



PRESS RELEASE

Relief and NeuroRx Conclude Enrollment in their Phase 2b/3 Trial of RLF-100™ for Critical COVID-19 with Respiratory Failure

Geneva, Switzerland and Radnor, Pa, USA, December 30, 2020 – RELIEF THERAPEUTICS Holding AG (SIX: RLF, OTCQB: RLFTF) ("Relief" or the "Company") and NeuroRx, Inc. today announced the conclusion of enrollment in the phase 2b/3 trial of ZYESAMI™ (previously RLF-100™: aviptadil) for the treatment of Respiratory Failure in patients with Critical COVID-19 (www.clinicaltrials.gov NCT04311697). No drug-related serious adverse events have been reported as of today. Enrollment was increased from 165 patients in order to amass as large a safety database as possible. Top line data are expected in late January - early February 2021.

"With FDA's authorization and the extraordinary dedication of our twelve clinical trial sites, we were able to take a drug not formulated or administered to patients in IV form since 2005 and advance it to the clinic in ten weeks. We hope that the highly encouraging results seen in the most critically-ill COVID-19 patients treated in our expanded access program can be replicated in patients who have Critical COVID-19 without an advanced comorbidity" said **Prof. Jonathan Javitt, CEO and founder of NeuroRx, Inc.** "The FDA did not agree to grant EUA, as applied for in September, based upon the open-label study reported earlier, but has advised us that they remain committed to working with us in the development of our product and will promptly review the forthcoming data from this randomized trial. Until that time, available stocks of ZYESAMITM (RLF-100TM) will continue to be administered under our Expanded Access Protocol and individual patient requests under *Right to Try* laws."

Raghuram (Ram) Selvaraju, Chairman of the Board of Relief, said: "We congratulate NeuroRx on the successful completion of patient enrollment in this crucial pivotal clinical study and look forward to the top line results. Our fervent hope is for RLF-100TM to bring benefit to critically ill patients suffering the consequences of COVID-19 infection."

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ABOUT VIP in COVID-19

Vasoactive Intestinal Polypeptide (VIP) was first discovered by the late Dr. Sami Said in 1970. Although first identified in the intestinal tract, VIP is now known to be produced throughout the body and to be primarily concentrated in the lungs. VIP has been shown in more than 100 peer-reviewed studies to have potent anti-inflammatory/anti-cytokine activity in animal models of respiratory distress, acute lung injury, and inflammation. Most importantly, 70% of the VIP in the body is bound to a rare cell in the lung, the alveolar type II cell (ATII), that is critical to transmission of oxygen to the body.

COVID-19-related respiratory failure is caused by selective infection of the ATII cell by the SARS-CoV-2 virus. They are vulnerable because of their (ACE2) surface receptors, which serve as the route of entry for the virus. These specialized cells manufacture surfactant that coats the lung and is essential for oxygen exchange. Loss of surfactant causes collapse of the air sacs (alveolae) in the lung and results in respiratory failure.

VIP is shown to block Coronavirus replication in the ATII cell, block cytokine synthesis, block viral-induced cell death (cytopathy), and upregulate surfactant production. Other than RLF-100™, no currently proposed treatments for COVID-19 specifically target these vulnerable Type II cells.





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ABOUT ZYESAMI™ (PREVIOUSLY RLF-100™: AVIPTADIL)

ZYESAMI™ (RLF-100TM: AVIPTADIL) IS A FORMULATION OF VASOACTIVE INTESTINAL POLYPEPTIDE (VIP) RLF-

100™ (Aviptadil) is a formulation of Vasoactive Intestinal Polypeptide (VIP) that was developed based on Prof. Sami Said's original work at Stony Brook University, for which Stony Brook was awarded an FDA Orphan Drug Designation in 2001. VIP is known to be highly concentrated in the lungs, where it inhibits coronavirus replication, blocks the formation of inflammatory cytokines, prevents cell death, and upregulates the production of surfactant. FDA has now granted IND authorization for intravenous and inhaled delivery of RLF-100™ for the treatment of COVID-19 and awarded Fast Track designation. RLF-100™ is being investigated in two placebo-controlled US phase 2b/3 clinical trials in respiratory deficiency due to COVID-19. Since July 2020, more than 300 patients with Critical COVID-19 and Respiratory Failure have been treated with RLF-100™ between the two FDA-approved protocols (randomized and expanded access). Information on the RLF-100™ Expanded Access program is at https://www.neurorxpharma.com/our-services/rlf-100.

ABOUT RELIEF

Relief focuses primarily on clinical-stage programs based on molecules of natural origin (peptides and proteins) with a history of clinical testing and use in human patients or a strong scientific rationale. Currently, Relief is concentrating its efforts on developing new treatments for respiratory disease indications. Its lead drug candidate RLF-100TM (aviptadil) is being investigated in two placebo-controlled U.S. phase 2b/3 clinical trials in respiratory deficiency due to COVID-19. Relief also holds a patent issued in the United States and various other countries covering potential formulations of RLF-100TM.

RELIEF THERAPEUTICS Holding AG is listed on the SIX Swiss Exchange under the symbol RLF and quoted in the U.S. on OTCQB under the symbol RLFTF.

www.relieftherapeutics.com

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ABOUT NEURORX, INC.

NeuroRx draws upon more than 100 years of collective drug development experience and by former senior executives of AstraZeneca, Eli Lilly, Novartis, Pfizer, and PPD. In addition to its work on ZYESAMI[™], which has been awarded FDA Fast Track designation (previously RLF-100[™]: Aviptadil), NeuroRx has been awarded Breakthrough Therapy Designation and a Special Protocol Agreement to develop NRX-101 in suicidal bipolar depression and is currently in Phase 3 trials. Its executive team is led by Prof. Jonathan C. Javitt, MD, MPH, who has served as a health advisor to four Presidential administrations and worked on paradigm-changing drug development projects for Merck, Allergan, Pharmacia, Pfizer, Novartis, and Mannkind, together with Robert Besthof, MIM, who served as the Global Vice President (Commercial) for Pfizer's Neuroscience and Pain Division. The Company has recently announced a plan to merge with Big Rock Partners Acquisition Corp (NASDAQ:BRPA), following which it plans to trade on the NASDAQ as NRXP.





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