## HIGHLIGHTS OF PRESCRIBING INFORMATION

These highlights do not include all the information needed to use AJOVY safely and effectively. See full prescribing information for AJOVY.

### AJOVY® (fremanezumab-vfrm) injection, for subcutaneous use

**INDICATIONS AND USAGE**

AJOVY is a calcitonin gene-related peptide antagonist indicated for the preventive treatment of migraine in adults.

### DOSAGE AND ADMINISTRATION

1. **Subcutaneous Administration**

   - **For subcutaneous use only**: (2.1, 2.2)
   - **Recommended Dosing**
     - Two subcutaneous dosing options of AJOVY are available to administer the recommended dosage:
       - 225 mg monthly, or
       - 675 mg every 3 months (quarterly).
   - **Dosage Regimen of AJOVY and At Least 2% Greater Than Placebo in Studies 1 and 2**
     - 225 mg Monthly
     - Placebo
     - %
     - AJOVY
     - Placebo

<table>
<thead>
<tr>
<th>Adverse Reaction</th>
<th>AJOVY (n=667)</th>
<th>Placebo (n=663)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Nausea</td>
<td>32%</td>
<td>26%</td>
</tr>
<tr>
<td>Headache</td>
<td>30%</td>
<td>26%</td>
</tr>
<tr>
<td>Dizziness</td>
<td>28%</td>
<td>24%</td>
</tr>
<tr>
<td>Fatigue</td>
<td>28%</td>
<td>28%</td>
</tr>
<tr>
<td>Constipation</td>
<td>24%</td>
<td>26%</td>
</tr>
</tbody>
</table>

2. **Loading Dose**

   - Administer the first loading dose of 675 mg as a single injection.
   - If a second loading dose is required, it may be administered at any time after the first dose.
   - Thereafter, AJOVY can be scheduled for the recommended dosing:

### CONTRAINDICATIONS

- AJOVY is contraindicated in patients with serious hypersensitivity to fremanezumab-vfrm or to any of the excipients.

### WARNINGS AND PRECAUTIONS

- **Hypersensitivity Reactions**: If hypersensitivity occurs, consider discontinuing AJOVY and institute appropriate therapy.

### ADVERSE REACTIONS

The most common adverse reactions (≥5% and greater than placebo) were injection site reactions.

- To report SUSPECTED ADVERSE REACTIONS, contact Teva Pharmaceuticals at 1-888-483-8279 or FDA at 1-800-FDA-1088 or www.fda.gov/medwatch.

### PATIENT COUNSELING INFORMATION

- Additional information is available at www.migraine.com.

### DOSAGE FORMS AND STRENGTHS

- AJOVY Injection: 225 mg/1.5 mL solution in a single-dose prefilled autoinjector.
- AJOVY Injection: 225 mg/1.5 mL solution in a single-dose prefilled syringe.

### ADVERSE REACTIONS

The following clinically significant adverse reactions are discussed in greater detail in other sections of the labeling:

- Hypersensitivity Reactions
  - [see Warnings and Precautions (5.1)]

### CLINICAL STUDIES

- Clinical Trials Experience

### HOW SUPPLIED/STORAGE AND HANDLING

- **Injection**:
  - 225 mg monthly or AJOVY 675 mg quarterly for at least 6 months, 775 patients for at least 12 months, and 138 patients for at least 15 months.
  - Placebo-controlled clinical trials (Studies 1 and 2), 662 patients received AJOVY 225 mg monthly for 12 weeks (with or without a loading dose of 675 mg), and 663 patients received AJOVY 675 mg quarterly for 12 weeks.

### CLINICAL PHARMACOLOGY

- **Immunogenicity**

### PATIENT COUNSELING INFORMATION

- Injection site reactions include multiple related adverse event terms, such as injection site pain, induration, and erythema.

### IMMUNOGENICITY

As with all therapeutic proteins, there is a potential for immunogenicity. The detection of antibody formation is highly dependent on sensitivity and specificity of the assay. Additionally, the observed incidence of antibody (including neutralizing antibody) positivity in an assay may be influenced by several factors, including assay methodology, sample handling, timing of sample collection, concomitant medications, and underlying disease. For these reasons, comparison of the incidence of antibodies to fremanezumab-vfrm in the studies described below with the incidence of antibodies to other products may be misleading.
Clinical immunogenicity of AJOVY was monitored by analyzing anti-drug antibodies (ADA) and neutralizing antibodies in drug-treated patients. The data reflect the percentage of patients whose test results were positive for antibodies to AJOVY in specific assays. In 3-month placebo-controlled studies, treatment-emergent ADA responses were observed in 6 out of 1701 (0.4%) AJOVY-treated patients. One of the 6 patients developed anti-AJOVY neutralizing antibodies at Day 84. In the ongoing long-term open-label study, ADA were detected in 16% of patients (30 out of 1888). Out of 30 ADA-positive patients, 17 had a neutralizing activity in their post-dose samples. Although these data do not demonstrate an impact of anti-fremazemab-vfrm antibody development on the efficacy or safety of AJOVY in these patients, the available data are too limited to make definitive conclusions.

8 USE IN SPECIFIC POPULATIONS

8.1 Pregnancy

Risk Summary

There are no adequate data on the developmental risk associated with the use of AJOVY in pregnant women. AJOVY has a long half-life [see Clinical Pharmacology (12.3)]. This should be taken into consideration for women who are pregnant or plan to become pregnant while using AJOVY. Administration of fremazemab-vfrm to rats and rabbits during the period of organogenesis or to rats throughout pregnancy and lactation at doses resulting in plasma levels greater than those expected clinically did not result in adverse effects on development [see Animal Data]. In the U.S. general population, the estimated background risk of major birth defects and miscarriage in clinically recognized pregnancies is 2-4% and 15-20%, respectively. The estimated rate of major birth defects (2.2-2.9%) and miscarriage (17%) among deliveries to women with migraine is similar to rates reported in women without migraine.

Clinical Considerations

12.1 Mechanism of Action

Fremazemab-vfrm is a fully humanized IgG2a/kappa monoclonal antibody specific for calcitonin gene-related peptide (CGRP) ligand. Fremazemab-vfrm is produced by recombinant DNA technology in Chinese hamster ovary (CHO) cells. The antibody consists of 1224 amino acids and has a molecular weight of approximately 148 kDa. When fremazemab-vfrm (0, 50, 100, or 200 mg/kg) was administered to male and female rats by weekly subcutaneous injection prior to and during mating and continuing in females throughout organogenesis, no adverse embryofetal effects were observed. The highest dose tested was associated with plasma exposures (AUC) approximately 2 times that in humans at a dose of 675 mg. Administration of fremazemab-vfrm (0, 10, 50, or 100 mg/kg) weekly by subcutaneous injection to pregnant rabbits throughout the period of organogenesis produced no adverse effects on embryofetal development. The highest dose tested was associated with plasma AUC approximately 3 times that in humans (675 mg). Administration of fremazemab-vfrm (0, 50, 100, or 200 mg/kg) weekly by subcutaneous injection to female rats throughout pregnancy and lactation resulted in no adverse effects on pre- and postnatal development. The highest dose tested was associated with plasma AUC approximately 2 times that in humans (675 mg).

8.2 Lactation

Risk Summary

There are no data on the presence of fremazemab-vfrm in human milk, the effects on the breastfed infant, or the effects on milk production. The developmental and health benefits of breastfeeding should be considered along with the mother’s clinical need for AJOVY and any potential adverse effects on the breastfed infant from AJOVY or from the underlying maternal condition.

8.4 Pediatric Use

Safety and effectiveness in pediatric patients have not been established.

8.5 Geriatric Use

Clinical studies of AJOVY did not include sufficient numbers of subjects aged 65 and over to determine whether they respond differently from younger subjects.

11 DESCRIPTION

Fremazemab-vfrm is a fully humanized IgG2a/kappa monoclonal antibody specific for calcitonin gene-related peptide (CGRP) ligand. Fremazemab-vfrm is produced by recombinant DNA technology in Chinese hamster ovary (CHO) cells. The antibody consists of 1224 amino acids and has a molecular weight of approximately 148 kDa. AJOVY (fremazemab-vfrm) injection is a sterile, preservative-free, clear to opalescent, colorless to slightly yellow solution for subcutaneous injection, supplied in a single-dose 225 mg/1.5 mL prefilled autoinjector and a single-dose 225 mg/1.5 mL prefilled syringe. Each prefilled autoinjector delivers 15 mL of solution containing 225 mg fremazemab-vfrm, disodium ethylenediaminetetraacetic acid dihydrate (EDTA) (0.204 mg), L-histidine (0.815 mg), L-histidine hydrochloride monohydrate (3.93 mg), polysorbate-80 (0.3 mg), sucrose (99 mg), and Water for Injection, and has a pH of 5.5.

12 CLINICAL PHARMACOLOGY

12.1 Mechanism of Action

Fremazemab-vfrm is a humanized monoclonal antibody that binds to calcitonin gene-related peptide (CGRP) ligand and blocks its binding to the receptor.

12.2 Pharmacodynamics

The relationship between the pharmacodynamic activity and the mechanism(s) by which fremazemab-vfrm exerts its clinical effects is unknown.

12.3 Pharmacokinetics

Absorption

After single subcutaneous (SC) administrations of 225 mg, 675 mg, and 900 mg fremazemab-vfrm, median time to maximum concentrations (tmax) was 5 to 7 days. Dose-proportionality, based on post-dose plasma PK, was observed between 225 mg to 900 mg. Steady state was achieved by approximately 168 days (about 6 months) following 225 mg SC monthly and 675 mg SC quarterly dosing regimens. Median accumulation ratio, based on once-monthly and once-quarterly dosing regimens, is approximately 2.3 and 1.2, respectively.

Distribution

Fremazemab-vfrm has an apparent volume of distribution of approximately 6 liters, suggesting minimal distribution to the extravascular tissues.

Metabolism

Similar to other monoclonal antibodies, fremazemab-vfrm is degraded by enzymatic proteolysis into small peptides and amino acids.

Elimination

Fremazemab-vfrm apparent clearance was approximately 0.141 L/day. Fremazemab-vfrm was estimated to have a half-life of approximately 31 days.

Specific Populations

A population PK analysis assessing effects of age, race, sex, and weight was conducted on data from 2287 subjects. No dose adjustments are recommended for AJOVY.

9 SAFETY

9.1 General Adverse Reactions

Adverse reactions were assessed as mild, moderate, or severe. Adverse reactions are summarized in Table 2.

<table>
<thead>
<tr>
<th>Study 1 Efficacy Endpoint</th>
<th>AJOVY 225 mg Monthly (N=297)</th>
<th>AJOVY 675 mg Quarterly (N=298)</th>
<th>Placebo (N=290)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Monthly migraine days (MMD)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Baseline migraine days</td>
<td>8.9</td>
<td>9.2</td>
<td>9.1</td>
</tr>
<tr>
<td>Change from baseline</td>
<td>-3.7</td>
<td>-3.4</td>
<td>-2.2</td>
</tr>
<tr>
<td>Difference from placebo</td>
<td>-1.5</td>
<td>-1.2</td>
<td></td>
</tr>
<tr>
<td>p-value</td>
<td>&lt;0.001</td>
<td>&lt;0.001</td>
<td></td>
</tr>
<tr>
<td>≤50% MDD responders</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>% responders</td>
<td>42.7%</td>
<td>44.4%</td>
<td>27.9%</td>
</tr>
<tr>
<td>Difference from placebo</td>
<td>19.8%</td>
<td>16.5%</td>
<td></td>
</tr>
<tr>
<td>p-value</td>
<td>&lt;0.001</td>
<td>&lt;0.001</td>
<td></td>
</tr>
</tbody>
</table>

continued
Figure 1 displays the mean change from baseline in the average monthly number of migraine days in Study 1.

Figure 1: Change from Baseline in Monthly Migraine Days in Study 1

![Change from Baseline in Monthly Migraine Days in Study 1](image)

Table 1: Efficacy Endpoint

<table>
<thead>
<tr>
<th>Study 1</th>
<th>Efficacy Endpoint</th>
<th>AJOVY 225 mg Monthly (N=377)</th>
<th>AJOVY 675 mg Quarterly (N=288)</th>
<th>Placebo (N=290)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Monthly acute headache medication days</td>
<td>Change from baseline</td>
<td>-3.0</td>
<td>-2.9</td>
<td>-1.6</td>
</tr>
<tr>
<td>Difference from placebo</td>
<td>-1.4</td>
<td>-1.3</td>
<td>-0.3</td>
<td></td>
</tr>
<tr>
<td>p-value</td>
<td>&lt;0.001</td>
<td>&lt;0.001</td>
<td>&lt;0.001</td>
<td></td>
</tr>
</tbody>
</table>

Figure 2 shows the distribution of change from baseline in monthly migraine days in Study 1 for both AJOVY and placebo groups.

Figure 2: Distribution of Change from Baseline in Monthly Migraine Days by Treatment Group in Study 1

![Distribution of Change from Baseline in Monthly Migraine Days by Treatment Group in Study 1](image)

Chronic Migraine

Study 2 (NCT 02621931) included adults with a history of chronic migraine. All patients were randomized (1:1:1) to receive subcutaneous injections of either AJOVY 675 mg starting dose followed by 225 mg monthly, 675 mg every 3 months (quarterly), or placebo monthly, over a 3-month treatment period. Patients were allowed to use acute headache treatments during the study. A subset of patients (21%) was allowed to use one additional concomitant, preventive medication.

The secondary endpoints were the mean change from baseline in the monthly average number of headache days of at least moderate severity at 4 weeks after the first dose. The mean change from baseline in monthly average number of headache days of at least moderate severity at month 3 in bins of 3 days by treatment group. A treatment benefit over placebo for both dosing regimens of AJOVY is seen across a range of changes from baseline in monthly migraine days.

Figure 3 displays the mean change from baseline in the average monthly number of headache days of at least moderate severity in Study 1 for both AJOVY and placebo groups.

Figure 3: Change from Baseline in Monthly Migraine Days in Study 1

![Change from Baseline in Monthly Migraine Days in Study 1](image)

Table 2: Efficacy Endpoints in Study 2

<table>
<thead>
<tr>
<th>Study 2</th>
<th>Efficacy Endpoint</th>
<th>AJOVY 225 mg* Monthly (N=375)</th>
<th>AJOVY 675 mg Quarterly (N=375)</th>
<th>Placebo (N=371)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Baseline headache days of any severity*</td>
<td>20.3</td>
<td>20.4</td>
<td>20.3</td>
<td></td>
</tr>
<tr>
<td>Baseline headache days of at least moderate severity*</td>
<td>12.8</td>
<td>13.2</td>
<td>13.3</td>
<td></td>
</tr>
<tr>
<td>Change from baseline in the monthly average number of headache days of at least moderate severity</td>
<td>-4.6</td>
<td>-4.3</td>
<td>-2.5</td>
<td></td>
</tr>
<tr>
<td>Difference from placebo</td>
<td>-2.1</td>
<td>-1.8</td>
<td>-1.7</td>
<td></td>
</tr>
<tr>
<td>p-value</td>
<td>&lt;0.001</td>
<td>&lt;0.001</td>
<td>&lt;0.001</td>
<td></td>
</tr>
</tbody>
</table>

Figure 4 shows the distribution of change from baseline in monthly headache days of at least moderate severity at month 3 in bins of 3 days by treatment group. A treatment benefit over placebo for both dosing regimens of AJOVY is seen across a range of changes from baseline in headache days.

Figure 4: Distribution of Mean Change from Baseline in Monthly Headache Days of At Least Moderate Severity in Study 2

![Distribution of Mean Change from Baseline in Monthly Headache Days of At Least Moderate Severity in Study 2](image)
AJOVY® (fremanezumab-vfrm) injection

AJOVY® (fremanezumab-vfrm) injection is not made with natural rubber latex.

AJOVY® is supplied as follows:

**Prefilled Autoinjector**
- Pack of 1 autoinjector: 225 mg/1.5 mL single-dose prefilled autoinjector NDC 57579-202-10
- Pack of 3 autoinjectors: 3 x 225 mg/1.5 mL single-dose prefilled autoinjectors NDC 57579-202-22

**Prefilled Syringe**
- Pack of 1 syringe: 225 mg/1.5 mL single-dose prefilled syringe NDC 57579-204-10

**Storage and Handling**
- Store refrigerated at 2°C to 8°C (36°F to 46°F) in the original outer carton to protect from light.
- If necessary, AJOVY may be kept in the original carton at room temperature up to 30°C (86°F) for a maximum of 7 days. After removal from the refrigerator, AJOVY must be used within 7 days or discarded. Once stored at room temperature, do not place back in the refrigerator.
- Do not freeze.
- Do not expose to extreme heat or direct sunlight.
- Do not shake.

## 17 PATIENT COUNSELING INFORMATION

Advises the patient and/or caregiver to read the FDA-approved patient labeling (Patient Information and Instructions for Use). Information on Preparation and Administration

Provide guidance to patients and caregivers on proper subcutaneous administration technique, including aseptic technique, and how to use the single-dose prefilled syringe [see Dosage and Administration (2.2)]. Instruct patients and/or caregivers to read and follow the Instructions for Use each time they use AJOVY.

Instruct patients prescribed the regimen of 675 mg every 3 months to administer the dosage as follows:
- If your prescribed dose is AJOVY 225 mg one time every month or AJOVY 675 mg one time every 3 months.
- If your prescribed dose is AJOVY 675 mg every 3 months, you must use 3 separate autoinjectors or 3 separate syringes.

Inform patients about the signs and symptoms of hypersensitivity reactions and that these reactions can occur up to 1 month after administration. Advise patients to contact their healthcare provider immediately if signs or symptoms of hypersensitivity reactions occur [see Warnings and Precautions (5.1)].

**Hypersensitivity Reactions**

Inform patients about the signs and symptoms of hypersensitivity reactions and that these reactions can occur up to 1 month after administration. Advise patients to contact their healthcare provider immediately if signs or symptoms of hypersensitivity reactions occur [see Warnings and Precautions (5.1)].

**Manufactured by:**
Teva Pharmaceuticals USA, Inc.
North Wales, PA 19454
US License No. 2016

AJOVY® (fremanezumab-vfrm), its use, or its process of manufacture, may be protected by one or more United States patents, including US 8,007,794, US 8,586,045 and US 9,896,502.

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### AJOVY® (fremanezumab-vfrm) injection

**Patient Information**

**AJOVY® (a-JO-vey)**
(fremanezumab-vfrm) injection
for subcutaneous use

**What is AJOVY?**

AJOVY® is a prescription medicine used for the preventive treatment of migraine in adults. It is not known if AJOVY is safe and effective in children.

**Who should not use AJOVY?**

Do not use AJOVY if you are allergic to fremanezumab-vfrm or any of the ingredients in AJOVY. See the end of this leaflet for a complete list of the ingredients in AJOVY.

**Before you use AJOVY, tell your healthcare provider if you:**
- are pregnant or plan to become pregnant. It is not known if AJOVY will harm your unborn baby.
- are breastfeeding or plan to breastfeed. It is not known if AJOVY passes into your breast milk. Talk to your healthcare provider about the best way to feed your baby while using AJOVY.

**Tell your healthcare provider about all the medicines you take,** including prescription and over-the-counter medicines, vitamins, and herbal supplements. Know the medicines you take. Keep a list of your medicines with you to show your healthcare provider and pharmacist when you get a new medicine.

**How should I use AJOVY?**

- See the detailed “Instructions for Use” for information on how to prepare and inject a dose of AJOVY.
- Use AJOVY exactly as your healthcare provider tells you to use it.
- AJOVY is given by injection under your skin (subcutaneously).
- Your healthcare provider should show you or your caregiver how to prepare and inject your dose of AJOVY before you or your caregiver give your AJOVY the first time.
- Your healthcare provider will tell you how much AJOVY to use and when to use it.
  - Your healthcare provider will tell you if you should use AJOVY 225 mg one time every month or AJOVY 675 mg one time every 3 months.
  - If your prescribed dose is AJOVY 675 mg every 3 months, you must use 3 separate autoinjectors or 3 separate syringes.
  - You will give 3 separate injections one time every 3 months.
- If you are giving 3 injections of AJOVY for your prescribed dose, you may use the same injection area for all 3 injections, but not the same spot.
- Do not inject AJOVY in the same injection site that you inject other medicine.
- If you are switching from using AJOVY one time every month to one time every 3 months or if you are switching from using AJOVY one time every 3 months to one time every month, give the first dose of AJOVY on the day it was due to be given on your old schedule.
- If you miss a dose of AJOVY, take it as soon as possible. If you need to take the dose late, you will need to change your schedule: if you take 225 mg of AJOVY, inject your next dose 1 month after the late dose. If you take 675 mg of AJOVY, inject your next dose 3 months after the late dose. If you have questions about your schedule, ask your healthcare provider.

**What are the possible side effects of AJOVY?**

AJOVY may cause serious side effects, including:
- **Allergic reactions.** Allergic reactions, including itching, rash, and hives, can happen within hours and up to 1 month after receiving AJOVY. Call your healthcare provider or get emergency medical help right away if you have any of the following symptoms of an allergic reaction:
  - swelling of your face, mouth, tongue, or throat
  - trouble breathing

**The most common side effects of AJOVY include:**

- injection site reactions
  Tell your healthcare provider if you have any side effect that bothers you or that does not go away.
  These are not all the possible side effects of AJOVY. For more information, ask your healthcare provider or pharmacist. Call your doctor for medical advice about side effects. You may report side effects to FDA at 1-800-FDA-1088.
How should I store AJOVY?

- Store AJOVY in the refrigerator between 36°F to 46°F (2°C to 8°C).
- Keep AJOVY in the carton it comes in to protect from light.
- If needed, AJOVY may be stored at room temperature up to 86°F (30°C) in the carton it comes in for up to 7 days. Do not use AJOVY if it has been out of the refrigerator for 7 days or longer. Throw away (dispose of) AJOVY in a sharps disposal or puncture-resistant container if it has been out of the refrigerator for 7 days or longer. Once stored at room temperature, do not place back in the refrigerator.
- Do not freeze. If AJOVY freezes, throw it away in a sharps disposal container.
- Keep AJOVY out of extreme heat and direct sunlight.
- Do not shake AJOVY.

Keep AJOVY prefilled autoinjector and AJOVY prefilled syringe out of the reach of small children.

General information about the safe and effective use of AJOVY.

Medicines are sometimes prescribed for purposes other than those listed in a Patient Information leaflet. Do not use AJOVY for a condition for which it was not prescribed. Do not give AJOVY to other people, even if they have the same symptoms that you have. It may harm them. You can ask your pharmacist or healthcare provider for information about AJOVY that is written for health professionals.

What are the ingredients in AJOVY?

Active ingredients: fremanezumab-vfrm

Inactive ingredients: disodium ethylenediaminetetraacetic acid dihydrate (EDTA), L-histidine, L-histidine hydrochloride monohydrate, polysorbate-80, sucrose, and Water for Injection.

AJOVY prefilled syringe and prefilled autoinjector are not made with natural rubber latex.

Manufactured by: Teva Pharmaceuticals USA, Inc., North Wales, PA 19454

US License No. 2016

AJ0PL-04

For more information, go to www.AJOVY.com or call 1-888-483-8279.

This Patient Information has been approved by the U.S. Food and Drug Administration. Revised: 10/2020

Instructions for Use

AJOVY® (a-JO-vee)

(fremanezumab-vfrm) injection

prefilled autoinjector, for subcutaneous use

For subcutaneous injection only.

Read and follow the Instructions for Use for your AJOVY prefilled autoinjector before you start using it and each time you get a refill.

Important:

- AJOVY prefilled autoinjector is for single-time (one-time) use only. Put AJOVY in a FDA-cleared sharps disposal or puncture-resistant container right away after use. Do not throw away (dispose of) your used sharps disposal container in your household trash.
- Before injecting, let AJOVY sit at room temperature for 30 minutes.
- Keep AJOVY prefilled autoinjector out of the reach of small children.
- After you remove the protective cap from AJOVY, to prevent infection, do not touch the needle.
- Do not inject AJOVY in your veins (intravenously).
- Do not re-use your AJOVY prefilled autoinjector as this could cause injury or infection.
- Do not share your AJOVY prefilled autoinjector with another person. You may give another person an infection or get an infection from them. You may give AJOVY yourself. If you feel uncomfortable, you should not get your first dose of AJOVY until you or your caregiver receive training from a healthcare provider on the right way to use AJOVY.

Storage Conditions:

- Store AJOVY in the refrigerator between 36°F to 46°F (2°C to 8°C).
- Keep AJOVY in the carton it comes in to protect from light.
- If needed, AJOVY may be stored at room temperature up to 86°F (30°C) in the carton it comes in for up to 7 days. Do not use AJOVY if it has been out of the refrigerator for 7 days or longer. Throw away (dispose of) AJOVY in a sharps disposal or puncture-resistant container if it has been out of the refrigerator for 7 days or longer. Once stored at room temperature, do not place back in the refrigerator.
- Do not freeze. If AJOVY freezes, throw it away in a sharps disposal container.
- Keep AJOVY out of extreme heat and direct sunlight.

Do not shake AJOVY.

AJOVY prefilled autoinjector (Before use). See Figure A.

AJOVY prefilled autoinjector (After use). See Figure B.

- The blue plunger moves down the viewing window during the injection. The blue plunger fills the window when the injection is complete. Note: When the blue plunger has filled the viewing window you will still be able to see the gray stopper, as shown in Figure B.
- When injecting AJOVY, hold the prefilled autoinjector so that your hand does not cover the viewing window.

Read this before you inject.

Step 1. Check the dose your healthcare provider has prescribed.

AJOVY comes as a single-dose (one time) prefilled autoinjector. Your healthcare provider will prescribe the dose that is best for you.

- If your healthcare provider has prescribed 225 mg of AJOVY each month for you, give 1 injection each month, using a 225 mg prefilled AJOVY autoinjector.
- If your healthcare provider has prescribed 675 mg of AJOVY every 3 months for you, give 3 separate injections, one after another, using a different 225 mg prefilled AJOVY autoinjector for each injection. Give these injections 1 time every 3 months.

Before you inject, always check the label of your single-dose prefilled autoinjector to make sure you have the correct medicine and the correct dose of AJOVY. If you are not sure of your dose, ask your healthcare provider.

How do I inject AJOVY?

Step 2. Remove the prefilled autoinjector from the carton.

- You may need to use more than 1 prefilled autoinjector depending on your prescribed dose.
- Remove the autoinjector from the carton (see Figure C).
- Do not shake the prefilled autoinjector at any time, as this could affect the way the medicine works.

Important: If there are any unused autoinjectors left in the carton, put the carton and unused autoinjectors back in the refrigerator.
AJOVY® (fremanezumab-vfrm) injection

Step 3. Gather the supplies you will need to inject AJOVY.
- Gather the following supplies (see Figure D) and the number of AJOVY 225 mg prefilled autoinjectors you will need to give your prescribed dose:
  - If your dose is 225 mg, you will need 1 AJOVY 225 mg prefilled autoinjector.
  - If your dose is 675 mg, you will need 3 AJOVY 225 mg prefilled autoinjectors.
- Alcohol swabs (not supplied).
- Gauze pads or cotton balls (not supplied).
- Sharps disposal or puncture-resistant container (not supplied).

Step 4. Let AJOVY reach room temperature.
- Place the supplies you have gathered on a clean, flat surface.
- Wait for 30 minutes to allow the medicine to reach room temperature.
- Do not leave the prefilled autoinjector in direct sunlight.
- Do not warm up the AJOVY prefilled autoinjector using a heat source such as hot water or a microwave.

Step 5. Wash your hands.
- Wash your hands with soap and water and dry well with a clean towel. Be careful not to touch your face or hair after washing your hands.

Step 6. Look closely at your AJOVY prefilled autoinjector.
Note: You may see air bubbles in the prefilled autoinjector. This is normal. Do not remove the air bubbles from the prefilled autoinjector before giving your injection. Injecting AJOVY with these air bubbles will not harm you.
- Check that the liquid medicine in the prefilled autoinjector viewing window is clear and colorless and slightly yellow before you give your injection. (See Figure E). If the liquid has any particles in it, or is discolored, cloudy, or frozen, do not use the prefilled autoinjector. Call your healthcare provider or pharmacist.
- Check that AJOVY appears on the prefilled autoinjector.
- Check the expiration date (EXP) printed on the prefilled autoinjector label.
- Do not use the prefilled autoinjector if it has any visible damage, such as cracks or leaks. See disposal instructions in Step 12.
- Do not use the prefilled autoinjector if it has the expiration date (EXP) has passed.
- Do not use if you have been given the wrong medicine.

Step 7: Choose your injection area.
- Choose an injection area from the following areas (see Figure F):
  - your stomach area (abdomen), avoid about 2 inches around the belly button.
  - the front of your thighs, an area that is at least 2 inches above the knee and 2 inches below the groin.
  - the back of your upper arms, in the fleshy area of the upper back portion.

Note: There are some injection areas on your body that are hard to reach (like the back of your arm). You may need help from someone who has been instructed on how to give your injection if you cannot reach certain injection areas.

Step 8. Clean your injection area.
- Clean the chosen injection area using a new alcohol swab. Let your skin dry.
- Do not inject AJOVY into an area that is tender, red, bruised, callused, tattooed, hard, or that has scars or stretch marks.
- Do not inject AJOVY in the same injection site that you inject other medicine.
- If you want to use the same injection area for the 3 separate injections needed for the 675 mg dose, make sure the second and third injections are not at the same spot you used for the other injections.

Step 9. Remove protective cap and do not replace.
- Pick up the prefilled autoinjector in 1 hand.
- Hold the prefilled autoinjector as shown in Figure G and pull the protective cap straight off with your other hand. Do not twist.
- Throw away the protective cap right away.
- Do not put the protective cap back on the prefilled autoinjector, to avoid injury and infection.

Step 10. Give your injection.
- 10.1 Place the prefilled autoinjector at a 90 degree angle against your skin at the injection site you have cleaned (see Figure H).
AJOVY® (fremanezumab-vfrm) injection

Instructions for Use

AJOVY® (a-JO-vee) (fremanezumab-vfrm) injection
prefilled syringe, for subcutaneous use

For subcutaneous injection only.

Read and follow the Instructions for Use for your AJOVY prefilled syringe before you start using it and each time you get a refill.

Important:

- AJOVY prefilled syringe is for single-time (one-time) use only. Put AJOVY in a FDA-cleared sharps disposal or puncture-resistant container right away after use. Do not throw away (dispose of) your used sharps disposal container in your household trash.
- Before injecting, let AJOVY sit at room temperature for 30 minutes.
- Keep AJOVY prefilled syringe out of the reach of small children.
- After you remove the needle cap from AJOVY, to prevent infection, do not touch the needle.
- Do not inject AJOVY in your veins (intravenously).
- Do not re-use your AJOVY prefilled syringe, as this could cause injury or infection.
- Do not share your AJOVY prefilled syringe with another person. You may give another person an infection or get an infection from them.

You may give AJOVY yourself. If you feel uncomfortable, you should not get your first dose of AJOVY until you or your caregiver receive training from a healthcare provider on the right way to use AJOVY.

Storage Conditions:

- Store AJOVY in the refrigerator between 36°F to 46°F (2°C to 8°C).
- Keep AJOVY in the carton it comes in to protect from light.
- If needed, AJOVY may be stored at room temperature up to 86°F (30°C) in the carton it comes in for up to 7 days. Do not use AJOVY if it has been out of the refrigerator for 7 days or longer. Throw away (dispose of) AJOVY in a sharps disposal or puncture-resistant container if it has been out of the refrigerator for 7 days or longer. Once stored at room temperature, do not place back in the refrigerator.
- Do not freeze. If AJOVY freezes, throw it away in a sharps disposal container.
- Keep AJOVY out of extreme heat and direct sunlight.
- Do not shake AJOVY.

10.2 Press down on the prefilled autoinjector and keep holding it down against the skin for about 30 seconds. Do not remove pressure until the 3 steps below are complete.

1. You hear the first “click” (this means the injection has started and the blue plunger starts to move).

2. You hear a second “click” (about 15 seconds after the first click. The plunger will be moving to the bottom of the viewing window as the medicine is being injected.)

3. You wait another 10 seconds (to make sure all the medicine is injected).

10.3 Check that the blue plunger has filled the viewing window and remove the autoinjector from the skin by lifting the prefilled autoinjector straight up (see Figure I).

Note: When the blue plunger has filled the viewing window you will be able to see the gray stopper.

As the prefilled autoinjector is lifted from the skin, the needle shield returns to the original (before use) position and locks into place, covering the needle.

Do not try to put the protective cap back on the used prefilled autoinjector as it is no longer needed.

Do not try to re-use the prefilled autoinjector.

Step 12. Dispose of your prefilled autoinjector right away.

- Put your used prefilled autoinjectors in a FDA-cleared sharps disposal container right away after use.
- Do not throw away (dispose of) prefilled autoinjectors in your household trash. Do not recycle your used sharps disposal container.
- If you do not have a FDA-cleared sharps disposal container, you may use a household container that is:
  - made of a heavy-duty plastic,
  - can be closed with a tight-fitting, puncture-resistant lid, without sharps being able to come out,
  - upright and stable during use,
  - leak-resistant, and
  - properly labeled to warn of hazardous waste inside the container.

- When your sharps disposal container is almost full, you will need to follow your community guidelines for the right way to dispose of your sharps disposal container. There may be state or local laws about how you should throw away used autoinjectors. For more information about safe sharps disposal, and for specific information about sharps disposal in the state that you live in, go to the FDA’s website at: http://www.fda.gov/safesharpsdisposal
- Do not dispose of your used sharps disposal container in your household trash unless your community guidelines permit this. Do not recycle your used sharps disposal container.

Injection Complete

This Instructions for Use has been approved by the U.S. Food and Drug Administration.

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AJOIFU-AI-004
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Figure I

Step 11. Apply pressure to the injection site.
- Use a clean, dry cotton ball, or gauze pad to gently press on the injection site for a few seconds.
- Do not rub the injection site.
- Do not re-use the prefilled autoinjector.

Storage Conditions:

- Keep AJOVY in the carton it comes in to protect from light.
- Store AJOVY in the refrigerator between 36°F to 46°F (2°C to 8°C).
- After you remove the needle cap from AJOVY, to prevent infection, do not touch the needle.
- Do not inject AJOVY in your veins (intravenously).
- Do not re-use your AJOVY prefilled syringe, as this could cause injury or infection.
- Do not share your AJOVY prefilled syringe with another person. You may give another person an infection or get an infection from them.

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Step 1. Check the dose your healthcare provider has prescribed.

AJOVY comes as a single-dose (one time) prefilled syringe. Your healthcare provider will prescribe the dose that is best for you.

- If your healthcare provider has prescribed 225 mg of AJOVY each month for you, give 1 injection each month using a 225 mg prefilled AJOVY syringe.
- If your healthcare provider has prescribed 675 mg of AJOVY every 3 months for you, give 3 separate injections, one after another, using a different 225 mg prefilled AJOVY syringe for each injection. Give these injections 1 time every 3 months.

Before you inject, always check the label of your single-dose prefilled syringe to make sure you have the correct medicine and the correct dose of AJOVY. If you are not sure of your dose, ask your healthcare provider.

How do I inject AJOVY?

Step 2. Remove the prefilled syringe from the carton.

- You may need to use more than 1 prefilled syringe depending on your prescribed dose.
- Hold the prefilled syringe (as shown in Figure C).
- Remove the syringe from the carton.
- Do not shake the prefilled syringe at any time, as this could affect the way the medicine works.

Step 3. Gather the supplies you will need to inject AJOVY.

- Gather the following supplies (see Figure D) and the number of AJOVY 225 mg prefilled syringes you will need to give your prescribed dose:
  - If your dose is 225 mg, you will need 1 AJOVY 225 mg prefilled syringe.
  - If your dose is 675 mg, you will need 3 AJOVY 225 mg prefilled syringes.
  - alcohol swabs (not supplied).
  - gauze pads or cotton balls (not supplied).
  - sharps disposal or puncture-resistant container (not supplied).

Step 4. Let AJOVY reach room temperature.

- Place the supplies you have gathered on a clean, flat surface.
- Wait for 30 minutes to allow the medicine to reach room temperature.
- Do not leave the prefilled syringe in direct sunlight.
- Do not warm up the AJOVY prefilled syringe using a heat source such as hot water or a microwave.

Step 5. Wash your hands.

- Wash your hands with soap and water and dry well with a clean towel. Be careful not to touch your face or hair after washing your hands.

Step 6. Look closely at your AJOVY prefilled syringe.

Note: You may see air bubbles in the prefilled syringe. This is normal. Do not remove the air bubbles from the prefilled syringe before giving your injection. Injecting AJOVY with these air bubbles will not harm you.

- Check that the liquid medicine in the prefilled syringe is clear and colorless to slightly yellow before you give your injection (see Figure E). If the liquid has any particles in it, or is discolored, cloudy, or frozen, do not use the prefilled syringe. Call your healthcare provider or pharmacist.
- Check that AJOVY appears on the prefilled syringe.
- Check the expiration date (EXP) printed on the prefilled syringe label.
- Do not use the prefilled syringe if it has any visible damage, such as cracks or leaks. See disposal instructions in Step 12.
- Do not use the prefilled syringe if the expiration date (EXP) has passed.

Step 7. Choose your injection area.

- Choose an injection area from the following areas (see Figure F):
  - your stomach area (abdomen), avoid about 2 inches around the belly button.
  - the front of your thighs, an area that is at least 2 inches above the knee and 2 inches below the groin.
  - the back of your upper arms, in the fleshy area of the upper back portion.
- Do not inject AJOVY into an area that is tender, red, bruised, callused, tattooed, hard, or that has scars or stretch marks.
- Do not inject AJOVY in the same injection site that you inject other medicine.

Step 8. Clean your injection area.

- Clean the chosen injection area using a new alcohol swab. Let your skin dry.
- Do not inject AJOVY into an area that is tender, red, bruised, callused, tattooed, hard, or that has scars or stretch marks.
- Do not inject AJOVY in the same injection site that you inject other medicine.
Step 9. **Remove needle cap and do not replace.**
- Pick up the body of the prefilled syringe with 1 hand.
- Pull the needle cap **straight off** with your other hand (see Figure G).
- **Do not** twist.
- **Throw away** the needle cap right away.
- **Do not** put the needle cap back on the prefilled syringe, to avoid injury and infection.

**Figure G**

**Step 10.** Give your injection following the 4 steps below.

1. **Use your free hand to gently pinch up** at least 1 inch of the skin that you have cleaned.
2. **Insert the needle into the pinched skin at a 45 to 90 degree angle.**
3. **When the needle is all the way into your skin, use your thumb to push the plunger.**
4. **Push the plunger slowly all the way down as far as it will go to inject all of the medicine.**

**Figure H**

**Step 11.** Remove the needle from your skin.
- After you have injected all of the medicine, **pull the needle straight out** (see Figure H).
- **Do not** recap the needle at any time to avoid injury and infection.

**Step 12.** Apply pressure to the injection site.
- Use a clean, dry cotton ball or gauze to **gently press on the injection site** for a few seconds.
- **Do not** rub the injection site.
- **Do not** re-use the prefilled syringe.

**Step 13.** Dispose of your prefilled syringe right away.
- Put your used prefilled syringes, needles, and sharps in a FDA-cleared sharps disposal container right away after use.
- **Do not throw away (dispose of) loose needles, syringes, or prefilled syringes in your household trash. Do not recycle your used sharps disposal container.**
- If you do not have a FDA-cleared sharps disposal container, you may use a household container that is:
  - made of a heavy-duty plastic,
  - can be closed with a tight-fitting, puncture-resistant lid, without sharp being able to come out,
  - upright and stable during use,
  - leak-resistant, and
  - properly labeled to warn of hazardous waste inside the container.