Press Release

Nicox’s Partner Fera Pharmaceuticals to Investigate Naproxcinod as Potential Covid-19 Adjuvant Treatment

December 11, 2020 – release at 7:30 am CET
Sophia Antipolis, France

Nicox SA (Euronext Paris: FR0013018124, COX), an international ophthalmology company, and Fera Pharmaceuticals, a privately-held, U.S. specialty pharmaceutical company, announced today that Fera will evaluate naproxcinod as a potential adjuvant treatment for patients with COVID-19 infection. Subject to successful completion of the ongoing manufacturing of naproxcinod test material, Fera plans to initiate pre-clinical proof-of-concept studies in models of COVID-19 infection in early 2021.

Naproxcinod, a Cyclooxygenase-Inhibiting Nitric Oxide (NO)-Donating (CINOD) naproxen, is a non-steroidal anti-inflammatory product candidate engineered to release NO and naproxen, originally discovered and developed by Nicox. Nicox and Fera entered into an agreement in December 2015 which granted Fera exclusive rights to develop and commercialize naproxcinod for the U.S. market. Nicox and Fera are amending their existing agreement to include COVID-19 as an indication, and Nicox will grant to Fera warrants1 to acquire 10,000 Nicox shares.

Michele Garufi, Chief Executive Officer and Chairman of Nicox, said: “There is a strong scientific rationale for using naproxcinod in the treatment of the inflammatory symptoms of COVID-19 infections, and potentially against the virus itself. In collaboration with Fera we will be testing this scientific rationale in relevant pre-clinical models and whilst this research is at an early stage, any potential future human trial would benefit from the extensive naproxcinod clinical data previously generated by Nicox for the treatment of signs and symptoms of osteoarthritis.”

Rationale for naproxcinod in COVID-19 treatment

Most outcomes of COVID-19 are associated with high levels of inflammation and dysfunction of the vascular system leading to thrombotic events2. Naproxcinod, a Cox-Inhibiting Nitric Oxide Donor (CINOD), would potentially treat multiple aspects of COVID-19 infection including fever, pain, inflammation and platelet aggregation, thus decreasing the risk of thrombus formation. In addition, NO donation might increase vasodilation and restore normal vascular functions. Moreover, NO has specifically been demonstrated to inhibit replication of the COVID-19 virus by two distinct mechanisms3. As an oral capsule formulation, naproxcinod could be easily administered to patients at the first signs of infection.

Once Fera has received the newly manufactured naproxcinod, they plan to initiate proof-of-concept pre-clinical tests in models of COVID-19 infection. Should the results of these studies prove positive, Fera plans to meet with the U.S. Food and Drug Administration (FDA) to identify the clinical trials that would be required to submit a New Drug Application (NDA) for naproxcinod in the treatment of COVID-19 infection.

1 The warrants will be issued at no cost. The subscription price of the new shares to be obtained by exercising the warrants will be equal to the VWAP calculated on the 3-days trading prior to the Board meeting that will decide on the issuance of the warrants.


www.nicox.com
Under the terms of the naproxcinod agreement with Nicox, Fera is responsible for all clinical development, manufacturing, regulatory and commercialization activities in the U.S. Nicox retains all rights to naproxcinod outside the U.S., subject to the payment of royalties to Fera, if intellectual property developed under the agreement is used outside the U.S.

Fera continues to review non-COVID-19 development options for naproxcinod, including addressing the U.S. FDA refusal letter concerning Fera’s application for Orphan Drug Designation (ODD) of naproxcinod in sickle-cell disease.

**About naproxcinod**

Naproxcinod is a nitric oxide (NO)-donating naproxen combining the cyclooxygenase (COX) inhibitory activity of naproxen with that of NO (COX-inhibiting NO donor, CINOD). While the inhibitory COX component provides the analgesic and anti-inflammatory efficacy, the NO part may play a significant role in maintaining vascular endothelial cell function and integrity, blood pressure homeostasis and microvascular circulation. A broad clinical package already exists for naproxcinod in osteoarthritis, including three phase 3 trials with over 2,700 patients.

**About Fera Pharmaceuticals**

Fera Pharmaceuticals is a privately held company. The company goal is to realize opportunities via acquisitions, in-licensing, developing and marketing abbreviated new drug applications (ANDAs), new drug applications (NDAs) and 505(b)(2) NDA products. Areas of interest include products that could benefit from lifecycle management with a special focus on niche markets. For more information visit [www.ferapharma.com](http://www.ferapharma.com).

**About Nicox**

Nicox S.A. is an international ophthalmology company developing innovative solutions to help maintain vision and improve ocular health. Nicox's lead program in clinical development is NCX 470, a novel, second-generation nitric oxide-donating bimatoprost analog, for lowering intraocular pressure in patients with glaucoma. The company is also developing NCX 4251, a proprietary formulation of fluticasone, for acute exacerbations of blepharitis. Nicox generates revenue from VYZULTA® in glaucoma, licensed exclusively worldwide to Bausch + Lomb, and ZERVIATE™ in allergic conjunctivitis, licensed in multiple geographies, including to EyeVance Pharmaceuticals, LLC, in the U.S. and Ocumen Therapeutics in the Chinese and in the majority of South East Asian markets.

Nicox is headquartered in Sophia Antipolis, France, is listed on Euronext Paris (Compartment B: Mid Caps; Ticker symbol: COX) and is part of the CAC Healthcare, CAC Pharma & Bio and Next 150 indexes.

For more information on Nicox, its products or pipeline, please visit: [www.nicox.com](http://www.nicox.com).

**Analyst coverage**

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Risks factors which are likely to have a material effect on Nicox’s business are presented (i) in the 3rd chapter of the ‘Document d’enregistrement universel, rapport financier annuel et rapport de gestion 2019’ filed with the French Autorité des Marchés Financiers (AMF) on March 6, 2020 which are available on Nicox’s website (www.nicox.com) and (ii) as restated in the 4th chapter of the half yearly financial report as of June 30, 2020, which is also available on Nicox’s website.