Nuvalent to Present Pivotal Data from ARROS-1 Clinical Trial of Zidesamtinib for TKI Pre-treated Patients with Advanced ROS1-positive NSCLC at WCLC 2025 Presidential Symposium

CAMBRIDGE, Mass., Aug. 13, 2025 /PRNewswire/ -- Nuvalent, Inc. (Nasdaq: NUVL), a clinical-stage biopharmaceutical company focused on creating *precisely* targeted therapies for clinically proven kinase targets in cancer, today announced that pivotal data for zidesamtinib, a novel ROS1-selective inhibitor, in TKI (tyrosine kinase inhibitor) pre-treated patients with advanced ROS1-positive non-small cell lung cancer (NSCLC) from the global ARROS-1 Phase 1/2 clinical trial, in addition to preliminary data for TKI-naïve patients, will be presented as part of the Presidential Symposium at the IASLC 2025 World Conference on Lung Cancer (WCLC 2025). The conference is hosted by the International Association for the Study of Lung Cancer and is being held September 6-9, 2025, in Barcelona, Spain.

"Since the inception of our ROS1 research program, collaboration with physician-scientists has been central to understanding the specific opportunities for a new therapeutic option to make a meaningful impact for patients living with ROS1-positive NSCLC. This collaboration helped to shape our vision for a future where physicians and their patients facing their next treatment decision do not have to choose between tolerability and efficacy," **said Christopher Turner, M.D., Chief Medical Officer of Nuvalent.** "We are highly encouraged by these pivotal results for zidesamtinib in a TKI pre-treated ROS1-positive NSCLC population as well as the preliminary results in a TKI-naïve population, and look forward to engaging with the global lung cancer community on this important milestone at WCLC."

Details of the presentation are as follows:
Title: Pivotal ARROS-1 efficacy and safety data: zidesamtinib in TKI pre-treated patients with advanced/metastatic ROS1+ NSCLC
Session: PL02 – Presidential Symposium 1

Session Date and Time: Sunday, September 7, 2025, 8:15 a.m. – 10:30 a.m. CEST
Presenting Author: Alexander Drilon, M.D. (Memorial Sloan Kettering Cancer Center, New York, USA)

The company has initiated its rolling NDA submission for zidesamtinib in TKI pre-treated patients with advanced ROS1-positive NSCLC. The FDA agreed to accept the NDA for participation in the Real-Time Oncology Review (RTOR) pilot program, which facilitates earlier submission of topline efficacy and safety results prior to the submission of the complete application to support an earlier start to the FDA's evaluation of the application. Completion of the NDA submission is targeted for the third quarter of 2025, and the company continues to engage with the FDA on potential opportunities for line-agnostic expansion.

About Zidesamtinib and the ARROS-1 Phase 1/2 Clinical Trial
Zidesamtinib is an investigational, novel brain-penetrant ROS1-selective inhibitor created with the aim to overcome limitations observed with currently available ROS1 inhibitors. Zidesamtinib is designed to remain active in tumors that have developed resistance to currently available ROS1 inhibitors, including tumors with treatment-emergent ROS1 mutations such as G2032R. In addition, zidesamtinib is designed for central nervous system (CNS) penetrance to improve treatment options for patients with brain metastases, and to avoid inhibition of the structurally related tropomyosin receptor kinase (TRK) family. Together, these characteristics have the potential to avoid TRK-related CNS adverse events seen with dual TRK/ROS1 inhibitors and to drive deep, durable responses for patients across all lines of therapy. Zidesamtinib has received breakthrough therapy designation for the treatment of patients with ROS1-positive metastatic non-small cell lung cancer (NSCLC) who have been previously treated with 2 or more ROS1 tyrosine kinase inhibitors and orphan drug designation for ROS1-positive NSCLC.

Zidesamtinib is currently being investigated in the ARROS-1 trial (NCT05118789), a first-in-human Phase 1/2 clinical trial for patients with advanced ROS1-positive NSCLC and other solid tumors. The completed Phase 1 portion enrolled ROS1-positive NSCLC patients who previously received at least one ROS1 TKI, or patients with other ROS1-positive solid tumors who had been previously treated. The Phase 1 portion of the trial was designed to evaluate the overall safety and tolerability of zidesamtinib, with additional objectives including determination of the recommended Phase 2 dose (RP2D), characterization of the pharmacokinetic profile, and evaluation of preliminary anti-tumor activity. The ongoing global, single arm, open label Phase 2 portion is designed with registrational intent for TKI-naïve and TKI pre-treated patients with advanced ROS1-positive NSCLC.

About Nuvalent
Nuvalent, Inc. (Nasdaq: NUVL) is a clinical-stage biopharmaceutical company focused on creating *precisely* targeted therapies for patients with cancer, designed to overcome the limitations of existing therapies for clinically proven kinase targets. Leveraging deep expertise in chemistry and structure-based drug design, we develop innovative small molecules that have the potential to overcome resistance, minimize adverse events, address brain metastases, and drive more durable responses. Nuvalent is advancing a robust pipeline with investigational candidates for ROS1-positive, ALK-positive, and HER2-altered non-small cell lung cancer, and multiple discovery-stage research programs.

Forward-Looking Statements
This press release contains forward-looking statements within the meaning of the Private Securities Litigation Reform Act of 1995, as amended, including, without limitation, implied and express statements regarding Nuvalent's strategy, business plans, and focus; the expected timing of data announcements and NDA submissions; the clinical development program for zidesamtinib; the potential benefits and effects of Nuvalent's product development candidates; the design of Nuvalent's clinical trials, including for ARROS-1 its intended pivotal registration-directed design; the potential of Nuvalent's pipeline programs, including zidesamtinib; Nuvalent's research and development programs for the treatment of cancer; and risks and uncertainties associated with drug development. The words "may," "might," "will," "could," "would," "should," "expect," "plan," "anticipate," "aim," "goal," "intend," "believe," "expect," "estimate," "seek," "predict," "future," "project," "potential," "continue," "target" or the negative of these terms and similar words or expressions are intended to identify forward-looking statements, although not all forward-looking statements contain these identifying words. Drug development and commercialization involve a high degree of risk, and only a small number of research and development programs result in commercialization of a product. You should not place undue reliance on these statements or the scientific data presented.

Any forward-looking statements in this press release are based on management's current expectations and beliefs and are subject to a number of risks, uncertainties, and important factors that may cause actual events or results to differ materially from those expressed or implied by any forward-looking statements contained in this press release, including, without limitation: risks that Nuvalent may not fully enroll its clinical trials or that enrollment will take longer than expected; unexpected concerns that may arise from additional data, analysis, or results obtained during preclinical studies or clinical trials; the risk that results of earlier clinical trials may not be predictive of the results of later-stage clinical trials; the risk that data from our clinical trials may not be sufficient to support registration and that Nuvalent may be required to conduct one or more additional studies or trials prior to seeking registration of our product candidates; the occurrence of adverse safety events; risks that the FDA may not approve our potential products on the timelines we expect, or at all; risks of unexpected costs, delays, or other unexpected hurdles; risks that Nuvalent may not be able to nominate drug candidates from its discovery programs; the direct or indirect impact of public health emergencies or global geopolitical circumstances on the timing and anticipated timing and results of Nuvalent's clinical trials, strategy, and future operations; the timing and outcome of Nuvalent's planned interactions with regulatory authorities; and risks related to obtaining, maintaining, and protecting Nuvalent's intellectual property. These and other risks and uncertainties are described in greater detail in the section entitled "Risk Factors" in Nuvalent's Quarterly Report on Form 10-Q for the quarter ended June 30, 2025, as well as any prior and subsequent filings with the Securities and

Exchange Commission. In addition, any forward-looking statements represent Nuvalent's views only as of today and should not be relied upon as representing its views as of any subsequent date. Nuvalent explicitly disclaims any obligation to update any forward-looking statements.

SOURCE Nuvalent, Inc.

Investor Contact:
Chelcie Lister
Nuvalent, Inc.
clister@nuvalent.com

Media Contact:
Josie Butler
1AB
josie@1abmedia.com
