

Patient information and treatment diary

To those of you who are being treated
with OPDIVO[®] (nivolumab) in combination
with YERVOY[®] (ipilimumab)

Introduction

This is a booklet intended as guidance for you who are being treated with OPDIVO combined with YERVOY. Here you will learn more about this medicine and what you can expect in terms of your treatment. The symptoms of possible side effects are described here, to which you should pay attention, as well as what to do should you experience such symptoms.

For complete information, please see the package leaflet for OPDIVO and YERVOY at www.fass.se.

About the treatment

The immune system consists of a number of different cells, including T-cells, which are part of the body's natural defence. T-cells find and destroy cells that the body considers to be alien, such as bacteria, viruses and cancer cells. Occasionally cancer cells can find a way to hide from the body's defence system which leads to the cancer being able to grow and spread.

OPDIVO helps by stopping the cancer cells' blocking mechanism towards the immune system's T-cells. YERVOY works in another way by increasing the activity of the T-cells so that they can attack and destroy the cancer cells.

Since this treatment works by activating the immune system it can result in side effects caused by the fact that your immune system becomes overactive and starts attacking healthy cells. It is important to contact the doctor or nurse treating you right away if you experience symptoms or signs of side effects. You can read more about side effects related to the treatment in the section "Side effects from the treatment" on pages 3–4.

Before the treatment starts

Before treatment, your doctor will check your general health. You will also give blood samples during your treatment.

Before treatment you and your doctor will go through whether:

- you have an autoimmune disease (a condition where the body attacks its own cells)
- you have melanoma in your eye
- you have previously been given YERVOY and had severe side effects from this medicine
- your cancer has spread to your brain
- you have previously had inflammation in your lungs
- you are taking or have taken other medicines

Inform your doctor if you are, or if you think you are, pregnant, if you plan to have a child or if you are breast-feeding. It is important to protect yourself against pregnancy.

How the treatment will be administered

OPDIVO in combination with YERVOY

OPDIVO and YERVOY will be administered via an infusion (drip) into the blood. The treatment will be administered in two phases.

Combination phase

During the first four treatments, OPDIVO will be administered, followed by YERVOY during the same day. The combination treatment will be administered every third week. The quantities of OPDIVO and YERVOY will be calculated on the basis of your body weight. Inform your treating doctor if you gain or lose weight.

Continued treatment

Afterwards, OPDIVO will be administered alone every two or four weeks. Your doctor will determine which is most suitable for you.

On the day that you receive treatment, you may experience a general sense of illness. The doctor and nurse treating you can inform you about how this can be treated.

Your doctor will continue to give you your treatment so long as you benefit from it or until you no longer tolerate it. It is your doctor who evaluates the treatment with, for example, an X-ray examination.

If you miss an appointment for treatment

It is very important that you come to your appointments to receive treatment. If you miss a treatment, your doctor will plan when to give you your next dose.

Side effects from the treatment

Like all medicines, this medicine can cause side effects, but not everybody experiences them. Your treatment affects your immune system and can cause inflammation in parts of the body. Inflammation can cause serious injury in the body and some inflammatory conditions can be life-threatening and may need treatment or may need you to discontinue your treatment.

Most side effects related to your treatment can be treated. For example, your doctor can decide to give you other medicines in order to decrease the symptoms and prevent complications, give your next dose at a later date or discontinue the treatment. Early measures to counteract side effects decrease the risk of having to discontinue the treatment. It is important that you contact your treating outpatient clinic early if you suspect a side effect or if you are not feeling well.

Side effects can occur at any time during treatment. Be attentive as side effects can occur weeks or months after your last dose.

It is important that you contact your treating outpatient clinic if you suspect an side effect. It is very important that side effects are detected early to allow treatment and thereby prevent your condition from deteriorating.

Immediately contact your doctor or nurse if you experience any of these signs or symptoms:



Problems with you liver: signs and symptoms can be yellow whites of your eyes or yellowish skin (jaundice), pain in the right part of your stomach or fatigue.



Diarrhoea: watery, thin or soft stools or other symptoms of inflammation in the bowels (colitis), like stomach pain and mucus or blood in the stools.



Problems with your kidneys: signs and symptoms can be decreased urine volume.



Problems with your lungs such as breathing difficulties or coughing. This can be a sign of inflammation of your lungs.



Inflammation in your muscles like myocarditis (inflammation of your heart muscle), myositis (inflammation of the muscles) and rhabdomyolysis (stiffness of the muscles and joints, muscular cramps). Signs and symptoms of this can be muscular pain, stiffness, weakness, chest pain or intense fatigue.



Problems with hormone-producing glands (including the thyroid gland and the adrenal glands): signs and symptoms that your glands do not work normally can be extreme fatigue, weight changes or headache and visual disturbances.



Diabetes (symptoms like exaggerated thirst, large increase in urine volume, increased appetite with weight loss, sensation of fatigue, drowsiness, weakness, depression, irritability and a general sensation of illness) or diabetic ketoacidosis (acid in the blood caused by the diabetes).



Inflammation in the skin which can cause serious skin reactions (known as toxic epidermal necrolysis and Stevens-Johnson syndrome). Signs and symptoms of serious skin reaction (sometimes lethal) can express itself as rash, itching and skin detachment.

Immediately contact the health care staff if you experience signs, symptoms described in this booklet, or if you have other adverse reactions. Receiving medical care at an early stage can prevent the problem from becoming serious. Do not try to treat your symptoms with other medicines on your own.

For a complete list of side effects, please see the package leaflet for OPDIVO and YERVOY at www.fass.se.

Common questions and answers about

OPDIVO in combination with YERVOY is a drug that helps the body's own immune system to attack and destroy cancer cells.

How does OPDIVO work in combination with YERVOY?

The immune system consists of a number of different cells, including T-cells, which are part of the body's natural defence. T-cells find and destroy cells that the body considers to be alien, such as bacteria, viruses and cancer cells. Occasionally cancer cells can find a way to hide from the body's defence system which leads to the cancer being able to grow and spread.

OPDIVO helps by stopping the cancer cells' blocking mechanism towards the immune system's T-cells. YERVOY works in another way by increasing the activity of the T-cells so that they can attack and destroy

the cancer cells. Your doctor will assess whether you should be given the combined treatment of OPDIVO and YERVOY.

As this treatment works by activating your immune system it can result in side effects caused by the fact that your immune system becomes overactive and starts attacking healthy cells. It is important to contact your doctor or nurse treating you early if you experience symptoms or signs of side effects. You can read more about side effects related to the treatment in the section “Side effects from the treatment” on pages 3-4.

How will I be given the treatment?

OPDIVO and YERVOY will be administered via an infusion (drip) into the blood. The treatment will be administered in two phases.

Combination phase: During the first four treatments, OPDIVO will be administered, followed by YERVOY during the same day. The combination treatment will be administered every third week. The quantities of OPDIVO and YERVOY will be calculated on the basis of your body weight. Tell us if you gain or lose weight.

Continued treatment: Afterwards, OPDIVO will be administered alone every two or four weeks. Your doctor will determine which is most suitable for you.

Where will I be given the treatment?

You will be administered the treatment at the hospital, under the supervision of your doctor.

How long will I be treated for?

Your doctor will continue to give you the treatment so long as you benefit from it or until you no longer tolerate it. It is your doctor who evaluates the treatment with, for example, an X-ray examination.

Will the treatment have to be stopped if I have side effects?

If you experience side effects, you must contact your doctor or nurse. Most side effects related to your treatment can be treated. For example, your doctor can decide to give you other medicines in order to decrease the symptoms and prevent complications, give your next dose at a later date or discontinue the treatment. Early measures to counteract side effects decrease the risk of having to discontinue the treatment.

What should I do if I experience side effects during the treatment?

It is important that you contact your doctor or nurse immediately so they can treat the side effect as early as possible, to prevent the side effect from getting worse. Side effects of the treatment can usually be alleviated with corticosteroids.

It is important that you contact your treating outpatient clinic early if you suspect a side effect or if you are not feeling well. Do not attempt to treat the symptom or side effect yourself. It is also important to know that side effects may occur at any time during your treatment and also several months after the treatment has finished.

Can I take other medicines during my treatment?

It is important that you tell your doctor about all other medicines that you are taking before you start the treatment. While treatment is ongoing, it is also important that you talk to your doctor before you start taking any other medicine.

What should I do if I miss a booked appointment time for treatment?

It is very important that you attend all treatments. If you miss a treatment, the unit handling your treatment will plan when to give you your next dose.

If you have any other questions about your treatment, contact your doctor or nurse.

Your patient card

Your doctor or your nurse will give you a patient card. It contains important safety information that you need to know before, during and after your treatment with the treatment. It also contains information that all medical staff need to know about your treatment. Therefore, it is important that you show your patient card to all other health care staff that you meet. This is also valid for your visit to a primary care centre or a another hospital.

If you have not been given a patient card or if you have lost it, ask the doctor or nurse treating you for a new one.

Always bring the patient card that you received from your doctor with you and show it if you need to visit another doctor, for example when you are travelling. It contains important information about symptoms that may need to be treated in agreement with your treating doctor.

Your treatment diary

This diary is created to make it easier for you, your doctor and your nurse to discover symptoms of side effects early on, which may occur with your treatment. If the symptoms are detected in time they can often be treated and can prevent the side effects from worsening.

Take time to fill in the diary and bring it to every visit to your treating doctor or nurse.

Date for treatment: day on /

..... day on/..... day on/..... day on/..... day on/..... day on/..... day on/..... day on/.....																	
How are you feeling?																							
Are you still able to perform your normal activities?																							
Are you nauseous and/or vomiting?																							
<input type="checkbox"/> YES <input type="checkbox"/> NO	<input type="checkbox"/> YES <input type="checkbox"/> NO	<input type="checkbox"/> YES <input type="checkbox"/> NO	<input type="checkbox"/> YES <input type="checkbox"/> NO	<input type="checkbox"/> YES <input type="checkbox"/> NO	<input type="checkbox"/> YES <input type="checkbox"/> NO	<input type="checkbox"/> YES <input type="checkbox"/> NO																	
Have you lost your appetite or do you feel less hungry than usually?																							
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How many defecations do you have each day?																							
Are your defecations more numerous than normally?																							
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Are your stools thin or watery?																							
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Have you seen blood or mucus in your stools?																							
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Are your defecations painful?																							
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Have you got a new cough or deteriorated cough?																							
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Do you have breathing problems?																							
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Have you had itching, skin rash, blisters, ulcers or peeling of the skin?																							
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Do you have a constant or unusual headache?																							
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Have you felt extremely tired?																							
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Have you had changes of your mood or your behaviour such as decreased sex drive, irritability or forgetfulness?																							
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Have you had numbness, pain or weakness in your muscles or difficulties walking?																							
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Do you have pain or numbness/swelling in joints or muscles?																							
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Have you had pain or redness in your eye, visual problems or blurred vision?																							
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Immediately contact the outpatient clinic if you feel ill during the treatment so that they can help you.

For complete information, please see the package leaflet for OPDIVO and YERVOY at www.fass.se

YERVOY® is an immunotherapy used to treat advanced melanoma, a type of skin cancer, in adults and adolescents aged 12 years and older. Yervoy in combination with Opdivo is used to treat advanced melanoma or advanced renal cell cancer in adults. It is important that you also read the package insert for Opdivo. Ask your doctor if you have questions about Opdivo.

The active substance in Yervoy is called ipilimumab.

▼ This medicine is subject to increased supervision

OPDIVO® is an immunotherapy that is used to treat adults with:

- advanced melanoma (a type of skin cancer)
- melanoma that has been completely removed with surgery (treatment after operation is called adjuvant treatment)
- advanced non-small cell lung cancer (a type of lung cancer)
- advanced renal cell cancer
- Hodgkin's lymphoma
- advanced head and neck cancer
- advanced urothelial cell cancer (cancer in the urine bladder and in the urinary tract)

Do not use Yervoy if you are allergic to nivolumab or any other excipient in this medicine.

You must not treat side effects that are related to Yervoy on your own and must always contact your doctor. Note that side effects may occur at any time during the treatment and sometimes weeks or months after your last dose.

Potency and package: Yervoy is a concentrate for infusion fluid with potency 5 mg/ml. The package

size is either 1 injection bottle at 10 ml or 1 injection bottle at 40 ml.

Further information: For complete information about Yervoy, carefully read the package leaflet contained in the package (it can also be found at www.fass.se).

Bristol Myers Squibb
Phone: 08-704 71 00
www.bms.com/se

Opdivo can be administered in combination with ipilimumab for advanced melanoma or advanced renal cell cancer. It is important that you also read the package leaflet for Yervoy. Ask your doctor if you have questions about Yervoy.

The active substance in Opdivo is called nivolumab.

Do not use Opdivo if you are allergic to nivolumab or any other excipient in this medicine.

You must not treat side effects that are related to Opdivo on your own and must always contact your doctor. Note that side effects may occur at any time during the treatment and sometimes weeks or months after your last dose.

Potency and package: Opdivo is a concentrate for infusion fluid with potency 10 mg/ml. The package size is either 1 injection bottle at 4 ml or 1 injection bottle at 10 ml or 1 injection bottle at 24 ml.

Further information: For complete information about Opdivo, carefully read the package leaflet contained in the package (it can also be found at www.fass.se).

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