

## II.D.2 Patient Understanding

### *Patient Comprehension Questions*

Program Steps	
One goal of the iPLEDGE Program is making sure pregnant women do not take isotretinoin	<ol style="list-style-type: none"> <li>1. True</li> <li>2. False</li> </ol>
You are in your third month of treatment, in order to get your isotretinoin prescription, you must	<ol style="list-style-type: none"> <li>1. Have a urine test for infection.</li> <li>2. Have a negative pregnancy test performed in a laboratory, discuss birth control with your doctor/prescriber and answer questions in the iPLEDGE Program system.</li> </ol>
The confidential iPLEDGE Program Pregnancy Registry collects information on pregnancies that happen during isotretinoin treatment or within 1 month after the last dose	<ol style="list-style-type: none"> <li>1. True</li> <li>2. False</li> </ol>
Each month you need to answer questions and:	<ol style="list-style-type: none"> <li>1. Enter 2 methods of birth control you are using.</li> <li>2. Do a home pregnancy test to show the pharmacist.</li> </ol>
For 1 month after your last dose, you must use 2 effective methods of birth control together correctly all the time	<ol style="list-style-type: none"> <li>1. True</li> <li>2. False</li> </ol>
Your doctor/prescriber tells you that you have to come back 1 month after your final dose. You return to the office to:	<ol style="list-style-type: none"> <li>1. Get a prescription for 30 days.</li> <li>2. Talk with your doctor/prescriber about birth control.</li> <li>3. Take your last pregnancy test.</li> </ol>
Your doctor/prescriber tells you that you need to have a pregnancy test each month. You would:	<ol style="list-style-type: none"> <li>1. Refuse the test because you know you are not pregnant.</li> <li>2. Agree to have a pregnancy test each month because it is important to know if you are pregnant because there is a very high chance of birth defects if an unborn baby's mother takes isotretinoin.</li> <li>3. Wait a week before getting the test done.</li> </ol>
You must keep your appointment every month because:	<ol style="list-style-type: none"> <li>1. It is important for you and your doctor/prescriber to interact before you get a maximum 30-day supply of isotretinoin each month.</li> <li>2. You need to sign a consent form each month.</li> <li>3. You need to tell your doctor where to call in your prescription.</li> </ol>

General Contraception Requirements	
Most women who got pregnant during isotretinoin treatment were using only 1 method of birth control.	1. True
	2. False
Alcohol and drugs can make it more difficult to use your birth control properly when having sex.	1. True
	2. False
You can change one primary method of birth control for another without talking with your doctor/prescriber first.	1. True
	2. False
You can use any 2 methods of birth control for the iPLEDGE Program	1. True
	2. False
You are using 2 effective methods of birth control in the month after your last dose. Your partner's male latex condom breaks. You would:	1. Forget about it since you have finished isotretinoin
	2. Call your doctor/prescriber to see if you might need emergency birth control.
	3. Get a home pregnancy test kit to see if you are pregnant.
While you are taking isotretinoin, you must use 2 effective methods of birth control together correctly all the time	1. True
	2. False
You should talk to your doctor/prescriber about birth control:	1. Each month during your office visit.
	2. Only when you have a problem.
	3. Only when you sign the second consent form.
You have finished your last dose of isotretinoin. Your doctor/prescriber has ordered a pregnancy test. For the next month, you:	1. Continue to use 2 effective methods of birth control together correctly all the time.
	2. Stop using your secondary method because you are not taking isotretinoin.
	3. Go for the pregnancy test at any time during the month.

Birth Defects and Pregnancy	
Which of the following are signs you might be pregnant?	1. You miss your menstrual period
	2. You have nausea (generally first thing in the morning), sometimes referred to as morning sickness.
	3. Your breasts feel tender, like at the beginning of a menstrual period.
	4. Any of the above is a sign you might be pregnant.
The risk for birth defects for a baby whose mother took isotretinoin is	1. So low that you do not have to worry.
	2. Low enough so you do not need birth control.
	3. Very high even if a female patient takes isotretinoin for only a short time.
You can possibly get pregnant if you have a uterus and ovaries.	1. True
	2. False
Which of the following birth defects may be caused by isotretinoin?	1. No ears
	2. Heart problems
	3. Small jaw and misshaped head
	4. A child could have any or all of these birth defects.
You think you may be pregnant and you have taken isotretinoin. You would:	1. Not worry because it was only a few doses.
	2. Stop isotretinoin and call your doctor/prescriber. Even the smallest amount of isotretinoin may cause birth defects.
It's okay to take isotretinoin when	1. You are using one effective method of birth control.
	2. You are using 2 effective methods of birth control together correctly all the time.
	3. You may be pregnant.
You cannot get pregnant if:	1. You never have sex with a man.
	2. You miss your hormone shots.
	3. You use only a male latex condom with spermicide.

Safety Information	
I must not give blood while taking isotretinoin and 1 month after stopping isotretinoin.	1. True 2. False
I will not share my isotretinoin with anyone.	1. Agree 2. Disagree
Your neighbor wants to try your isotretinoin. You tell her or him:	1. I must not share isotretinoin. 2. He or she must see a doctor/prescriber who can prescribe isotretinoin. 3. You tell her or him both of these reasons.
The Red Cross calls you to give blood right after your last dose. You tell them:	1. I can never give blood again. 2. I cannot give blood until 1 month after my last dose of isotretinoin.
The Red Cross calls you to give blood during your first month after your last dose. You tell them:	1. I cannot give blood until 1 month after my last dose of isotretinoin. 2. I can never give blood again.

Obtaining a Prescription	
Your pharmacist can fill and dispense your prescription only after:	1. You phone it in. 2. Checking with the iPLEDGE Program system to see if you can get isotretinoin. 3. You email it in.
You can get a prescription for isotretinoin only if:	1. Your pregnancy test is negative. 2. You answered the questions correctly. 3. You entered your 2 methods of birth control and they match the birth control methods entered by your doctor/prescriber. 4. Your doctor/prescriber entered your 2 methods of birth control. 5. All of these.
You had a pregnancy test on Tuesday for your next prescription. What is the last day you can obtain your prescription?	1. Later that same day (7 hours later) 2. By Friday before the pharmacy closes 3. On Monday of the next week
You can obtain your prescription any time you want	1. True 2. False
Your doctor/prescriber has entered your negative pregnancy test result and your two methods of birth control into the iPLEDGE Program system and gives you a prescription. Before you can obtain isotretinoin at a pharmacy you must:	1. Answer your comprehension questions in the iPLEDGE Program system. 2. Schedule your next appointment with your doctor/prescriber 3. Sign your informed consent each month.

Contraception Choices	
You missed 2 days taking your birth control pills earlier this week. You would:	<ol style="list-style-type: none"> <li>1. You should call your doctor/prescriber as soon as you realize it.</li> <li>2. Have sex whenever you want.</li> </ol>
I have been using birth control pills for 5 years and have not gotten pregnant. Why do I need another method of birth control now?	<ol style="list-style-type: none"> <li>1. Any form of birth control can fail</li> <li>2. The birth defects are too severe to risk getting pregnant</li> <li>3. Isotretinoin may change how well some methods of birth control work</li> <li>4. All of these</li> </ol>
Birth control pills work best when you take them about the same time every day and do not miss any pills.	<ol style="list-style-type: none"> <li>1. True</li> <li>2. False</li> </ol>
You can switch to the progesterone-only mini-pill for your primary method of birth control while you are taking isotretinoin.	<ol style="list-style-type: none"> <li>1. True</li> <li>2. False</li> </ol>
Birth control pills may not work as well when you are taking isotretinoin. It is important to:	<ol style="list-style-type: none"> <li>1. Take two birth control pills every day.</li> <li>2. Remember to take your birth control pill about the same time every day and always use a second effective method correctly to help prevent pregnancy.</li> <li>3. Take your birth control pill only before having sex</li> </ol>
I'm using my primary method of birth control exactly as prescribed and my partner uses male latex condoms as the second method. My partner does not have any male latex condoms with him tonight. We would:	<ol style="list-style-type: none"> <li>1. Have sex anyway.</li> <li>2. Wait until he can get a male latex condom.</li> <li>3. Have sex but use withdrawal.</li> </ol>
I'm using my primary method of birth control exactly as prescribed and my partner uses male latex condoms with spermicide as the second method. My partner's male latex condom breaks. I would:	<ol style="list-style-type: none"> <li>1. Not worry about getting pregnant.</li> <li>2. Worry that I may become pregnant but not do anything.</li> <li>3. Call my doctor/prescriber to talk about emergency birth control.</li> </ol>
My partner has only one male latex condom with him. We would:	<ol style="list-style-type: none"> <li>1. Have sex more than once and reuse the male latex condom.</li> <li>2. Have sex once and wait until he can get more male latex condoms.</li> <li>3. Have sex but use withdrawal.</li> </ol>
For the iPLEDGE Program, male latex condoms can be used with or without a spermicide.	<ol style="list-style-type: none"> <li>1. True</li> <li>2. False</li> </ol>
If you use lubricant with a male latex condom, it should be a water-based lubricant.	<ol style="list-style-type: none"> <li>1. True</li> <li>2. False</li> </ol>
Your partner should put his male latex condom on:	<ol style="list-style-type: none"> <li>1. Any time during sex.</li> <li>2. As soon as he gets an erection, because waiting too long lets sperm leak out.</li> <li>3. Only when he remembers.</li> </ol>

You find a rip/tear in your diaphragm after you had sex. You should:	1. Forget about it since you were near your period. 2. Call your doctor/prescriber to see if you might need emergency birth control.
For the iPLEDGE Program, diaphragms can be used with or without a spermicide.	1. True 2. False
After sex, your diaphragm should stay in place for:	1. 24 hours or more. 2. 2 to 4 hours 3. At least 6 hours and up to 24 hours.
You and your partner want to have sex a second time after you put in your diaphragm. You must:	1. Put spermicide in your vagina again before you have sex. 2. Have sex again without using more spermicide for up to 24 hours at a time. 3. Take out your diaphragm and clean it before having sex again.
Your gynecologist or family doctor needs to check how your diaphragm fits:	1. Every time you have sex. 2. Every 2 years, or if you gain or lose 10 pounds, or after childbirth or an abortion. 3. Every 2 months.
The diaphragm with a spermicide can help prevent:	1. Pregnancy 2. The spread of HIV/AIDS 3. The spread of other sexually transmitted infections (STIs)
I'm using my primary method of birth control exactly as prescribed and a cervical cap with a spermicide as my second method. I forgot my cap at home. I would:	1. Have sex anyway. 2. Wait until I can use the cervical cap with a spermicide. 3. Have sex but use withdrawal.
For the iPLEDGE Program cervical caps can be used with or without a spermicide.	1. True 2. False
You cannot use the cervical cap if there is any vaginal bleeding, such as during your menstrual period.	1. True 2. False
You and your partner want to have sex a second time after you put in your cervical cap. You do not need to put in more spermicide.	1. True 2. False
After sex, your cervical cap should stay in place for:	1. 24 hours. 2. 4 to 8 hours. 3. At least 6 hours and up to 48 hours.

The cervical cap with spermicide can help prevent:	1. Pregnancy.
	2. The spread of HIV/AIDS
	3. The spread of other sexually transmitted infections (STIs)
I'm using my primary method of birth control exactly as prescribed and use a vaginal sponge as my second method. I ran out of sponges tonight. My partner and I would:	1. Wait until I can buy more sponges
	2. Have sex anyway.
	3. Have sex but use withdrawal.
I'm using my primary method of birth control exactly as prescribed, but I forgot to put in my vaginal sponge this one time. I would:	1. Not worry that I could have gotten pregnant.
	2. Worry that I may become pregnant, but do nothing.
	3. Call my doctor/prescriber to talk about emergency birth control.
The vaginal sponge can help prevent:	1. The spread of HIV/AIDS
	2. The spread of other sexually transmitted infections (STIs)
	3. Pregnancy
You should insert your sponge:	1. Any time during sex.
	2. Any time up to 30 hours before you have sex.
	3. Only when you remember.
After sex, your sponge should stay in place for:	1. 24 hours or more.
	2. 2 to 4 hours.
	3. At least 6 hours.
If you and your partner want to have sex a second time after you put in your sponge:	1. You must put spermicide in your vagina again before you have sex.
	2. You can have sex again without using a new sponge for up to 30 hours at a time.
	3. You must take out your sponge and out in a new one before having sex.
I have been using the hormonal skin patch perfectly for over a year and have not gotten pregnant. Why do I need another method of birth control now?	1. Any method of birth control can fail.
	2. Using two methods of birth control together all the time drastically reduces the chance that you will get pregnant.
	3. Most female patients who got pregnant during isotretinoin treatment were using only 1 method of birth control.
	4. All of these.
I have been using the hormonal skin patch, but I was 2 days late changing it. If I have sex my chances of getting pregnant are:	1. Increased
	2. Decreased

I am using the hormonal skin patch exactly as my doctor/prescriber tells me. I should not put it on:	1. My upper outer arm.
	2. My stomach or upper body.
	3. My breasts.
The hormonal skin patch works best when you change it every week on the same day each week for 3 weeks.	1. True
	2. False
You can get pregnant using the hormonal patch if:	1. It becomes loose or falls off for more than 24 hours.
	2. You leave the same hormonal patch on more than one week.
	3. Both 1 and 2.
My hormonal vaginal ring slipped out and I did not replace it within 3 hours, I:	1. Could get pregnant.
	2. Could have sex when I want to, my partner's male latex condom with spermicide is enough.
I have been using a hormonal vaginal ring for 1 year and have not gotten pregnant. Why do I need another method of birth control now?	1. Any method of birth control can fail.
	2. Using two methods of birth control all the time drastically reduces the chance that you will get pregnant.
	3. Most female patients who got pregnant during isotretinoin treatment were using only 1 method of birth control.
	4. All of these.
I am using the hormonal vaginal ring as my primary method. I can use it with which of the following barrier methods?	1. Cervical cap with spermicide.
	2. Male latex condom with or without spermicide.
	3. Diaphragm with spermicide.
The hormonal ring works best when you leave it in place for 3 weeks and insert a new ring every 4 weeks on the same day of the week.	1. True
	2. False
Pregnancy can happen when you are using a hormonal vaginal ring if:	1. The unopened package containing the hormonal vaginal ring is put into direct sunlight or gets hot.
	2. It slips out of your vagina and you do not replace it within 3 hours.
	3. It does not stay in your vagina for 3 weeks.
	4. You leave it in your vagina for more than 3 weeks.
	5. All of these reasons.

I have been getting hormone shots for 1 year and have not gotten pregnant. Why do I need another method of birth control now?	1. Any method of birth control can fail.
	2. Using two methods of birth control all the time drastically reduces the chance that you will get pregnant.
	3. Most female patients who got pregnant during isotretinoin treatment were using only 1 method of birth control.
	4. All of these.
You get hormone shots:	1. Every 12 weeks
	2. Your dermatologist, gynecologist, family doctor or pharmacist (in some states) can give you a shot.
	3. Both are true.
You are using hormonal shots as your primary method of birth control, you should remember:	1. You are not protected against HIV/AIDS
	2. You need to see your dermatologist, gynecologist, family doctor or pharmacist (in some states) to get the hormonal shot
	3. They might delay your ability to get pregnant after you stop using hormonal shots.
	4. All of the above.
One advantage of hormone shots is that they also protect against sexually transmitted infections (STIs).	1. True
	2. False
You have chosen hormone shots as your primary method of birth control. Another acceptable method for you to use would be:	1. Female condoms
	2. Hormonal IUD
	3. Withdrawal
	4. Male latex condoms
You have received a prescription for isotretinoin from your doctor/prescriber but have not had the prescription filled. It is OK to have unprotected sex since you have had a negative pregnancy test.	1. True
	2. False
I have been using an IUD for 3 years and have not gotten pregnant. Why do I need another method of birth control now?	1. Any method of birth control can fail.
	2. Using two methods of birth control all the time drastically reduces the chance that you will get pregnant.
	3. Most female patients who got pregnant during isotretinoin treatment were using only 1 method of birth control.
	4. All of these.
Your IUD needs to be checked by your doctor/prescriber:	1. Within 3 months after you had it inserted.
	2. If your weight stays the same.

You have chosen an IUD as your primary method of birth control. Another acceptable method for you to use would be:	1. Female condoms
	2. Hormonal IUD
	3. Withdrawal
	4. Male Latex Condoms
I had tubal sterilization (blocking my tubes) 5 years ago and have not gotten pregnant. Why do I need another method of birth control now?	1. Any form of birth control can fail, including tubal sterilization (blocking my tubes).
	2. Using two methods of birth control all the time drastically reduces the chance that you will get pregnant.
	3. Most female patients who got pregnant during isotretinoin treatment were using only 1 method of birth control.
	4. All of these.
My primary method of birth control is tubal sterilization (blocking my tubes). I also need to use a secondary method or another primary method of birth control while I am taking isotretinoin:	1. True
	2. False
Important information to know about tubal sterilization (blocking your tubes) is:	1. Tubal sterilization (blocking your tubes) does not protect against sexually transmitted infections (STIs).
	2. Tubal sterilization (blocking your tubes) does not require surgery.
	3. It is easy to re-open the tubes.
Tubal sterilization (blocking your tubes) is a highly effective method of birth control and does not require use of another effective method of birth control while taking isotretinoin:	1. True
	2. False
You have chosen tubal sterilization (blocking your tubes) as your primary method of birth control. Another acceptable method for you to use would be:	1. Female condoms
	2. Hormonal IUD
	3. Withdrawal
	4. None of the above
My only partner had a male vasectomy 5 years ago and I have not gotten pregnant. Why do I need another method of birth control now?	1. Using two methods of birth control all the time drastically reduces the chance that you will get pregnant.
	2. Any method of birth control can fail.
	3. Most female patients who got pregnant during isotretinoin treatment were using only 1 method of birth control.
	4. All of these.
My primary method of birth control is my only partner's male vasectomy. I also need to use a secondary method or another primary method of birth control while I am taking isotretinoin.	1. True
	2. False

Important information to know about your partner's male vasectomy is:	1. Your partner's male vasectomy does not protect against sexually transmitted infections (STIs).
	2. Your partner's male vasectomy does not require surgery.
	3. It is easy to open the tubes again if your partner wants a child later.
Your partner's male vasectomy is a highly effective method of birth control and does not require use of another effective method of birth control while taking isotretinoin:	1. True
	2. False
You have chosen your partner's male vasectomy as your primary method of birth control. Another acceptable method for you to use would be:	1. Female condoms
	2. IUD
	3. Withdrawal
	4. None of the above
I have had implanted hormones for 2 years and have not gotten pregnant. Why do I need another method of birth control now?	1. Using two methods of birth control all the time drastically reduces the chance that you will get pregnant.
	2. Any method of birth control can fail.
	3. Most female patients who got pregnant during isotretinoin treatment were using only 1 method of birth control.
	4. All of these.
My primary method of birth control is implanted hormones. I also need to use a secondary method or another primary method of birth control while I am taking isotretinoin.	1. True
	2. False
My implantable hormone will only be effective through the first month while I am taking isotretinoin, after that I only need to use one effective method of birth control for the rest of my treatment.	1. True
	2. False
You have chosen implantable hormones as your primary method of birth control. Another acceptable method for you to use would be:	1. Female condoms
	2. Hormonal IUD
	3. Withdrawal
	4. None of the above
Abstinence means that you commit to not having sex or sexual contact with any male 24 hours a day, 7 days a week for 1 month before, during, and for 1 month after your isotretinoin treatment.	1. True
	2. False
One of the most common causes of unplanned pregnancy is not being able to avoid sex (failing to maintain abstinence).	1. True
	2. False

<p>You planned not to have sex while taking isotretinoin, so you do not use any birth control. If you decide to have sex, you need to:</p>	1. Tell the doctor/prescriber who prescribes your isotretinoin before you engage in sexual activity and make a plan to start your birth control and be sure you are not pregnant. .
	2. Stop having sex until you use your new birth control method for at least 1 month and have a negative pregnancy test.
	3. Start using three methods of effective birth control, just to be sure.
	4. 1 and 2 are correct.
<p>You must not take isotretinoin if you cannot follow the birth control requirements of the iPLEDGE Program.</p>	1. True
	2. False
<p>Important information to know about birth control pills is:</p>	1. You cannot get pregnant while using birth control pills even if you skip pills.
	2. Birth control pills do not protect against sexually transmitted infections (STIs) or HIV/AIDS
	3. There are no side effects from using birth control pills.
<p>The male latex condom can help prevent:</p>	1. Pregnancy.
	2. The spread of HIV/AIDS
	3. The spread of other sexually transmitted infections (STIs)
	4. All of the above.
<p>For 1 month after your last dose, you must use 2 effective methods of birth control together all the time.</p>	1. True
	2. False



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Committed to Pregnancy Prevention

## Wholesaler Registration Letter

Dear wholesaler or chain drug executive,

The iPLEDGE Program requires annual registration of all wholesalers distributing isotretinoin. For the purposes of the iPLEDGE Program, the term wholesaler refers to a wholesaler and each of its individual distribution centers, a distributor and each of its individual distribution centers, and/or each warehousing chain pharmacy distribution center.

A Registration Form is enclosed with this letter.

### **Key Elements of the iPLEDGE Program, Which Relate to Your Business Practices:**

- The program covers all isotretinoin products (brand and generic)
- To be registered, each wholesaler must agree to meet all iPLEDGE Program requirements for wholesale distribution of isotretinoin products by completing the enclosed agreement
- Wholesalers registered in the iPLEDGE Program will only ship isotretinoin to:
  - Wholesalers registered in the iPLEDGE Program with prior written consent from the manufacturer, or
  - Pharmacies registered with and activated in the iPLEDGE Program
- The registration of wholesalers that do not abide by the terms of the agreement will be revoked after an investigation process, and manufacturers of FDA-approved isotretinoin products will not continue to provide them with isotretinoin for distribution
- Registration in the iPLEDGE Program expires in 12 months and requires re-registration annually.

### **To Register in the iPLEDGE Program, Mail the Completed Agreement to:**

iPLEDGE Program  
P.O. Box 29094  
Phoenix, AZ 85038

OR

Fax the completed form, using the following iPLEDGE fax number: 1-866-495-0660

We thank you for your support in complying with the iPLEDGE Program.

Sincerely,

iPLEDGE Program Sponsors



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## Wholesaler Registration (continued)

### SAFETY NOTICE

Isotretinoin must not be used by female patients who are or may become pregnant. There is an extremely high risk that severe birth defects will result if pregnancy occurs while taking isotretinoin in any amount, even for a short period of time. Potentially any fetus exposed during pregnancy can be affected. There are no accurate means of determining whether an exposed fetus has been affected. Because of this toxicity, isotretinoin can only be marketed under a special restricted distribution program. This program is called iPLEDGE<sup>®</sup>. Under this program, prescribers must be registered and activated with the iPLEDGE Program and can prescribe isotretinoin only to registered patients who meet all the requirements of iPLEDGE. Isotretinoin can be dispensed only by a pharmacy registered and activated with iPLEDGE. Registered and activated pharmacies can only receive isotretinoin from wholesalers registered with iPLEDGE.

Patients on isotretinoin have been known to become depressed or to develop other serious mental health problems. Some people have had thoughts of hurting themselves or putting an end to their own lives. Some people tried to end their own lives and some have ended their own lives. There have been reports that people on isotretinoin were aggressive or violent. No one knows if isotretinoin caused these problems or behaviors, or if they would have happened even if the person did not take isotretinoin.

Isotretinoin use has been associated with pseudotumor cerebri, a condition caused by increased pressure on the brain. This condition may occur more often in patients also taking tetracycline. Patients should be aware of other serious side effects, including problems with the skin, pancreas, liver, stomach, bones, muscles, hearing, vision, lipids, allergic reactions, blood sugar, or red and white blood cells. The most common, less serious adverse events include dry skin, chapped lips, dry eyes, and dry nose that may lead to nosebleeds. Patients should be advised about these adverse events and routinely monitored by a doctor during treatment with isotretinoin.



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iPLEDGE Program  
P.O. Box 29094  
Phoenix, AZ 85038

**Wholesaler Registration Form On Behalf of the Wholesaler Listed Below, I Acknowledge That:**

For the purpose of the iPLEDGE Program, the term wholesaler refers to a wholesaler and each of its individual distribution centers, a distributor and each of its individual distribution centers, and/or each warehousing chain pharmacy distribution center. To distribute isotretinoin, wholesalers must be registered with the iPLEDGE Program and agree to meet all iPLEDGE Program requirements for wholesaler distribution of isotretinoin products. Wholesalers must register with the iPLEDGE Program by signing and returning this agreement that affirms they will comply with all iPLEDGE Program requirements for distribution of isotretinoin. The registration of wholesalers that do not abide by the terms of the agreement will be revoked after an investigation process, and manufacturers of FDA approved isotretinoin products will not continue to provide them with isotretinoin for distribution. Each distribution center operated by a wholesaler must register if they are to distribute isotretinoin. These requirements include:

- Registering prior to distributing isotretinoin and registering annually thereafter
- Distributing only FDA approved isotretinoin product obtained directly from the isotretinoin manufacturers (or delegate) or another registered wholesaler
- Beginning November 1, 2005 only ship isotretinoin to:
  - Wholesalers registered in the iPLEDGE Program with prior written consent from the manufacturer, or
  - Pharmacies licensed in the US and registered and activated in the iPLEDGE Program
- Notifying immediately the isotretinoin manufacturer (or delegate) of any nonregistered and/or nonactivated pharmacy or unregistered wholesaler that attempts to order isotretinoin
- Complying with inspection of wholesaler records by the isotretinoin manufacturer (or delegate) for verification of compliance with the iPLEDGE Program
- Returning to the manufacturer (or delegate) any undistributed product if registration is Permanently Deactivated by the iPLEDGE Program or if the wholesaler chooses to not reregister annually.

An agreement must be completed for each distributor center.

I am authorized to execute this agreement on behalf of the wholesaler (and its distribution centers, if applicable)



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Wholesaler Name \_\_\_\_\_  
(Type or Print)

Distribution Center's DEA# \_\_\_\_\_

Distribution Center's Address \_\_\_\_\_

Distribution Center's City \_\_\_\_\_

Distribution Center's State \_\_\_\_\_ Distribution Center's ZIP \_\_\_\_\_

Distribution Center's Phone Number \_\_\_\_\_

Distribution Center's Fax Number \_\_\_\_\_

Authorized Representative \_\_\_\_\_  
(Print First, MI, Last)

Title \_\_\_\_\_

E-mail Address for Key Contact \_\_\_\_\_

E-mail Address for Pharmacy Eligibility File Delivery\* \_\_\_\_\_

Authorized Representative's Signature \_\_\_\_\_

Date \_\_\_\_/\_\_\_\_/\_\_\_\_  
Month Day Year

\*The list of registered and activated pharmacies will be e-mailed to this address daily.

***This agreement expires 12 months from agreement date. Annual registration is required.***

[www.ipledgeprogram.com](http://www.ipledgeprogram.com) | 1-866-495-0654



**iPLEDGE**<sup>®</sup>  
Committed to Pregnancy Prevention

## Wholesaler Product Return Letter

### Isotretinoin Return Customer Instruction Guide - Wholesaler

Wholesaler Name  
Address 1  
Address 2  
City, State Postal Code

Dear Wholesaler:

You are no longer registered in the iPLEDGE Program for the following reason:

- Your facility is closing, and will no longer be a wholesaler in the iPLEDGE Program.

Therefore, you are required to return all isotretinoin inventories in stock **effective immediately**.

This action is pursuant to the FDA approval of the iPLEDGE Program through the special restricted distribution program. If you have questions about your status in the iPLEDGE Program, please call 1-866-495-0654 or visit [www.ipledgeprogram.com](http://www.ipledgeprogram.com) for more information.

#### **Action required by a wholesaler no longer registered in the iPLEDGE Program:**

- Immediately remove all isotretinoin from your stock.
- If you have isotretinoin inventory, please call **1-866-495-0654** for further instructions on returning this product to the manufacturer.

Sincerely,

iPLEDGE Program Sponsors

[www.ipledgeprogram.com](http://www.ipledgeprogram.com) | 1-866-495-0654



## Wholesaler Product Return Letter (continued)

### SAFETY NOTICE

Isotretinoin must not be used by female patients who are or may become pregnant. There is an extremely high risk that severe birth defects will result if pregnancy occurs while taking isotretinoin in any amount, even for a short period of time. Potentially any fetus exposed during pregnancy can be affected. There are no accurate means of determining whether an exposed fetus has been affected. Because of this toxicity, isotretinoin can only be marketed under a special restricted distribution program. This program is called iPLEDGE®. Under this program, prescribers must be registered and activated with the iPLEDGE Program and can prescribe isotretinoin only to registered patients who meet all the requirements of iPLEDGE. Isotretinoin can be dispensed only by a pharmacy registered and activated with iPLEDGE. Registered and activated pharmacies can only receive isotretinoin from wholesalers registered with iPLEDGE.

Patients on isotretinoin have been known to become depressed or to develop other serious mental health problems. Some people have had thoughts of hurting themselves or putting an end to their own lives. Some people tried to end their own lives and some have ended their own lives. There have been reports that people on isotretinoin were aggressive or violent. No one knows if isotretinoin caused these problems or behaviors or if they would have happened even if the person did not take isotretinoin.

Isotretinoin use has been associated with pseudotumor cerebri, a condition caused by increased pressure on the brain. This condition may occur more often in patients also taking tetracycline. Patients should be aware of other serious side effects, including problems with the skin, pancreas, liver, stomach, bones, muscles, hearing, vision, lipids, allergic reactions, blood sugar, or red and white blood cells. The most common, less serious adverse events include dry skin, chapped lips, dry eyes, and dry nose that may lead to nosebleeds. Patients should be advised about these adverse events and routinely monitored by a doctor during treatment with isotretinoin.

iPLEDGE, an enhanced pregnancy risk management program designed to minimize fetal exposure to isotretinoin, has been approved by the FDA through a special restricted distribution program.

[www.ipledgeprogram.com](http://www.ipledgeprogram.com) | 1-866-495-0654

# [Name of Manufacturer]

## {Product Name and Generic Designation} Wholesaler to Wholesaler Shipment Request Form

### REQUESTING PARTY INFORMATION: (Please print or type)

Requester Name: \_\_\_\_\_ Telephone: \_\_\_\_\_

E-Mail Address: \_\_\_\_\_ Date of Request: \_\_\_\_\_

Wholesaler Name: \_\_\_\_\_ DEA: \_\_\_\_\_

Address: \_\_\_\_\_ City: \_\_\_\_\_ State: \_\_\_\_\_ Zip: \_\_\_\_\_

*Requesting Party represents and warrants that the Receiving Party listed below is registered with the iPLEDGE Program.*

Requester Signature: \_\_\_\_\_ Date: \_\_\_\_\_

### RECEIVING PARTY INFORMATION

Wholesaler Name: \_\_\_\_\_ DEA: \_\_\_\_\_

Address: \_\_\_\_\_ City: \_\_\_\_\_ State: \_\_\_\_\_ Zip: \_\_\_\_\_

Ship to Address: \_\_\_\_\_ City: \_\_\_\_\_ State: \_\_\_\_\_ Zip: \_\_\_\_\_

### MANUFACTURER'S CONSENT

Shipments of [Product Name and Generic Designation] between wholesalers\* must be in compliance with the iPLEDGE Program. Labeling for [Product Name] states that wholesalers registered in the iPLEDGE Program may only ship to other registered wholesalers with prior written consent from the manufacturer. Pursuant to the labeling for [Product Name], the authorized signature below shall serve as written consent from the manufacturer provided that the requesting and receiving party's registration is verified prior to each shipment. Registration will be verified by [Manufacturer Name] upon receipt of shipment request.

*\*The term wholesaler refers to a wholesaler and each of its individual distribution centers, a distributor and each of its individual distribution centers, and/or each warehousing chain pharmacy distribution center.*

NOTE: This Wholesaler to Wholesaler Shipment Request Form is not required when shipping isotretinoin between distribution centers that are under the same parent company.

[Name of Manufacturer]

By: \_\_\_\_\_ Title: \_\_\_\_\_

Print Name: \_\_\_\_\_ Date: \_\_\_\_\_

**Please return completed forms to [Insert Appropriate Information Including Fax Number, Email URL, Telephone Number, etc.]**

**Please note that this approval is for a one time shipment. Further shipments require consent from [Manufacturer].**