BioCryst Announces Publication of Data from Open-label Extension of the APeX-2 Pivotal Trial of ORLADEYO® (berotralstat)

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– Results published in JACI: In Practice show rapid and sustained reductions in HAE attacks and improved quality of life over 96 weeks of treatment with ORLADEYO –

RESEARCH TRIANGLE PARK, N.C., Dec. 19, 2023 (GLOBE NEWSWIRE) -- BioCryst Pharmaceuticals, Inc. (Nasdaq: BCRX) today announced that data from the open-label extension (OLE) of the APeX-2 trial of oral, once-daily ORLADEYO® (berotralstat) for the prophylactic treatment of hereditary angioedema (HAE) in patients 12 years and older have been published online by the *Journal of Allergy and Clinical Immunology: In Practice* (*JACI: In Practice*).

"As detailed in this manuscript, long-term prophylaxis with ORLADEYO has enabled patients with HAE to better manage their condition, including reducing the number of HAE attacks they experience and demonstrating clinically meaningful improvement in their quality of life. I have seen firsthand in my practice evidence that the longer patients remain on ORLADEYO, the better their outcomes are," said William R. Lumry, M.D., clinical professor of internal medicine at the University of Texas Southwestern Medical School.

"We are pleased to share long-term efficacy and safety data from APeX-2 as published in *JACI: In Practice*. These data further illustrate the potential lasting outcomes that can be appreciated by patients who are treated with oral, once-daily ORLADEYO. We continue to see long-term safety and effectiveness data that reinforce ORLADEYO as an important treatment option for patients with HAE, and we look forward to sharing additional real-world evidence at upcoming medical congresses," said Dr. Ryan Arnold, chief medical officer of BioCryst.

APeX-2 was a Phase 3, double-blind, placebo-controlled, parallel-group, three-part study evaluating ORLADEYO versus placebo for the prevention of HAE attacks in 121 patients with HAE Type I or Type II. In part 3 of the study (weeks 49-96), all patients (n=81) were treated with open-label ORLADEYO at 150 mg. The primary endpoint of the OLE was long-term safety and tolerability, while secondary endpoints included HAE attack rates and quality of life (QoL).

The authors concluded that ORLADEYO was generally well tolerated, provided rapid and sustained reductions in HAE attacks and improved QoL over the study duration of 96 weeks. The results from part 3 of APeX-2 were first presented at the European Academy of Allergy and Clinical Immunology (EAACI) Congress in July 2021.

Additional details can be found in the manuscript, which is available online at jaci-inpractice.org. *JACI: In Practice* is an official journal of the American Academy of Allergy, Asthma, and Immunology (AAAAI).

About ORLADEYO® (berotralstat)

ORLADEYO® (berotralstat) is the first and only oral therapy designed specifically to prevent attacks of hereditary angioedema (HAE) in adult and pediatric patients 12 years and older. One capsule of ORLADEYO per day works to prevent HAE attacks by decreasing the activity of plasma kallikrein.

U.S. Indication and Important Safety Information

INDICATION

ORLADEYO® (berotralstat) is a plasma kallikrein inhibitor indicated for prophylaxis to prevent attacks of hereditary angioedema (HAE) in adults and pediatric patients 12 years and older.

Limitations of use

The safety and effectiveness of ORLADEYO for the treatment of acute HAE attacks have not been established. ORLADEYO should not be used for the treatment of acute HAE attacks. Additional doses or dosages of ORLADEYO higher than 150 mg once daily are not recommended due to the potential for QT prolongation.

IMPORTANT SAFETY INFORMATION

An increase in QT prolongation was observed at dosages higher than the recommended 150 mg once-daily dosage and was concentration dependent.

The most common adverse reactions (≥10% and higher than placebo) in patients receiving ORLADEYO were abdominal pain, vomiting, diarrhea, back pain, and gastroesophageal reflux disease.

A reduced dosage of 110 mg taken orally once daily with food is recommended in patients with moderate or severe hepatic impairment (Child-Pugh B or C).

Berotralstat is a substrate of P-glycoprotein (P-gp) and breast cancer resistance protein. P-gp inducers (eg, rifampin, St. John's wort) may decrease berotralstat plasma concentration, leading to reduced efficacy of ORLADEYO. The use of P-gp inducers is not recommended with ORLADEYO.

ORLADEYO at a dose of 150 mg is a moderate inhibitor of CYP2D6 and CYP3A4. For concomitant medications with a narrow therapeutic index that are predominantly metabolized by CYP2D6 or CYP3A4, appropriate monitoring and dose titration is recommended. ORLADEYO at a dose of 300 mg

is a P-gp inhibitor. Appropriate monitoring and dose titration is recommended for P-gp substrates (eg, digoxin) when coadministering with ORLADEYO.

The safety and effectiveness of ORLADEYO in pediatric patients <12 years of age have not been established.

There are insufficient data available to inform drug-related risks with ORLADEYO use in pregnancy. There are no data on the presence of berotralstat in human milk, its effects on the breastfed infant, or its effects on milk production.

To report SUSPECTED ADVERSE REACTIONS, contact BioCryst Pharmaceuticals, Inc. at 1-833-633-2279 or FDA at 1-800-FDA-1088 or www.fda.gov/medwatch.

Please see full Prescribing Information.

About BioCryst Pharmaceuticals
BioCryst Pharmaceuticals is a global biotechnology company with a deep commitment to improving the lives of people living with complement-mediated and other rare diseases. BioCryst leverages its expertise in structure-guided drug design to develop first-in-class or best-in-class oral small-molecule and protein therapeutics to target difficult-to-treat diseases. BioCryst has commercialized ORLADEYO® (berotralstat), the first oral, once-daily plasma kallikrein inhibitor, and is advancing a pipeline of small-molecule and protein therapies. For more information, please visit www.biocryst.com or follow us on LinkedIn.

Forward-Looking Statements

This press release contains forward-looking statements, including statements regarding future results, performance or achievements. These statements involve known and unknown risks, uncertainties and other factors which may cause actual results, performance or achievements to be materially different from any future results, performance or achievements expressed or implied by the forward-looking statements. These statements reflect our current views with respect to future events and are based on assumptions and are subject to risks and uncertainties. Given these uncertainties, you should not place undue reliance on these forward-looking statements. Some of the factors that could affect the forward-looking statements contained herein include: the ongoing COVID-19 pandemic, which could create challenges in all aspects of BioCryst's business, including without limitation delays, stoppages, difficulties and increased expenses with respect to BioCryst's and its partners' development, regulatory processes and supply chains, negatively impact BioCryst's ability to access the capital or credit markets to finance its operations, or have the effect of heightening many of the risks described below or in the documents BioCryst files periodically with the Securities and Exchange Commission; BioCryst's ability to successfully implement its commercialization plans for, and to commercialize, ORLADEYO, which could take longer or be more expensive than planned; the commercial viability of ORLADEYO, including its ability to achieve market acceptance; the FDA or other applicable regulatory agency may require additional studies beyond the studies planned for products and product candidates, may not provide regulatory clearances which may result in delay of planned clinical trials, may impose certain restrictions, warnings, or other requirements on products and product candidates, may impose a clinical hold with respect to product candidates, or may withhold, delay, or withdraw market approval for products and product candidates; BioCryst's ability to successfully manage its growth and compete effectively; risks related to the international expansion of BioCryst's business; and actual financial results may not be consistent with expectations, including that revenue, operating expenses and cash usage may not be within management's expected ranges. Please refer to the documents BioCryst files periodically with the Securities and Exchange Commission, specifically BioCryst's most recent Annual Report on Form 10-K, Quarterly Reports on Form 10-Q, and Current Reports on Form 8-K, which identify important factors that could cause the actual results to differ materially from those contained in BioCryst's forward-looking statements.

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Contact:
John Bluth
+1 919 859 7910
jbluth@biocryst.com