PATIENT PACKAGE INSERT IN ACCORDANCE WITH THE PHARMACISTS' REGULATIONS (PREPARATIONS) - 1986
The medicine is dispensed with a doctor's prescription only

Yuflyma
40 mg
80 mg
Solution for injection in a prefilled pen

Active ingredient and its concentration: adalimumab 100 mg/ml
Each 40 mg Yuflyma prefilled pen contains:
adalimumab 40 mg/0.4 ml
Each 80 mg Yuflyma prefilled pen contains:
adalimumab 80 mg/0.8 ml
Inactive and allergenic ingredients in the preparation – see section 6 in this leaflet.
Read the leaflet carefully in its entirety before using the medicine. This leaflet contains concise information about the medicine. If you have further questions, refer to the doctor or pharmacist.
This medicine has been prescribed to treat your ailment/you. Do not pass it on to others. It may harm them even if it seems to you that their ailment/medical condition is similar.
Yuflyma is a biosimilar preparation. For additional information on biosimilar preparations, refer to the Ministry of Health website:
https://www.health.gov.il/UnitsOffice/HD/MT/Drugs/Registration/Pages/Biosimilars.aspx

In addition to the leaflet, Yuflyma has a 'Patient safety information card'. This card includes important safety information which you should know before starting and during the treatment with Yuflyma and act accordingly. Read the 'Patient safety information card' and the patient leaflet before starting to use the preparation. Keep the card for further reading, if necessary.

Please note that it is important that each time you receive the medicine from the pharmacy, be sure that you receive the medicine that was prescribed for you by the specialist treating you. If the medicine you received looks different than the one you usually receive, or if the instructions for use have changed, please refer immediately to the pharmacist and make sure that you received the correct medicine. Any change or change in the dosage of a medicine containing adalimumab (the active ingredient in the medicine) must be made by the attending specialist only. Please check that the tradename of the preparation prescribed for you by the specialist in the prescription, is identical to the name of the medicine you received from the pharmacist.

1. WHAT IS THE MEDICINE INTENDED FOR?
Rheumatoid arthritis
Yuflyma in combination with methotrexate, is indicated for:
- the treatment of moderate to severe, active rheumatoid arthritis in adult patients when the response to disease-modifying anti-rheumatic drugs (DMARDs), including methotrexate, has been inadequate.
- the treatment of severe, active and progressive rheumatoid arthritis in adults not previously treated with methotrexate.

Yuflyma can be given as monotherapy in case of intolerance to methotrexate or when continued treatment with methotrexate is inappropriate.
Adalimumab has been shown to reduce the rate of progression of joint damage as measured by X-ray and to improve physical function, when given in combination with methotrexate.
Polyarticular juvenile idiopathic arthritis
Yuflyma in combination with methotrexate is indicated for the treatment of active polyarticular juvenile idiopathic arthritis in patients from the age of 2 years, weighing at least 30 kg, who have had an inadequate response to one or more disease-modifying anti-rheumatic drugs (DMARDs). Yuflyma can be given as monotherapy in case of intolerance to methotrexate or when continued treatment with methotrexate is inappropriate. Adalimumab has not been studied in patients aged less than 2 years.
Enthesitis-related arthritis
Yuflyma is indicated for the treatment of active enthesitis-related arthritis, in patients 6 years of age and older, weighing at least 30 kg, who have had an inadequate response to, or who are intolerant of, conventional therapy.
Ankylosing spondylitis (AS)
Yuflyma is indicated for the treatment of adults with severe active ankylosing spondylitis who have had an inadequate response to conventional therapy.
Axial spondyloarthritis without radiographic evidence of AS
Yuflyma is indicated for the treatment of adults with severe axial spondyloarthritis without radiographic evidence of AS, but with objective signs of inflammation by radiological and/or laboratory tests including MRI and serum CRP levels, who have had an inadequate response to, or are intolerant to non-steroidal anti-inflammatory drugs (NSAIDs).
Psoriatic arthritis
Yuflyma is indicated for the treatment of active and progressive psoriatic arthritis, in adults, when the response to previous disease-modifying anti-rheumatic drug therapy has been inadequate. Adalimumab has been shown to reduce the rate of progression of peripheral joint damage, as measured by X-ray in patients with polyarticular symmetrical subtypes of the disease and to improve physical function.
Psoriasis
Yuflyma is indicated for the treatment of moderate to severe chronic plaque psoriasis in adult patients who are candidates for systemic therapy.
Pediatric plaque psoriasis
Yuflyma is indicated for the treatment of severe chronic plaque psoriasis in children and adolescents from 4 years of age, weighing at least 30 kg, who have had an inadequate response to or are inappropriate candidates for topical therapy and phototherapies.
Hidradenitis suppurativa (HS)
Yuflyma is indicated for the treatment of moderate to severe hidradenitis suppurativa (acne inversa) in adults and adolescents from 12 years of age, weighing at least 30 kg, with an inadequate response to conventional systemic HS therapy.
Crohn's disease
Yuflyma is indicated for reducing signs and symptoms and inducing and maintaining clinical remission in adult patients with moderately to severely active Crohn's disease, who have had an inadequate response to conventional therapy. Yuflyma is indicated for reducing signs and symptoms and inducing clinical remission in these patients if they have also lost response to or are intolerant to infliximab.
Pediatric Crohn's disease
Yuflyma is indicated for the treatment of moderately to severely active Crohn's disease in paediatric patients from 6 years of age, weighing at least 40 kg, who have had an inadequate response to conventional therapy including primary nutrition therapy and corticosteroid and/or an immunomodulator, or who are intolerant to or have contraindications for such therapies.
Ulcerative colitis
Yuflyma is indicated for treatment of moderately to severely active ulcerative colitis in adult patients, who have had an inadequate response to conventional therapy including corticosteroids and 6-mercaptopurine (6-MP) or azathioprine (AZA), or who are intolerant to or have medical contraindications for such therapies.
Pediatric ulcerative colitis
Yuflyma is indicated for the treatment of moderately to severely active ulcerative colitis in paediatric patients (from 6 years of age), who have had an inadequate response to conventional therapy including corticosteroids and/or 6-mercaptopurine (6-MP) or azathioprine (AZA), or who are intolerant to or have medical contraindications for such therapies.
Uveitis
Yuflyma is indicated for the treatment of non-infectious intermediate, posterior and panuveitis in adult patients, who have had an inadequate response to corticosteroids, in patients in need of corticosteroid-sparing, or in whom corticosteroid treatment is inappropriate.
Pediatric uveitis
Yuflyma is indicated for the treatment of chronic non-infectious uveitis in paediatric patients from 2 years of age, weighing at least 30 Kg, who have had an inadequate response to or are intolerant to conventional therapy, or in whom conventional therapy is inappropriate.
Intestinal Behcet's disease
Yuflyma is indicated for the treatment of intestinal Behcet's disease, in patients who have had an inadequate response to conventional therapy.
Yuflyma contains the active ingredient adalimumab.
The active ingredient in Yuflyma, adalimumab, is a human monoclonal antibody.
Monoclonal antibodies are proteins that attach to specific targets. The target of adalimumab is a protein called tumor necrosis factor (TNFa), which is involved in the immune (defense) system and is present at increased levels in the inflammatory diseases listed above. By attaching to TNFa, Yuflyma reduces the inflammatory process in these diseases.
There is no information regarding use of Yuflyma in children under two years of age.
Therapeutic group: TNF blocker.

2. BEFORE USING THE MEDICINE
Do not use the medicine if:
- you are sensitive (allergic) to the active ingredient, or to any of the additional ingredients included in the medicine (see section 6 "Further information").
- you have active tuberculosis or other severe infections, such as sepsis and opportunistic infections (see "Special warnings regarding use of the medicine"). It is important that you tell your doctor if you have symptoms of infection, for example fever, wounds, feeling tired and dental problems.
- you have moderate or severe heart failure. It is important to tell your doctor if you have or have had a serious heart problem (see "Special warning regarding use of the medicine").

Special warnings regarding use of the medicine
Before treatment with Yuflyma, inform your doctor:
Allergic reactions
- If you suffer from allergic reactions with symptoms such as chest tightness, wheezing, dizziness, swelling or rash, do not inject more Yuflyma and contact your doctor immediately since, in rare cases, these reactions can be life-threatening.

Infections
- If you have an infection, including prolonged infection or a localized infection (for example, leg ulcer) consult your doctor before using Yuflyma. If you are uncertain, refer to your doctor.
- You might get infections more easily while receiving Yuflyma treatment. This risk may increase if you have a problem with your lungs. These infections may be serious and include:
  - tuberculosis
  - infections caused by viruses, fungi, parasites or bacteria
  - severe infection in the blood (sepsis)

  In rare cases, these infections can be life-threatening. It is important to tell your doctor if you get symptoms such as fever, wounds, feeling tired or dental problems. Your doctor may recommend that you temporarily stop treatment with Yuflyma.
- Consult your doctor if you live or travel in regions where fungal infections (for example, histoplasmosis, coccidioidomycosis or blastomycosis) are very common.
- Consult your doctor if you have had recurrent infections or other conditions that increase the risk of infections.
- If you are over the age of 65 years, you may be more likely to get infections during treatment with Yuflyma. You and your doctor should pay special attention to signs of infection during the course of treatment with Yuflyma. It is important to tell your doctor if you have symptoms of infections, such as fever, wounds, feeling tired or dental problems.

Tuberculosis
- It is very important that you tell your doctor if you have ever had tuberculosis, or if you have been in close contact with someone who has had tuberculosis. If you have active tuberculosis, do not use Yuflyma.
  - As cases of tuberculosis have been reported in patients treated with adalimumab, your doctor will check you for signs or symptoms of tuberculosis before you start Yuflyma treatment. This will include a thorough medical evaluation, including patient history and appropriate screening tests (for example, chest X-ray and a tuberculin test). The conduct and results of these tests should be recorded on your **'Patient Safety Information Card'**.
  - Tuberculosis can develop during therapy, even if you have received treatment for the prevention of tuberculosis.
  - If symptoms of tuberculosis (for example, persistent cough, weight loss, lack of energy, mild fever), or of any other infection appears during or after treatment with Yuflyma, refer to your doctor immediately.

Hepatitis B
- Tell your doctor if you are a carrier of the hepatitis B virus (HBV), if you have active HBV or if you think you might be at risk of contracting HBV.
  - Your doctor should perform an HBV test. In people who carry HBV, Yuflyma can cause its reactivation.
  - In rare cases, especially if you are taking other medicines that suppress the immune system, reactivation of HBV can be life-threatening.

Dental procedure or surgery
- If you are due to undergo dental procedures or surgery, inform your doctor that you are taking Yuflyma. Your doctor may recommend temporary discontinuation of Yuflyma.

Demyelinating diseases
- If you have or develop a demyelinating disease (a disease that affects the insulating layer around the nerves, such as multiple sclerosis), your doctor will decide if you should receive or continue to receive Yuflyma. Tell your doctor immediately if you experience symptoms like changes in your vision, weakness in your arms or legs or numbness or tingling in any part of the body.

Vaccinations
- Certain vaccines may cause infections and you should not receive them during the course of treatment with Yuflyma.
  - Consult your doctor before you receive any vaccine.
  - It is recommended that children, if possible, be given all the vaccinations scheduled for their age before starting treatment with Yuflyma.
  - If you were treated with Yuflyma during pregnancy, your baby may be at higher risk for an infection for approximately five months after the last dose you received during pregnancy. It is important that you tell your baby's doctors and healthcare providers that you were treated with Yuflyma during pregnancy so they can decide when your baby can receive vaccines.

Heart failure
- If you have mild heart failure and are being treated with Yuflyma, your heart failure status must be closely monitored by your doctor. It is important to tell your doctor if you have or have had a serious heart problem. If new heart failure symptoms develop or if the existing symptoms worsen (e.g., shortness of breath, or swelling of the feet), refer to your doctor immediately. Your doctor will decide if you should receive Yuflyma.

Fever, bruising, bleeding or pallor
- In some patients, the body fails to produce a sufficient amount of the blood cells that fight infections or that help stop bleeding. Your doctor may decide to stop treatment with Yuflyma if a fever that does not go away, mild bruises or if you bleed very easily or look very pale, refer to your doctor immediately.

Cancer
- Very rare cases of certain kinds of cancer in adults and children treated with adalimumab or other TNF blockers have been described.
  - People with more serious rheumatoid arthritis who have had the disease for a long time, may be at higher than average risk of getting lymphoma (a cancer that affects the lymphatic system) and leukemia (a cancer that affects the bone marrow and the blood).
  - If you being treated with Yuflyma, the risk of getting lymphoma, leukemia, or other cancers may increase. On rare occasions, an uncommon and severe type of lymphoma, has been seen in patients taking adalimumab. Some of those patients were also treated with azathioprine or 6-mercaptopurine.
  - Tell your doctor if you are taking azathioprine or 6-mercaptopurine together with Yuflyma.
  - Cases of non-melanoma skin cancer have been observed in patients taking adalimumab.
  - Tell your doctor if new skin lesions appear or if there is a change in existing lesions during or after therapy.
- Cases of non-lymphoma cancers have been reported in patients who took a different TNF blocker and who have a certain type of lung disease called chronic obstructive pulmonary disease (COPD). If you have COPD, or are a heavy smoker, consult with your doctor on whether treatment with a TNF blocker is appropriate for you.

Autoimmune disease
- On rare occasions, treatment with Yuflyma could result in lupus-like syndrome. Refer to your doctor if symptoms such as persistent unexplained rash, fever, joint pain or tiredness occur.

Smoking
If you are a heavy smoker, consult with your attending doctor on whether treatment with a TNF blocker is appropriate for you (see additional information in section "Special warnings regarding use of the medicine").
Children and adolescents
Vaccinations: If possible, children should receive all the necessary vaccinations before starting treatment with Yuflyma.
Drug interactions
If you are taking, or have recently taken, other medicines, including non-prescription medicines and nutritional supplements, tell the doctor or pharmacist.
Do not take Yuflyma with medicines containing the following active ingredients, due to increased risk of serious infections:
- anakinra.
- abatacept.

These medicines are used to treat rheumatoid arthritis.
Yuflyma can be taken together with:
- methotrexate
- certain disease-modifying anti-rheumatic agents for treatment of arthritis (for example, sulfasalazine, hydroxychloroquine, leflunomide and injectable gold preparations)
- steroids or pain medications, including non-steroidal anti-inflammatory drugs (NSAIDs).

If you have questions, ask your doctor.
Pregnancy and breastfeeding
- You should consider the use of adequate contraception to prevent pregnancy and continue its use for at least 5 months after the last Yuflyma treatment.
- If you are pregnant, think you may be pregnant or are planning a pregnancy, consult with your doctor about taking this medicine.
- Yuflyma should only be used during a pregnancy if needed.
- In a study that assessed its use in pregnant women, there was no higher risk of birth defects when the mother had received adalimumab during pregnancy compared with mothers with the same disease who did not receive adalimumab.
- Yuflyma can be taken during breastfeeding.
- If you received Yuflyma during pregnancy, your baby may be at higher risk for developing infections.
- Before your baby receives any vaccine, it is important to inform the pediatrician treating your baby, as well as the healthcare providers in the clinic and Family Health Center ("Tipat Chalav"), that you took Yuflyma during the pregnancy. For more information on vaccines, see section "Special warnings regarding use of the medicine".

Driving and using machinery
Yuflyma may have a negligible effect on the ability to drive, ride a bicycle or operate machinery. After treatment with Yuflyma, you may have a sensation of dizziness and visual disturbances.

3. HOW SHOULD YOU USE THE MEDICINE?
Always use the preparation according to the doctor's instructions.
Check with the doctor or pharmacist if you are uncertain regarding the dosage and treatment regimen of the preparation.
The dosage and treatment regimen will be determined by the doctor only.
Do not exceed the recommended dose.
Method of administration:
Yuflyma solution for injection in a prefilled pen is injected under the skin (subcutaneous use). Yuflyma is provided in a package of 40 mg or 80 mg. Thus, Yuflyma can not be used for children who require a dosage smaller than a full 40 mg dose. If such a dosage is required, other medicines containing adalimumab should be used.
Detailed instructions on how to inject Yuflyma are provided in section 7 'Instructions for use'.
If you accidentally took a higher dosage
If you accidentally injected Yuflyma more frequently than instructed by your doctor or pharmacist, call your doctor or pharmacist and report this to them. Always bring the package of the medicine with you, even if it is empty.
If you forgot to inject Yuflyma
If you forgot to inject Yuflyma, inject the next dose as soon as you remember. The next dose should be taken as originally scheduled had you not forgotten a dose. Adhere to the treatment regimen as recommended by the doctor. Even if there is an improvement in your health, do not stop treatment with the medicine without consulting the doctor.
If you stop taking Yuflyma
Discuss discontinuation of Yuflyma with your doctor. Your symptoms may recur when you stop taking Yuflyma.
Do not take medicines in the dark! Check the label and the dose each time you take medicine. Wear glasses if you need them.
If you have further questions regarding use of the medicine, consult the doctor or pharmacist.

4. SIDE EFFECTS
As with any medicine, use of Yuflyma may cause side effects in some users. Do not be alarmed when reading the list of side effects. You may not suffer from any of them.

Most side effects are mild to moderate. However, some side effects may be serious and require treatment. Side effects may occur up to at least 4 months after the last Yuflyma treatment.
Refer to your doctor immediately if you notice any of the following symptoms:
- severe rash, hives or other signs of allergic reaction
- swelling in the face, hands, feet
- breathing difficulties, swallowing difficulties
- shortness of breath with physical activity or upon lying down or swelling of the feet

Refer to your doctor as soon as possible if you notice any of the following symptoms:
- signs suggestive of infection such as fever, nausea, wounds, dental problems, burning upon urination
- feeling weak or tired
- cough
- tingling
- numbness
- double vision
- arm or leg weakness
- a bump or open sore that does not heal
- signs and symptoms suggestive of blood disorders such as persistent fever, bruising, bleeding, paleness

The symptoms described above can be signs of the following side effects, which have been observed with Yuflyma.
Very common side effects (effects that occur in more than 1 user in 10):
- injection site reactions (including pain, swelling, redness or itching)
- respiratory tract infections (including cold, runny nose, sinus infection, pneumonia)
- headache
- abdominal pain
- nausea and vomiting
- rash
- musculoskeletal pain

Common side effects (effects that occur in 1-10 in 100 users):
- serious infections (including blood poisoning and influenza)
- intestinal infections (including gastroenteritis)
- skin infections (including cellulitis, shingles)
- ear infections
- oral infections (including tooth infections and cold sores)
- reproductive tract infections
- urinary tract infection
- fungal infections
- joint infections
- benign tumors
- skin cancer
- allergic reactions (including seasonal allergy)
- dehydration
- mood swings (including depression)
- anxiety
- sleeping difficulties
- sensation disorders such as sense of tingling, prickling or numbness
- migraine
- nerve root compression (including lower back pain and leg pain)
- vision disturbances
- eye inflammation
- inflammation of the eyelid and eye swelling
- vertigo (feeling of dizziness or spinning)
- sensation of rapid heart beats
- high blood pressure
- flushing
- hematoma (collection of blood outside of blood vessels)
- cough
- asthma
- shortness of breath
- gastrointestinal bleeding
- dyspepsia (indigestion, bloating, heartburn)
- acid reflux disease
- sicca syndrome (including dry eyes and dry mouth)
- itching
- itchy rash
- bruising
- inflammation of the skin (such as eczema)
- breaking of fingernails and toenails
- increased sweating
- hair loss
- new onset or worsening of psoriasis
- muscle spasms
- blood in the urine
- kidney problems
- chest pain
- edema (swelling)
- fever
- reduction in blood platelets which increases the risk of bleeding or bruising
- impaired healing

Uncommon side effects (effects that occur in 1-10 in 1,000 users):
- opportunistic infections (which include tuberculosis and other infections that occur when resistance of the body to disease is lowered)
- neurological infections (including viral meningitis)
- eye infections
- bacterial infections
- diverticulitis (infection and inflammation of the large intestine)
- cancer
- cancer that affects the lymphatic system
- melanoma
- immune system disorders that could affect the lungs, skin and lymph nodes (most commonly presenting as sarcoidosis)
- inflammation of blood vessels (vasculitis)
- tremor
- disorder of the nerves (neuropathy)
- stroke
- hearing loss, buzzing
- sensation of heart beating irregularly, such as skipping a beat
- heart problems that can cause shortness of breath or ankle swelling
- heart attack
- a sac in the wall of a major artery, inflammation and clot of a vein, blockage of a blood vessel
- lung diseases causing shortness of breath (including inflammation)
- pulmonary embolism (blockage in an artery of the lung)
- abnormal collection of fluid in the pleural space (pleural effusion)
- inflammation of the pancreas which causes severe pain in the abdomen and back
- difficulty in swallowing
- facial edema (swelling of the face)
- gallbladder inflammation, gallbladder stones
- fatty liver
- night sweats
- scarring
- abnormal breakdown of muscle tissue
- systemic lupus erythematosus (including inflammation of skin, heart, lung, joints and other organ systems)
- sleep interruptions
- impotence
- inflammations

Rare side effects (effects that occur in 1-10 in 10,000 users):
- leukemia (cancer affecting the blood and bone marrow)
- severe allergic reaction with shock
- multiple sclerosis
- nerve disorders (such as eye nerve inflammation and Guillain-Barré syndrome that may cause muscle weakness, abnormal sensations, tingling in the arms and upper body)
- heart stops pumping
- scarring of the lung (pulmonary fibrosis)
- intestinal perforation (hole in the intestine)
- hepatitis
- reactivation of hepatitis B
- autoimmune hepatitis (inflammation of the liver caused by the body's own immune system)
- inflammation of blood vessels in the skin (cutaneous vasculitis)
- Stevens-Johnson syndrome (early symptoms include malaise, fever, headache and rash)
- facial edema (swelling of the face) associated with allergic reactions
- erythema multiforme (inflammatory skin rash)
- lupus-like syndrome
- angioedema (localized swelling of the skin)
- lichenoid skin reaction (itchy reddish-purple skin rash)

Side effects of unknown frequency (the frequency can not be estimated from the available data):
- hepatosplenic T-cell lymphoma (a rare blood cancer that is often fatal)
- Merkel cell carcinoma (a type of skin cancer)
- Kaposi's sarcoma, a rare cancer related to infection with human herpes virus 8. Kaposi's sarcoma most commonly appears as purple lesions on the skin.
- liver failure
- worsening of dermatomyositis (looks like a skin rash accompanied by muscle weakness)
- weight gain (most patients had a minor weight gain)

Some of the side effects, observed when using the preparation, have no symptoms and can only be detected by blood tests. These include:
Very common side effects (effects occurring in more than 1 user in 10):
- low levels of white blood cells
- low levels of red blood cells
- increased lipids in the blood
- elevated liver enzymes

Common side effects (effects occurring in 1-10 in 100 users):
- high levels of white blood cells
- low levels of platelets
- increased uric acid in the blood
- abnormal levels of sodium in the blood
- low levels of calcium in the blood
- low levels of phosphate in the blood
- high blood sugar
- high levels of the lactate dehydrogenase enzyme in the blood
- presence of autoantibodies in the blood
- low levels of potassium in the blood

Uncommon side effects (effects occurring in 1-10 in 1,000 users):
- high bilirubin values (liver function blood test)

Rare side effects (effects occurring in 1-10 in 10,000 users):
- low levels of white blood cells, red blood cells and platelets

If a side effect occurs, if one of the side effects worsens, or if you are suffering from a side effect not mentioned in the leaflet, consult the doctor.
Side effects can be reported to the Ministry of Health by clicking on the link "Report Side Effects of Drug Treatment" found on the Ministry of Health homepage (www.health.gov.il), that directs you to the online form for reporting side effects or by entering the link: https://sideeffects.health.gov.il.
In addition, they can be reported to Padagis via the following address: Padagis.co.il

5. HOW SHOULD THE MEDICINE BE STORED?
- Avoid poisoning! This medicine and any other medicine must be kept in a safe place out of the reach and sight of children and/or infants to avoid poisoning. Do not induce vomiting unless explicitly instructed to do so by a doctor.
- Do not use the medicine after the expiry date (exp. date) that appears on the package and label. The expiry date refers to the last day of that month.
- Store refrigerated 2°C-8°C.
- Do not freeze.
- Store the pen in the original outer package to protect from light.

Alternative storage conditions:
- When needed, a **single** Yuflyma prefilled pen may be stored at a maximum temperature of up to 25°C for up to 31 days. The prefilled pen must be protected from light and discarded if not used within 31 days of taking it out of the refrigerator.
- Record date of removal from refrigerator on the carton package.
- Do not dispose of medicines via wastewater or household waste. Ask your doctor or pharmacist how to dispose of medicines you no longer need. These measures will help protect the environment.

6. FURTHER INFORMATION
- In addition to the active ingredient, the medicine also contains:
Glycine, Polysorbate 80, Sodium acetate trihydrate, Acetic acid, Water for injection

What does the medicine look like and what are the contents of the package?
- The package contains Yuflyma solution for injection in a prefilled pen for single use. The solution is colorless to light brown, clear to slightly cloudy.
The solution is a sterile solution of adalimumab in the following volumes:
40 mg/0.4 ml
80 mg/0.8 ml
Yuflyma (40 mg) solution for injection in a prefilled pen (Auto Injector-A)):
- 1 prefilled pen (0.4 ml sterile solution), with 2 alcohol pads.
- 2 prefilled pens (0.4 ml sterile solution), with 2 alcohol pads.
- 4 prefilled pens (0.4 ml sterile solution), with 4 alcohol pads.
- 6 prefilled pens (0.4 ml sterile solution), with 6 alcohol pads.

Yuflyma (80 mg) solution for injection in a prefilled pen (Auto Injector-A)):
- 1 prefilled pen (0.8 ml sterile solution), with 2 alcohol pads.
- 3 prefilled pens (0.8 ml sterile solution), with 4 alcohol pads.

Not all package sizes may be marketed.
- **Manufacturer and its address:** Celltrion Ltd., Incheon, South Korea.
- **Registration holder and its address:** Padagis Israel Agencies Ltd., 1 Rakefet St., Shoham, Israel.
- Revised in July 2024.
- Registration number of the medicine in the National Drug Registry of the Ministry of Health: 17196.37092
- For the sake of simplicity and ease of reading, this leaflet was written as intended for you. Although, the medicine is intended for you or for your child.

7. INSTRUCTIONS FOR USE
- The following instructions explain how to give yourself a subcutaneous injection of Yuflyma using the prefilled pen.
First, read all the instructions carefully and then follow them step by step.
- Your doctor or nurse will teach you the technique of self-injection.
- **Do not** attempt to self-inject the medicine until you are sure that you understand how to prepare and inject the medicine.
- After proper training, you can inject the medicine yourself or allow someone else to give you the injection, such as a family member or friend.
- Use each prefilled pen for one injection only.

Yuflyma prefilled pen

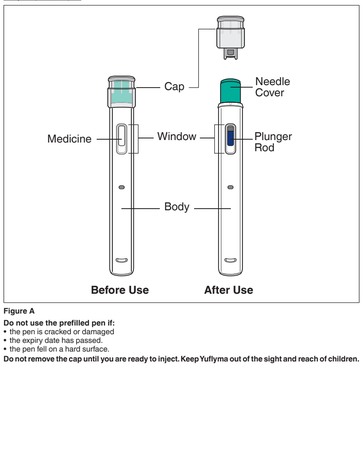

Figure A

Do not use the prefilled pen if:
- the pen is cracked or damaged
- the expiry date has passed.
- the pen fell on a hard surface.

Do not remove the cap until you are ready to inject. Keep Yuflyma out of the sight and reach of children.

1. Gather the supplies for the injection
a. Prepare a clean, flat surface, such as a table or work surface, in a well-lit area.
b. Remove one prefilled pen from the carton package stored in the refrigerator.
c. Make sure you have the following supplies:
- Prefilled pen
- one alcohol pad

Not included in the carton package:
- Cotton ball or gauze
- Adhesive bandage
- Sharps disposal container

2. Inspect the prefilled pen
a. Ensure you have the correct medicine (Yuflyma) and the correct dosage.
b. Look at the prefilled pen and make sure it is not cracked or damaged.
c. Check the expiry date on the label of the prefilled pen.

Do not use the prefilled pen if:
- it is cracked or damaged.
- the expiry date has passed.
- it has been dropped onto a hard surface.

Figure B

3. Inspect the medicine.
a. Look through the window and make sure that the liquid is clear, colorless to pale brown, and free of particles.
- **Do not** use the prefilled syringe if the liquid is discolored (yellow or dark brown), cloudy or contains particles in it.
- You may see air bubbles in the liquid. This is normal.

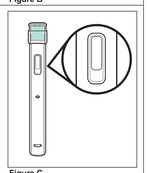

Figure C

4. Wait 15 to 30 minutes
a. Leave the prefilled pen at room temperature for 15 to 30 minutes to allow the solution to warm up.
- **Do not** warm the prefilled pen using heat sources such as hot water or a microwave.

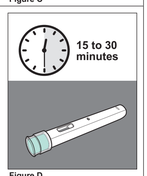

Figure D

5. Choose an appropriate injection site
a. You may inject into:
- the front area of the thighs.
- the abdomen except for 5 cm around the navel.
- the outer area of the upper arm (will be done by a caregiver or healthcare professional **ONLY**).

- **Do not** inject into skin in the area within 5 cm of the navel, or into red, hard, tender, damaged, bruised, or scarred skin.
- If you have psoriasis, do not inject directly into any raised, thick, red or scaly skin patches or lesions on your skin.
- **Do not** inject through clothes.

b. Rotate the injection site each time you give an injection. Each new injection site should be at least 3 cm away from the previous injection site.

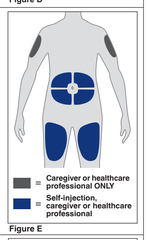

Figure E

6. Wash your hands
a. Wash your hands with soap and water and dry them thoroughly.

Figure F

7. Clean the injection site
a. Clean the injection site with an alcohol pad using a circular motion.
b. Let the skin dry before injecting.
- **Do not** blow on or touch the injection site again before giving the injection.

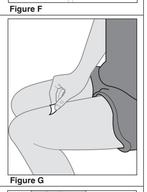

Figure G

8. Remove the cap
a. With one hand, hold the prefilled pen by the injector body with the cap facing upwards. Gently pull the cap straight off with the other hand.
- **Do not** remove the cap until you are ready to inject.
- **Do not** touch the needle or needle cover. Doing so may result in a needle stick injury.
- **Do not** recap the prefilled pen. Dispose of the cap immediately into a special container.
- It is normal to see a few drops of liquid come out of the needle.

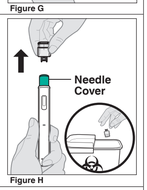

Figure H

9. Place the prefilled pen over the injection site.
a. Hold the prefilled pen such that you can see the window.
b. Without pinching or stretching the skin, place the prefilled pen over the injection site at a 90-degree angle.

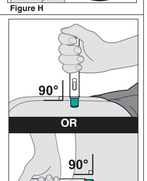

Figure I

10. Begin the injection
a. Press the prefilled pen **firmly** against the skin. When the injection starts, you will hear the first loud "click" and the blue plunger rod will begin to fill the window.
b. Keep holding the prefilled pen firmly against the skin and listen for the second loud "click."
c. After you hear the second loud "click," continue to hold the prefilled pen firmly against the skin and count slowly to 5 to make sure you inject the full dose.
- **Do not** change the position of the prefilled pen after the injection has started.

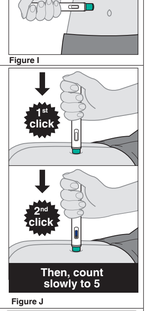

Figure J

11. Remove the prefilled pen from the injection site and care for the injection site
a. Look at the prefilled pen and make sure that the blue plunger rod with the gray top fills the window completely.
b. Remove the prefilled pen from your skin.
- After you remove the prefilled pen from the injection site, the needle will be automatically covered.
Do not recap the pen.
- If the window has not turned completely blue or if the medicine is still injecting, this means you have not received a full dose. Call your doctor or healthcare provider immediately.

c. Treat the injection site by gently pressing, without rubbing, a cotton ball or gauze on the injection site and apply an adhesive bandage, if necessary. Some bleeding may occur.
- **Do not** reuse the prefilled pen.
- **Do not** rub the injection site.

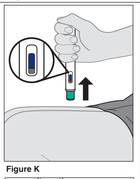

Figure K

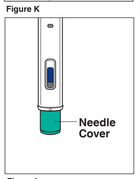

Figure L

12. Dispose of the prefilled pen
a. Throw away the used prefilled pen in a special container as instructed by your doctor, nurse or pharmacist.
b. The alcohol pad and packaging may be disposed of in the household waste.
- **Always** keep the prefilled pen and the special container out of the reach and sight of children.

Figure M

08/07/2024