

## INSTRUCTIONS FOR PREPARING AND GIVING AN INJECTION OF XYNTHA IN THE PREFILLED DUAL CHAMBER SYRINGE

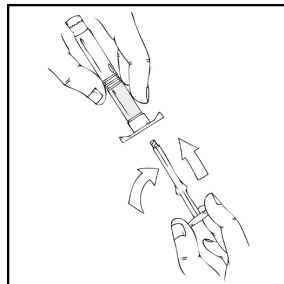
XYNTHA is given by intravenous (IV) injection after reconstituting (mixing) the freeze-dried powder with the diluent (0.9% sodium chloride solution). Both the XYNTHA powder and the diluent are supplied within the prefilled dual chamber syringe.

### Preparation

1. Always wash your hands before doing the following procedures.
2. Aseptic technique (clean and germ-free) should be used during the reconstitution procedure.
3. All components used in the reconstitution and administration of this product should be used as soon as possible after opening their sterile containers to minimise unnecessary exposure to the air.

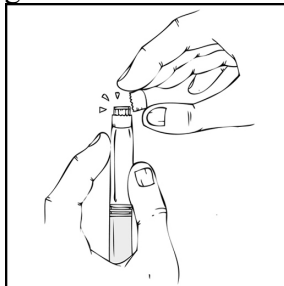
### Reconstitution

1. Allow the prefilled dual chamber syringe of freeze-dried XYNTHA to reach room temperature.
2. Remove the contents of the XYNTHA prefilled dual chamber syringe kit and place on a clean surface, making sure you have all the supplies you will need.
3. Grasp the plunger rod as shown in the following diagram. Avoid contact with the shaft of the plunger rod. Screw the plunger rod firmly into the opening in the finger rest of the XYNTHA prefilled dual chamber syringe by pushing and turning firmly until resistance is felt (approximately 2 turns).

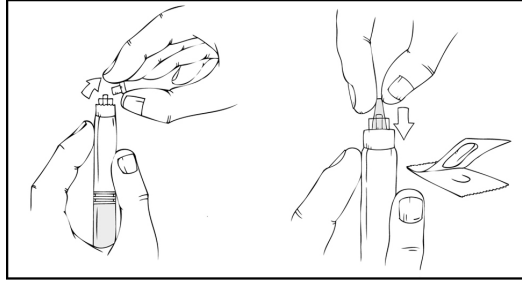


**Note:** Throughout the reconstitution process it is important to keep the XYNTHA prefilled dual chamber syringe upright (with the white powder above the clear solution) to prevent possible leakage.

4. Holding the XYNTHA prefilled dual chamber syringe upright, remove the white tamper-evident seal by bending the seal right to left (or a gentle rocking motion) to break the perforation of the cap and expose the grey rubber tip cap of the XYNTHA prefilled dual chamber syringe.

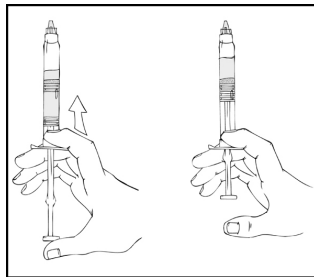


5. Remove the protective blue vented sterile cap from its package. While holding the XYNTHA prefilled dual chamber syringe upright, remove the grey rubber tip cap and replace it with the protective blue vented cap (prevents pressure build-up). Avoid touching the open end of both the syringe and the protective blue vented cap.

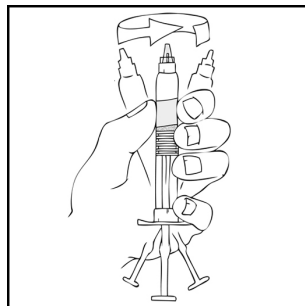


6. **Gently and slowly** advance the plunger rod by pushing until the two stoppers inside the XYNTHA prefilled dual chamber syringe meet, and all the diluent is transferred to the chamber containing the XYNTHA powder.

**Note:** To prevent the escape of fluid from the tip of the syringe, the plunger rod should not be pushed with excessive force.



7. With the XYNTHA prefilled dual chamber syringe remaining upright, swirl **gently** several times until the powder is dissolved.



**Note:** Look at the final solution in the XYNTHA prefilled dual chamber syringe. The solution should be clear to slightly opalescent and colourless. If it is not, the solution should be discarded and a new kit should be used.

8. Holding the XYNTHA prefilled dual chamber syringe in an upright position, slowly advance the plunger rod until most, but not all, of the air is removed from the drug product chamber.

XYNTHA should be administered within 3 hours after reconstitution or removal of the grey tip from the prefilled dual-chamber syringe. If you are not going to administer the XYNTHA

solution immediately, you should store the prefilled dual chamber syringe at 2°C to 8°C in an upright position, with the protective blue vented cap on the syringe until you are ready to administer it.

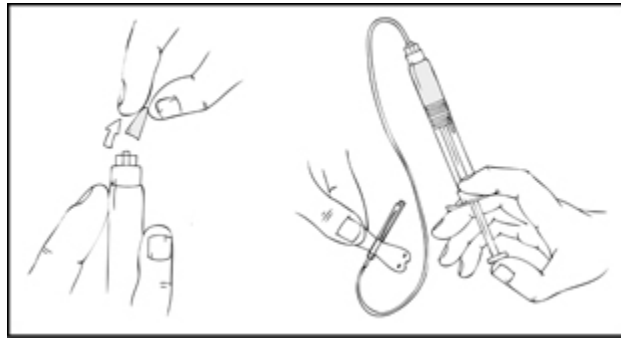
**If you have not used it in 3 hours, throw it away.**

### **Administration (Intravenous Injection)**

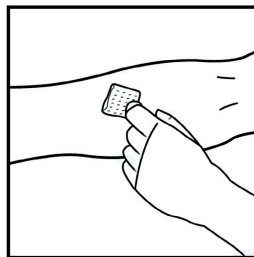
Your doctor or healthcare professional should teach you how to infuse XYNTHA. Once you learn how to self-infuse, you can follow the instructions in this insert.

**Administer XYNTHA, as provided in the prefilled dual chamber syringe by intravenous (IV) injection using the infusion set provided in the kit, unless otherwise advised by your doctor or healthcare professional.**

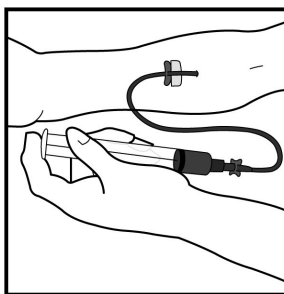
1. Remove the protective blue vented cap and firmly attach the intravenous infusion set provided in the kit onto the XYNTHA prefilled dual chamber syringe.



2. Apply a tourniquet and prepare the injection site by wiping the skin well with an alcohol swab provided in the kit.



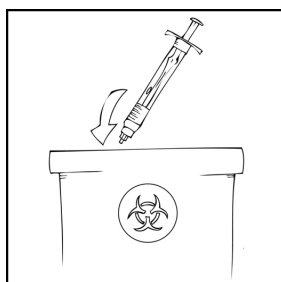
3. Remove the protective needle cover and insert the butterfly needle of the infusion set tubing into your vein as instructed by your doctor or healthcare professional. Remove the tourniquet. Infuse the reconstituted XYNTHA product over several minutes. Your doctor may change your recommended infusion rate to make the infusion more comfortable.



**Reconstituted XYNTHA must not be administered in the same tubing or container with other medicinal products.**

4. After infusing XYNTHA, remove the infusion set and discard. The amount of drug product left in the infusion set will not affect your treatment.

**Note:** Dispose of all unused solution, the empty prefilled dual chamber syringe and other used medical supplies in an appropriate container for medical waste. Medicines must not be disposed of down the toilet or via household waste. These measures help protect other people and the environment.



It is recommended to record the lot number from the XYNTHA prefilled dual chamber syringe label every time you use XYNTHA. You can use the peel-off label found on the XYNTHA prefilled dual chamber syringe to record the lot number.

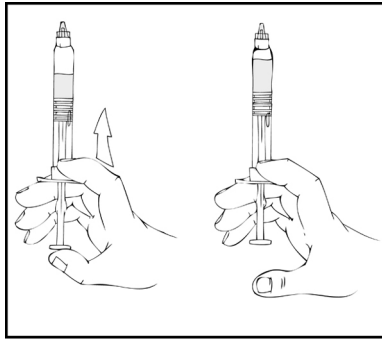
#### **Additional Instructions:**

#### **Multiple XYNTHA Prefilled Dual Chamber Syringe Reconstitution to a 10 mL or Larger Luer Lock syringe (10 mL or larger luer lock syringes not provided)**

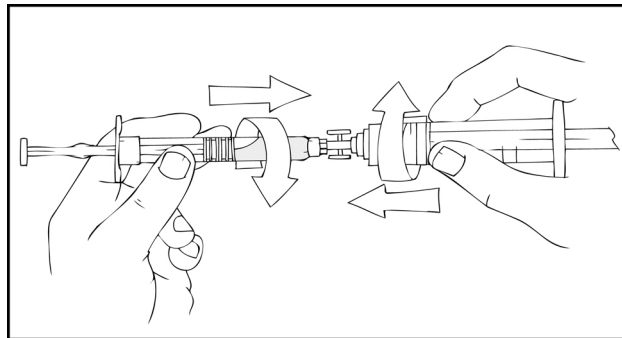
The instructions below are for the use of multiple XYNTHA prefilled dual chamber syringe kits with a 10 mL or larger luer lock syringe.

1. Reconstitute all XYNTHA prefilled dual chamber syringes according to instructions shown above in the reconstitution directions (see Reconstitution and Administration).

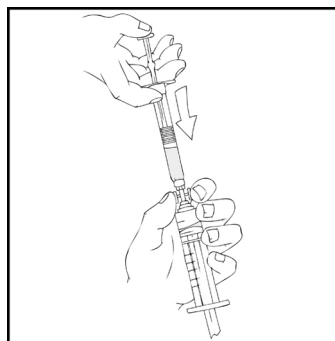
Holding the XYNTHA prefilled dual chamber syringe in an upright position, slowly advance the plunger rod until most, but not all, of the air is removed from the drug product chamber.



2. Remove the luer-to-luer syringe connector from its package (luer-to-luer syringe connectors are not provided).
3. After removing the protective blue vented cap, connect a sterile 10 mL or larger luer lock syringe to one opening (port) in the syringe connector and the XYNTHA prefilled dual chamber syringe to the remaining open port on the opposite end.

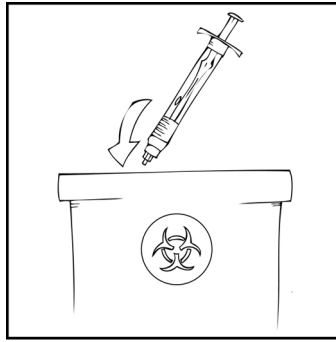


4. With the XYNTHA prefilled dual chamber syringe on top, slowly depress the plunger rod until the contents empty into the 10 mL or larger luer lock syringe.



5. Remove the empty XYNTHA prefilled dual chamber syringe and repeat procedures 3 and 4 above for any additional reconstituted syringes.
6. Remove the luer-to-luer syringe connector from the 10 mL or larger luer lock syringe and attach the infusion set, as described above in the directions for administration of the prefilled dual chamber syringe [see administration (Intravenous Injection)].

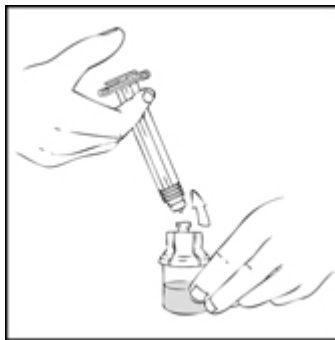
**Note:** Dispose of all unused solution, the empty prefilled dual chamber syringe, and the used medical supplies in an appropriate container for medical waste.



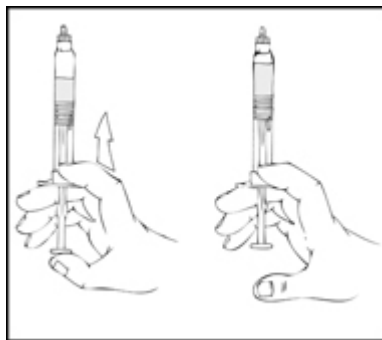
### **Use of a XYNTHA Vial Kit and a XYNTHA Prefilled Dual Chamber Syringe Kit**

These instructions are for the use of only one XYNTHA vial kit and one XYNTHA prefilled dual chamber syringe kit.

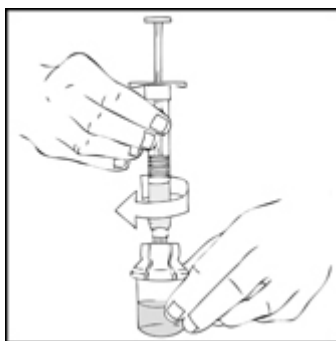
1. Reconstitute the XYNTHA vial using the instructions included with the kit. Detach the empty diluent syringe from the vial adapter by gently turning and pulling the syringe counterclockwise, leaving the contents in the vial and the vial adapter in place.



2. Reconstitute the XYNTHA prefilled dual chamber syringe using the instructions included with the kit, remembering to remove most, but not all, of the air from the drug product chamber.



3. After removing the protective blue vented cap, connect the XYNTHA prefilled dual chamber syringe to the vial adapter by inserting the tip into the adapter opening while firmly pushing and turning the syringe clockwise until secured.



4. Slowly depress the plunger rod of the XYNTHA prefilled dual chamber syringe until the contents empty into the XYNTHA vial. The plunger rod may move back slightly after release.

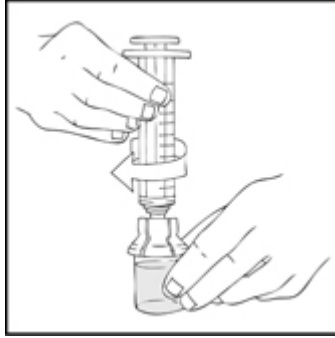


5. Detach and discard the empty XYNTHA prefilled dual chamber syringe from the vial adapter.

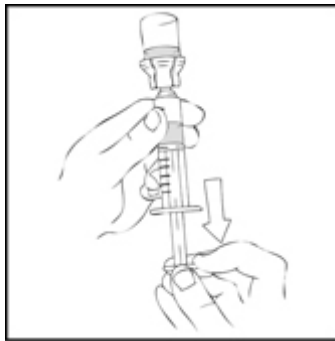
**Note:** If the syringe turns without detaching from the vial adapter, grasp the white collar and turn.



6. Connect a sterile 10 mL or larger luer lock syringe to the vial adapter. You may want to inject some air into the vial to make withdrawing the vial contents easier.



7. Invert the vial and slowly draw the solution into the 10 mL or larger luer lock syringe.



8. Detach the syringe from the vial adapter by gently turning and pulling the syringe counterclockwise. Discard the vial with the adapter attached.
9. Attach the infusion set to the 10 mL or larger luer lock syringe as described above in the directions for administration of the prefilled dual chamber syringe [see administration (Intravenous Injection)].

**Note:** Dispose of all unused solution, the empty prefilled dual chamber syringe, and the used medical supplies in an appropriate container for medical waste.