



3-13-03-Final Draft submitted by HLR

MEDICATION GUIDE

Roferon[®]-A

(Interferon alfa-2a, recombinant)

Solution for Injection - Vials

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Before you start taking Roferon-A (ro-FER-on), please read this Medication Guide carefully. Read this Medication Guide each time you refill your prescription in case new information has been added. This information does not take the place of talking with your healthcare provider.

What is the most important information I should know about Roferon-A?

Roferon is used to treat people with hepatitis C, hairy cell leukemia, AIDS-related Kaposi's sarcoma, and Philadelphia chromosome positive chronic myelogenous leukemia (CML). However, Roferon-A can cause some serious side effects that may cause death in rare cases. Before starting Roferon-A, you should talk with your healthcare provider about the possible benefits and the possible side effects of treatment, to decide if Roferon-A is right for you. While taking Roferon-A, you will need to see your healthcare provider regularly for medical examinations and blood tests to make sure your treatment is working and to check for side effects.

The most serious possible side effects of Roferon-A treatment include:

1. **Mental health problems:** Roferon-A may cause some patients to develop mood or behavioral problems. Signs of these problems include irritability (getting easily upset), depression (feeling low, feeling bad about yourself or feeling hopeless), and anxiety. Some patients may have aggressive behavior and think about hurting others. Some patients may develop thoughts about ending their lives (suicidal thoughts) and may attempt to do so. A few patients have even ended their lives. Former drug addicts may fall back into drug addiction or overdose. You must tell your healthcare provider if you are being treated for a mental illness or have a history of mental illness or if you are or have ever been addicted to drugs or alcohol. Call your healthcare provider immediately if you develop any of these problems while on Roferon-A treatment.
2. **Heart problems:** Roferon-A may cause some patients to experience high blood pressure, a fast heartbeat, chest pain, and very rarely a heart attack. Tell your healthcare provider if you have or have had any heart problems in the past.
3. **Blood problems:** Many patients taking Roferon-A have had a drop in the number of their white blood cells and their platelets. If the numbers of these blood cells are too low, you could be at risk for infections or bleeding.

37 **Stop taking Roferon-A and call your healthcare provider immediately if you**
38 **develop any of these symptoms:**

- 39 • **You become very depressed or think about suicide**
- 40 • **You have severe chest pain**
- 41 • **You have trouble breathing**
- 42 • **You have a change in your vision**
- 43 • **You notice unusual bleeding or bruising**
- 44 • **High fever**
- 45 • **Severe stomach pain. If the pain is in the lower part of your stomach area it**
46 **could mean that your bowels are inflamed (colitis)**
- 47

48 *For more information on possible side effects with Roferon-A therapy, please read the*
49 *section on "What are the possible side effects of Roferon-A?" in this Medication Guide.*

50 **What is Roferon-A?**

51 Roferon-A is a treatment that is used for some people who are infected with the hepatitis
52 C virus, hairy cell leukemia, AIDS-related Kaposi's sarcoma, and Philadelphia
53 chromosome positive chronic myelogenous leukemia (CML). Patients with hepatitis C
54 have the virus that causes hepatitis in their blood and liver. Patients with hairy cell
55 leukemia produce abnormal white blood cells that travel to the spleen where they trap and
56 destroy normal blood cells. In AIDS-related Kaposi's sarcoma, the virus affects the
57 immune system causing many patients to develop lesions on the skin, in the nose, mouth,
58 and sometimes in the lungs, bowels, and lymph nodes. In CML, your body produces too
59 many of certain blood cells. Roferon-A works in these conditions by reducing the amount
60 of virus in the body, destroying cells that may be harmful to your body and keeping the
61 body from producing too many cells.

62 **Who should not take Roferon-A?**

63 Do not use Roferon-A if:

- 64 • **You are pregnant or breast-feeding or are planning to become pregnant**
- 65 • **You are allergic to alpha interferons, *Escherichia coli*-derived products or any**
66 **component of Roferon-A (see ingredients listed at the end of this Medication Guide)**
- 67 • **You have autoimmune hepatitis (hepatitis caused by your immune system attacking**
68 **your liver)**
- 69

70 **If you have or have had any of the following conditions or serious medical problems,**
71 **discuss them with your doctor before taking Roferon-A:**

- 72 • History of or current severe mental illness (such as depression or anxiety)
- 73 • Previous heart attack or heart problems
- 74 • Sleep problems
- 75 • High blood pressure
- 76 • Autoimmune disease (where the body's immune system attacks the body's own
- 77 cells), such as vasculitis, psoriasis, systemic lupus erythematosus, rheumatoid arthritis
- 78 • Kidney problems
- 79 • Blood disorders-Low blood counts or bleeding problems
- 80 • You take a medicine called theophylline
- 81 • Diabetes (high blood sugar)
- 82 • Thyroid problems
- 83 • Liver problems, other than hepatitis C
- 84 • Hepatitis B infection
- 85 • HIV infection (the virus that causes AIDS)
- 86 • Problems with your vision
- 87 • Colitis
- 88 • Body organ transplant and are taking medicine that keeps your body from rejecting
- 89 your transplant (suppresses your immune system)
- 90 • Alcoholism
- 91 • Drug abuse or addiction

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93 If you have any doubts about your health condition or about taking Roferon-A, talk to
94 your healthcare provider.

95 **What should I avoid while taking Roferon-A?**

- 96 • Avoid becoming pregnant while taking Roferon-A. Roferon-A may harm your
- 97 unborn child or cause you to lose your baby (miscarry)
- 98 • You should not breast-feed your baby while taking Roferon-A

99 **How should I take Roferon-A?**

100 To get the most benefit from this medicine, it is important to take Roferon-A exactly as
101 your healthcare provider tells you.

102 Your healthcare provider will tell you how much medicine to take and how often to take
103 it. Once you start treatment with Roferon-A, do not switch to another brand of interferon
104 without talking to your doctor. Other interferons may not have the same effect on the
105 treatment of your disease. Switching brands will also require a change in your dose. Your
106 healthcare provider will tell you how long you need to use Roferon-A.

107 Over time, your healthcare provider may change your dose of Roferon-A. Do not change
108 your dose unless your doctor tells you to change it.

109 Roferon-A is supplied in vials as a ready-to-use solution and in prefilled syringes. This
110 Medication Guide describes how to use the vials. Whether you give yourself the injection
111 or another person gives the injection to you, it is important to follow the instructions in
112 this Medication Guide (see the section, "How do I inject Roferon-A?").

113 If you miss a dose of Roferon-A, take the missed dose as soon as possible during the
114 same day or the next day, then continue on your regular dosing schedule. If several days
115 go by after you miss a dose, check with your doctor about what to do. Do not double the
116 next dose or take more than one dose a day unless your doctor tells you to. Call your
117 doctor right away if you take more than your prescribed Roferon-A dose. Your doctor
118 may wish to examine you more closely and take blood for testing.

119 You must get regular blood tests to help your healthcare provider check how the
120 treatment is working and to check for side effects.

121 Tell your doctor if you are taking or planning to take other prescription or non-
122 prescription medicines, including vitamins and mineral supplements and herbal
123 medicines.

124 **What are the possible side effects of Roferon-A?**

125 Possible, serious side effects include:

- 126 • **Mental health problems including suicide, suicidal thoughts, heart problems,**
127 **and blood problems:** See "What is the most important information I should know
128 about Roferon-A?"
- 129 • **Other body organ problems:** Some patients may experience lung problems (such as
130 difficulty breathing or pneumonia) and vision problems
- 131 • **New or worsening autoimmune disease:** Some patients may develop an
132 autoimmune disease (a disease where the body's own immune system begins to attack
133 itself) while on Roferon-A therapy. These diseases can include vasculitis (an
134 inflammation of your blood vessels), rheumatoid arthritis or lupus erythematosus,
135 psoriasis or thyroid problems. In some patients who already have an autoimmune
136 disease, the disease may worsen while on Roferon-A therapy

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138 Common, but less serious, side effects include:

- 139 • **Flu-like symptoms:** Most patients who take Roferon-A have flu-like symptoms that
140 usually lessen after the first few weeks of treatment. Flu-like symptoms may include
141 unusual tiredness, fever, chills, muscle aches, and joint pain. Taking acetaminophen
142 or ibuprofen before you take Roferon-A can help with these symptoms. You can also
143 try taking Roferon-A at night. You may be able to sleep through the symptoms
- 144 • **Extreme fatigue (tiredness):** Many patients may become extremely tired while on
145 Roferon-A therapy
- 146 • **Upset stomach:** Nausea, taste changes, diarrhea, and loss of appetite occur

- 147 commonly
- 148 • **Blood sugar problems:** Some patients may develop a problem with the way their
 - 149 body controls their blood sugar and may develop diabetes
 - 150 • **Thyroid problems:** Some patients may develop changes in their thyroid function.
 - 151 Symptoms of these changes may include feeling hot or cold all the time, trouble
 - 152 concentrating, changes in your skin (your skin may become very dry), and changes in
 - 153 your weight
 - 154 • **Skin reactions:** Some patients may develop a rash, dry or itchy skin, and redness and
 - 155 swelling at the site of injection
 - 156 • **Sleep disturbances and headache:** Trouble sleeping and headaches may also occur
 - 157 during Roferon-A therapy
 - 158 • **Hair thinning:** Hair loss is not uncommon while using Roferon-A. This hair loss is
 - 159 temporary and hair growth should return after you stop taking Roferon-A

160
161 These are not all of the side effects of Roferon-A. Your doctor or pharmacist can give
162 you a more complete list.

163 Talk to your healthcare provider if you are worried about side effects or find them very
164 bothersome.

165 **How do I inject Roferon-A?**

166 The instructions that follow will help you learn how to use Roferon-A vials, ~~and prefilled~~
167 ~~syringes~~. Please read all of these directions before trying to take your medicine. It is
168 important to follow these directions carefully. Talk to your healthcare provider if you
169 have any concerns about how to use Roferon-A. Whether you are giving yourself an
170 injection or if you are giving this injection to someone else, a healthcare provider must
171 teach you how to inject. If you are giving this injection to someone else, a healthcare
172 provider must teach you how to avoid needle-sticks. Being stuck by a used needle can
173 pass diseases on to you.

174 **General advice about prescription medicines**

175 Medicines are sometimes prescribed for purposes other than those listed in a Medication
176 Guide. If you have any concerns or questions about Roferon-A, contact your healthcare
177 provider. Do not use Roferon-A for a condition or person other than that for which it is
178 prescribed. If you want to know more about Roferon-A, your healthcare provider or
179 pharmacist will be able to provide you with detailed information that is written for
180 healthcare providers.

181 **Ingredients:** interferon alfa-2a recombinant, sodium chloride, polysorbate 80, benzyl
182 alcohol, and ammonium acetate.

This Medication Guide has been approved by the U.S. Food and Drug Administration.

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APPENDIX: Use of Roferon-A Solution in Vials

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187 Your doctor will tell you whether you will be injecting Roferon-A under the surface of
188 the skin (subcutaneous) yourself or if your caregiver will be giving you injections into
189 your muscle (intramuscular). The size of the syringe and needle you will use depends on
190 where you will be getting your injections.

191 Find a well-lit, clean, flat working surface like a table and remove a vial of Roferon-A
192 from the refrigerator. You should check the date on the vial and make sure that it has not
193 expired. **Do not use a vial if the date on the label has passed.**

1. Collect all the supplies you will need to give yourself or get an injection:

- one vial of Roferon-A

196

- the correct sterile disposable syringe and needle (as directed by your doctor or
197 healthcare provider)

197

- two alcohol swabs

199

- puncture-resistant disposable container

200

201 2. Check the vial of Roferon-A to see if it is clear or colorless to a light yellow in color
202 and check the date on the vial to make sure it has not expired.

- Do not use Roferon-A if:

the medicine is cloudy

the medicine has particles floating in it

the medicine is any color besides clear or colorless to light yellow

it has passed the expiration date

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209 3. Gently warm the refrigerated medicine by slowly rolling the vial in the palms of your
210 hands for about one minute.

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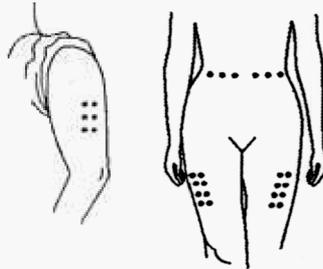
4. Wash your hands with soap and warm water. This step is very important to help
212 prevent infection.

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5. Pick a site for the injection.

214 A. For injections under the skin (subcutaneous) the following sites may be used

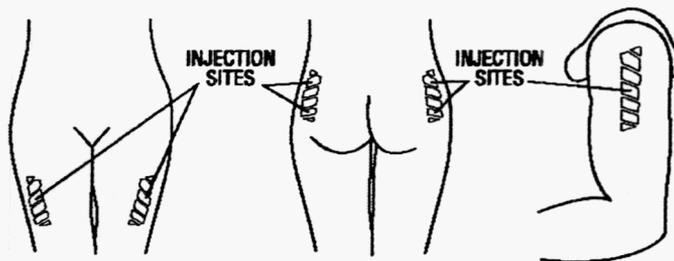
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- abdomen, avoiding the navel and waistline area
 - If someone else is giving you an injection you can also use the outer surface of your upper arm



- 219
- 220
- You should choose a different spot each time you give or receive an injection

B. For injections into the muscle (intramuscular) the following sites may be used:

- 222
- 223
- 224
- If someone else is giving you an injection you can use the upper, outer quadrant of the buttocks and the upper outer arm
 - If you are giving yourself the injection you can use the upper thigh



- 227
- 228
- Again, you should choose a different injection site each time you give or receive an injection

6. Preparing the injection site:

- 230
- 231
- Clean the skin where the injection will be given with an alcohol swab and leave the swab on the site

7. Preparing your dose of Roferon-A:

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- 234
- A. Using your thumb, flip off the plastic top covering the top of vial. Clean the rubber stopper on the top of the vial with an alcohol swab



B. Remove the syringe from the package

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C. Remove the cover from the needle. Do not use the syringe if the needle cover is not on the needle securely

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D. Pull the syringe plunger back to the mark that corresponds to the dose of Roferon-A prescribed by your doctor. This draws air into the syringe barrel



E. Push the needle through the center of the stopper on the vial

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F. Slowly inject all the air from the syringe into the space above the solution. Do not inject air into the Roferon-A fluid



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248 G. Keep needle in vial and turn both upside down

249 H. Holding the vial and syringe straight up, while keeping the tip of the needle in the
250 fluid, slowly pull the plunger back to the mark on syringe barrel ordered by your
251 doctor

252 I. To avoid air bubbles from forming in the syringe, make sure the needle tip always
253 stays in the liquid. If air bubbles appear, gently tap the syringe barrel then slowly
254 push the solution back into vial and withdraw the solution again



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J. Leave the needle in the vial and lay the vial and needle down on your clean working surface

K. Withdraw the needle from vial

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L. Hold syringe and needle straight up, check again for air bubbles. If there are bubbles, lightly tap the side of syringe with your thumb and finger to bring any air bubbles to the top.

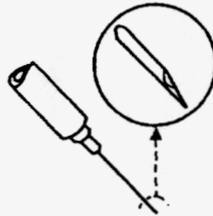
M. Press plunger slightly to push the air out through the needle.

8. For injections under the skin (subcutaneous) follow the instructions below:

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- Remove the alcohol swab from the site you cleaned for the injection. Allow the site to dry for 10 seconds
- Position the bevel of the needle so that it is (at point of needle) facing up



- Pinch an area of skin firmly between the thumb and forefinger



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- Hold the needle like a pencil or a dart, at a 45° to 90° degree angle to skin and in one quick motion insert the needle as far as it will go



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- Once inserted, draw back slowly on the syringe. If blood appears in the syringe, the needle has entered a blood vessel

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Do not inject Roferon-A at that site and discard the filled syringe and needle. Prepare a new syringe for the injection and use a different injection site.

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- If blood does not appear in the syringe, then slowly push the plunger all the way down so that you get all of your medicine

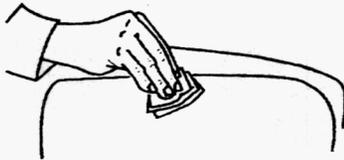
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- Withdraw needle at same angle it was inserted

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- When you are finished, place an alcohol swab over the injection site and press slightly

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- Place the syringe and needle in a puncture-resistant container

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- If you are using a Single-Use vial discard any remaining solution. If you are using the 18 million IU Multi-Dose vial, return Roferon-A to the refrigerator for storage

291

292

9. For injections into the muscle (intramuscular), follow the instructions below:

- With one hand, stretch the skin away from the injection site so it is tight
- Hold syringe in the other hand like a pencil, and insert the needle at a 90° angle with a quick dart-like thrust. You should feel some resistance

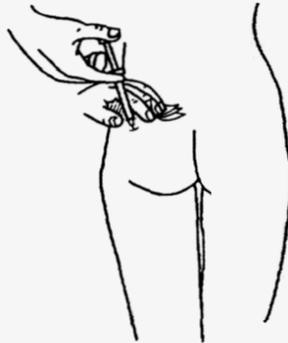
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- Once inserted, release the stretched skin and slowly inject the medication by pushing the plunger down



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- Withdraw the needle at the same angle it was inserted
- When you are finished, place an alcohol swab over the site and press lightly



- Apply light pressure over site
- Place the syringe and needle in a puncture-resistant container

- 308 ● If you are using a Single-Use vial discard any remaining solution. If you are
309 using the 18 million IU Multi-Dose vials return Roferon-A to the refrigerator
310 for storage.

311 **How should I dispose of materials used to inject Roferon-A?**

- Do not recap the needle
- Place the entire syringe and needle in a puncture-resistant container. A home “Sharps Container” may be purchased at your pharmacy or you can use a hard plastic container with a screw top or a coffee can with a plastic lid. You should talk to your healthcare provider about how to properly dispose of a full container of used syringes. There may be special state or local laws about disposing used syringes and needles, so please check with your physician, nurse or pharmacist for instructions. DO NOT throw the filled container in the household trash and DO NOT recycle
- The needle cover and alcohol swabs can be thrown in the regular trash. You should always keep your syringes and disposal container out of the reach of children

322 **How should I store Roferon-A?**

323 Roferon-A must be stored in the refrigerator at a temperature of 36°F to 46°F (2°C to
324 8°C). Do not leave Roferon-A outside of the refrigerator for more than 24 hours. Do not
325 freeze Roferon-A. Keeping Roferon-A at temperatures outside the recommended range
326 can destroy the medicine.

Do not shake Roferon-A. Shaking can destroy Roferon-A so that it will not work.

Protect Roferon-A from light during storage.

Keep this and all other medicines out of the reach of children.

R_x only



Pharmaceuticals

Hoffmann-La Roche Inc.
340 Kingsland Street
Nutley, New Jersey 07110-1199

332 U.S. Govt. Lic. No.0136

333 27898099

334

335 Issued: March 2003

336 Printed in USA

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