

XIAFLEX REMS Video (Peyronie’s Disease)

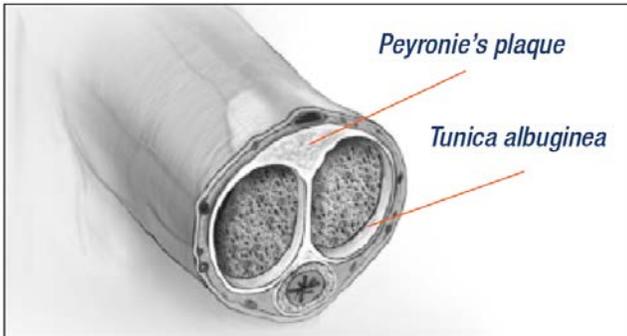
VISUAL	AUDIO
<p>TITLE FRAME</p>  <p>ON-SCREEN COPY: REMS training video for administering Xiaflex for Peyronie’s disease.</p> <p>VISUAL: REMS description scrolls</p> <p>A REMS is a strategy to manage known or potential risks associated with a drug, and is required by the FDA to ensure that the benefits of the drug outweigh its risks.</p> <p>Xiaflex, or collagenase clostridium histolyticum, is indicated for the treatment of adult men with Peyronie’s disease with a palpable plaque and curvature deformity of at least thirty degrees at the start of therapy</p> <p>Xiaflex is only available under a restricted distribution program called the Xiaflex REMS because of the risks of penile fracture and other serious penile injury associated with using Xiaflex in treating Peyronie’s disease.</p> <p>This training video is part of the Xiaflex REMS program. This video discusses:</p> <ul style="list-style-type: none"> • the steps necessary to prepare and administer Xiaflex • the in-office penile modeling procedure that is part of each Xiaflex treatment cycle • the daily, at-home penile modeling activities 	<p>VO:</p> <p>Welcome to the REMS training video for administering Xiaflex for Peyronie’s disease. A REMS is a strategy to manage known or potential risks associated with a drug, and is required by the FDA to ensure that the benefits of the drug outweigh its risks. Xiaflex, or collagenase clostridium histolyticum, is indicated for the treatment of adult men with Peyronie’s disease with a palpable plaque and curvature deformity of at least thirty degrees at the start of therapy [Ref: PI/p.3/§1/¶ 2].</p> <p>Xiaflex is only available under a restricted distribution program called the Xiaflex REMS because of the risks of penile fracture and other serious penile injury associated with using Xiaflex in treating Peyronie’s disease.</p> <p>This training video is part of the Xiaflex REMS program. This video discusses:</p> <ul style="list-style-type: none"> • the steps necessary to prepare and administer Xiaflex • the in-office penile modeling procedure that is part of each Xiaflex treatment cycle • the daily, at-home penile modeling activities that are performed for approximately 6 weeks after each treatment cycle • counseling your patient with the Xiaflex Patient Counseling Tool <p>Xiaflex is also indicated for the treatment of adult patients with Dupuytren’s contracture with a palpable cord. [Ref: PI/p.3/§1/¶ 1] Use of Xiaflex for Dupuytren’s contracture is covered under separate REMS requirements. Information on Dupuytren’s contracture is available by visiting www.XIAFLEX.com or by calling 1-877-313-1235.</p>

Introduction—Mechanism of Action

<p>that are performed for approximately 6 weeks after each treatment cycle</p> <ul style="list-style-type: none">• counseling your patient with the Xiaflex Patient Counseling Tool <p>Xiaflex is also indicated for the treatment of adult patients with Dupuytren’s contracture with a palpable cord. [Ref: PI/p.3/§1/¶ 1] Use of Xiaflex for Dupuytren’s contracture is covered under separate REMS requirements. Information on Dupuytren’s contracture is available by visiting www.Xiaflex.com or by calling 1-877-313-1235.</p> <p>Xiaflex® (collagenase clostridium histolyticum)</p> <p>© 2013 Auxilium Pharmaceuticals, Inc. All rights reserved RMX-00008</p>	
<p>Frame 1</p> <div data-bbox="131 1010 761 1346" data-label="Image">The image shows the Xiaflex logo, which consists of the word "XIAFLEX" in a bold, blue, sans-serif font with a registered trademark symbol. Below it, the words "collagenase clostridium histolyticum" are written in a smaller, grey, sans-serif font. A thin red horizontal line is positioned below the logo.</div> <p>VISUAL: REMS headline fades away and Xiaflex logo moves to center of screen.</p> <p>ON-SCREEN COPY: Xiaflex® collagenase clostridium histolyticum</p> <p>www.XiaflexREMS.com.</p>	<p>VO:</p> <p>For the treatment of Peyronie’s disease, Xiaflex should be administered by a healthcare provider experienced in the treatment of male urological diseases who has completed the required training. [Ref: PI/p.6/§2.2/¶ 1]</p> <p>This video demonstrates the steps necessary to prepare and administer Xiaflex. It also describes the in-office penile modeling procedure that is part of each Xiaflex treatment cycle, as well as the daily, at-home penile modeling activities that are performed for approximately six weeks after each cycle. [Ref: PI/p.9/§2.2/Penile Modeling for PD/¶1; PI/p.9/§2.2/Penile Modeling for PD/¶2] It also includes directions for utilizing the Xiaflex Patient Counseling Tool. The full Prescribing Information and Medication Guide are available at www.XiaflexREMS.com.</p>

Introduction—Mechanism of Action

Frame 2



VISUAL: Cut to cross-section illustration of penis with callouts for Peyronie's plaque and tunica albuginea

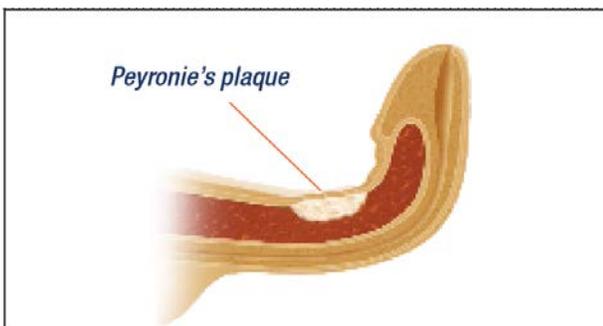
ON-SCREEN COPY: Peyronie's plaque; Tunica albuginea

VO:

Peyronie's disease is a localized connective tissue disorder characterized by changes in collagen composition in the tunica albuginea. [Ref: Hellstrom, p. 397/col 1/¶2/lines 1-4] These changes cause an abnormal scar formation known as Peyronie's plaque, which is typically a palpable bump under the skin. [Ref: Ralph, p 1, col 1, ¶1, lines 6-7; Ref: Moreland, p.408/col 2/¶1; Ref: Bella/p. 1529/col 1/¶ 2/lines 10-11] The Peyronie's plaque is composed predominantly of collagen, and replaces the normally elastic fibers of the tunica albuginea. [Ref: CSR/p.14/¶1/lines 2-3]

Microvascular trauma resulting from excessive bending or injury to the erect penis (possibly during sexual activity) is thought to be an important trigger for the inflammatory response and plaque development characteristic of Peyronie's disease. [Ref 3:Moreland, p.407/col 2/¶2; p.408/col.1/¶1] Genetic predisposition and autoimmunity may also play a role in its development. [Moreland, p.406/col 2/¶2/lines 3-8; lines 21-31]

Frame 3



VISUAL: Animation changes perspective to illustrate penis profile with Peyronie's plaque causing curvature

ON-SCREEN COPY: Peyronie's plaque

VO:

One of the hallmarks of Peyronie's disease is curvature deformity. Peyronie's disease may also cause other types of deformities, including narrowing, indentation, and shortening of the penis. [Ref 4: Bella/p. 1527/col 1/¶1/lines 4-8]

Mechanism of Action

Frame 4

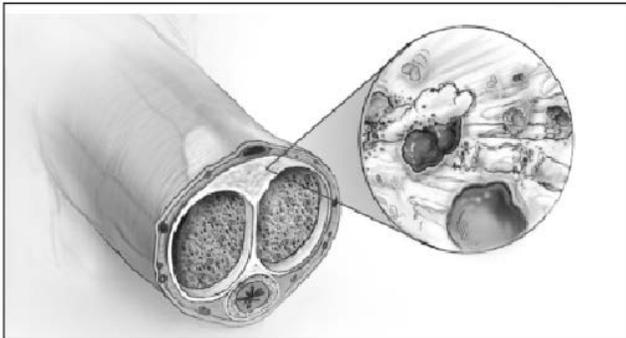


ON-SCREEN COPY: Xiaflex®
collagenase clostridium histolyticum
Mechanism of Action

VO:

Xiaflex contains two different types of purified collagenase clostridium histolyticum in a defined mass ratio. [\[Ref: PI/p.19/§11/¶1\]](#)

Frame 5

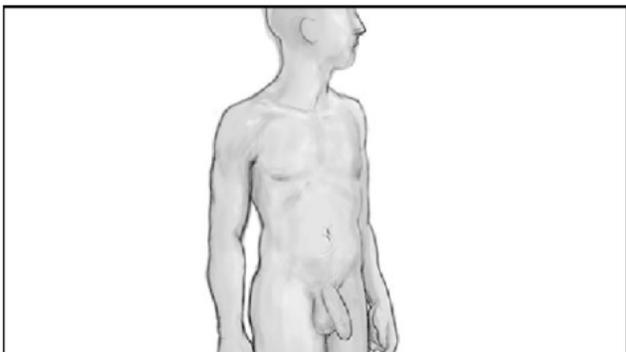


VISUAL: Animation illustrates enzymatic disruption of the collagen found in Peyronie's plaque

VO:

Injection of Xiaflex into a Peyronie's plaque, which is composed mostly of collagen, may result in enzymatic disruption of the collagen found in Peyronie's plaque. [\[Ref: PI/p.20/§12.1/¶3, lines 1-3\]](#)

Frame 6



VISUAL: Transition to full male figure

VO:

Following this disruption of the collagen-containing plaque, penile curvature deformity may improve and Patient-Reported Bother may be reduced. [\[Ref: PI/p.20/12.1/¶3/lines 2-4\]](#)

PREPARATION FOR ADMINISTRATION

Frame 7



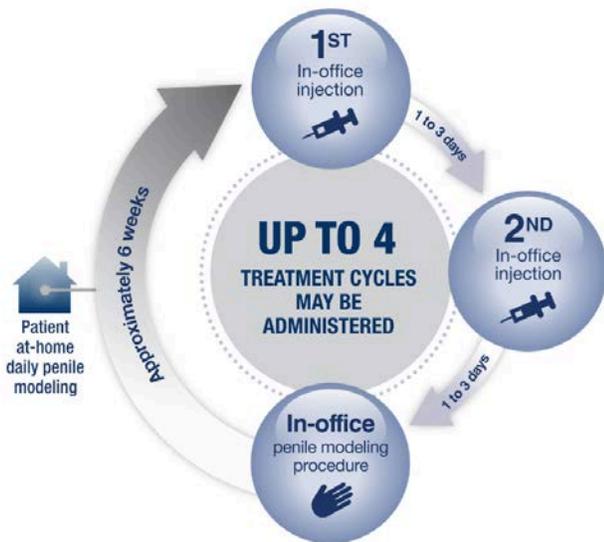
ON-SCREEN COPY: Xiaflex®
collagenase clostridium histolyticum
Treatment Overview

VO:

Xiaflex should be administered by a healthcare provider experienced in the treatment of male urological diseases who has completed the required training. [Ref: PI/p.6/§2.2/¶1]

Xiaflex, supplied as a lyophilized powder, must be reconstituted with the provided diluent prior to use. [Ref: PI/p.7/§2.2/¶2/lines 1-2] The dose of Xiaflex is 0.58 mg per injection administered into a Peyronie’s plaque. If more than one plaque is present, inject into the plaque causing the curvature deformity. [Ref: PI/p.7/§2.2/¶2/lines 2-4]

Frame 8



VISUAL: Treatment cycle diagram; Treatment Cycle 1 circles are highlighted in dark blue to match the VO

ON-SCREEN COPY: One Xiaflex Treatment Cycle

Treatment Cycle 1; 1st in-office injection procedure → 1 to 3 days → 2nd in-office injection procedure → 1 to 3 days → In-office penile modeling procedure
approximately 6 weeks; At-home penile modeling

VO:

A treatment course consists of a maximum of four treatment cycles. Each treatment cycle consists of two Xiaflex injection procedures and one in-office penile modeling procedure. The second Xiaflex injection procedure occurs one to three days after the first. The in-office penile modeling procedure is performed one to three days after the second injection of the treatment cycle. [Ref: PI/p.7/§2.2/¶3/lines 1-5] It is necessary to identify the treatment area prior to each treatment cycle.

[Ref: PI/p.8/§2.2/ID of Tx Area for PD, bullet a/line 1] Healthcare providers must counsel patients on the risks of penile fractures or other serious injuries of the penis and how to perform the at-home penile modeling activities as appropriate. After the third office visit of each treatment cycle, the patient performs approximately six weeks of daily, at-home penile modeling activities. [Ref: PI/p.9/§2.2/Penile Modeling for PD/¶2]

Up to four treatment cycles (for a total of eight injection procedures and four modeling procedures) may be administered per plaque causing the curvature deformity. If the curvature deformity is less than 15 degrees

PREPARATION FOR ADMINISTRATION

activities→Treatment Cycle 2; approximately 6 weeks; At-home penile modeling activities→Treatment Cycle 3; approximately 6 weeks; At-home penile modeling activities→Treatment Cycle 4; approximately 6 weeks; At-home penile modeling activities

Up to 4 treatment cycles may be administered

If the curvature deformity is less than 15 degrees after the 1st, 2nd or 3rd Treatment Cycle, or if you determine that further treatment is not clinically indicated, then subsequent treatment cycles should not be administered.

The safety of more than one treatment course of Xiaflex is not known.

after the first, second or third treatment cycle, or if you determine that further treatment is not clinically indicated, then the subsequent treatment cycles should not be administered. The safety of more than one treatment course of Xiaflex is not known. [Ref: PI/p.7/§2.2/¶1-3]

Frame 9



ON-SCREEN COPY: Xiaflex®
collagenase clostridium histolyticum
Preparation for Administration

VO:

In this section, we will explain the procedure for reconstitution of the lyophilized powder of Xiaflex.

Frame 10



VISUAL: Xiaflex vial

VO:

Xiaflex, supplied as a lyophilized powder, must be reconstituted with the provided diluent prior to use. Xiaflex is supplied in single-use glass vials containing 0.9 mg of collagenase clostridium histolyticum as a sterile, lyophilized powder for reconstitution. [Ref: PI/p.10/§3]

PREPARATION FOR ADMINISTRATION

Frame 11



VISUAL: Close-up of sterile diluent vial

VO:

Sterile diluent for reconstitution is provided in the package in a single-use glass vial containing 3 mL of 0.3 mg/mL calcium chloride dihydrate in 0.9% sodium chloride. Xiaflex must be reconstituted with the provided diluent prior to use. [\[Ref: PI/p.10/§3\]](#)

Frame 12



VISUAL: Xiaflex vial; Sterile diluent vial; temperature gauge illustrating 2°C to 8°C and 36°F to 46°F

ON-SCREEN COPY: Prior to Reconstitution
Store in refrigerator

VO:

The vials of lyophilized Xiaflex powder and sterile diluent should be stored in a refrigerator at two to eight degrees Celsius or thirty-six to forty-six degrees Fahrenheit. Do not freeze. [\[Ref: PI/p.31/§16/¶1\]](#)

Frame 13



VISUAL: Xiaflex vial; Sterile diluent vial

VO:

Before use, remove the vial containing the lyophilized powder of Xiaflex and the vial containing the diluent for reconstitution from the refrigerator and check the labels on both the diluent vial and the lyophilized powder vial to make sure they have not expired. Allow the two vials to stand at room temperature for at least fifteen minutes and no longer than sixty minutes [\[Ref: PI/p.7/§2.2/bullet a, lines 1-4\]](#)

PREPARATION FOR ADMINISTRATION

ON-SCREEN COPY: After refrigeration, let the vial containing Xiaflex and the vial containing diluent stand at room temperature for at least 15 minutes before use
No longer than 60 minutes

Frame 14



VISUAL: Two Xiaflex vials one intact, one eroded

ON-SCREEN COPY: INTACT; ERODED
If eroded, call 1-877-663-0412

VO:

Visually inspect the vial containing Xiaflex. The cake of lyophilized powder should be intact and white in color. If the cake has been eroded, it should not be used, and should be reported to Auxilium by calling 1-877-663-0412.

[Ref: PI/p.7/§2.2/bullet a, lines 4-5]

Frame 15



VISUAL: Gloved hands remove flip-off cap from Xiaflex vial

VO:

After removing the flip-off cap from each vial, using aseptic technique, swab the rubber stopper and surrounding surface of the vial containing Xiaflex and the vial containing the diluent for reconstitution with sterile alcohol.

[Ref: PI/p.8/§2.2/bullet b]

PREPARATION FOR ADMINISTRATION

Frame 16

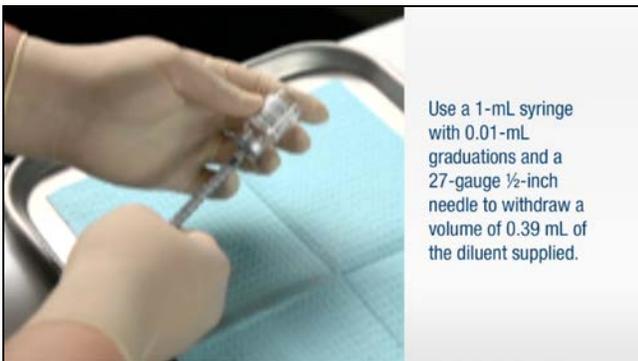


VISUAL: Gloved hands remove flip-off cap from diluent vial

VO:

No other antiseptics should be used. [\[Ref: PI/p.8/§2.2/bullet b\]](#) Use only the supplied diluent for reconstitution. The diluent contains calcium, which is required for the activity of Xiaflex. [\[Ref: PI/p.8/§2.2/bullet c\]](#)

Frame 17



VISUAL: Gloved hands hold 1-mL syringe with 0.01-mL graduations and a 27-gauge ½-inch needle to withdraw diluent

ON-SCREEN COPY: Use a 1-mL syringe with 0.01-mL graduations and a 27-gauge ½-inch needle to withdraw a volume of 0.39 mL of the **diluent supplied**

VO:

Using a 1-mL syringe with 0.01-mL graduations with a twenty-seven-gauge one-half-inch needle, which is not supplied, withdraw a volume of 0.39 mL of the **diluent supplied**. [\[Ref: PI/p.8/§2.2/bullet d\]](#)

PREPARATION FOR ADMINISTRATION

Frame 18



VISUAL: Gloved hands hold 1-mL syringe and inject diluent slowly into sides of Xiaflex vial

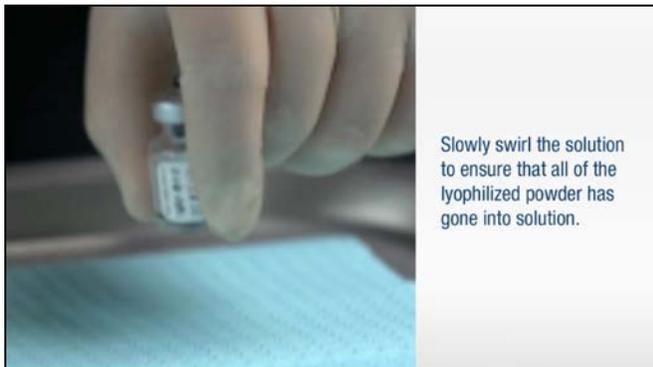
ON-SCREEN COPY: Inject the diluent slowly into the sides of the Xiaflex vial. Do not invert the vial or shake the solution

VO:

Inject the diluent slowly into the sides of the vial containing the lyophilized powder of Xiaflex.

[Ref: PI/p.8/§2.2/bullet e/lines 1-2]

Frame 19



VISUAL: Gloved hands slowly swirl the solution

ON-SCREEN COPY: Slowly swirl the solution to ensure that all of the lyophilized powder has gone into solution

VO:

Do not invert the vial or shake the solution. Slowly swirl the solution to ensure that all of the lyophilized powder has gone into solution. Do not use if opaque particles, discoloration, or other foreign particles are present. *[Ref: PI/p.8/§2.2/bullet e/lines 2-3]*

PREPARATION FOR ADMINISTRATION

Frame 20



VISUAL: Gloved hand holds vial of reconstituted Xialflex solution

VO:

The Xialflex solution is now ready for injection.

Frame 21



VISUAL: Vial with reconstituted Xialflex; clock illustrating 15-minute duration; clock animates to 60 minutes, illustrating the VO “and no longer than 60 minutes”

ON-SCREEN COPY: Reconstituted Xialflex
After refrigeration, let stand at room temperature for 15 minutes before use

VO:

The reconstituted Xialflex solution can be kept at room temperature of twenty to twenty-five degrees Celsius, or sixty-eight to seventy-seven degrees Fahrenheit, for up to one hour; or refrigerated at two to eight degrees Celsius, or thirty-six to forty-six degrees Fahrenheit, for up to four hours prior to administration. If the reconstituted Xialflex solution is refrigerated, allow this solution to return to room temperature for approximately fifteen minutes before use and no longer than sixty minutes.

[\[Ref: PI/p.8/§2.2/bullet f\]](#)

PREPARATION FOR ADMINISTRATION

Frame 22



VISUAL: Gloved hand discards syringe

VO:

As a final step, discard the syringe, needle and diluent used for reconstitution using medical waste disposal procedures. [\[Ref: PI/p.8/§2.2/bullet g\]](#)

Frame 23



ON-SCREEN COPY: Xiaflex®
collagenase clostridium histolyticum

Self-Test Questions

VO:

This completes the section on Xiaflex preparation for administration. To confirm understanding of the key points of this section, please answer the following self-test questions. After answering these questions correctly, you may continue to the next section.

INJECTION PROCEDURE

<p>Frame 24</p> <div data-bbox="136 275 760 604" style="border: 1px solid black; padding: 10px; text-align: center;">  <hr style="border: 0.5px solid orange; margin: 5px 0;"/> <p style="color: orange; font-weight: bold;">Self-Test Questions</p> </div> <p>ON-SCREEN COPY: Xiaflex® collagenase clostridium histolyticum Self-Test Questions</p>	<p>VO: Self-Test Questions</p>
<p>Frame 25 VISUAL: Question appears on-screen and answers appear sequentially, corresponding to VO ON-SCREEN COPY: Before use, for how long should the vials containing Xiaflex and the diluent be left to stand at room temperature? a) 5 to 10 minutes b) At least 15 but no more than 60 minutes c) 60 to 90 minutes d) At least 2 hours</p>	<p>VO: Before use, for how long should the vials containing Xiaflex and the diluent be left to stand at room temperature? a) Five to ten minutes b) At least fifteen but no more than sixty minutes c) Sixty to ninety minutes d) At least two hours</p>
<p>Frame 25A VISUAL: The correct answer is highlighted. ON-SCREEN COPY: Before use, for how long should the vials containing Xiaflex and the diluent be left to stand at room temperature? a) 5 to 10 minutes b) At least 15 but no more than 60 minutes c) 60 to 90 minutes d) At least 2 hours</p>	<p>VO: b) At least fifteen but no more than sixty minutes [<i>Ref: PI/p.8/§2.2/bullet f</i>]</p>
<p>Frame 26 VISUAL: Question appears on-screen and answers appear sequentially, corresponding to VO ON-SCREEN COPY: The amount of diluent that should be used for reconstituting the lyophilized powder of Xiaflex is: a) 0.15 mL</p>	<p>VO: The amount of diluent that should be used for reconstituting the lyophilized powder of Xiaflex is: a) 0.15 mL b) 0.25 mL c) 0.31 mL</p>

INJECTION PROCEDURE

<p>b) 0.25 mL c) 0.31 mL d) 0.39 mL</p>	<p>d) 0.39 mL</p>
<p>Frame 26A VISUAL: The correct answer is highlighted. ON-SCREEN COPY: The amount of diluent that should be used for reconstituting the lyophilized powder of Xiaflex is: a) 0.15 mL b) 0.25 mL c) 0.31 mL d) 0.39 mL</p>	<p>VO: d) 0.39 mL [<i>Ref: PI/p.8/§2.2/bullet d</i>]</p>
<p>Frame 27 VISUAL: Question appears on screen and answers appear sequentially, corresponding to VO ON-SCREEN COPY: The reconstituted Xiaflex solution can be kept at room temperature for up to 1 hour or refrigerated for up to: a) 2 hours b) 3 hours c) 4 hours d) 5 hours</p>	<p>VO: The reconstituted Xiaflex solution can be kept at room temperature for up to 1 hour or refrigerated for up to: a) Two hours b) Three hours c) Four hours d) Five hours</p>
<p>Frame 27A VISUAL: The correct answer is highlighted. ON-SCREEN COPY: The reconstituted Xiaflex solution can be kept at room temperature for up to 1 hour or refrigerated for up to: a) 2 hours b) 3 hours c) 4 hours d) 5 hours</p>	<p>VO: c) four hours [<i>Ref: PI/p.8/§2.2/bullet f</i>]</p>

INJECTION PROCEDURE

Frame 28



ON-SCREEN COPY: Xiaflex®
collagenase clostridium histolyticum

Identifying the Treatment Area and Injecting
Xiaflex

VO:

We will now discuss the proper procedures for identifying the treatment area and injecting the reconstituted Xiaflex (collagenase clostridium histolyticum) solution into the Peyronie's plaque.

Prior to administering Xiaflex and as part of every treatment-related visit, use the Patient Counseling Tool to discuss important information with each patient.

Frame 29



VISUAL: Gloved hand identifies the treatment area and induces an erection

VO:

Prior to each treatment cycle, identify the treatment area as follows: Induce a penile erection. A single intracavernosal injection of ten micrograms or twenty micrograms of alprostadil may be used for this purpose. Apply antiseptic at the site of injection and allow the skin to dry prior to the intracavernosal injection. [\[Ref: PI/p.8/§2.2/ID of Tx Area for PD, bullet a, sub-bullet 1\]](#)

Frame 30



VISUAL: Gloved hands locate the plaque at point of maximum concavity and mark the focal point with a surgical marker

VO:

Locate the plaque at the point of maximum concavity, or focal point, in the bend of the penis. [\[Ref: PI/p.8/§2.2/ID of Tx Area for PD, bullet a, sub-bullet 2\]](#)

Mark the point with a surgical marker. This indicates the target area in the plaque for Xiaflex deposition. [\[Ref: PI/p.8/§2.2/ID of Tx Area for PD, bullet a, sub-bullet 3\]](#)

INJECTION PROCEDURE

Frame 31



VISUAL: Gloved hand holds up vial for inspection

VO:

The reconstituted Xiaflex solution should be clear. Inspect the solution visually for particulate matter and discoloration prior to administration. If the solution contains particulates, is cloudy, or is discolored, do not inject the reconstituted solution. [\[Ref: PI/ p.8/§2.2/Inj Procedure for PD/bullet a\]](#)

Frame 32

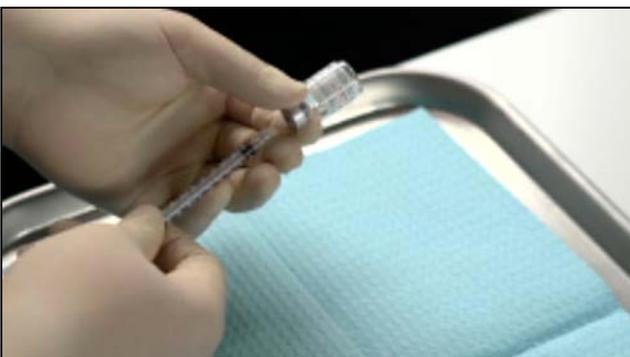


VISUAL: Gloved hands apply antiseptic

VO:

Apply antiseptic at the site of the injection and allow the skin to dry. If desired, administer suitable local anesthetic. [\[Ref: PI/ p.8/§2.2/Inj Procedure for PD/bullets b,c\]](#)

Frame 33



VISUAL: Gloved hands with hubless 0.01-mL syringe withdraw fluid

VO:

Using a new hubless syringe containing 0.01-mL graduations with a permanently fixed twenty-seven-gauge half-inch needle, which is not supplied, withdraw a volume of 0.25 mL of **reconstituted solution containing 0.58 mg of Xiaflex**. There will be reconstituted solution remaining in the vial. [\[Ref: PI/ p.8/§2.2/Inj Procedure for PD/bullet d\]](#)

INJECTION PROCEDURE

Frame 34

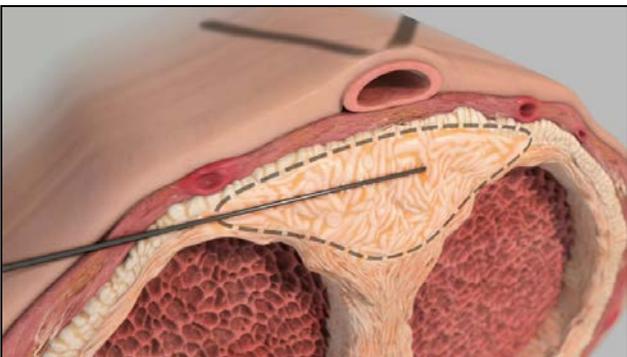


VISUAL: Gloved hands orient the needle so that it enters the plaque from the side

VO:

The penis should be in a flaccid state before injecting Xiaflex. Place the needle tip on the side of the target plaque in alignment with the point of maximal concavity. Orient the needle so that it enters the plaque from the side, NOT downward or perpendicularly toward the body of the corpora cavernosum. [\[Ref: PI/ p.8/§2.2/Inj Procedure for PD/bullet e\]](#)

Frame 35



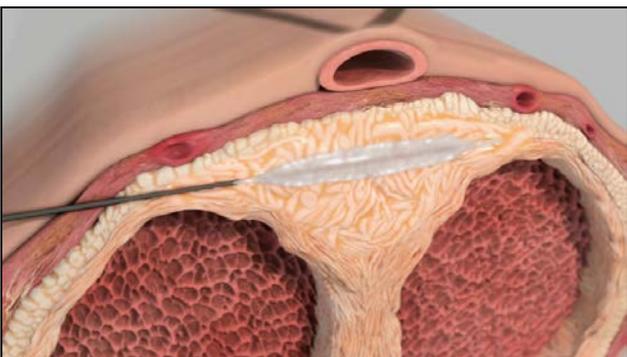
VISUAL: Cut to needle insertion animation

ON-SCREEN COPY: Image not to scale

VO:

Insert and advance the needle transversely through the width of the plaque, toward the opposite side of the plaque and without passing completely through it. Proper needle position is confirmed by carefully noting resistance to minimal depression of the syringe plunger. [\[Ref: PI/ p.9/§2.2/Inj Procedure for PD/bullet f\]](#)

Frame 36



VO:

With the tip of the needle placed within the plaque, initiate the injection, maintaining steady pressure to slowly inject the drug into the plaque. Withdraw the needle slowly so as to deposit the full dose along the needle track within the plaque. For plaques that are only a few millimeters in width, the distance of withdrawal of the syringe may be very minimal. The goal is always to deposit the full dose entirely within the plaque. [\[Ref: PI/ p.9/§2.2/Inj Procedure for PD/bullet g\]](#)

INJECTION PROCEDURE

<p>VISUAL: Needle withdrawal animation; arrow indicates direction</p> <p>ON-SCREEN COPY: Image not to scale</p>	
<p>Frame 37</p>  <p>VISUAL: A gloved hand applies pressure at the injection site</p>	<p>VO: Upon complete withdrawal of the needle, apply gentle pressure at the injection site. Apply a dressing as necessary. [Ref: PI/ p.9/§2.2/Inj Procedure for PD/bullet h]</p> <p>Discard the unused portion of the reconstituted solution and diluent after each injection. Do not store, pool, or use any vials containing unused reconstituted solution or diluent. [Ref: PI/ p.9/§2.2/Inj Procedure for PD/bullet i]</p>
<p>Frame 38</p>  <p>VISUAL: Gloved hand holds penis in place; copy and x's appear, indicating injection sites 2-3 mm apart</p> <p>ON-SCREEN COPY: 2-3 mm apart, but still within the plaque</p>	<p>VO: The second injection of each treatment cycle should be made approximately two to three millimeters apart from the first injection and within the plaque. [Ref: PI/ p.9/§2.2/Inj Procedure for PD/bullet j]</p>

INJECTION PROCEDURE

Frame 39

ON-SCREEN COPY: Xiaflex®
collagenase clostridium histolyticum

At each patient visit, counsel the patient as appropriate on the following:

- The risks of corporal rupture (penile fracture) and other serious injury to the penis
- That their penis may appear bruised and/or swollen
- That they may have mild-to-moderate penile pain that can be relieved by taking over-the-counter pain medications
- To promptly contact their physician if, at any time, they have any of these symptoms:
 - A popping sound or sensation in an erect penis
 - Sudden loss of the ability to maintain an erection
 - Severe purple bruising and swelling of the penis
 - Difficulty urinating or blood in the urine,
 - Severe pain in the penis
- These symptoms may be accompanied by a popping or cracking sound from the penis
- To return to their healthcare provider's office when directed for further injection(s) and/or penile modeling procedure(s)
- **To wait two weeks after the second injection of each treatment cycle before resuming sexual activity, provided pain and swelling have subsided**

VO:

At each patient visit, counsel the patient as appropriate on the following:

- The risks of corporal rupture (penile fracture) and other serious injury to the penis
- That their penis may appear bruised and/or swollen [\[Ref: PI/p.32/§17.2/bullet 2\]](#)
- That they may have mild-to-moderate penile pain that can be relieved by taking over-the-counter pain medications [\[Ref: PI/p.32/§17.2/bullet 3\]](#)
- To promptly contact their physician if, at any time, they have any of these symptoms:
 - a popping sound or sensation in an erect penis
 - sudden loss of the ability to maintain an erection
 - severe purple bruising and swelling of the penis
 - difficulty urinating or blood in the urine,
 - severe pain in the penis

These symptoms may indicate penile fracture, and may require surgery [\[Ref: PI/p.32/§17.2/bullet 4\]](#)

- To return to their healthcare provider's office when directed for further injection(s) and/or penile modeling procedure(s) [\[Ref: PI/p.32/§17.2/bullet 5\]](#)
- To wait two weeks after the second injection of each treatment cycle before resuming sexual activity, provided pain and swelling have subsided [\[Ref: PI/p.32/§17.2/bullet 6\]](#)

INJECTION PROCEDURE

Frame 40

XIAFLEX[®]
collagenase clostridium histolyticum

Self-Test Questions

ON-SCREEN COPY: Xiaflex[®]
collagenase clostridium histolyticum

Self-Test Questions

VO:

This completes the section on identifying the treatment area and injecting Xiaflex. To confirm understanding of the key points of this section, please answer the following self-test questions. After answering these questions, you may continue to the next section.

INJECTION PROCEDURE—SELF-TEST

<p>Frame 41</p> <div style="border: 1px solid black; padding: 10px; text-align: center;">  <p style="color: orange; margin-top: 20px;">Self-Test Questions</p> </div> <p>ON-SCREEN COPY: Xiaflex® collagenase clostridium histolyticum</p> <p>Self-Test Questions</p>	<p>VO:</p> <p>SELF-TEST QUESTIONS</p>
<p>Frame 42</p> <p>VISUAL: Question appears on-screen and answers appear sequentially, corresponding to VO</p> <p>ON-SCREEN COPY: The proper site of injection for Xiaflex is:</p> <ul style="list-style-type: none"> a) Laterally into the distal two-thirds of the penis b) At the point of minimal concavity in the bend of the penis c) At the point of maximal concavity in the bend of the penis d) 2 mm from the base of the erect penis 	<p>VO:</p> <p>The proper site of injection for Xiaflex is:</p> <ul style="list-style-type: none"> a) Laterally into the distal two-thirds of the penis b) At the point of minimal concavity in the bend of the penis c) At the point of maximal concavity in the bend of the penis d) Two millimeters from the base of the erect penis
<p>Frame 42A</p> <p>VISUAL: The correct answer is highlighted. ON-SCREEN COPY: The proper site of injection for Xiaflex is:</p> <ul style="list-style-type: none"> a) Laterally into the distal two-thirds of the penis b) At the point of minimal concavity in the bend of the penis c) At the point of maximal concavity in the bend of the penis d) 2 mm from the base of the erect penis 	<p>VO:</p> <p>c) At the point of maximal concavity in the bend of the penis <i>[Ref: PI/p.8/§2.2/ID of Tx Area for PD, bullet a, sub-bullet 2]</i></p>
<p>Frame 43</p> <p>VISUAL: Question appears on-screen and answers appear sequentially, corresponding to VO</p> <p>ON-SCREEN COPY: The amount of reconstituted Xiaflex that should be injected into the Peyronie’s plaque is:</p> <ul style="list-style-type: none"> a) 0.20 mL b) 0.25 mL c) 0.31 mL 	<p>VO:</p> <p>The amount of reconstituted Xiaflex that should be injected into the Peyronie’s plaque is:</p> <ul style="list-style-type: none"> a) 0.20 mL b) 0.25 mL c) 0.31 mL d) 0.39 mL

INJECTION PROCEDURE—SELF-TEST

<p>d) 0.39 mL</p>	
<p>Frame 43A VISUAL: The correct answer is highlighted. ON-SCREEN COPY: The amount of reconstituted Xiaflex that should be injected into the Peyronie’s plaque is: a) 0.20 mL b) 0.25 mL c) 0.31 mL d) 0.39 mL</p>	<p>VO: b) 0.25 mL [Ref: PI/ p.8/§2.2/Inj Procedure for PD/bullet d]</p>
<p>Frame 44 VISUAL: Question appears on-screen and answers appear sequentially, corresponding to VO ON-SCREEN COPY: The penis should be _____ when marking the treatment area and _____ when injecting Xiaflex. a) Erect; erect b) Flaccid; flaccid c) Erect; flaccid d) Flaccid; erect</p>	<p>VO: When marking the treatment area the penis should be (a) flaccid or (b) erect. When injecting Xiaflex the penis should be: (a) flaccid or (b) erect.</p>
<p>Frame 44A VISUAL: Answers disappear and the correct answers are inserted into the question and highlighted. ON-SCREEN COPY: The penis should be erect when marking the treatment area and flaccid when injecting Xiaflex.</p>	<p>VO: The penis should be erect when marking the treatment area and flaccid when injecting Xiaflex. [Ref: PI/p.8/§2.2/ID of Tx Area for PD/bullet a/sub-bullet 1] [Ref: PI/ p.8/§2.2/Inj Procedure for PD/bullet e/line 1]</p>
<p>ALT FRAME 44 VISUAL: Question appears on-screen and answers appear sequentially, corresponding to VO ON-SCREEN COPY: When marking the treatment area should the penis be (a) flaccid or (b) erect?</p>	<p>VO: When marking the treatment area should the penis be (a) flaccid or (b) erect?</p>

INJECTION PROCEDURE—SELF-TEST

<p>ALT FRAME 44A VISUAL: The correct answer is highlighted. ON-SCREEN COPY: When marking the treatment area should the penis be (a) flaccid or (b) erect</p>	<p>VO: When marking the treatment area the penis should be erect. [Ref: PI/p.8/§2.2/ID of Tx Area for PD/bullet a/sub-bullet 1]</p>
<p>ALT FRAME 44B VISUAL: Question appears on-screen and answers appear sequentially, corresponding to VO ON-SCREEN COPY: When injecting Xiaflex should the penis be: (a) flaccid or (b) erect?</p>	<p>VO: When injecting Xiaflex should the penis be: (a) flaccid or (b) erect?</p>
<p>ALT FRAME 44C VISUAL: Answers disappear and the correct answers are inserted into the question and highlighted. ON-SCREEN COPY: When injecting Xiaflex should the penis be: (a) flaccid or (b) erect?</p>	<p>VO: When injecting Xiaflex should the penis be: flaccid [Ref: PI/ p.8/§2.2/Inj Procedure for PD/bullet e/line 1]</p>
<p>Frame 45 VISUAL: Question appears on-screen and answers appear sequentially, corresponding to VO ON-SCREEN COPY: The needle needs to be inserted _____ the plaque a) Perpendicular to b) Transversely through the width of c) Parallel to d) Adjacent to, but not within</p>	<p>VO: The needle needs to be inserted <i>in which direction into</i> the plaque? a) Perpendicular to b) Transversely through the width of c) Parallel to d) Adjacent to, but not within</p>
<p>Frame 45A VISUAL: Answers disappear and the correct answer is inserted into the question and highlighted. ON-SCREEN COPY: The needle needs to be inserted <u>transversely through the width of</u> the plaque</p>	<p>VO: The needle needs to be inserted transversely through the width of the plaque? [Ref: PI/ p.9/§2.2/Inj Procedure for PD/bullet f/lines 1-2]</p>

INJECTION PROCEDURE—SELF-TEST

<p>Frame 46 VISUAL: Question appears on-screen and answers appear sequentially, corresponding to VO ON-SCREEN COPY: The goal of injection is always to deposit the full dose of drug: a) Entirely within the plaque b) Mostly within the plaque c) Entirely outside of the plaque d) Both inside and outside of the plaque</p>	<p>VO: The goal of injection is always to deposit the full dose of drug... a) Entirely within the plaque b) Mostly within the plaque c) Entirely outside of the plaque d) Both inside and outside of the plaque</p>
<p>Frame 46A VISUAL: The correct answer is highlighted. ON-SCREEN COPY: The goal of injection is always to deposit the full dose of drug: a) Entirely within the plaque b) Mostly within the plaque c) Entirely outside of the plaque d) Both inside and outside of the plaque</p>	<p>VO: The goal of injection is always to deposit the full dose of drug entirely within the plaque <i>[Ref: PI/p.9/\$2.2/Inj Procedure for PD/bullet g/lines 5-6]</i></p>

PENILE MODELING (IN-OFFICE AND AT-HOME)

Frame 47



ON-SCREEN COPY: Xiaflex®
collagenase clostridium histolyticum
Penile Modeling (In-Office and At-Home)

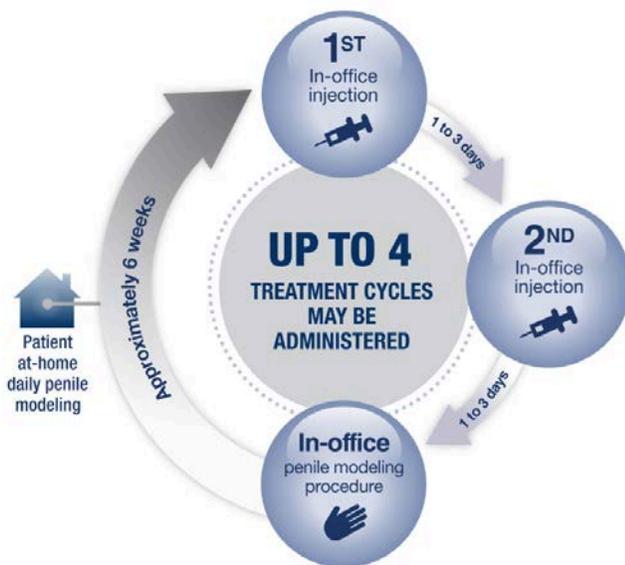
VO:

In conjunction with Xiaflex, penile modeling helps improve curvature deformity and straighten the penile shaft. In this section, we will review the in-office penile modeling procedure that is performed one to three days following the second injection of Xiaflex in each treatment cycle. [\[Ref: PI/ p.9/§2.2/Penile Modeling Procedure/¶ 1\]](#) We will also review the at-home penile modeling activities that patients should be instructed to do daily.

[\[Ref: PI/ p.9/§2.2/Penile Modeling for PD/¶ 2\]](#)

Prior to administering Xiaflex and as part of every treatment-related visit, use the Patient Counseling Tool to discuss important information with each patient.

Frame 48



VISUAL: “In-office penile modeling procedure” is highlighted to correspond with VO

ON-SCREEN COPY: One Xiaflex Treatment Cycle

Treatment Cycle 1; 1st in-office injection procedure → 1 to 3 days → 2nd in-office injection procedure → 1 to 3 days → In-office penile modeling

VO:

At a follow-up visit one to three days after the second injection of each treatment cycle, perform the in-office penile modeling procedure on the flaccid penis in order to stretch and elongate the treated plaque. [\[Ref: PI/ p.9/§2.2/Penile Modeling for PD/¶ 1\]](#)

PENILE MODELING (IN-OFFICE AND AT-HOME)

procedure

approximately 6 weeks; At-home penile modeling activities→Treatment Cycle 2; approximately 6 weeks; At-home penile modeling activities→Treatment Cycle 3; approximately 6 weeks; At-home penile modeling activities→Treatment Cycle 4; approximately 6 weeks; At-home penile modeling activities

Up to 4 treatment cycles may be administered

If the curvature deformity is less than 15 degrees after the 1st, 2nd or 3rd Treatment Cycle, or if you determine that further treatment is not clinically indicated, then subsequent treatment cycles should not be administered.

The safety of more than one treatment course of Xiaflex is not known.

Frame 49



VISUAL: Gloved hands grasp the plaque or indurated portion of the flaccid penis about 1 cm proximal and distal to the injection site.

VO:

If desired, administer suitable local anesthetic.
[\[Ref: PI/ p.9/§2.2/Penile Modeling Procedure/bullet 1\]](#)

Wearing gloves, grasp the plaque or indurated portion of the flaccid penis about one centimeter proximal and distal to the injection site. Avoid direct pressure on the injection site.
[\[Ref: PI/ p.9/§2.2/Penile Modeling Procedure/bullet 2\]](#)

SELF-TEST — PENILE MODELING (IN-OFFICE AND AT-HOME)

Frame 50



VISUAL: Gloved hands apply firm, steady pressure to elongate and stretch the plaque, creating bending opposite to the patient's penile curvature

ON-SCREEN COPY:

- Use target plaque as fulcrum point
- Apply firm, steady pressure
- Gradually bend the shaft in the opposite direction of the curvature

VO:

Using the target plaque as a fulcrum point, use both hands to apply firm, steady pressure to elongate and stretch the plaque. The goal is to gradually create bending opposite to the patient's penile curvature, with stretching to the point of moderate resistance.

[\[Ref: PI/ p.9/§2.2/Penile Modeling Procedure/bullet 3\]](#)

Hold pressure for thirty seconds then release.

[\[Ref: PI/ p.9/§2.2/Penile Modeling Procedure/bullet 3\]](#)

Frame 51



VISUAL: Gloved hands repeat the penile modeling technique

ON-SCREEN COPY:

- Rest for 30 seconds
- Repeat a total of 3 times
- Hold for 30 seconds each time

VO:

After a thirty-second rest period, repeat the penile modeling technique for a total of three modeling attempts at thirty seconds each.

[\[Ref: PI/ p.9/§2.2/Penile Modeling Procedure/bullet 4\]](#)

Frame 52

VO:

For the approximately six weeks following each treatment cycle, patients will need to perform the following penile modeling activities at home daily to help reduce penile curvature [\[Ref:](#)

[PI/p.9/§2.2/Penile Modeling for PD/¶12\]](#)

SELF-TEST — PENILE MODELING (IN-OFFICE AND AT-HOME)



ON-SCREEN COPY: Xiaflex®
collagenase clostridium histolyticum
At-home Penile Modeling Activities

Frame 52A



VO:
There are two types of home penis activities. One is a stretching activity. The other is a straightening activity. Be sure to instruct patients on exactly when to start the activities and how long to continue performing them. [\[Ref: PI/p.9/§2.2/Penile Modeling for PD/¶2\]](#)

The stretching activity should be performed three times daily and when the penis is not erect. Instruct patients to grasp the tip of the penis with the fingers of one hand and hold the base of the penis with the fingers of the other. Then, **gently** pull the penis away from the body to its full length. Hold the stretch for thirty seconds. Then let go and allow the penis to return to its normal unstretched length. [\[Ref: PI/p.10/§2.2/Penile Modeling for PD/bullet 2\]](#)

The penile straightening activity is performed a maximum of once per day on an erection unrelated to sexual activity. If the patient does not have a spontaneous erection, he should not attempt the penis straightening. Instruct patients to **gently** attempt to bend the shaft of the erect penis in the opposite direction of the curve, but not so forcefully as to produce significant pain or discomfort. Patients should hold the penis in this more straightened position for thirty seconds, then let go. [\[Ref: PI/p.10/§2.2/Penile Modeling for PD/bullet 1\]](#)

SELF-TEST — PENILE MODELING (IN-OFFICE AND AT-HOME)

 <p>VISUAL: Animation demonstrates at-home patient stretching (first of three images above) and straightening (second and third images above) activities.</p>	
<p>Frame 53</p> <p>ON-SCREEN COPY: Provide instructions on at-home penile modeling activities:</p> <ul style="list-style-type: none"> • On flaccid penis: Stretch 3 times daily for 30 seconds at a time • On erect penis: Gently straighten and hold for 30 seconds, once daily 	<p>VO: In summary, penis stretching is performed on a non-erect penis, three times a day for thirty seconds each time. Penis straightening is performed no more than once a day on a spontaneous erection unrelated to sexual activity. Discuss with patients the best time to perform these activities. <i>[Ref: PI/p.10/§2.2/Penile Modeling for PD/bullets 1-2]</i></p>
<p>Frame</p> <div data-bbox="168 1283 764 1602" style="border: 1px solid black; padding: 10px; text-align: center;">  </div> <p>54</p> <p>ON-SCREEN COPY: Xiaflex® collagenase clostridium histolyticum Self-Test Questions</p>	<p>VO: This completes the section on the Xiaflex penile modeling, both in-office and at-home. To confirm understanding of the key points of this section, please answer the following self-test questions. After answering these questions, you may continue to the next section.</p>
<p>Frame 55</p> <p>VISUAL: Question appears on-screen and answers</p>	<p>VO: How soon should the in-office penile modeling</p>

SELF-TEST — PENILE MODELING (IN-OFFICE AND AT-HOME)

<p>appear sequentially, corresponding to VO</p> <p>ON-SCREEN COPY: In-office penile modeling procedure should be performed _____ after the second injection of each treatment cycle:</p> <ul style="list-style-type: none">a) Immediatelyb) 15 to 60 minutesc) 1 to 3 daysd) 5 to 7 days	<p>procedure be performed after the second injection of each treatment cycle:</p> <ul style="list-style-type: none">a) Immediatelyb) Fifteen to sixty minutesc) One to three daysd) Five to seven days
<p>Frame 55A</p> <p>VISUAL: The correct answer is highlighted.</p> <p>ON-SCREEN COPY: In-office penile modeling procedure should be performed _____ after the second injection of each treatment cycle:</p> <ul style="list-style-type: none">a) Immediatelyb) 15 to 60 minutesc) 1 to 3 daysd) 5 to 7 days	<p>VO:</p> <p>In-office penile modeling procedure should be performed one to three days after the second injection of each treatment cycle: [Ref: PI/p.9/§2.2/Penile Modeling for PD/¶1]</p>
<p>Frame 56</p> <p>VISUAL: Question appears on-screen and answers appear sequentially, corresponding to VO</p> <p>ON-SCREEN COPY: When performing in-office penile modeling procedure, hold the pressure for 30 seconds and rest for 30 seconds for a total of:</p> <ul style="list-style-type: none">a) 2 timesb) 3 timesc) 5 timesd) 10 times	<p>VO:</p> <p>When performing in-office penile modeling procedure, hold the pressure for thirty seconds and rest for thirty seconds for a total of:</p> <ul style="list-style-type: none">a) Two timesb) Three timesc) Five timesd) Ten times
<p>Frame 56A</p> <p>VISUAL: The correct answer is highlighted.</p> <p>ON-SCREEN COPY: When performing in-office penile modeling procedure, hold the pressure for 30 seconds and rest for 30 seconds for a total of:</p> <ul style="list-style-type: none">a) 2 timesb) 3 timesc) 5 timesd) 10 times	<p>VO:</p> <p>When performing in-office penile modeling procedure, hold the pressure for thirty seconds and rest for thirty seconds for a total of: three times. [Ref: PI/ p.9/§2.2/Penile Modeling Procedure/bullets 3 and 4]</p>

SUMMARY

<p>Frame 57 VISUAL: Question appears on-screen and answers appear sequentially, corresponding to VO ON-SCREEN COPY: The patient should be instructed to perform at-home penile <i>straightening</i> activity on a spontaneous <u>erection</u> unrelated to sexual activity no more than once daily for 30 seconds. The patient should also be instructed to perform the <i>stretching</i> activity on the <u>flaccid penis</u>:</p> <ul style="list-style-type: none">a) At no timeb) Once daily for a total of 1 minutec) 5 times daily for 30 seconds at a timed) 3 times daily for 30 seconds at a time	<p>VO: The patient should be instructed to perform at-home penile <i>straightening</i> activity on a spontaneous <u>erection</u> unrelated to sexual activity no more than once daily for thirty seconds. How often should the patient perform the <i>stretching</i> activity on the <u>flaccid penis</u>:</p> <ul style="list-style-type: none">a) At no timeb) Once daily for a total of one minutec) Five times daily for thirty seconds at a timed) Three times daily for thirty seconds at a time
<p>Frame 57A VISUAL: The correct answer is highlighted. ON-SCREEN COPY: The patient should be instructed to perform at-home penile <i>straightening</i> activity on an <u>erection</u> unrelated to sexual activity once daily for 30 seconds. The patient should also be instructed to perform the <i>stretching</i> activity on the <u>flaccid penis</u>:</p> <ul style="list-style-type: none">a) At no timeb) Once daily for a total of 1 minutec) 5 times daily for 30 seconds at a timed) 3 times daily for 30 seconds at a time	<p>VO: Instruct the patient to perform the <i>stretching</i> activity on the <u>flaccid penis</u> three times daily for thirty seconds at a time. [Ref: PI/p.10/§2.2/Penile Modeling for PD/bullet 2]</p>
<p>Frame 58 VISUAL: Title screen with Xiaflex logo and copy below. ON-SCREEN COPY: Xiaflex® collagenase clostridium histolyticum Instructions for Using the Xiaflex Patient Counseling Document</p>	<p>VO: Instructions for using the Xiaflex Patient Counseling Document.</p>

SUMMARY

<p>Frame 59 VISUAL: Copy appears and scrolls as VO is heard</p> <p>ON-SCREEN COPY: Prior to initiating treatment with Xiaflex, and as part of each treatment-related visit, discuss the following information included in the Patient Counseling Document with each patient:</p> <ul style="list-style-type: none">• The risks of corporal rupture (penile fracture) and other serious penile injury• Precautions related to the patient’s role in reducing the risks of these adverse outcomes (for example, advising patients to wait two weeks until after the second injection of each treatment cycle before resuming sexual activity)• Conditions under which patients should promptly contact their healthcare provider• Clear instructions on at-home penile modeling activities• Important information regarding the safe use of Xiaflex in treating Peyronie’s disease <p>The patient must be given a copy of the Patient Counseling Document to take home.</p> <p>In addition, provide a Medication Guide to each patient prior to each injection of Xiaflex.</p>	<p>VO:</p> <p>Prior to initiating treatment with Xiaflex, and as part of each treatment-related visit, discuss the following information included in the Patient Counseling Tool with each patient:</p> <ul style="list-style-type: none">• The risks of corporal rupture (penile fracture) and other serious penile injury• Precautions related to the patient’s role in reducing the risk of these adverse outcomes (for example, advising patients to wait two weeks until after the second injection of each treatment cycle before resuming sexual activity)• Conditions under which patients should promptly contact their healthcare provider• Clear instructions on at-home penile modeling activities• Important information regarding the safe use of Xiaflex in treating Peyronie’s disease <p>The patient must be given a copy of the Patient Counseling Document to take home.</p> <p>In addition, provide a Medication Guide to each patient prior to each injection of Xiaflex.</p>
<p>Frame 60</p> <p>VISUAL: Scrolling copy stops and fades to new copy</p> <p>ON-SCREEN COPY: To obtain copies of the Patient Counseling Document:</p> <ul style="list-style-type: none">• Visit www.XiaflexREMS.com• Call 1-877-XIAFLEX (1-877-942-3539)• Or contact your Xiaflex sales representative	<p>VO:</p> <p>To obtain copies of the Patient Counseling Document:</p> <ul style="list-style-type: none">• Visit www.XiaflexREMS.com• Call 1-877-XIAFLEX (1-877-942-3539)• Or contact your Xiaflex sales representative

SUMMARY

<p>Frame 61</p> <div data-bbox="136 285 764 625" style="border: 1px solid black; padding: 10px; text-align: center;"><p>XIAFLEX[®] collagenase clostridium histolyticum</p><hr/><p>Summary</p></div> <p>ON-SCREEN COPY: Xiaflex[®] collagenase clostridium histolyticum Summary</p>	<p>VO:</p> <p>Thank you for taking the time to review the REMS training video for using Xiaflex in treating Peyronie’s disease. Here are some key points to remember.</p>
<p>Frame 62</p> <p>ON-SCREEN COPY:</p> <p>Storage and handling</p> <ul style="list-style-type: none">• Keep vials of Xiaflex and sterile diluent refrigerated between 2° and 8° C or 36° to 46° F• Before use, allow vials to stand at room temperature for at least 15 minutes and no longer than 60 minutes• Reconstituted solution can be kept at room temperature for up to 1 hour or refrigerated for up to 4 hours prior to administration. If refrigerated, allow to return to room temperature for 15 minutes before use	<p>VO:</p> <p>STORAGE AND HANDLING</p> <p>Prior to reconstitution, the vials of lyophilized powder of Xiaflex and sterile diluent should be stored in a refrigerator at two to eight degrees Celsius or thirty-six to forty-six degrees Fahrenheit. [Ref: PI/p.31/§16/¶1]</p> <ul style="list-style-type: none">• Before preparing Xiaflex, allow vials to stand at room temperature for at least fifteen minutes and no longer than sixty minutes. [Ref: PI/p.7/§2.2/bullet a, lines 1-4]• The reconstituted Xiaflex solution can be kept at room temperature of twenty to twenty-five degrees Celsius or sixty-eight to seventy-seven degrees Fahrenheit for up to one hour or refrigerated at two to eight degrees Celsius or thirty-six to forty-six degrees Fahrenheit for up to four hours prior to administration. [Ref: PI/p.8/§2.2/bullet f, lines 1-3]
<p>Frame 63</p>	<p>VO:</p> <p>If the reconstituted Xiaflex solution is refrigerated, allow this solution to return to room temperature for approximately fifteen minutes before use.</p> <p>[Ref: PI/p.8/§2.2/bullet f, lines 3-5]</p>

SUMMARY



VISUAL: Vial with reconstituted Xiaflex; clock illustrating 15-minute interval

ON-SCREEN COPY: Reconstituted Xiaflex

After refrigeration, let stand at room temperature for 15 minutes before use

Frame 64



VISUAL: Gloved hands mark injection site with surgical marker.

ON-SCREEN COPY: Xiaflex®
collagenase clostridium histolyticum

Identifying the Treatment Area

- Induce an erection
- Locate the point of maximum concavity
- Mark the area with a marker

VO:

Identifying the Treatment Area

After inducing an erection, locate the point of maximum concavity, or focal point in the bend of the penis, which is where the plaque and the curvature intersect. *[Ref: PI/p.8/§2.2/ID of Tx Area for PD, bullet a, sub-bullets 1-2]*

Frame 65

VO:

INJECTION PROCEDURE

Withdraw 0.25 mL of reconstituted Xiaflex solution. *[Ref: PI/p. 8/§2.2/Inj Procedure for PD/bullet d]*

The penis should be in a flaccid state before injecting Xiaflex. Place the needle tip on the side of the target plaque in alignment with

SUMMARY



VISUAL: Two gloved hands inject penis with syringe; copy bullets appear, corresponding with VO

ON-SCREEN COPY: Injection Procedure

- Withdraw 0.25 mL of reconstituted Xiaflex solution
- Penis should be flaccid
- Place needle tip on the side of the plaque in alignment with the point of maximal concavity
- Insert and advance the needle transversely
- Confirm proper positioning by noting resistance in syringe plunger
- Deposit full dose within the plaque

the point of maximal concavity. Orient the needle so that it enters the plaque from the side, NOT downward or perpendicularly toward the body of the corpora cavernosum.

[\[Ref: PI/p. 8/§2.2/Inj Procedure for PD/bullet e\]](#)

Insert and advance the needle transversely through the width of the plaque, toward the opposite side of the plaque without passing completely through it. Proper needle position is confirmed by carefully noting resistance to minimal depression of the syringe plunger.

[\[Ref: PI/p. 9/§2.2/Inj Procedure for PD/bullet f\]](#)

Withdraw the needle slowly, so as to deposit the full dose along the needle track within the plaque. The goal is always to deposit the full dose entirely within the plaque. [\[Ref: PI/p. 9/§2.2/Inj Procedure for PD/bullet g\]](#)

Frame 66



VISUAL: Gloved hands bending penis opposite to penile curvature, with stretching to the point of moderate resistance.

ON-SCREEN COPY:

- Create a reverse bend
- Stretch to point of moderate resistance
- Hold for 30 seconds

IN-OFFICE PENILE MODELING

VO:

PENILE MODELING (IN-OFFICE AND AT-HOME)

In-office penile modeling procedure should be performed one to three days after the second injection of each treatment cycle. [\[Ref: PI/p.9/§2.2/Penile Modeling Procedure/¶ 1\]](#)

Using the target plaque as a fulcrum point, use both hands to apply firm, steady pressure to elongate and stretch the plaque. The goal is to gradually create bending opposite to the patient's penile curvature, with stretching to the point of moderate resistance.

Hold pressure for thirty seconds and then release. [\[Ref: PI/p.9/§2.2/Penile Modeling Procedure/bullet 3\]](#)

After a thirty-second rest period, repeat the penile modeling technique for a total of three times at thirty seconds each.

SUMMARY

	<p>[Ref: PI/ p.9/§2.2/Penile Modeling Procedure/bullet 4]</p>
<p>Frame 67</p> <p>At-home penile modeling activities:</p> <div data-bbox="136 380 641 590"><p>Stretching</p><p>On flaccid penis: Stretch 3 times daily for 30 seconds at a time</p><p>Straightening</p><p>On erect penis: Gently straighten and hold for 30 seconds, once daily</p></div> <p>VISUAL: Illustrations of stretching and straightening technique for at-home penile modeling</p> <p>ON-SCREEN COPY: At-home penile modeling activity:</p> <ul style="list-style-type: none">• On flaccid penis: Stretch 3 times daily for 30 seconds at a time• On erect penis: Gently straighten and hold for 30 seconds once per day	<p>At-home penile modeling activities involve a gentle stretching and straightening of the penis.</p> <p>[Ref: PI/p.9/§2.2/Penile Modeling for PD/¶12]</p> <p>The stretching activity should be performed three times daily and when the penis is not erect. Stretching should last for thirty seconds.</p> <p>[Ref: PI/p.10/§2.2/Penile Modeling for PD/bullet 2]</p> <p>The straightening activity is performed for thirty seconds, no more than once per day on a spontaneous erection unrelated to sexual activity. [Ref: PI/p.10/§2.2/Penile Modeling for PD/bullet 1]</p> <p>Remember, prior to initiating treatment and as part of each treatment-related visit, use the Xiaflex Patient Counseling Tool to discuss important information with each patient.</p>

CONTACT US

FRAME 68

XIAFLEX[®]
collagenase clostridium histolyticum

Contact Us
1-877-663-0412

ON-SCREEN COPY: Xiaflex[®]
collagenase clostridium histolyticum
Contact Us
1-877-663-0412

VO:

If you have product-related questions or to report adverse events, please contact the Auxilium Drug Information Call Center at 1-877-663-0412.

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