

INDICATION

DAWNZERA™ (donidalorsen) is a prescription medicine used to prevent hereditary angioedema (HAE) attacks in adults and children 12 years of age and older.

It is not known if DAWNZERA is safe and effective in children under 12 years of age.

SELECT IMPORTANT SAFETY INFORMATION

Do not use DAWNZERA if you have had a serious allergic reaction, including anaphylaxis to donidalorsen or any of the ingredients in DAWNZERA.

Before using DAWNZERA, tell your healthcare provider about all of your medical conditions, including if you: are pregnant or plan to become pregnant, are breastfeeding or plan to breastfeed. It is not known if DAWNZERA can harm your unborn baby, or if it passes into your breast milk and if it can harm your baby.



It's time to

REDEFINE WHAT'S "NORMAL" FOR HAE

About three-fourths of people with hereditary angioedema (HAE) reported that they make trade-offs in their lives due to the unpredictability of attacks.*

While everyone's experience with HAE is different, you may recognize some similarities in what others have shared.



9 OUT 10

people reported that they frequently missed or avoided events because of their HAE.*

^{*}Online survey conducted by The Harris Poll of 150 US adults diagnosed with HAE by a health care provider and 228 board-certified allergists and immunologists practicing in the US, sponsored by Ionis.



93%

of people reported an **HAE attack in the past year**, but doctors estimated only 64%, highlighting a gap in awareness.*



TWO-THIRDS

of people reported they felt that they still **hadn't found the best preventative treatment** for them.*

If HAE attacks are holding you back, it may be time to consider something new

*Online survey conducted by The Harris Poll of 150 US adults diagnosed with HAE by a health care provider and 228 board-certified allergists and immunologists practicing in the US, sponsored by Ionis.

Looking for a sign to SWITCH?

Or, are you looking to start your first preventative treatment? Meet DAWNZERA

DAWNZERA offers the **convenience of a self-administered autoinjector**. No mixing or syringes, and although refrigeration is recommended, it can be stored at room temperature for 6 weeks.



Dosing every 4 or 8 weeks

Recommended dosing for DAWNZERA is once every 4 weeks, but every 8 weeks may also be considered.



Autoinjector

DAWNZERA is delivered through an autoinjector. DAWNZERA contains no citric acid, which is associated with injection-site pain.*



Flexible storage

DAWNZERA autoinjectors can be stored in its original carton at room temperature (up to 86 °F [30 °C]) for up to 6 weeks.

Please note: Do not expose your DAWNZERA autoinjector to heat, do not freeze it, and protect it from direct light.





How to Use DAWNZERA

Watch the How to Use DAWNZERA training video at **DAWNZERA.com/taking-dawnzera** for step-by-step instructions on how to **place**, **push**, **and hold** the autoinjector.

*In clinical studies, injection site reactions, including injection site pain, were the most frequently reported adverse reactions.

SELECT IMPORTANT SAFETY INFORMATION

Tell your healthcare provider about all the medicines you take, including prescription and over-the-counter medicines, vitamins, and herbal supplements.

What are the possible side effects of DAWNZERA?

DAWNZERA can cause serious side effects including allergic reactions. Allergic reactions can include rash, trouble breathing, chest pain, fainting, dizziness, feeling lightheaded, swelling of the face, lips or tongue, and itching. Stop using DAWNZERA and call your healthcare provider or get emergency help right away if you have any of these symptoms.

DAWNZERA works differently BY USING RNA TECHNOLOGY

to prevent HAE attacks

RNA technology has been used across many types of diseases for almost 30 years and works by reducing, enhancing, or modifying specific proteins.

DAWNZERA is designed to work in 3 steps:

DAWNZERA is specifically targeted to the liver.



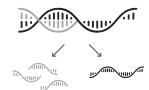
DAWNZERA finds and directs the breakdown of the message (called mRNA) that tells your body to make plasma PKK.



By targeting mRNA, **DAWNZERA** helps to control the pathway that leads to swelling.











DAWNZERA has the precision of RNA technology without changing your DNA.



WHY DOES DAWNZERA TARGET THE LIVER?

Plasma PKK, a protein that plays a key role in swelling, is made in the liver. Therefore, DAWNZERA works in the liver to lessen plasma PKK in the blood and help to control the pathway that leads to swelling

DNA=deoxyribonucleic acid; mRNA=messenger RNA; PKK=prekallikrein; RNA=ribonucleic acid.



LASTING REDUCTIONS

in HAE attacks

In clinical studies, there was a **steady reduction over time** in the average of HAE attacks for those who took DAWNZERA every 4 weeks.

Primary Study



fewer attacks on average at 6 months

vs placebo (primary endpoint)*



fewer attacks on average at 6 months from the 2nd dose

vs placebo (secondary endpoint)*

Long-term Safety Study



fewer attacks at 1 year

on average vs their attack rate before starting treatment (secondary endpoint)

About 90% fewer moderate-to-severe HAE attacks, HAE attacks requiring rescue medication, and HAE attacks requiring ER visits[†]

ER=emergency room.

*Reductions in HAE attacks were measured 24 weeks after their first dose and 20 weeks after their second dose.

†Average percentage (%) observed in people using DAWNZERA every 4 weeks. Moderate-to-severe HAE attacks and rescue medication use were secondary endpoints, and ER visits due to HAE attacks were an exploratory endpoint.

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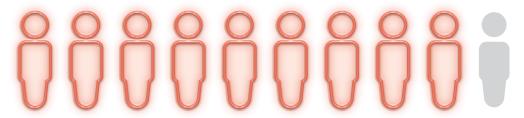
Tell your healthcare provider about all the medicines you take, including prescription and over-the-counter medicines, vitamins, and herbal supplements.

In the open-label study,

The majority of people SWITCHED SAFELY without an overall increase in HAE attacks

DAWNZERA was evaluated in people who switched from lanadelumab, berotralstat, or a C1-INH.*

HAE mean attack reduction was seen regardless of prior preventative treatment.[†]





people in the study who used the AECT reported being **well-controlled** after switching to DAWNZERA (secondary endpoint interim analysis at Week 16)[‡]



HOW MANY PEOPLE SURVEYED PREFERRED DAWNZERA?

84% of people surveyed preferred DAWNZERA over their previous preventive treatment, citing better disease control, less time required for treatment, and reduced injection pain as reasons for their preference[§]

*These are not head-to-head data. Findings are from an open-label, uncontrolled safety study in patients who wanted to switch to DAWNZERA, and was not powered for any comparisons between prior treatment groups; thus, these observations cannot be generalized to other patients on prior long-term preventative treatments.

AECT=Angioedema Control Test; C1-INH=C1 esterase inhibitor.

Previous on-treatment monthly HAE attack rate established in the 10-week screening period.

¹The AECT is a simple, 4-question survey that helps you and your doctor understand how well your swelling symptoms have been managed over the past month.

§More than 1 reason was permitted. Exploratory endpoint interim analysis at Week 16.



SAFETY AND SIDE EFFECTS

seen with DAWNZERA

The most common side effects of DAWNZERA were injection-site reactions.*

Adverse reaction, n (%)	DAWNZERA every 4 weeks (N=45)	DAWNZERA every 8 weeks (N=23)	Placebo (N=22)
Injection-site reactions [†]	11 (24)	1 (4)	1 (5)
Upper respiratory tract infection	4 (9)	2 (9)	1 (5)
Urinary tract infection	4 (9)	2 (9)	0
Abdominal discomfort	3 (7)	0	0

Redness was the most common injection-site reaction.

In clinical trials, hypersensitivity reactions, including anaphylaxis, have occurred. Symptoms included generalized rash, shortness of breath, chest pain, and swelling around the mouth.

These are not the only possible side effects of DAWNZERA. Tell your doctor if you have any side effect that bothers you or that does not go away while taking DAWNZERA.

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Tell your healthcare provider about all the medicines you take, including prescription and over-the-counter medicines, vitamins, and herbal supplements.

^{*}Injection site reactions include: erythema (redness), discoloration, pain, pruritus (itching), induration (hardening or thickening of the skin), bruising, hematoma (localized bleeding under the skin), hypersensitivity, swelling, reaction, and urticaria (hives).

†All injection-site reactions were mild, nonserious, and the majority of them resolved without receiving any treatment.

Let's take this step by step, TOGETHER



Ionis Every Step is here for you every step of the way, from getting your prescribed medicine to learning how to use it and ways to help follow your doctor's treatment plan.

Signing up for Ionis Every Step gives you access to:



A Patient Education Manager*

You'll be connected with a dedicated Patient Education Manager—your primary Ionis Every Step partner from the start and throughout your treatment journey.



Insurance Navigation[†]

Ionis Every Step works with you and your doctor to help get insurance approval for your medication and provide updates along the way.



Affordability Programs[‡]

Ionis Every Step is committed to helping you access your Ionis medication. There are many financial assistance options available regardless of your insurance.



DAWNZERA Direct

Your digital companion for managing your DAWNZERA treatment journey. Reach out to your Patient Education Manager to get started.

TAKE YOUR NEXT STEP with Ionis Every Step.
Call 1-844-444-4305, Monday to Friday, 8 AM to 8 PM ET, to sign up



^{*}Patient Education Managers cannot give medical advice and do not replace your medical team. Contact your doctor for any medical concerns.

*Insurance approval is not guaranteed. Ionis Every Step offers financial assistance programs for patients who are uninsured or denied coverage for their Ionis medication.

[‡]Terms and conditions apply. Programs subject to change or discontinue without notice, including in specific states.

READ THE SIGNS

ASK FOR DAWNZERA

Use the information in this guide to help you talk with your doctor about what option may best fit your life and treatment goals.

Why DAWNZERA?



Lasting reductions in HAE attacks, with the majority of side effects being mild or moderate



The **longest time between doses** available for
HAE attack prevention*



A new targeted way of working to prevent HAE attacks



Visit DAWNZERA.com to hear from people taking DAWNZERA, learn more about treatment, and access important resources for getting started

*Available in every-4-week and every-8-week dosing.

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The most common side effects of DAWNZERA include injection site reactions (such as redness or pain at the injection site), upper respiratory tract infection, urinary tract infection, abdominal discomfort.

These are not all the possible side effects of DAWNZERA. Tell your healthcare provider or treatment team about any side effects you may have.

Ready for a NEW TREATMENT?

Tear off this page and check any of the statements that apply to you. You can also use this checklist to help you talk with your doctor about whether DAWNZERA could be a good fit.

I feel dissatisfied with how my current treatment is working

I feel like preparing my treatment is more complicated than I'd like

I wish I didn't have to worry about taking my treatment every day

I want a more effective treatment, but I'm scared of needles

I'm looking for more 1:1 support with my next treatment

I want the option to have up to 2 months between doses



LOOKING FOR OTHER OPTIONS?

Living with HAE can be hard to explain to others, even doctors

If you're thinking about a new treatment for HAE, this template can help you start a conversation with your doctor. Feel free to personalize it based on your experiences.

HI DOCTOR,

I've been thinking about how my HAE has been affecting my life. I learned about the treatment DAWNZERA (pronounced: dawn-ZAIR-ah) and I'd like to talk with you to see if it might be a good option for me.

I read that DAWNZERA offers:

- The longest time between doses available for HAE attack prevention*
- A technology for treating HAE attacks called RNA technology
- Lasting HAE prevention with a well-tolerated safety profile

*Available in every-4-week and every-8-week dosing.

RNA=ribonucleic acid.

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DAWNZERATM (donidalorsen) is a prescription medicine used to prevent hereditary angioedema (HAE) attacks in adults and children 12 years of age and older.

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Please see Important Safety Information throughout, and accompanying full Prescribing Information and Patient Information for DAWNZERA.

IONIS

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