The first and only DHE autoinjector. 1,2



Is your headache medicine providing you long lasting* relief for the acute treatment of migraine or cluster headaches?

Hospital-strength headache relief — now in your hands^{1,3-5}



ASK YOUR HEALTHCARE PROVIDER ABOUT BREKIYA AUTOINJECTOR

DHE, dihydroergotamine

*Lasting represents a duration of approximately 24-72 hours.^{3,4}

[†]Cluster headache results reported using intravenous (IV) administration.⁴

Indication

Brekiya autoinjector is a prescription medicine used for the acute treatment of migraine with or without aura and acute cluster headaches in adults.

Brekiya autoinjector is not used to prevent migraine or used to treat other types of headaches such as hemiplegic migraines (that make you unable to move on one side of your body) or basilar migraines (rare form of migraine with aura). It is not known if Brekiya autoinjector is safe and effective in children.

IMPORTANT SAFETY INFORMATION

Serious or potentially life-threatening reductions in blood flow to the brain or extremities due to interactions between dihydroergotamine (the active ingredient in Brekiya autoinjector) with strong CYP3A4 inhibitors (such as protease inhibitors and macrolide antibiotics) have been reported rarely. As a result, these medications should not be taken together.

Please see Boxed Warning above, Important Safety Information throughout this brochure, and Full Prescribing Information and Medication Guide in the pocket.

Relief shouldn't require the emergency room

Headache is the **fourth most common reason** for emergency department visits. But the ER is often the last place you want to be during an attack⁶⁻⁹:



Migraine attacks are often worsened by **bright lights**, **loud sounds**, **and strong smells** — all common triggers found in emergency rooms⁷



Cluster headache attacks are often misdiagnosed in emergency settings and **may be mistaken for less severe conditions.** Because they can last between 15 minutes to 3 hours, the attack may be over before a patient is seen^{8,9}



DID YOU KNOW?

Cluster headache is a type of primary headache disease that causes repeated attacks of sudden, severe pain, usually around one eye or on one side of the head.

It's often described as one of the most painful conditions a person can experience.¹⁰

IMPORTANT SAFETY INFORMATION (continued)

Do not use Brekiya autoinjector if you:

- are taking medicines known as strong CYP3A4 inhibitors, such as: ritonavir, nelfinavir, indinavir, erythromycin, clarithromycin, ketoconazole, or itraconazole.
- have heart problems or a history of heart problems, or uncontrolled high blood pressure.
- have narrowing of blood vessels in your legs, arms, stomach, or kidneys (peripheral vascular disease).
- have sepsis, had vascular surgery, severe liver or kidney problems.
- are allergic to dihydroergotamine, ergot alkaloids, latex, or any of the ingredients in Brekiya autoinjector.

Please see Important Safety Information throughout this brochure, including Boxed Warning on cover, and Full Prescribing Information and Medication Guide in the pocket.

A medicine that has been trusted for decades

For more than 70 years, D.H.E. 45° has been a trusted acute treatment option for migraine with or without aura and cluster headache^{5,11}

HOW DHE WORKS

DHE is believed to work in multiple ways to help ease migraine with or without aura and cluster headache symptoms. It regulates multiple pain and inflammatory neurotransmitters in the brain.¹²



Ready when you need it most

DHE can be used at any point during an attack and, in hospital settings, is typically given intravenously (IV), often when other treatments haven't worked.^{1,5,11}

DHE refers to dihydroergotamine mesylate, the active ingredient in D.H.E. 45°.11



IMPORTANT SAFETY INFORMATION (continued)

Do not use Brekiya autoinjector if you: (continued)

- have taken any of the following medicines in the last 24 hours: sumatriptan, frovatriptan, ergotamine or ergotamine-type medicines, almotriptan, naratriptan, zolmitriptan, eletriptan, or rizatriptan.
- have taken any medicines that constrict your blood vessels or raise your blood pressure.

Before you take Brekiya autoinjector, tell your healthcare provider about all your medical conditions, including if you:

• have high blood pressure, liver problems, or kidney problems.



DHE has been studied in both migraine and cluster headache

It has been shown to help relieve pain and may help keep it from coming back^{3,4}



of people with migraine treated with subcutaneous D.H.E. 45° had **pain relief 24 hours after their first dose**—compared to 77% treated with subcutaneous sumatriptan, a common treatment for migraine^{3,‡}



of people who received subcutaneous D.H.E. 45° for migraine with or without aura did not experience another headache within 24 hours^{3,‡}

Note that these studies did not include Brekiya autoinjector. Brekiya autoinjector contains DHE.^{1,3,4}

IMPORTANT SAFETY INFORMATION (continued)

Before you take Brekiya autoinjector, tell your healthcare provider about all your medical conditions, including if you: (continued)

- have risk factors for heart disease, such as: high blood pressure, high cholesterol levels, smoke, are overweight, diabetes, family history of heart disease.
- are pregnant or plan to become pregnant. Brekiya autoinjector may cause preterm labor. Brekiya autoinjector should be avoided during pregnancy. Talk to your healthcare provider right away if you are pregnant or want to become pregnant.
- are breastfeeding or plan to breastfeed. Brekiya autoinjector may reduce breast milk supply and pass into your breast milk. Brekiya autoinjector may be harmful to your baby. Do not breastfeed your baby while taking Brekiya autoinjector and for 3 days after you use Brekiya autoinjector. Talk with your healthcare provider about the best way to feed your baby if you take Brekiya autoinjector.

Please see Important Safety Information throughout this brochure, including Boxed Warning on cover, and Full Prescribing Information and Medication Guide in the pocket.



of people with episodic cluster headache didn't experience attacks for 3 months after intravenous D.H.E. 45° treatment^{4,§}

Note that these studies did not include Brekiya autoinjector. Brekiya autoinjector contains DHE.^{1,3,4}

[†]In a study of 295 patients who took under-the-skin (subcutaneous) injections for either D.H.E 45® or sumatriptan to treat their migraines. They reported results within 24 hours. Pain relief was defined as a change from moderate or severe pain, to mild or none.³

§In a study of 54 patients with cluster headaches, treated in hospital with intravenous DHE with dosing adjustments until they were headache-free for 24 hours.⁴

IMPORTANT SAFETY INFORMATION (continued)

Tell your healthcare provider about all the medicines you take, including prescription and over-the-counter medicines, vitamins, and herbal supplements. Especially tell your healthcare provider if you take: sumatriptan, ergot-type medicine, saquinavir, nefazodone, fluconazole, grapefruit juice, zileuton, nicotine, propranolol or other medicines that can lower your heart rate, any medicines that can increase your blood pressure, selective serotonin reuptake inhibitors. These are not all of the medicines that could affect how Brekiya autoinjector works. Your healthcare provider can tell you if it is safe to take Brekiya autoinjector with other medicines.

brekiya® dihydroergotamine mesylate) 1 mg/mL injection autoinjector

DHE has been shown to help across a range of different migraine and cluster headache types



It may be an option worth exploring if any of these sound familiar. 1.1

- Acute migraine attacks with or without aura¹
- Migraine attacks lasting >72 hours (status migrainosus)^{13,14}
- Recurring migraine attacks⁵
- Migraine not responding to other treatments, such as triptans and gepants^{5,13,14}

- Migraine attacks that come on quickly⁵
- Migraine upon awakening¹⁵
- Migraine linked to medication overuse^{13,14}
- Cluster headache attacks¹

The active ingredient in Brekiya autoinjector is DHE.1

Aura is a series of sensory disturbances or disruptions that usually happens shortly before a migraine attack.¹⁶

These results are based on studies of DHE given in different ways, including as a nasal spray, intravenous (IV) injection, and under-the-skin (subcutaneous) injection.^{1,5,13-15}

¹Brekiya provides the same amount of DHE in the body as a subcutaneous injection of D.H.E. 45[®]. ¹

IMPORTANT SAFETY INFORMATION (continued)

How should I use Brekiya autoinjector?

- Certain people should take their first dose of Brekiya autoinjector in their healthcare provider's office or in another medical setting.
- Brekiya autoinjector is for injection under the skin (subcutaneous) only.
- Use Brekiya autoinjector exactly as your healthcare provider tells you to use it.
- Each autoinjector contains one dose (1 mg).
- If your headache comes back after the first complete dose, you may give yourself up to 2 more doses as needed. Wait at least one hour between doses.
- **Do not** inject more than 3 doses (3 mg) of Brekiya autoinjector in a 24-hour period or 6 doses (6 mg) in a 1-week (7 day) period.

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Is DHE right for me?

Ask your healthcare provider if **Brekiya autoinjector** could be a good option for managing attacks like these

IMPORTANT SAFETY INFORMATION (continued)

What are the possible side effects of Brekiya autoinjector? Brekiya autoinjector can cause serious side effects, including:

- Heart attack and other heart problems. Heart problems may lead to death. Stop taking
 Brekiya autoinjector and get emergency medical help right away if you have any symptoms
 of a heart attack, such as: numbness or tingling in your fingers and toes, muscle
 pain or cramps in your arms and legs, weakness in your legs, temporary speeding or slowing
 of your heart rate, or swelling or itching. Brekiya autoinjector is not for
 people with risk factors for heart disease unless a heart exam is done
 and shows no problem.
- **Stroke.** Stop using Brekiya autoinjector and get emergency medical help right away if you have any of the symptoms of a stroke.



Brekiya autoinjector contains the same medicine trusted in hospitals — now in your hands^{1,5}





Is the **first and only autoinjector** to deliver DHE for lasting# headache relief in a disposable, ready-to-use, single-dose autoinjector injected into the thigh 1-5,**



Can be used at any point during an attack and may help reduce headache recurrence^{1,3,4}

*Lasting represents a duration of approximately 24-72 hours. 3,4

**Cluster headache results reported using intravenous (IV) administration.4

IMPORTANT SAFETY INFORMATION (continued)

What are the possible side effects of Brekiya autoinjector? Brekiya autoinjector can cause serious side effects, including: (continued)

- Changes in color or sensation in your fingers and toes (Raynaud's syndrome).
- Stomach and intestinal problems.
- Increased blood pressure.
- Medicine overuse headache. Some people who use too much Brekiya autoinjector may make their headaches worse. If your headaches get worse, your healthcare provider may decide to stop your treatment with Brekiya autoinjector.
- Preterm labor.

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May be helpful for people who struggle with oral treatments because it is given subcutaneously⁵



Designed so you can use it on your own, with **no need for refrigeration**, assembly, or preparation of the autoinjector¹

IMPORTANT SAFETY INFORMATION (continued)

What are the possible side effects of Brekiya autoinjector? Brekiva autoiniector can cause serious side effects, including: (continued)

• Tissue changes (fibrotic complications). Inflammation and fiber-like tissue that is not normal (fibrosis) can occur around the lungs and stomach. Tell your healthcare provider if you have any side effect that bothers you or that does not go away. These are not all of the possible side effects of Brekiya autoinjector. Call your healthcare provider for medical advice about side effects.



How to use Brekiya autoinjector



Designed so you can use it on your own, with no need for refrigeration, assembly, or preparation of the autoinjector.¹ It's important to use Brekiya autoinjector exactly as your healthcare provider tells you.

DOSING INFORMATION

- Each autoinjector contains one dose (1 mg)¹
- If your headache comes back after the first complete dose, you may give yourself up to 2 more doses as needed. Wait at least one hour between doses¹
- **Do not** inject more than 3 doses (3 mg: 3 doses of 1 mg) of Brekiya autoinjector in a 24-hour period or 6 doses (6 mg) in a 1-week (7 day) period¹
- Brekiya autoinjector should not be used for chronic daily administration.



For illustration purposes only. Not actual size.

IMPORTANT SAFETY INFORMATION (continued)

How should I store Brekiya autoinjector?

- Store at room temperature between 68°F to 77°F (20°C to 25°C). Do not refrigerate or freeze.
- · Protect Brekiya autoinjector from light.
- Keep Brekiya autoinjector in the original pack until ready to use.

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Scan the QR code to watch a step-by-step video on how to use Brekiya autoinjector safely and confidently

100%

of patients were able to use Brekiya autoinjector on their own, when following instructions for use^{2,††}

^{††}According to a human factor study of 30 people, living with migraine and/or cluster headache, who had no prior training for using the autoinjector.²

References: 1. Brekiya [package insert]. Bridgewater, NJ: Amneal Pharmaceuticals LLC; 2025. 2. Data on file, Amneal Pharmaceuticals LLC. 3. Winner P, Ricalde O, Le Force B, Saper J, Margul B. Arch Neurol. 1996;53(2):180-184. doi:10.1001/archneur.1996.00550020092020 4. Mather PJ, Silberstein SD, Schulman EA, Hopkins MM. Headache. 1991;31(8):525-532. doi:10.1111/j.1526-4610.1991.hed3108525.x 5. Silberstein SD, Shrewsbury SB, Hoekman J. Headache. 2020;60(1):40-57. doi:10.1111/head.13700 6. Phelan A, DeMera R, Halker Singh RB, Hentz JG, Spears RC. Headache. 2021;61(10):1499-1507. doi:10.1111/head.14194 7. American Migraine Foundation. Understanding migraine treatment in the emergency room. July 12, 2018. Accessed June 18, 2025. https://americanmigrainefoundation.org/resource-library/understanding-migraine-treatment-in-the-emergency-room/8. Hasirci Bayir BR, Nazli E, Ulutas C. Medicina. 2025;61(3):437. doi:10.3390/medicina61030437 9. Wei DY-T, Ong JJY, Goadsby PJ. Ann Indian Acad Neurol. 2018;21(Suppl 1):S3-S8. doi:10.4103/aian. AIAN_349_17 10. Leroux E, Ducros A. Orphanet J Rare Dis. 2008;3:20. doi:10.1186/1750-1172-3-20 11. D.H.E. 45® [package insert]. Aliso Viejo, CA: Valeant Pharmaceuticals North America; 2008 12. Silberstein SD, McCrory DC. Headache. 2003;43(2):144-166. doi:10.1046/j.1526-4610.2003.03034.x 13. Shafqat R, Flores-Montanez Y, Delbono V, Nahas SJ. J Pain Res. 2020;13:859-864. doi:10.2147/JPR.S203650 14. Silberstein SD, Kori SH. CNS Drugs. 2013;27(5):385-394. doi:10.1007/s40263-013-0061-2 15. Tepper SJ. Headache. 2013; 53. doi:10.1111/head.13700.



Eligible, commercially-insured patients may pay as little as \$40 per carton^{‡‡,§§}

HOW DO I GET MY PRESCRIPTION?

- 1. Your healthcare provider will submit your prescription to a specialty pharmacy, and that pharmacy will call or text^{III} you within 2 business days.
- The specialty pharmacy will arrange delivery to your preferred location.



888-618-4126



888-282-5166

SCAN THIS QR CODE OR VISIT BREKIYA.COM TO LEARN MORE



IMPORTANT SAFETY INFORMATION (continued)

Do not use Brekiya autoinjector if you:

- are taking medicines known as strong CYP3A4 inhibitors, such as: ritonavir, nelfinavir, indinavir, erythromycin, clarithromycin, ketoconazole, or itraconazole.
- have heart problems or a history of heart problems, or uncontrolled high blood pressure.
- have narrowing of blood vessels in your legs, arms, stomach, or kidneys (peripheral vascular disease).
- have sepsis, had vascular surgery, severe liver or kidney problems.
- are allergic to dihydroergotamine, ergot alkaloids, latex, or any of the ingredients in Brekiya autoinjector.
- have taken any of the following medicines in the last 24 hours: sumatriptan, frovatriptan, ergotamine or ergotamine-type medicines, almotriptan, naratriptan, zolmitriptan, eletriptan, or rizatriptan.
- · have taken any medicines that constrict your blood vessels or raise your blood pressure.

To report SUSPECTED ADVERSE REACTIONS, contact Amneal Specialty, a division of Amneal Pharmaceuticals LLC at 1-877-835-5472, or FDA at 1-800-FDA-1088 or www.fda.gov/medwatch.

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[#]Restrictions and quantity limits apply. See Brekiva.com for more information.

^{§§} Each carton contains 4 single dose autoinjectors.

Standard text messaging rates apply.