Ad hoc announcement pursuant to Art. 53 LR

Relief Therapeutics Announces IRB Approval and Initiation of an Investigator Initiated Trial of Nexodyn for Epidermolysis Bullosa at Ann & Robert H. Lurie Children's Hospital of Chicago

Professor Amy Paller, M.D., will serve as Principal Investigator to evaluate Nexodyn for the management of colonized dystrophic and junctional Epidermolysis Bullosa wounds, an important unmet need in this group of rare inherited skin diseases that affects approximately 250,000 patients worldwide.

Geneva, Switzerland, September 22, 2022 – RELIEF THERAPEUTICS Holding SA (SIX: RLF, OTCQB: RLFTF, RLFTY) ("**Relief**"), a biopharmaceutical company identifying, developing and commercializing novel, patent protected products in selected specialty, rare and ultra-rare disease areas on a global basis, announced today that an Investigator Initiated Trial with Nexodyn for Epidermolysis Bullosa (EB) has received IRB approval and will shortly enroll participants at Ann & Robert H. Lurie Children's Hospital of Chicago with Professor Amy Paller, M.D., Chair, Department of Dermatology, Feinberg School of Medicine, Northwestern University, serving as Principal Investigator. This trial represents an important step in the clinical development pathway of Relief's APR-TD011, a pharmaceutical grade hypochlorous acid topical spray being evaluated for decolonization, symptom alleviation, and healing of EB wounds.

Professor Amy Paller stated "EB is a rare, inherited skin disease characterized by widely distributed, chronic wounds that easily become infected with a risk of sepsis and death. As there is no cure for EB, a crucial element of patient management involves proper and timely wound care. This pilot clinical study will evaluate how the bactericidal activity of this differentiated hypochlorous acid wound spray that has been shown to kill methicillin-sensitive and methicillin-resistant Staphylococcus aureus, as well as Pseudomonas aeruginosa, could reduce wound colonization, thus improving the microbiome. Future studies will determine the impact of APR-TD011 on infection control, avoidance of chronic antibiotic use, accelerated wound healing, and quality of life for patients living with EB."

Dr. Nermeen Varawalla, Chief Medical Officer of Relief Therapeutics, noted, "This proof-of-concept clinical trial, to be conducted in 15 participants over a 12-week study period, will be most valuable for the swift, effective and efficient execution of Relief's clinical development plan for APR-TD011. The clinical data from this Investigator Initiated Trial will facilitate the design and conduct of follow-on, multi-center, pivotal clinical trials for future, potential U.S. Food and Drug Administration (FDA) and European Medicines Agency (EMA) approval of APR-TD011 as an effective, convenient, well tolerated, anti-microbial wound management solution for EB."

ABOUT Nexodyn and APR-TD011

Nexodyn, engineered by the TEHCLO® proprietary technology, is a highly pure and stabilized hypochlorous acid (HClO >95% of free chlorine species), with pH between 2.5 - 3.0 and high Reduction-Oxidation Potential (ORP 1.000 - 1.200 mV). It is presented as a self-administered sprayable solution intended for use in the debridement, irrigation, cleansing, and moistening of acute and chronic wounds and is certified in the EU as a Class III medical device and, in the US, as a 510(k) cleared unclassified device. The spray formulation enables wound application whilst avoiding skin contact and cross-contamination, of a solution that has been consistently shown to accelerate wound closure with reduced infection rates.

APR-TD011, manufactured with the company's proprietary TECHLO® technology, is a GMP grade pharmaceutical hypochlorous acid wound solution that has been granted an FDA Orphan Drug Designation for EB. If approved, it would be the first product specifically indicated to prevent or reduce colonization and infections in EB wounds, and via modulation of the wound microbiome accelerate wound healing and closure, while reducing antibiotic use. This, along with its anti-inflammatory action, could provide symptom relief and wound healing.

EB is a group of rare, genetic, life-threatening connective tissue disorders characterized by fragile skin and mucous membrane with severe blistering throughout the body. There are currently an estimated 250,000 patients with EB worldwide, with an estimated 30,000 patients in the European Union and 20,000 patients in the U.S.

ABOUT RELIEF THERAPEUTICS

Relief is a Swiss, commercial-stage, biopharmaceutical company focused on identification, development and commercialization of novel, patent protected products intended for the treatment of rare and ultra-rare diseases including metabolic disorders, pulmonary diseases, and connective tissue disorders. Relief's diversified pipeline consists of assets that have the potential to effectively address significant unmet medical needs, including PKU GOLIKE®, engineered with the proprietary Physiomimic technology, which is the first prolonged-release amino acid product commercialized for the dietary management of phenylketonuria ("PKU"). Relief has a Collaboration and License Agreement with Acer Therapeutics for the worldwide development and commercialization of ACER-001 (sodium phenylbutyrate) for the treatment of various inborn errors of metabolism, including Urea Cycle Disorders ("UCDs") and Maple Syrup Urine Disease ("MSUD"). Relief also continues to develop aviptadil for several rare pulmonary indications. Further, Relief is undertaking the clinical development of APR-TD011, a differentiated acid oxidizing solution of hypochlorous acid intended for the treatment of epidermolysis bullosa ("EB"), a group of rare, genetic, life-threatening connective tissue disorders; APR-TD011 has been granted Orphan Drug Designation by the FDA. Finally, Relief is commercializing several legacy products via licensing and distribution partners.

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

For more information, visit www.relieftherapeutics.com. Follow Relief on LinkedIn.

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