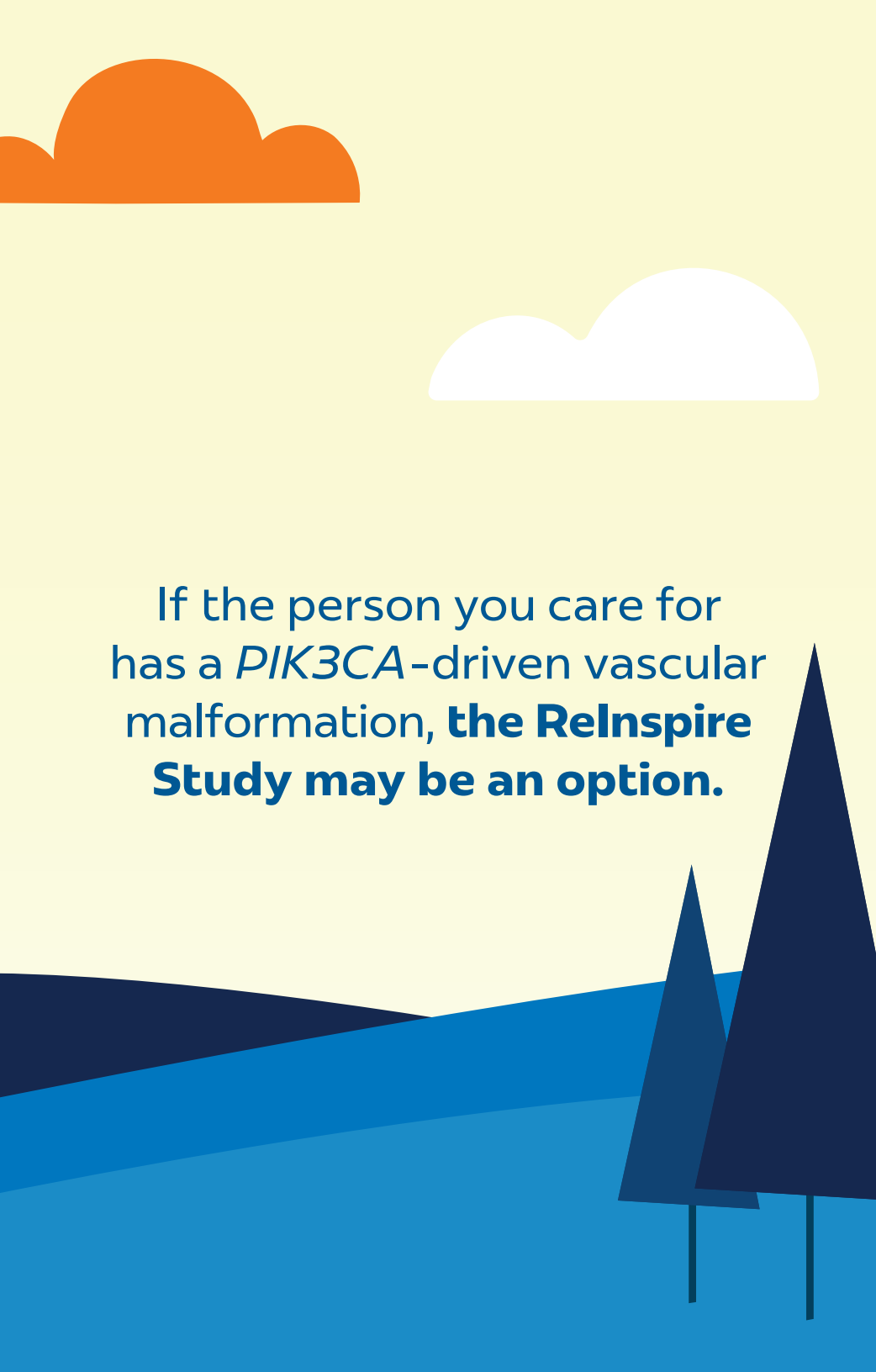


A stylized landscape illustration with a winding blue path leading through a field of purple flowers towards a sunset over rolling hills. The sky is a gradient of yellow and orange, and the hills are in shades of orange and red. Dark blue evergreen trees are scattered throughout the scene.

Could a more  
**selective treatment**  
mean brighter days?

Consider clinical research for individuals with *PIK3CA*-driven vascular malformations including Lymphatic Malformations and *PIK3CA*-Related Overgrowth Spectrum (PROS).



If the person you care for has a *PIK3CA*-driven vascular malformation, **the ReInspire Study may be an option.**

## What Are *PIK3CA*-Driven Vascular Malformations?

*PIK3CA*-driven vascular malformations may include lymphatic malformations and *PIK3CA*-related overgrowth spectrum (PROS). PROS consists of syndromes such as Klippel-Trenaunay, CLOVES, FAVA, MCAP, and more.

This clinical research study is designed to learn about a potential new treatment option for *PIK3CA*-driven vascular malformations. These types of malformations are caused by mutations or changes in the *PIK3CA* gene, which create a malfunctioning form of the protein PI3K $\alpha$ . When mutated, PI3K $\alpha$  can cause vascular malformations and/or overgrowths throughout the body, along with other symptoms. Although everyone has a unique experience, common symptoms include pain, swelling, infections, and impaired physical functioning, which can significantly impact quality of life.

## About the ReInspire Study

The ReInspire Study is evaluating an investigational study drug for the treatment of *PIK3CA*-driven vascular malformations. "Investigational" means that the study drug has not yet been approved by the US Food and Drug Administration (FDA) or any other regulatory agency, such as the European Commission in the EU, and is not available outside of a clinical research study.



## About the Investigational Study Drug

The study drug being evaluated is called RLY-2608. It is a drug taken orally that is designed to specifically hinder the mutated PI3K $\alpha$  protein that can cause *PIK3CA*-driven vascular malformations. By selectively targeting the mutated proteins and limiting impact on the non-mutated proteins, the study drug aims to reduce side effects and provide more effective treatment over non-mutant-selective therapies.

## Study Parts

The ReInspire Study is made up of three parts. Participants will only be enrolled in one part.


**Part 1** will determine the dose(s) of RLY-2608 that provides the most benefit with the least side effects.

**Part 2** will continue to collect data on the safety and effectiveness of the recommended dose(s) found during Part 1.


**Part 3** will open after Parts 1 and 2 and will further confirm the treatment effect of the drug.


## Study Design

To enter the study, the person you care for will first need to give their consent, or permission, by reviewing and signing an informed consent form. No matter what part of the ReInspire Study the person you care for is enrolled in, their participation will consist of three periods:

 **Screening:** A study doctor will evaluate the person you care for to see if the study is a good match.

- During this period, the person you care for will have a single or multiple screening visits to complete the tests or procedures such as a physical exam, routine labs, MRI, and others, to determine if they meet the requirements to participate.

 **Treatment:** If the study is a good match for the person you care for and they agree to take part, they will begin taking the study drug and attend monthly study site visits. Their treatment period will last as long as the person you care for and their doctor determine it is in their best interest, or until the study ends.

 **Follow-up:** If the person you care for stops taking the study drug, their study doctor will continue to monitor their health for a period of time.

## Who Can Participate?

**To participate in this study, the person you care for must:**

- Have a diagnosis of a *PIK3CA*-driven vascular malformation, such as lymphatic malformation or PROS (includes syndromes such as Klippel-Trenaunay, CLOVES, FAVA, MCAP, and more)
- NOT have another genetic driver that is known to cause their disease

Additional requirements will apply. Discuss with the doctor of the person you care for whether the ReInspire Study may be an option.

**To learn more, visit [ReInspireStudy.com](https://ReInspireStudy.com) or email [clinicaltrials@relaytx.com](mailto:clinicaltrials@relaytx.com).**



## What You Should Know About Clinical Research Studies

Clinical research studies, also called clinical trials, aim to answer specific questions about how drugs work in the participants who take them. You and the person you care for should feel fully informed about what to expect from their participation in a clinical research study.

### Researchers use clinical research studies to:

- Learn about the safety and effects of investigational medicines
- Help find new ways of using certain medications
- Identify new treatment options
- Answer specific health questions

Participation in any clinical research study is voluntary (the choice of you and the person you care for). The clinical research study team will inform the person you care for of the potential risks and benefits of participation, as well as possible side effects. To make an informed decision, talk to the person you care for and their healthcare providers about any questions you both may have.

### All clinical research studies are:

- Developed to protect the rights, safety, and well-being of participants
- Conducted according to strict scientific and ethical principles
- Reviewed and approved by an institutional review board (IRB) or ethics committee (EC)