NEWS RELEASE

# Nature Medicine Publishes Two Studies Highlighting Signatera's Clinical Utility in Patients with Gastrointestinal Malignancies

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AMPLIFY-201, a phase I trial evaluating treatment response to the ELI-002 vaccine, is the first study to read out using ctDNA to enroll and monitor patients with solid tumors

PANDA study shows ctDNA status in patients with gastric and gastroesophageal cancers is strongly associated with recurrence risk and pathologic response

AUSTIN, Texas--(BUSINESS WIRE)-- **Natera, Inc.** (NASDAQ: NTRA), a global leader in cell-free DNA testing, today announced the publication of two new studies in Nature Medicine evaluating Natera's personalized and tumor-informed molecular residual disease (MRD) test, Signatera.™

The **first study** shows results from the single-arm, **phase I AMPLIFY-201** trial evaluating the ELI-002 cancer vaccine. The study enrolled a total of 25 patients (20 pancreatic, 5 colorectal), 21 of whom were Signatera-positive after locoregional treatment. Signatera dynamics were also used as a secondary endpoint to assess tumor biomarker response, together with serum tumor antigens. This study is the first of its kind to have utilized circulating tumor DNA (ctDNA)-positivity in conjunction with other tumor biomarkers for enrollment, highlighting the use of Signatera for detecting early evidence of anti-tumor activity of a drug candidate to help assess for a dose response.

The study reported that tumor biomarker responses were observed in 21/25 patients (84%), and ctDNA clearance was observed in 6/25 patients (24%). In contrast, serum tumor antigens declined but did not clear, highlighting the potential utility of ctDNA to be used as a reliable surrogate biomarker for treatment efficacy.

“We are excited about our partnership with Natera, which has resulted in a successful prospective, phase I clinical trial focused on trial enrichment by enrolling Signatera-positive patients and monitoring therapy response in patients receiving a cancer vaccine,” said Christopher Haqq, MD, PhD, Elicio Therapeutic, Inc’s executive vice president, head of research and development, and chief medical officer. “We are optimistic about the potential for this study to improve outcomes for patients with pancreatic and colorectal cancer, who face clinical challenges and are often incurable when ctDNA is detected after treatment.”

A **second study** published in Nature Medicine, called PANDA, is a single-arm, open-label, phase II study that investigated preoperative atezolizumab plus chemotherapy in patients with resectable, non-metastatic gastric and gastroesophageal junction adenocarcinoma. The study included 20 patients who underwent surgery and were evaluated for safety, pathologic response, and survival endpoints, and Signatera was used to detect and monitor ctDNA dynamics in all 20 patients.

The PANDA study found that ctDNA clearance after neoadjuvant therapy correlated with pathologic response in 11/11 patients, while 3/6 patients with poor pathologic response remained ctDNA-positive (P=0.029). None of the patients with complete pathologic response (pCR) were ctDNA-positive, and among the nonresponders who were ctDNA negative (n=3), the study reported superior long-term outcomes. Additionally, ctDNA-positivity at the MRD and follow-up time points was associated with a recurrence rate of 100%.

“We’re pleased to see these excellent results from the PANDA study which demonstrate the strong correlation between ctDNA and pathologic response pre-surgery and the therewith associated recurrence risk, highlighting the potential utility of Signatera in gastric and gastroesophageal cancers,” said Myriam Chalabi MD, PhD, medical oncologist, Netherlands Cancer Institute, and principal investigator of the PANDA study.

“PANDA and AMPLIFY-201 add to the growing number of studies supporting the use of Signatera across various GI indications, showing the promise of tumor-informed ctDNA testing to help personalize treatment strategies,” said Alexey Aleshin, general manager of oncology and early cancer detection at Natera. “These studies also point to the valuable insights longitudinal ctDNA monitoring can provide in the evaluation of novel therapies.”

## About Signatera

**Signatera** is a personalized, tumor-informed, molecular residual disease test for patients previously diagnosed with cancer. Custom-built for each individual, Signatera uses circulating tumor DNA to detect and quantify cancer left in the body, identify recurrence earlier than standard-of-care tools, and help optimize treatment decisions. The test is available for clinical and research use and is covered by Medicare for patients with colorectal cancer, breast cancer (stage IIb and higher) and muscle invasive bladder cancer, as well as for immunotherapy monitoring of any solid

tumor. Signatera has been clinically validated across multiple cancer types and indications, with published evidence in more than 50 peer-reviewed papers.

## About Natera

Natera™ is a global leader in cell-free DNA testing, dedicated to oncology, women's health, and organ health. We aim to make personalized genetic testing and diagnostics part of the standard of care to protect health, and inform earlier, more targeted interventions that help lead to longer, healthier lives. Natera's tests are validated by more than 180 peer-reviewed publications that demonstrate high accuracy. Natera operates ISO 13485-certified and CAP-accredited laboratories certified under the Clinical Laboratory Improvement Amendments (CLIA) in Austin, Texas and San Carlos, California. For more information, visit **www.natera.com**.

## Forward-Looking Statements

All statements other than statements of historical facts contained in this press release are forward-looking statements and are not a representation that Natera's plans, estimates, or expectations will be achieved. These forward-looking statements represent Natera's expectations as of the date of this press release, and Natera disclaims any obligation to update the forward-looking statements. These forward-looking statements are subject to known and unknown risks and uncertainties that may cause actual results to differ materially, including with respect to whether the results of clinical or other studies will support the use of our product offerings, the impact of results of such studies, our expectations of the reliability, accuracy and performance of our tests, or of the benefits of our tests and product offerings to patients, providers and payers. Additional risks and uncertainties are discussed in greater detail in "Risk Factors" in Natera's recent filings on Forms 10-K and 10-Q and in other filings Natera makes with the SEC from time to time. These documents are available at **www.natera.com/investors** and **www.sec.gov**.

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