines**
- erythromycin, clarithromycin, telithromycin - these are **antibiotics**
- ritonavir - this is **an antiviral medicine**
- phenytoin, carbamazepine, phenobarbital - these are **treatments for epilepsy**
- rifampicin - this is a treatment for **tuberculosis**
- famotidine, omeprazole - these are medicines that **block the secretion of stomach acid**
- St. John's Wort - an herbal preparation obtained without a prescription used to treat **depression** and other conditions (also known as *Hypericum perforatum*).

Do not take medicines that neutralize stomach acids (antacids such as aluminum hydroxide / magnesium hydroxide) within 2 hours before or after taking DASATINIB NEAPOLIS.

Tell your doctor if you are taking medicines that thin the blood or prevent blood clots.

DASATINIB NEAPOLIS with food and drinks

Do not take DASATINIB NEAPOLIS with grapefruit or grapefruit juice.

Pregnancy and breastfeeding

If you are pregnant or think you may be pregnant, tell your doctor immediately.

DASATINIB NEAPOLIS should not be used during pregnancy unless clearly needed. Your doctor will discuss with you the potential risks of taking DASATINIB NEAPOLIS during pregnancy.

It is recommended for men and women treated DASATINIB NEAPOLIS to use effective contraception throughout the treatment period.

If you are breastfeeding, tell your doctor. You should stop breastfeeding while taking DASATINIB NEAPOLIS.

Driving and using machines

Take special care in driving and using machines if you suffer from side effects such as dizziness or blurred vision.

DASATINIB NEAPOLIS contains lactose.

If your doctor has told you that you have an intolerance to certain sugars, tell your doctor before taking DASATINIB NEAPOLIS.

3. How to take DASATINIB NEAPOLIS ?

DASATINIB NEAPOLIS can only be prescribed for you by a doctor who has experience in the treatment of leukemia. Always take this medicine exactly as your doctor has told you. Check with your doctor or pharmacist if in doubt. DASATINIB NEAPOLIS is prescribed for adults and children aged at least 1 year.

The recommended starting dose for adult patients with chronic phase CML is 100 mg once daily.

The recommended starting dose for adult patients with accelerated phase or blast crisis with CML or for patients with Ph + ALL is 140 mg once daily.

The dosage in children with chronic phase CML or Ph + ALL is based on weight. DASATINIB NEAPOLIS is taken orally once a day as DASATINIB NEAPOLIS tablets. DASATINIB NEAPOLIS tablets are not recommended for patients weighing less than 10 kg. Powder for oral suspension should be used for patients weighing less than 10 kg and patients who cannot swallow the tablets.

A change in dosage may be necessary when switching from one presentation to another (Tablets and powder for oral suspension), so you should not switch between them.

The doctor will decide on the proper presentation and dosage depending on your weight, potential side effects and treatment response. The starting dose of DASATINIB NEAPOLIS for children is calculated based on body weight as indicated below :

Body weight (kg)	Daily dosage (mg)
from 10 to less than 20 kg	40 mg
from 20 to less than 30 kg	60 mg
from 30 to less than 45	70 mg
at least 45 kg	100 mg

The tablet is not recommended for patients weighing less than 10 kg ;

There is no dosage recommendation for DASATINIB NEAPOLIS in children less than 1 year of age. Depending on your response to treatment, your doctor may suggest increasing or decreasing the doses or briefly stopping treatment. For higher or lower doses, you may need to take a combination of different dosages.

How to take DASATINIB NEAPOLIS

Take your tablets at the same time every day. Swallow the tablets as they are. Do not crush, cut or chew them. Do not take scattered tablets. You may not be sure that you are getting the correct dose if you crush, cut or scatter the tablets. DASATINIB NEAPOLIS tablets can be taken with or without food.

Special handling instructions for DASATINIB NEAPOLIS

The tablets are unlikely to break. However, if this happens, people other than the patient himself should wear gloves when handling DASATINIB NEAPOLIS.

Duration of treatment with DASATINIB NEAPOLIS

Take DASATINIB NEAPOLIS daily until your doctor tells you to stop. Make sure you take DASATINIB NEAPOLIS for as long as you will be prescribed.

If you take more DASATINIB NEAPOLIS than you should

If you accidentally take more tablets than necessary, tell your doctor **immediately**. You may need medical supervision.

If you forget to take DASATINIB NEAPOLIS

Do not take a double dose to make up for the dose you forgot to take. Take the next dose at the usual time.

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

4. Possible side effects ?

Like all medicines, this medicine can cause side effects, although not everybody gets them.

• The following symptoms may be manifestations of serious side effects

- If you have chest pain, difficulty breathing, if you cough or if you faint
- If you are **bleeding unexpectedly or if you have bruises** without injury
- If you notice blood in vomit, stools or urine, or if you have black stools
- If you have **signs of infection** such as fever or severe chills
- If you have fever, pain in the mouth or throat, blisters or peeling of the skin and / or mucous membranes

Tell your doctor immediately if you notice any of these effects.

Very common side effects (may affect more than 1 in 10 people)

- **Infections** (including bacterial, viral and fungal infections)
- **Heart and lungs** : difficulty breathing
- **Digestive disorders** : diarrhea, feeling nauseous (nausea, vomiting)
- **Skin, hair, eyes, general** : rash, fever, swelling of the face, hands and feet, headache, fatigue or weakness, bleeding
- **Pain** : pain in the muscles (during or after stopping treatment), stomach pain (abdominal)

DASATINIB NEAPOLIS®

DASATINIB 20 mg-50mg-70mg-140mg

Film-coated tablet - DASATINIB

- **Examinations may reveal** : low platelet count, low white blood cell count

(neutropenia), anemia, fluid around the lungs.

Common side effects (may affect up to 1 in 10 people)

- **Infections** : pneumonia, viral herpes infection (including cytomegalovirus - CMV), upper respiratory tract infection, severe blood or tissue infection (some uncommon cases with fatal outcome)
- **Heart and lung** : palpitations, irregular heartbeat, congestive heart failure, weakness of the heart muscle, high blood pressure, increased blood pressure in the lungs, cough
- **digestive disorders** : disorders of appetite, taste disturbance, swelling and stomach distension (abdomen), inflammation of the colon, constipation, heartburn, mouth ulceration, weight gain, weight loss, gastritis
- **Skin, hair, eyes, general** : tingling of the skin, itching, dry skin, acne, inflammation of the skin, persistent noise in the ears, hair loss, excessive sweating, visual disturbances (including blurred vision and visual discomfort), dry eyes, bruising, depression, insomnia, hot flushes, dizziness, bruising (blue), anorexia, drowsiness, generalized edema
- **Pain** : joint pain, muscle weakness, chest pain, pain around the hands and feet, chills, stiffness in muscles or joints, muscle spasm
- **The examinations can reveal** : fluid around the heart, fluid in the lungs, arrhythmia, febrile neutropenia, gastrointestinal bleeding, high levels of uric acid in the blood

Uncommon side effects (may affect up to 1 in 100 people)

- **Heart and lung** : heart attack (some fatal), inflammation of the membrane surrounding the heart, irregular heartbeat, chest pain due to lack of blood supply to the heart (angina), low blood pressure, narrowing of the bronchi which may cause difficulty breathing, asthma, increased blood pressure in the arteries (blood vessels) of the lungs
- **Digestive disorders** : inflammation of the pancreas, peptic ulcer, inflammation of the digestive tract, bloating of the stomach (abdomen), tearing of the skin of the anal canal, difficulty swallowing, inflammation of the gallbladder, obstruction of the bile ducts, gastroesophageal reflux disease (reflux acid and other stomach contents in the throat)

• **Skin, hair, eyes, general** : allergic reaction including a pressure pain, sensitive and red nodules on the skin (erythema nodosum), anxiety, confusion, mood changes, decreased libido, fainting, tremor, Inflammation of the eye may cause redness or pain, a skin disease characterized by sensitivity, redness, and well-defined spots with abrupt onset of fever and increased levels of white blood cells (neutrophilic dermatosis), hearing loss, sensitivity to light, visual disturbances, lacrimation, change in skin color, inflammation of fatty tissue under the skin, skin ulcer, blisters on the skin, changes in nails, hair disorders, hand-foot syndrome, renal impairment, frequent urination, enlarged breasts in men, menstrual disorders, uneasiness and general weakness, decreased thyroid function, loss of balance while walking, osteonecrosis (disease that reduces blood flow to bones, which can lead to bone loss and death of bone tissue), arthritis, swelling of the skin anywhere on the body

- **Pain** : inflammation of the veins which can lead to redness, sensitivity and swelling, inflammation of the tendon
- **Brain** : memory loss

• **Parameters can show** : abnormal blood test results and possible impairment of renal function due to the waste produced by the dying tumor (tumor lysis syndrome), low level of albumin in the blood, low level of lymphocytes (a type of white blood cell) in the blood, high cholesterol in the blood, swollen lymph nodes, bleeding in the brain, irregularity in the electrical activity of the heart, increase in the size of the heart, inflammation of the liver, protein in the urine, increased creatine phosphokinase (an enzyme mainly found in the heart, brain, and skeletal muscles), increased troponin (an enzyme found mainly in the heart and skeletal muscles), increased level of gamma-glutamyltransferase (an enzyme found mainly in the liver)

Rare side effects (may affect up to 1 in 1,000 people)

- **Heart and lung** : increase in the volume of the right ventricle of the heart, inflammation of the heart muscle, range of events leading to blockage of blood supply to the heart muscle (acute coronary syndrome), cardiac arrest (stopping blood flow from the heart), coronary artery (heart) disease, inflammation of the tissue covering the heart and lungs, blood clots, blood clots in the lungs
- **Digestive disorders** : loss of vital nutrients such as protein via the digestive tract, intestinal obstruction, anal fistula (abnormal opening of the anus to the skin around the anus), impaired kidney function, diabetes
- **Skin, hair, eyes, general** : seizure, inflammation of the optic nerve which may cause partial or complete loss of vision, purplish-blue spots on the skin, abnormally high thyroid function, inflammation of the thyroid, ataxia (a condition associated with poor muscle coordination), difficulty walking, miscarriage, inflammation of the blood vessels in the skin, fibrosis
- **Brain** : stroke, temporary episode of neurological dysfunction caused by decreased blood flow, facial nerve paralysis, dementia
- **Immune system** : severe allergic reaction
- **Musculoskeletal and subcutaneous tissue** : delayed fusion of the rounded ends that form the joints (epiphyses) ; slower or delayed growth

Other side effects which have been reported with an unknown frequency (which cannot be estimated from the available data)

- Lung inflammation
- Bleeding in the stomach or intestines, which can be fatal
- Hepatitis B reappears (reactivates) if you have had hepatitis B in the past (liver infection)
- A reaction with fever, blisters on the skin and mucosal ulceration
- Kidney disease with symptoms including edema and abnormal lab test results such as protein in the urine and low protein in the blood
- Injury of blood vessels called thrombotic microangiopathy (TMA), including decreased red blood cell count, decreased number of platelets and blood clots

Your doctor **will** check for some of these effects during your treatment.

Reporting side effects

If you get any side effects **talk to your doctor or pharmacist**. This also applies to any side effects that are not mentioned in this leaflet. You can also report side effects directly via the national reporting system : CNPV (National Center of Pharmacovigilance). By reporting side effects you can help provide more information on the safety of this medicine.

5. How to store DASATINIB NEAPOLIS ?

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

Store at a temperature not exceeding 30 ° C in the original outer packaging.

Do not use this medicine after the expiry date which is stated on the blister or the outer carton after EXP.

The expiration date refers to the last day of that month.

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

6. Content of the packaging and other information

What DASATINIB NEAPOLIS contains

DASATINIB NEAPOLIS 20mg, film-coated tablet	:	Dasatinib	20mg Per tablet
DASATINIB NEAPOLIS 50mg, film-coated tablet	:	Dasatinib	50mg Per tablet
DASATINIB NEAPOLIS 70mg, film-coated tablet	:	Dasatinib	70mg Per tablet
DASATINIB NEAPOLIS 140mg, film-coated tablet	:	Dasatinib	140mg Per tablet

The other components are :

Tablet core : Lactose monohydrate, microcrystalline cellulose, Croscarmellose sodium, Hydroxypropylcellulose, Magnesium stearate

Film Coating Agent : Polyvinyl Alcohol, Titanium Dioxide, Polyethylene Glycol / Macrogol, Talc

What does DASATINIB NEAPOLIS look like and content of the outer packaging

DASATINIB NEAPOLIS 20 mg : box of 60 white to off-white, biconvex, round film coated-tablets with "N21" debossed on one side in Aluminum / Aluminum blisters.

DASATINIB NEAPOLIS 50 mg : box of 60 white to off-white, biconvex, oval-shaped film coated-tablets in Aluminum / Aluminum blisters.

DASATINIB NEAPOLIS 70 mg : box of 60 white to off-white, biconvex, round film coated-tablets in Aluminum / Aluminum blisters.

DASATINIB NEAPOLIS 140 mg : box of 30 and 60 white to off-white, biconvex, round film coated-tablets with "N24" debossed on one side in Aluminum / Aluminum blisters

Prescription and supply condition : Table A / list 1

M. A. N° :

DASATINIB NEAPOLIS 20 mg film-coated tablet box of 60 :

DASATINIB NEAPOLIS 50 mg film-coated tablet box of 60 :

DASATINIB NEAPOLIS 70 mg film-coated tablet box of 60 :

DASATINIB NEAPOLIS 140 mg film-coated tablet box of 30 :

DASATINIB NEAPOLIS 140 mg film-coated tablet box of 60 :

Marketing authorization holder and manufacturer :

Neapolis Pharma

Road of Tunis Km 7 - PB 206 Nabeul 8000 - Tunisia.

Tel : +216. 31.833.400 / Fax : +216.72.235.016 / +216.31.338.421

E.mail : neapolispharma@neapolispharma.com

This leaflet was last reviewed : june 2020

THIS IS A MEDICINE

- A medicine is not like other consumer goods.
- A medication is a product that affects your health and its consumption without compliance to the prescription exposes you to danger.
- You should strictly follow your doctor's prescription and directions for use, as well as the advice of your pharmacist.
- Your doctor and pharmacist are familiar with the medication, its indications and contraindications.
- Do not stop treatment on your own during the prescribed period.
- Do not resume, do not increase doses without consulting your doctor.

KEEP MEDICINES OUT OF THE REACH OF CHILDREN



NEAPOLIS
PHARMA