

PATIENT INFORMATION

RASUVO® (ruh-SOO-voh) (methotrexate) injection, for subcutaneous use

What is Rasuvo?

Rasuvo is a single-dose manually-triggered auto-injector containing a prescription medicine, methotrexate. Methotrexate is used to:

- treat certain adults with severe, active rheumatoid arthritis (RA), and children with active polyarticular juvenile idiopathic arthritis (pJIA), after treatment with other medicines including non-steroidal anti-inflammatory (NSAIDs) have been used and did not work well.
- control the symptoms of severe, resistant, disabling psoriasis in adults when other types of treatment have been used and did not work well.

Rasuvo is available in doses of 7.5, 10, 12.5, 15, 17.5, 20, 22.5, 25, 27.5 and 30 mg. Your doctor will prescribe a different way to take methotrexate if you need to take methotrexate by mouth or in some other way. Your doctor may also change your prescription if your dose does not match the available Rasuvo doses, such as doses of less than 7.5 mg, more than 30 mg, or doses in between the available Rasuvo doses.

Rasuvo should not be used for the treatment of cancer.

Rasuvo should not be used for the treatment of children with psoriasis.

What is the most important information I should know about Rasuvo?

Rasuvo can cause serious side effects that can lead to death, including:

1. Organ system toxicity. People who use methotrexate for the treatment of cancer, psoriasis, or rheumatoid arthritis, have an increased risk of death from organ toxicity. Types of organ toxicity can include:

- | | |
|--------------------|-----------|
| ○ gastrointestinal | ○ nerve |
| ○ bone marrow | ○ lung |
| ○ liver | ○ kidneys |
| ○ immune system | ○ skin |

Your doctor will do blood tests and other types of tests before you take and while you are taking Rasuvo to check for signs and symptoms of organ toxicity. Call your doctor right away if you have any of the following symptoms of organ toxicity:

- | | |
|---------------|------------------------------|
| ○ vomiting | ○ neck stiffness |
| ○ diarrhea | ○ paralysis |
| ○ mouth sores | ○ irritability |
| ○ fever | ○ sleepiness |
| ○ confusion | ○ problems with coordination |
| ○ weakness | ○ dry cough |

- temporary blindness
- seizures
- headache
- back pain
- trouble breathing
- severe skin rash
- infection

2. Women who are pregnant are at increased risk for death of the baby and birth defects. Women who are pregnant or who plan to become pregnant **must not take Rasuvo**. A pregnancy test should be performed before starting Rasuvo.

Contraception should be used by both females and males while taking Rasuvo. Pregnancy should be avoided if either partner is receiving Rasuvo:

- for a minimum of 3 months after treatment with Rasuvo for males.
- during and for at least 1 menstrual cycle after treatment with Rasuvo for females.

Who should not take Rasuvo?

Do not take Rasuvo if you:

- are pregnant or planning to become pregnant. See **“What is the most important information I should know about Rasuvo?”**
- are breastfeeding.
- Rasuvo can pass into your breast milk and may harm your baby. **Do not** breastfeed while taking Rasuvo. Talk to your doctor about the best way to feed your baby if you take Rasuvo.
- have alcohol problems (alcoholism)
- have liver problems
- have problems fighting infection (immunodeficiency syndrome)
- have been told you have (or think you have) a blood disorder such as low levels of white blood cells, red blood cells (anemia), or platelets.
- have had an allergy to methotrexate or any of the ingredients in Rasuvo. See the end of this leaflet for a complete list of ingredients in Rasuvo.

Talk to your doctor before taking this medicine if you have any of these conditions.

What should I tell my doctor before taking Rasuvo?

Before you take Rasuvo, tell your doctor if you have any other medical conditions.

Tell your doctor about all of the medicines you take, including prescription, over-the-counter medicines, vitamins, and herbal supplements.

Rasuvo may affect how other medicines work, and other medicines may affect how Rasuvo works causing side effects.

Ask your doctor or pharmacist for a list of medicines if you are not sure.

Know the medicines you take. Keep a list of them to show your doctor and pharmacist when you get a new medicine.

How should I take Rasuvo?

- **Read the Instructions for Use that come with Rasuvo.**
- Take Rasuvo exactly as your doctor tells you to take it.
- Inject Rasuvo **only 1 time each week. Do not** take Rasuvo every day.
- Taking Rasuvo every day may cause death from toxicity.
- Your doctor will show you or your caregiver how to inject Rasuvo. You should not inject Rasuvo until you have been trained on the right way to use it.
- Check Rasuvo before you inject it. Rasuvo should be yellow to brown in color and should not have any lumps or particles in it.
- Rasuvo should be injected under the skin of the abdomen or thigh.
- **Do not** inject Rasuvo within 2 inches of the belly button (navel)
- Use a different site each time you inject. This may help to decrease any reactions at the injection site.
- **Do not** inject Rasuvo in the arms or any other areas of the body.
- **Do not** inject Rasuvo in areas where the skin is tender, bruised, red, scaly, hard, or has scars or stretch marks.
- If you are not sure if Rasuvo was injected, or if you have hard time giving the injection, **do not** inject another dose. Call your pharmacist or doctor right away.
- If you inject too much Rasuvo, call your doctor or go to the nearest hospital emergency room right away.

What should I avoid while taking Rasuvo?

- Do not drink alcohol while taking Rasuvo. Drinking alcohol can increase your chances of getting serious side effects.
- Rasuvo can cause dizziness and tiredness. Do not drive a car, operate machinery, or do anything that needs you to be alert until you know how Rasuvo affects you.
- Certain vaccinations should be avoided while taking Rasuvo. Talk to your doctor before you or members of your household receive any vaccines.

What are the possible side effects of Rasuvo?

Rasuvo may cause serious side effects, including:

See **“What is the most important information I should know about Rasuvo?”**

- **fertility problems.** Methotrexate, the active ingredient in Rasuvo, may affect your ability to have a baby. Males may have a decreased sperm count, and females may have changes to their menstrual cycle. This can happen while taking Rasuvo and for a short period of time after you stop.
- **certain cancers.** Some people who have taken methotrexate have had a certain type of cancer called Non-Hodgkin’s lymphoma and other tumors. Your doctor may tell you to stop taking Rasuvo if this happens.
- **tissue and bone problems.** Taking methotrexate while having radiation therapy may increase the risk of your tissue or bone not receiving enough blood. This may lead to death of the tissue or bone.

Common side effects of Rasuvo include:

- o nausea
- o stomach pain
- o indigestion (dyspepsia)
- o mouth sores
- o rash
- o stuffy or runny nose and sore throat
- o diarrhea
- o abnormal liver function tests
- o vomiting
- o headache
- o bronchitis
- o low red, white, and platelet blood cell count
- o hair loss
- o dizziness
- o sensitivity to light
- o burning skin lesions
- o lung problems

Tell your doctor if you have any side effect that bothers you or that does not go away.

These are not all the possible side effects of Rasuvo. For more information, ask your doctor or pharmacist.

Call your doctor for medical advice about side effects. You may report side effects to FDA at 1-800-FDA-1088.

How should I dispose of Rasuvo?

- **Do not throw away in the household trash.** Put used Rasuvo in an FDA-cleared sharps disposal container right away after use.
- If you do not have an FDA-cleared sharps disposal container, you may use a household container that is:
 - o made of a heavy-duty plastic
 - o can be closed with a tight-fitting, puncture-resistant lid, without sharps being able to come out
 - o upright stable during use
 - o leak-resistant
 - o properly labeled to warn of hazardous waste inside the container
- When your sharps disposal container is almost full, you will need to follow your community guidelines for the right way to dispose of your sharps disposal container. There may be state or local laws about how you should throw away used needles and syringes. For more information about the safe sharps disposal, and for specific information about sharps disposal in the state that you live in, go to the FDA's website at: <http://www.fda.gov/safesharpsdisposal>.
- Do not dispose of your used sharps disposal container in your household trash unless your community guidelines permit this. Do not recycle your used sharps disposal container.
- Safely dispose of Rasuvo that is out of date or is no longer needed.

How should I store Rasuvo?

Store Rasuvo at room temperature between 68°F to 77°F (20°C to 25°C)

- Do not freeze
- Keep Rasuvo out of the light.

Keep Rasuvo and all medicines out of the reach of children.

General information about the safe and effective use of Rasuvo.

Methotrexate is sometimes prescribed for purposes other than those listed in the Patient Information leaflet. Do not use Rasuvo for a condition for which it was not prescribed. Do not give Rasuvo to other people, even if they have the same symptoms that you have. It may harm them.

This Patient Information leaflet summarizes the most important information about Rasuvo. If you would like more information, talk to your doctor. You can ask your doctor or pharmacist for information about Rasuvo that is written for health professionals.

For more information, please contact Medac Pharma, Inc. at our number 1-855-336-3322.

What are the ingredients in Rasuvo?

Active ingredient: methotrexate

Inactive ingredients: sodium chloride, sodium hydroxide and water for injection, USP, and if necessary hydrochloric acid, USP.

This Patient Information has been approved by the U.S. Food and Drug Administration.

Manufactured for:

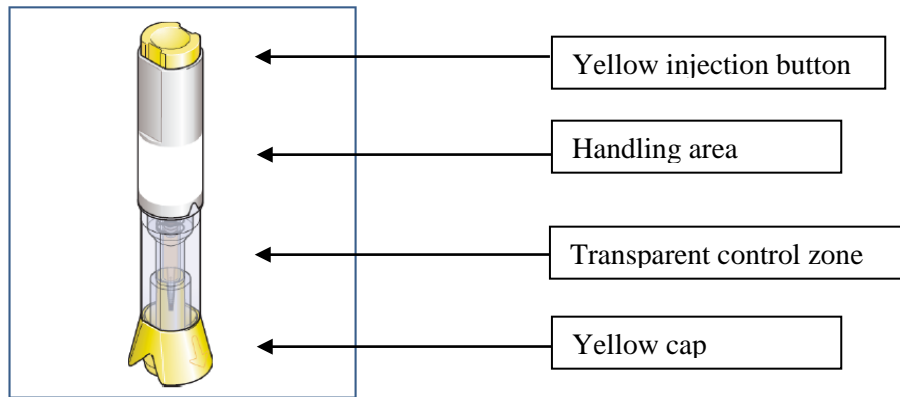
Medac Pharma, Inc.

29 N Wacker Drive, Suite 704
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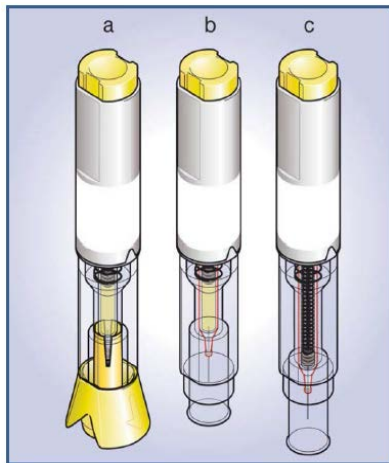
Issued: 11/2014

**Read this Instructions for Use before using
Rasuvo® (ruh-SOO-voh) (methotrexate) injection, for subcutaneous use
Follow these instructions each time you use Rasuvo®.**

Parts of your Rasuvo auto-injector



(Figure A)



(Figure B)

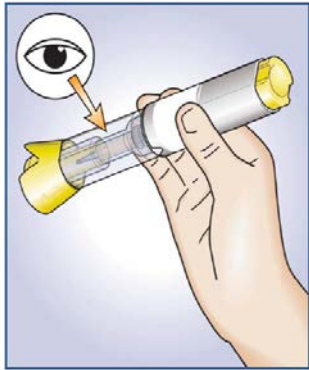
- Pre-filled auto-injector with cap before injection
- Pre-filled auto-injector after cap removal before injection
- Pre-filled auto-injector after injection

Prepare to Use Rasuvo

- Wash your hands well with soap and warm water.

- Select a clean, well-lit, flat work surface, such as a table.
- Place the Rasuvo carton containing the auto-injector on your flat work surface.
- Be sure that the dose, either 7.5, 10, 12.5, 15, 17.5, 20, 22.5, 25, 27.5 or 30 mg, stated on the carton is the same as the dose prescribed by your doctor.
- Check the expiration date on the label. **Do not** use if expired.
- Remove one Rasuvo auto-injector from the packaging.
- If the Rasuvo appears to be damaged **do not** use it. Use another Rasuvo.
- In addition to Rasuvo, you will need the following items: one alcohol swab and one cotton ball or gauze and small adhesive bandage strip, if desired.

Check the Liquid



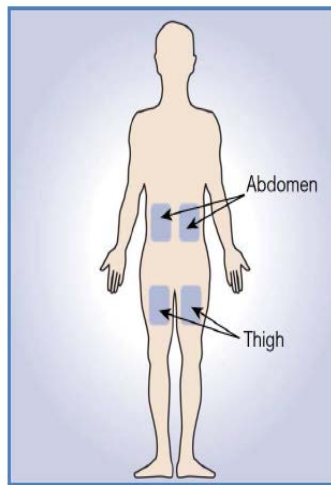
(Figure C)

- Look at the transparent control zone (see Figure C). The prefilled syringe is visible within the transparent control zone. Examine the contents of the syringe carefully. If the syringe is cracked or broken, **do not** use it. Use another auto-injector.
- The liquid should be clear and yellow to brown in color and should not have any lumps or particles in it. **Do not** use Rasuvo if the liquid is cloudy, discolored or contains particles.
- You may see an air bubble. This is normal.
- If you are not able to see or to check the Rasuvo auto-injector correctly prior to injection, ask a caretaker for assistance.
- **Do not** remove the yellow cap from the auto-injector until you are ready to use Rasuvo.

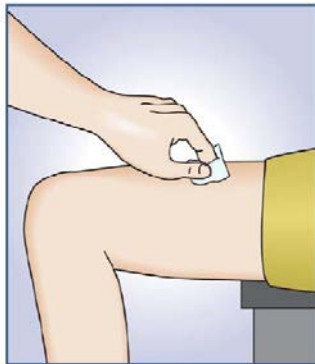
Choose an Injection Site

- Rasuvo should be injected into the stomach (abdomen) or the upper thigh. **Do not** inject Rasuvo within 2 inches of the belly button (navel) (see Figure D).
- **Do not** inject Rasuvo in the arms or any other areas of the body.

- **Do not** inject Rasuvo in areas where the skin is tender, bruised, red, scaly, hard, or has scars or stretch marks.
- Use a different site each time you inject. This may help to decrease any reactions at the injection site.
- Wipe the area with an alcohol swab (**see Figure E**).
- Allow the skin to dry and **do not** touch this area again before giving Rasuvo. **Do not** fan or blow on the clean area.



(Figure D)



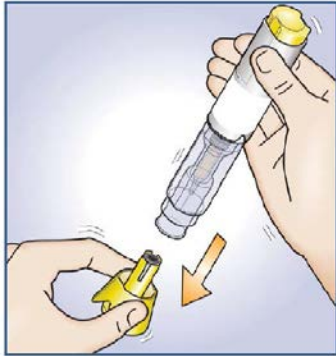
(Figure E)

Give your Injection

STEP 1: Remove the Yellow Cap (see Figure F)

- Hold the Rasuvo auto-injector with one hand in the handling area.

- Use your other hand to pull the yellow cap straight off (**see Figure F**). **Do not** twist the cap. If you are unable to remove the cap, ask a caretaker for assistance.
- You may notice 1 or 2 drops of medicine. This is normal.



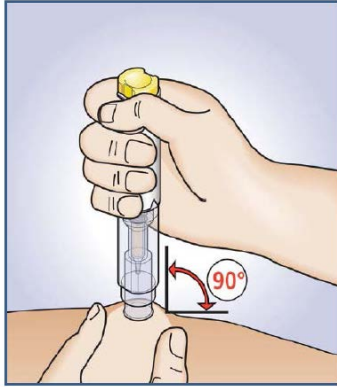
(Figure F)

Caution!

- **Do not** touch the needle end with your hands or fingers.
- This could inject the medicine into your hand.
- To avoid any injury, **never insert your fingers in the opening of the protect tube** covering the needle.
- **Do not** replace the cap after it has been removed.
- After the cap is removed Rasuvo must be used without delay or disposed of safely.
- **Do not** press the yellow injection button until you are ready to inject Rasuvo.

STEP 2: Prepare the Injection

- Pinch a pad of skin surrounding the cleaned injection site with your thumb and forefinger of your free hand by gently squeezing. Patients with rheumatoid arthritis who are unable to pinch the skin can inject directly into the thigh without pinching if needed.
- Hold the skin pinched until Rasuvo is removed from the skin after the injection.
- Position the uncapped transparent end of the Rasuvo auto-injector perpendicular (at a 90 degree angle) to the skin (**see Figure G**).
- Without pressing the button, **push Rasuvo firmly onto your skin until you feel the stop point** in order to unlock the yellow injection button (**see Figure G**).
- If you are unable to push Rasuvo to the stop-point, ask a caretaker for assistance.



(Figure G)

STEP 3: Inject Rasuvo

- While still holding Rasuvo firmly against the skin, **press the yellow injection button** with your thumb (see Figure H).
- You will hear a click which indicates the start of the injection. **Hold Rasuvo against the skin until all of the medicine is injected. This can take up to 5 seconds** (slowly count 1, 2, 3, 4, 5). **To avoid an incomplete injection**, do not remove the Rasuvo from the skin before the end of the injection.
- **Look at the transparent control zone while you are injecting to make sure that the entire dose is injected.** When the movement stops, the injection is completed.
- If you have problems with your hearing, slowly count to **5 seconds** from the moment you have pressed the button.
- It is not necessary to keep the button of the Rasuvo pressed down with your thumb after the injection has begun.

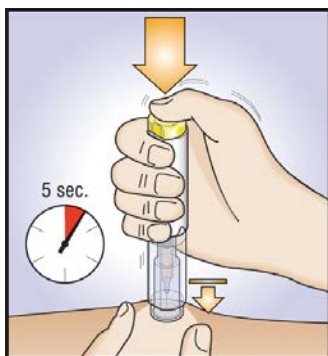


Figure H

- **After completing the injection, remove Rasuvo from the injection site** by pulling straight up (perpendicular to the skin).
- The protective needle shield automatically moves into place and locks over the needle.

- Put a small adhesive bandage strip over the injection site, if desired.

STEP 4: Check the Transparent Control Zone

Check visually to make sure **that there is no liquid left in the** syringe inside **the transparent control zone.**

- If there is liquid left, not all of the medicine has been injected correctly. Consult your doctor or health care professional immediately. **Do not** use another Rasuvo, unless advised by your doctor.

STEP 5: Dispose of Rasuvo

- Each Rasuvo can be used only 1 time.
- **Do not throw away in the household trash.** Put used Rasuvo in an FDA-cleared sharps disposal container right away after use.
- If you do not have an FDA-cleared sharps disposal container, you may use a household container that is:
 - made of a heavy-duty plastic
 - can be closed with a tight-fitting, puncture-resistant lid, without sharps being able to come out
 - upright and stable during use
 - leak-resistant
 - properly labeled to warn of hazardous waste inside the container
- When your sharps disposal container is almost full, you will need to follow your community guidelines for the right way to dispose of your sharps disposal container. There may be state or local laws about how you should throw away used needles and syringes. For more information about the safe sharps disposal, and for specific information about sharps disposal in the state that you live in, go to the FDA's website at: <http://www.fda.gov/safesharpsdisposal>.
- Do not dispose of your used sharps disposal container in your household trash unless your community guidelines permit this. Do not recycle your used sharps disposal container.
- Safely dispose of Rasuvo that is out of date or no longer needed.

How should I store Rasuvo?

- Store Rasuvo at room temperature between 68°F to 77°F (20°C to 25°C). Do not freeze.
- Keep Rasuvo out of the light.
- Always keep your Rasuvo out of the reach of children.

This Instructions for Use has been approved by the U.S. Food and Drug Administration.

Manufactured for:
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