

FRUSTRATED BY PAINFUL KNEES?

Give your knees
the cushioning they need.



*SYNVISC® (hylan G-F 20) lubricates and cushions your knee, and can provide up to six months of osteoarthritis knee pain relief.**

* As shown in a medical study comparing Synvisc to an injection of salt water.

SYNVISC is used to relieve knee pain due to osteoarthritis (OA). It is for patients who do not get enough relief from simple painkillers such as acetaminophen, or from exercise and physical therapy.

Important Safety Information

Before trying SYNVISC, tell your doctor if you have had an allergic reaction, such as swelling of the face, tongue or throat, respiratory difficulty, rash, itching or hives to SYNVISC or any hyaluronan-based products.

[Please see Important Safety Information on page 10.](#)
[Please see Important Patient Information on page 11.](#)

SYNVISC
HYLAN G-F 20

YOU DON'T HAVE TO SETTLE FOR OSTEOARTHRITIS KNEE PAIN

It's time to find the right treatment option for you

Talking to your doctor is the best way to start exploring your treatment options. And when you do, keep in mind:

Osteoarthritis knee pain affects everyone differently

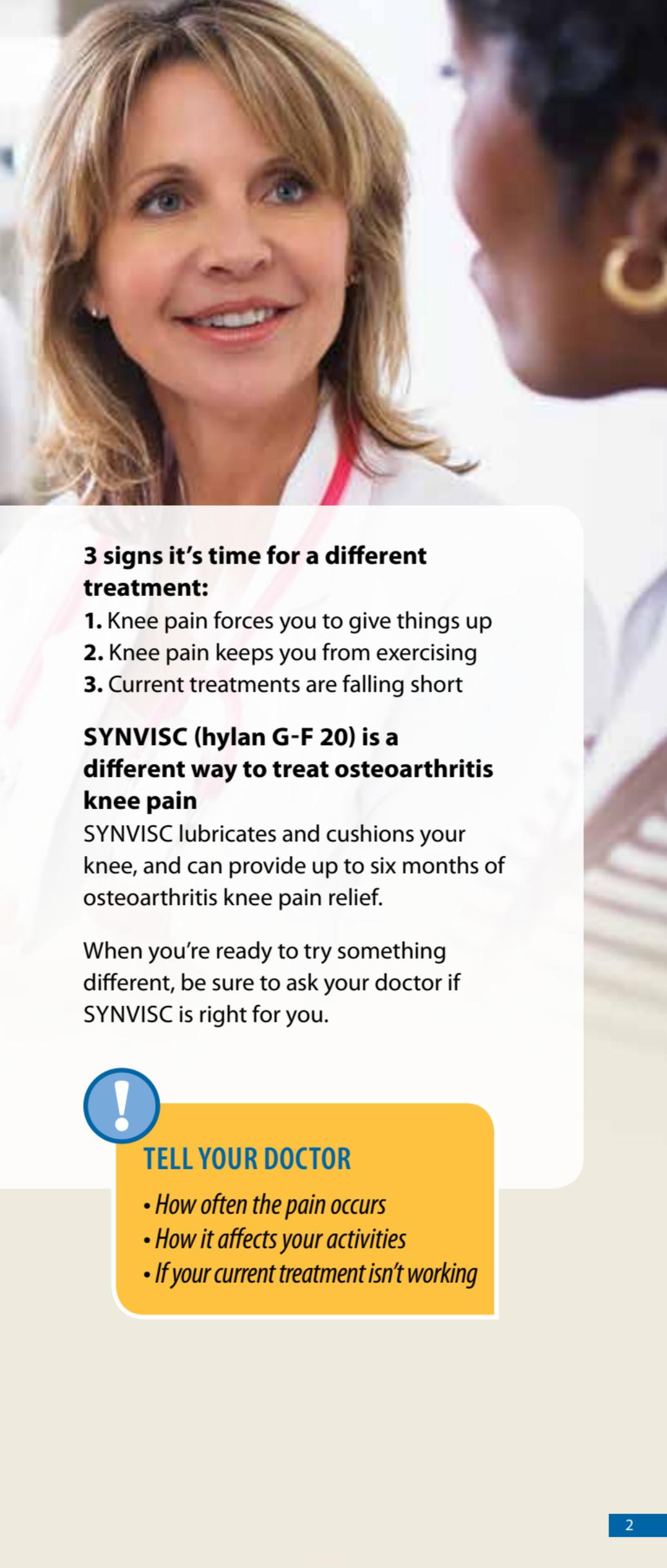
An X-ray can show your grade or stage of osteoarthritis, but your level of pain isn't necessarily connected to it. Your osteoarthritis could be mild with a great deal of pain or severe with little pain. Your doctor is relying on you to give a clear description of how your knee feels.

It's caused by more than just cartilage loss

In knees with osteoarthritis, the cartilage protecting the ends of the bones gradually deteriorates, joint fluid loses its shock-absorbing qualities, and bones may begin to rub against each other—all of which may cause the osteoarthritis knee pain you're feeling.

Please see Important Safety Information on page 10.

Please see Important Patient Information on page 11.



3 signs it's time for a different treatment:

1. Knee pain forces you to give things up
2. Knee pain keeps you from exercising
3. Current treatments are falling short

SYNVISC (hylan G-F 20) is a different way to treat osteoarthritis knee pain

SYNVISC lubricates and cushions your knee, and can provide up to six months of osteoarthritis knee pain relief.

When you're ready to try something different, be sure to ask your doctor if SYNVISC is right for you.



TELL YOUR DOCTOR

- *How often the pain occurs*
- *How it affects your activities*
- *If your current treatment isn't working*

ASK ABOUT SYNVISC® *(hylan G-F 20)*

A natural way to
treat osteoarthritis knee pain

What is SYNVISIC?

SYNVISC is a series of three injections, given one week apart, that supplement the fluid in your knee to help lubricate and cushion the joint, and can provide up to six months of osteoarthritis knee pain relief.

What is SYNVISIC made from?

The injection contains a gel-like mixture made from a substance called hyaluronan (pronounced hy-al-u-ROE-nan) that comes from chicken combs.

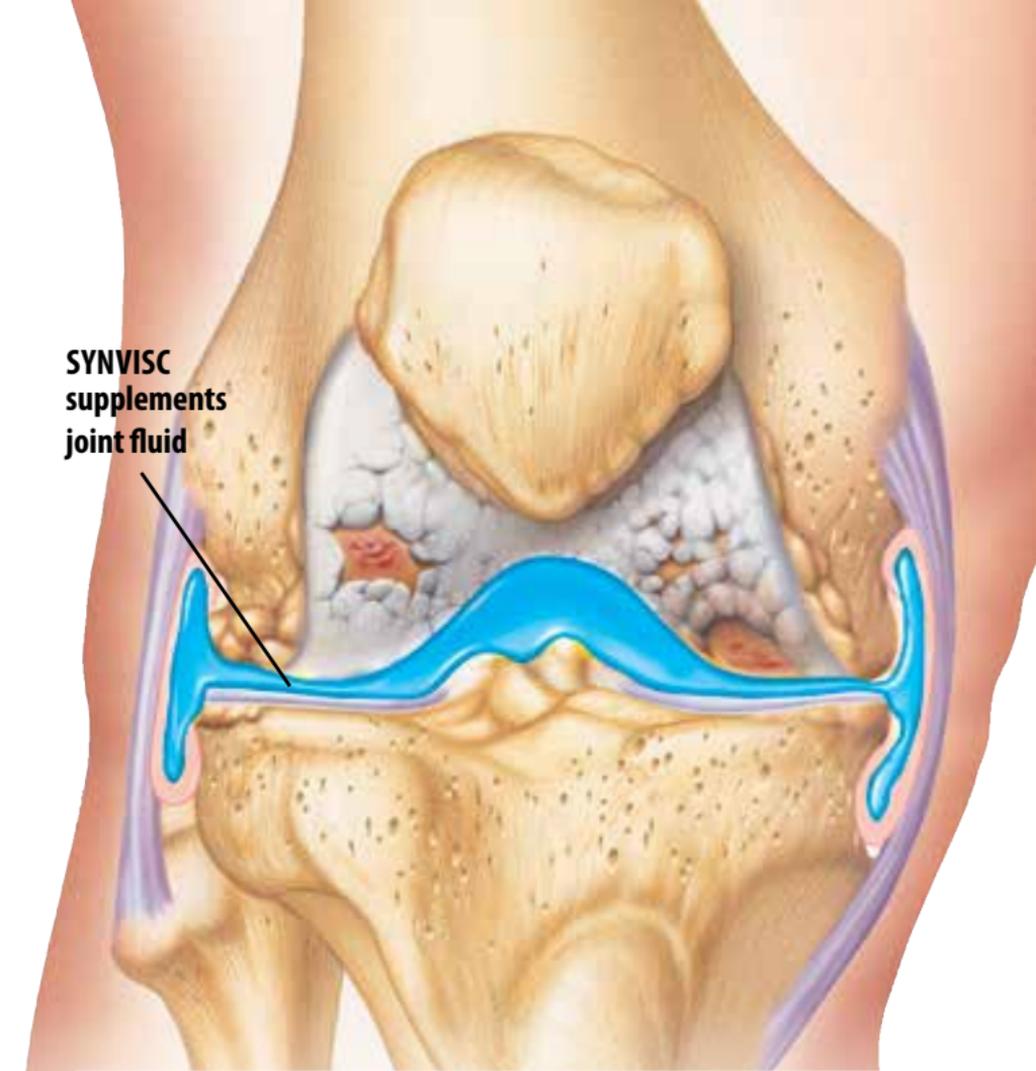
Hyaluronan is a natural substance found in the body and is present in very high amounts in joints. The body's own hyaluronan acts like a lubricant and a shock absorber in the joint and is needed for the joint to work properly.

When should I consider SYNVISIC?

SYNVISC is for people with knee osteoarthritis who have not received enough pain relief from diet, exercise and over-the-counter pain medication. If you've tried these options and are still feeling osteoarthritis knee pain, tell your doctor and ask if SYNVISIC could help.

Please see Important Safety Information on page 10.

Please see Important Patient Information on page 11.



**SYNVISC
supplements
joint fluid**

SYNVISC supplements your joint fluid to relieve pain and improve lubrication and cushioning in your knee.

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What are the side effects?

Side effects from SYNVISC are infrequent. In clinical trials, about 7% of patients experienced pain, swelling and/or fluid buildup around the knee.



ASK YOUR DOCTOR ABOUT A NATURAL SOLUTION

SYNVISC is a product made from a natural substance that lubricates and cushions your knee joint.

PREPARE FOR YOUR APPOINTMENTS

Be informed about SYNVISIC®
(hylan G-F 20) injections

How is SYNVISIC given?

SYNVISIC is injected directly into your knee over the course of three doctor visits, one week apart.

What is the procedure like?

Each injection takes a few minutes and can be done in your doctor's office.

Will the injection hurt?

Your doctor will likely apply a numbing agent before your SYNVISIC injections. You may feel some pressure, but it shouldn't be painful.

A white lab coat with buttons and a teal surgical drape.

Is it covered by insurance?

SYNVISC is covered by Medicare and most insurance plans.

What happens after the injections?

You can resume normal day-to-day activities immediately after the injections, but you should avoid any strenuous activities for about 48 hours.

KNOW WHAT TO EXPECT

What happens after your SYNVISIC® (hylan G-F 20) injections

In the first 48 hours after your SYNVISIC injection:

- You should be able to resume your normal day-to-day activities
- You should avoid activities that put excessive strain on your knee such as jogging, lifting or prolonged standing
- If you have any mild pain or swelling at the injection site, place an ice pack on your knee for 10 minutes or as recommended by your doctor

Please see Important Safety Information on page 10.
Please see Important Patient Information on page 11.



In the coming months:

- Some patients may begin to feel osteoarthritis knee pain relief after the first injection, but you need to have all three injections to get the maximum benefit
- Although everyone responds differently, most patients feel the greatest relief 8 to 12 weeks after the first injection
- SYNVISC can provide up to six months of knee pain relief

JOIN the ONTRACK PROGRAM

Get timely treatment resources

Sign up to receive the latest information on knee exercises, nutrition and activities.

Join today for your FREE Knee Exercise Program Guide

With step-by-step instructions for exercises designed specifically for patients with knee osteoarthritis.



JOIN **ONTRACK**

Two easy ways to enroll

- 1 Call 1-866-480-8912 or
- 2 Visit SynviscOne.com/ontrack

Please see Important Safety Information on page 10.
Please see Important Patient Information on page 11.

Indication

SYNVISC® (hylan G-F 20) is used to relieve knee pain due to osteoarthritis (OA). It is for patients who do not get enough relief from simple painkillers such as acetaminophen, or from exercise and physical therapy.

Important Safety Information

Before trying SYNVISIC, tell your doctor if you have had an allergic reaction, such as swelling of the face, tongue or throat, respiratory difficulty, rash, itching or hives to SYNVISIC or any hyaluronan-based products. Should not be used in patients with an infected knee joint, skin disease or infection around the area where the injection will be given, or circulatory problems in the legs.

SYNVISC is only for injection into the knee, performed by a doctor or other qualified health care professional. SYNVISIC has not been tested to show pain relief in joints other than the knee. Tell your doctor if you are allergic to products from birds—such as feathers, eggs or poultry—or if your leg is swollen or infected.

SYNVISC has not been tested in children (≤ 21 years old), pregnant women or women who are nursing. You should tell your doctor if you think you are pregnant or if you are nursing a child.

Talk to your doctor before resuming strenuous weight-bearing activities after treatment.

The most commonly occurring side effects reported after SYNVISIC include pain, swelling, heat, redness, and/or fluid build-up in or around the knee. Tell your doctor if you experience any side effects after treatment with SYNVISIC.



IMPORTANT PATIENT INFORMATION

SYNVISC® (HYLAN G-F 20)

Be sure to read the following important information carefully. This information does not take the place of your doctor's advice. If you do not understand this information or want to know more, ask your doctor.

Glossary of Terms

Hyaluronan (pronounced hy-al-u-ROE-nan): is a natural substance that is present in very high amounts in joints. It acts like a lubricant and a shock absorber in the joint and is needed for the joint to work properly.

Non-steroidal anti-inflammatory drugs: also known as "NSAIDs"; medication used to treat pain or swelling. There are many examples of NSAIDs, including (but not limited to) aspirin and ibuprofen. Some of these are over-the-counter drugs, and some can only be obtained by prescription.

Osteoarthritis (pronounced OS-te-o-arth-RI-tis): (OA) is a type of arthritis that involves the wearing down of cartilage (the protective covering on the ends of your bones) and loss of cushioning fluid in the joint.

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What is the SYNVIS[®] product?

SYNVISC is a gel-like mixture that is made up of hylan A fluid, hylan B gel, and salt water. Hylan A and hylan B are made from a substance called hyaluronan (pronounced hy-al-u-ROE-nan), also known as sodium hyaluronate that comes from chicken combs. Hyaluronan is a natural substance found in the body and is present in very high amounts in joints. The body's own hyaluronan acts like a lubricant and a shock absorber in the joint and is needed for the joint to work properly. Osteoarthritis (pronounced os-TE-o-ar-THRI-tis) (OA) is a type of arthritis that involves the wearing down of cartilage (the protective covering on the ends of your bones). In OA, there may not be enough hyaluronan, and there may be a decrease in the quality of the hyaluronan in the joint. SYNVIS[®] comes in syringes containing 2 mL (half a teaspoon) of product. SYNVIS[®] is injected directly into your knee.

How is the SYNVIS[®] product used? (Indications)

The FDA-approved indication for SYNVIS[®] is: SYNVIS[®] is indicated for the treatment of pain in osteoarthritis (OA) of the knee in patients who have failed to respond adequately to conservative non-pharmacologic therapy and simple analgesics, e.g., acetaminophen.

How is the SYNVIS[®] product given?

Your doctor will inject SYNVIS[®] into your knee.

Are there any reasons why I should not receive SYNVIS[®] injections? (Contraindications)

Your doctor will determine if there is any reason why you are not an appropriate candidate for SYNVIS[®]. You should be aware that SYNVIS[®]:

- Should not be used in patients who have had any prior allergic reactions to SYNVIS[®], Synvisc-One[®] or any hyaluronan-based products. Signs of an allergic reaction may include swelling of your face, tongue, or throat; difficulty breathing or swallowing; shortness of breath; wheezing; chest pain; a tightness in your throat; sleepiness; rash; itching; hives; flushing; and/or fever.
- Should not be used in patients with a knee joint infection, skin disease or infection around the area where the injection will be given, or circulatory problems in the legs.

What should my doctor warn me about?

The following are important treatment considerations for you to discuss with your doctor and understand in order to help avoid unsatisfactory results and complications:

- SYNVIS[®] is only for injection into the knee, performed by a doctor or other qualified health care professional. SYNVIS[®] has not been tested to show pain relief in joints other than the knee.
- SYNVIS[®] has not been tested to show better pain relief when combined with other injected medicines.
- Tell your doctor if you are allergic to products from birds such as feathers, eggs, and poultry.
- Tell your doctor if you have significant swelling or blood clots in the leg.
- SYNVIS[®] has not been tested in pregnant women, or women who are nursing. You should tell your doctor if you think you are pregnant, or if you are nursing a child.
- SYNVIS[®] has not been tested in children (≤ 21 years of age).

What are the risks of getting SYNVIS[®] injections?

The side effects (also called reactions) sometimes seen when SYNVIS[®] is injected into the knee as a first or repeat set of injections were pain, swelling, heat, redness, and/or fluid build-up around the knee. These reactions were generally mild and did not last long. If you have a reaction where the swelling is extensive and painful you should notify your doctor. The reactions seemed to occur more often when SYNVIS[®] was injected into the knee as a repeat set of injections than when SYNVIS[®] was injected as a first set of injections. Reactions are generally treated by resting and applying ice to the injected knee. Sometimes it is necessary to give pain relievers by mouth such as acetaminophen or NSAIDs, or to give injections of steroids, or to remove fluid from the knee joint. Patients rarely undergo arthroscopy (a surgical inspection of the knee joint) or other medical procedures related to these reactions.

Other less common side effects have been: rashes, hives, itching, muscle pain/cramps, flushing and/or swelling of your face, fast heart beat, nausea (or feeling sick to your stomach), dizziness, fever, chills, headache, difficulty breathing, swelling in your arms and/or legs, prickly feeling of your skin, and in rare cases a low number of platelets in the blood (platelets are a type of blood cell that are needed to help clot your blood when you are cut or injured). Before you are given SYNVISIC, tell your doctor if something like this has ever happened to you after receiving an injection of SYNVISIC or other hyaluronan products. If any of the above symptoms or signs appear after you are given SYNVISIC, or if you have any other problems, you should call your doctor. Rare cases of knee joint infection have been reported after SYNVISIC injections.

What are the benefits of getting SYNVISIC® injections?

As shown in medical studies of patients with osteoarthritis (OA) of the knee, where approximately half received a single injection of SYNVISIC and the other half either had fluid removed from the knee and/or received injections of the same volume of salt water (a "Saline Control" injection), the major benefits of SYNVISIC are pain relief and improvement in other symptoms related to OA of the knee.

What do I need to do after I get a SYNVISIC® injection?

It is recommended you avoid strenuous activities (for example, high impact sports such as soccer, tennis or jogging) or prolonged weight bearing activities for approximately 48 hours following the injection. You should consult your doctor regarding the appropriate time to resume such activities.

What other treatments are available for OA?

If you have OA, there are other things you can do besides getting SYNVISIC. These include:

Non-drug treatments

- Avoiding activities that cause knee pain
- Exercise or physical therapy
- Weight loss
- Removal of excess fluid from your knee

Drug therapy

- Pain relievers such as acetaminophen and narcotics
- Drugs that reduce inflammation (signs of inflammation are swelling, pain or redness), such as aspirin and other nonsteroidal anti-inflammatory drugs (NSAIDs, for example ibuprofen and naproxen)
- Steroids that are injected directly into your knee.

What did the clinical studies show?

Two medical studies involving a total of 132 patients were done in Germany. The patients in these studies were at least 40 years old and had knee pain due to OA. The patients were placed in one of two groups. One group was given an injection of SYNVISIC into one or both knees once a week for three weeks. The second group was given an injection of salt water once a week for three weeks. As part of the study, knee joint pain was measured for 26 weeks. Also, patients and doctors were asked to judge the success of the treatment for 26 weeks. Patients with OA knee pain, who did not get pain relief with other medicines, got pain relief with SYNVISIC. The patients given SYNVISIC had more pain relief than the patients

given salt water. Some patients started to feel pain relief after the first week of SYNVISIC treatment. The most pain relief and the greatest amount of treatment success was seen 8 to 12 weeks after SYNVISIC treatment started.

A medical study done in the United States involved 90 patients. The patients were at least 40 years old and had knee pain due to OA. Patients were placed into one of two groups. One group was given SYNVISIC once a week for three weeks. The second group had a needle inserted into the knee to have any fluid removed (this procedure is called arthrocentesis [pronounced AR-thro-sen-TEE-sis]) once a week for three weeks.

Patients improved after SYNVISIC treatment, but not more than patients who had arthrocentesis. This study was different from the German studies because the last time the two groups were compared was only two weeks after the last SYNVISIC injection. The study was also different in other ways, including length of time that patients had to stop taking medicines before they could start treatment. The length of time patients had to stop taking medicines was two weeks in the German studies and four weeks in the U.S. study.

What adverse events were observed in the clinical studies?

The side effects (also called reactions) sometimes seen when SYNVISIC is injected into the knee as a first or repeat set of injections were pain, swelling, heat, redness, and/or fluid build-up around the knee. These reactions were generally mild and did not last long.

How do I get more information about the SYNVISIC® product? (User Assistance)

If you have any questions or would like to find out more about SYNVISIC, you may call Genzyme Biosurgery at 1-888-3-SYNVISIC (1-888-379-6847) or visit www.synvisic.com.

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