A Phase 2 Study of Mutant-selective PI3Kα Inhibitor, RLY-2608, in Adults and Children With PIK3CA Related Overgrowth Spectrum and Malformations Driven by PIK3CA Mutation (The ReInspire Study)

Status: RECRUITING

Eligibility Criteria

Sex: ALL
Age: 2 years and over
Healthy Volunteers: This study is NOT accepting healthy volunteers
Key Inclusion Criteria:

- The participant must have a clinical diagnosis of PROS or a malformation within the ISSVA classification.
- One or more documented activating PIK3CA mutation(s) that are targeted by selective PI3Kα inhibitors in lesional tissue and/or cell-free DNA from the lesion or blood. Some participants may be eligible without a documented PIK3CA mutation, with the sponsor's approval, as long as no other genetic driver has been documented.
- Lansky (<16 yo) or Karnofsky (≥16 yo) performance status of ≥50.
- Agree to provide archived lesional fluid and/or tissue or be willing to undergo pretreatment lesional biopsy (if considered safe and medically feasible) to assess PIK3CA status.

Key Exclusion Criteria:

- Known hypersensitivity to RLY-2608.
- Any factors that increase the risk of QTc prolongation or risk of arrhythmic events
- Clinically significant, uncontrolled cardiovascular disease
- Received disease-directed therapy prior to the first dose of study drug:

1. Systemic therapy or antibody within 5 half-lives of the therapy.
2. Local therapy including radiation, surgery, or other procedures within 28 days; lesion(s) must have demonstrated progression after the procedure.

Conditions & Interventions

Interventions:
DRUG: RLY-2608, DRUG: Placebo
Conditions:
PIK3CA-Related Overgrowth Spectrum (PROS), Lymphatic Malformations, Vascular Malformations, PIK3CA Mutation, CLOVES Syndrome, Klippel Trenaunay Syndrome, Megalencephaly-capillary Malformation Polymicrogyria Syndrome (MCAP), Vascular Anomalies

More Information

Description:
This is a 3-part Phase 2 randomized study evaluating the safety and efficacy of the mutant-selective PI3Kα inhibitor, zovegalisib (RLY-2608), in adults and children with PIK3CA Related Overgrowth Spectrum (PROS) and malformations driven by PIK3CA mutation. Part 1 is a dose selection, Part 2 is a basket design with exploratory single-arm cohorts for various subpopulations of participants, and Part 3 is randomized, double-blinded study vs placebo.

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