

OPDIVO + YERVOY
+ CHEMOTHERAPY

Patient information and treatment diary

For patients being treated with
OPDIVO® (nivolumab) in combination with
YERVOY® (ipilimumab) and chemotherapy
for non-small cell lung cancer

OPDIVO[®]
(nivolumab)



YERVOY[®]
(ipilimumab)

Introduction

This booklet is intended as a guide for patients being treated with OPDIVO in combination with YERVOY and two treatment cycles of platinum-based chemotherapy. Here you will learn more about OPDIVO and YERVOY and what you can expect in conjunction with 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 on www.fass.se.

This booklet does not include information specifically on chemotherapy. It is therefore important to consult the package leaflets for these medicines. Ask your doctor if you have questions about your chemotherapy, or read the information in the package leaflet for these medicines on 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 immune 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 your treating doctor or nurse right away if you experience symptoms or signs of side effects. You can read more information about side effects related to treatment in 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 previously received OPDIVO or YERVOY and experienced serious side effects from these medicines
- your cancer has spread to the brain
- you have previously had inflammation in the lungs
- you are taking or have recently taken other medicines

Inform your doctor if you are or think you may be pregnant, if you are planning to have a baby or if you are breastfeeding. It is important to protect yourself against pregnancy in conjunction with the treatment.

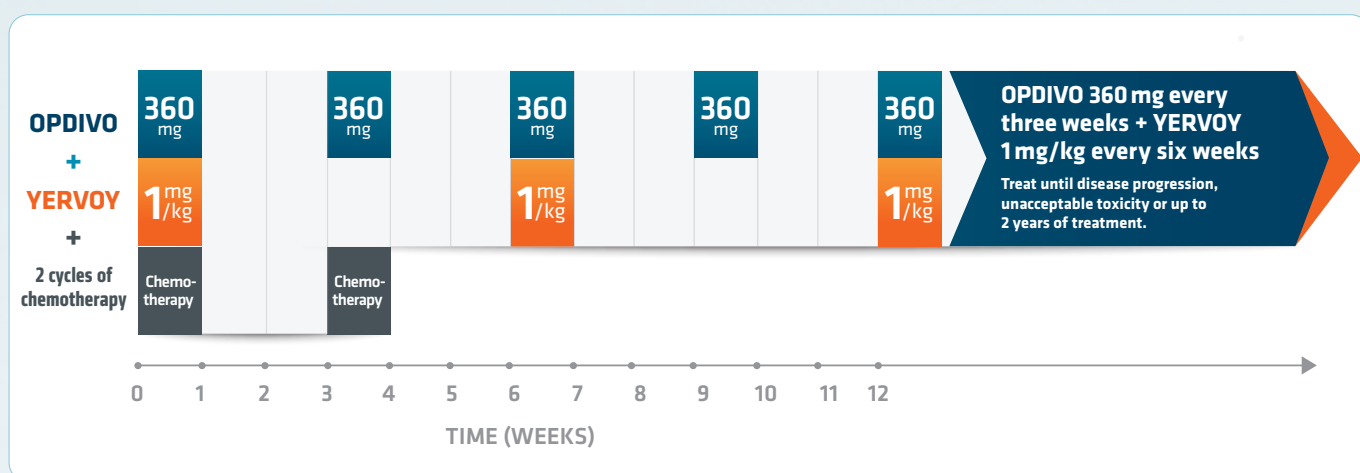
How the treatment will be administered

OPDIVO in combination with YERVOY and chemotherapy

OPDIVO, YERVOY and chemotherapy will be administered to you via an infusion (drip) into the blood. The infusions are given over 30 minutes each with at least 30 minutes of rest between treatments. The time for treatment with chemotherapy depends on which chemotherapy is being received.

In the first treatment session, OPDIVO is given followed by YERVOY and then chemotherapy. After three weeks, OPDIVO is given followed by the last dose of chemotherapy.

After this point, OPDIVO is given every three weeks and YERVOY every six weeks. This means that every six weeks, treatment with both OPDIVO and YERVOY is given, while the treatment round in between only includes treatment with OPDIVO.



On the day that you receive treatment you may experience a general sense of illness. Your treating doctor and nurse can in such case inform you of how this can be treated. The amount of YERVOY will be calculated on the basis of your body weight. Inform your doctor or nurse if you gain or lose weight.

Your doctor will continue giving you your treatment for up to 2 years or until you no longer tolerate it. Your doctor will evaluate the treatment, e.g. by X-ray examination.

If you miss a scheduled 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, these medicines 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 you may need to discontinue your treatment.

The vast majority of side effects related to your treatment can be counteracted. For example, your doctor can decide to give you other medicines in order to decrease the symptoms and prevent complications, give you 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 doctor or nurse early on 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 also weeks or months after your last dose.

It is important that you contact your treating doctor or nurse if you suspect a 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 your liver: signs and symptoms can be yellow whites of your eyes or yellowish skin (jaundice), pain in the right part of your digestive tract 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 manifest as 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 are not working 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 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 fatal) can manifest as rash, itching and skin detachment.

Contact the health care staff immediately if you experience signs or 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 on www.fass.se.

Common questions and answers about your treatment

OPDIVO in combination with YERVOY are medicines that help the body's own immune system 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, however, cancer cells can find a way to get around 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 determine whether you will receive the combined treatment with OPDIVO and YERVOY.

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 that you contact your treating doctor or nurse early on if you experience symptoms or signs of side effects. You can read more information about side effects related to treatment in section "Side effects from the treatment" on pages 3-4.

How will I be given the treatment?

OPDIVO, YERVOY and chemotherapy will be administered to you via an infusion (drip) into the blood. The infusions are given over 30 minutes each with at least 30 minutes of rest between treatments. The time for treatment with chemotherapy depends on which chemotherapy is being received.

In the first treatment session, OPDIVO is given followed by YERVOY and then chemotherapy. After three weeks, OPDIVO is given followed by the last dose of chemotherapy.

After this point, OPDIVO is given every three weeks and YERVOY every six weeks. This means that every six weeks treatment with both OPDIVO and YERVOY is given, while the treatment round in between only includes treatment with OPDIVO.

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 giving you your treatment for up to 2 years or until you no longer tolerate it. Your doctor will evaluate the treatment, e.g. by X-ray examination.

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

If you experience side effects, you must contact your doctor or nurse. The vast majority of side effects related to your treatment can be treated. Your doctor can decide to give you other medicines in order to decrease the symptoms and prevent complications, give you 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 from treatment with OPDIVO and YERVOY can usually be alleviated with corticosteroids.

It is important that you contact your treating doctor or nurse early on 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 even several months after completion of treatment.

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 OPDIVO and YERVOY. 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 also applies when you visit a primary care centre or a 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 with you and show it if you need to visit another doctor, e.g. if you are on holiday. It contains important information about symptoms that may need to be treated in consultation with your treating doctor.

Your treatment diary

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

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

Treatment date day, the /

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How are you feeling?

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Are you still able to perform your normal activities?

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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
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Have you lost your appetite or do you feel less hungry than usual?

<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
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How many defecations do you have each day?

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Are your defecations more numerous than normal?

<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
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Are your stools thin or watery?

<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
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Have you seen blood or mucus in your stools?

<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
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Are your defecations painful?

<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
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Have you got a new cough or worsened cough?

<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
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Do you have breathing problems?

<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
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Have you had itching, skin rash, blisters, ulcers or peeling of the skin?

<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
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Do you have a constant or unusual headache?

<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
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Have you felt extremely tired?

<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
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Have you had changes of your mood or your behaviour such as decreased sex drive, irritability or forgetfulness?

<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
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Have you had numbness, pain or weakness in your muscles or difficulties walking?

<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
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Do you have pain or numbness/swelling in joints or muscles?

<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
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Have you had pain or redness in your eye, visual problems or blurred vision?

<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
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Immediately contact your treating doctor or nurse if you feel ill during the treatment so that they can help you.

For more information about OPDIVO or OPDIVO in combination with YERVOY, visit the website www.opdivopatient.se

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

- advanced melanoma (a type of skin cancer)
- Melanoma that has been completely removed through surgery (treatment after surgery is called adjuvant treatment)
- advanced non-small cell lung cancer (a type of lung cancer)
- advanced renal cell carcinoma
- Hodgkins lymphoma
- advanced head and neck cancer
- advanced urothelial cell cancer (cancer in the urine bladder and the urinary tract)
- advanced oesophageal cancer

The active substance in OPDIVO is called nivolumab. OPDIVO can be administered in combination with YERVOY for advanced melanoma or advanced renal cell carcinoma. OPDIVO can be administered in combination with YERVOY and chemotherapy for advanced non-small cell lung cancer. It is important that you also read the package leaflet for YERVOY. Ask your doctor if you have questions about YERVOY.

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. 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.

Strength and package: OPDIVO is a 10 mg/ml concentrate for solution for infusion. The package size is either 1 injection bottle of 4 ml or 1 injection bottle of 10 ml or 1 injection bottle of 24 ml.

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

Bristol Myers Squibb
Tel: 08-704 71 00
www.bms.com/se
Version 4

YERVOY is an immunotherapy used to treat advanced melanoma, a type of skin cancer, in adults and adolescents aged 12 years and older. YERVOY can be administered in combination with OPDIVO for advanced melanoma or advanced renal cell carcinoma in adults. YERVOY can be administered in combination with OPDIVO and chemotherapy for advanced non-small cell lung 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.

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. 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.

Strength and package: YERVOY is a 5 mg/ml concentrate for solution for infusion. The package size is either 1 injection bottle containing 10 ml or 1 injection bottle containing 40 ml.

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

Bristol Myers Squibb
Tel: 08-7047100
www.bms.com/se
Version 2