



ORYZON GENOMICS, S.A.

Pursuant to the provisions of article 227 of the Restated Text of the Securities Market Act approved by Royal Legislative Decree 4/2015 of 23 October, ORYZON GENOMICS, S.A. ("**ORYZON**" or the "**Company**") hereby gives notice of the following

OTHER RELEVANT INFORMATION

ORYZON announces that it has received approval from the Spanish Drug Agency (AEMPS) to conduct a Phase II clinical trial with vafidemstat in severely ill COVID-19 patients.

The pressrelease that will be distributed today is attached

Madrid, 24 April 2020

ORYZON to initiate ESCAPE: a Phase II clinical trial to test efficacy of vafidemstat in severely ill COVID-19 patients

- ❖ **CTA approval, the European IND equivalent, granted by the Spanish Drug Agency via an accelerated procedure**
- ❖ **Primary endpoint: To investigate the efficacy of vafidemstat, in combination with standard of care treatment, to prevent Acute Respiratory Distress Syndrome (ARDS) in adult severely ill COVID-19 patients**

MADRID, SPAIN and CAMBRIDGE, MA, UNITED STATES, April 24th, 2020 - Oryzon Genomics, S.A. (ISIN Code: ES0167733015, ORY), a public clinical-stage biopharmaceutical company leveraging epigenetics to develop therapies in diseases with strong unmet medical need, announced today that under certain urgency provisions related to the COVID-19 pandemic it has received approval from the Spanish Drug Agency (AEMPS) to conduct a Phase II clinical trial with vafidemstat in seriously ill COVID-19 patients.

The study, named ESCAPE (*Efficacy and Safety of a Combined treatment with vafidemstat to prevent ARDS in adult Patients with severe COVID-19*) (EudraCT No.: 2020-001618-39), is an open-label, randomized, double-arm Phase II trial to assess the efficacy and tolerability of vafidemstat in combination with standard of care treatment, to prevent progression to Acute Respiratory Distress Syndrome (ARDS). Initially, it is planned to include 20 patients in each arm of the trial. In principle, initially two hospitals will participate (H. Valle de Hebrón and H. del Mar, both in Barcelona, Spain), while more centers can be added if necessary.

ESCAPE's aim is to explore a therapeutic intervention to prevent progression to ARDS, one of the main causes of death in severe COVID-19 patients. ARDS is the result of a so-called "cytokine storm", a violent systemic reaction to the infection that frequently leads to a multi-organ failure. As was the case in the previous MERS-CoV epidemic, in the COVID-19 pandemic it has been observed that the cytokines Interleukin-6 (IL-6) and Interleukin-1B (IL-1B) are central to triggering this cytokine storm. In acute preclinical models of inflammation vafidemstat has been shown to produce a rapid and strong decrease in IL-6, IL-1B and other relevant immunomodulatory inflammatory cytokines such as TNF- α and IFN- γ . In a recent clinical study with vafidemstat in the elderly population (ETHERAL Study in Alzheimer's disease) it has been shown to be very safe in long-term treatments, as well as demonstrating a significant decrease in a relevant marker of brain inflammation.

Carlos Buesa, President and CEO of Oryzon, said: "While we are not infectious diseases specialists, in these troubling times we feel compelled to initiate this study to potentially find a way to help these critically ill patients, both in Spain and abroad. We believe we have a solid rationale for vafidemstat in this indication. We wish to warmly thank the AEMPS and the heads of infectious disease departments of

various Spanish hospitals for their scientific and clinical advice, which has allowed us to design in a record time a protocol with such a strong scientific rationale. Provided the results of the ESCAPE study are encouraging, we should be able to quickly increase the number of participating patients.”

About Oryzon

Founded in 2000 in Barcelona, Spain, Oryzon (ISIN Code: ES0167733015) is a clinical stage biopharmaceutical company considered as the European champion in Epigenetics. Oryzon has one of the strongest portfolios in the field. Oryzon’s LSD1 program has rendered two compounds, vafidemstat and iadademstat, in clinical trials. In addition, Oryzon has ongoing programs for developing inhibitors against other epigenetic targets. Oryzon has a strong technological platform for biomarker identification and performs biomarker and target validation for a variety of malignant and neurological diseases. Oryzon has offices in Spain and the United States. For more information, visit www.oryzon.com

About Vafidemstat

Vafidemstat (ORY-2001) is an oral, CNS optimized LSD1 inhibitor. The molecule acts on several levels: it reduces cognitive impairment, including memory loss and neuroinflammation, and at the same time has neuroprotective effects. In animal studies vafidemstat not only restores memory but reduces the exacerbated aggressiveness of SAMP8 mice, a model for accelerated aging and Alzheimer’s disease (AD), to normal levels and also reduces social avoidance and enhances sociability in murine models. In addition, vafidemstat exhibits fast, strong and durable efficacy in several preclinical models of multiple sclerosis (MS). Oryzon has performed a Phase IIa clinical trial in aggressiveness in patients with different psychiatric disorders (REIMAGINE) and in aggressive/agitated patients with moderate or severe AD (REIMAGINE-AD), with positive preliminary clinical results reported. Additional Phase IIa clinical trials with vafidemstat are ongoing in patients with Mild to Moderate AD (ETHERAL), where a significant reduction of the inflammatory biomarker YKL40 has been observed after 6 months of treatment, and in Relapse-Remitting and Secondary Progressive MS (SATEEN).

FORWARD-LOOKING STATEMENTS

This communication contains, or may contain, forward-looking information and statements about Oryzon, including financial projections and estimates and their underlying assumptions, statements regarding plans, objectives and expectations with respect to future operations, capital expenditures, synergies, products and services, and statements regarding future performance. Forward-looking statements are statements that are not historical facts and are generally identified by the words “expects,” “anticipates,” “believes,” “intends,” “estimates” and similar expressions. Although Oryzon believes that the expectations reflected in such forward-looking statements are reasonable, investors and holders of Oryzon shares are cautioned that forward-looking information and statements are subject to various risks and uncertainties, many of which are difficult to predict and generally beyond the control of Oryzon that could cause actual results and developments to differ materially from those expressed in, or implied or projected by, the forward-looking information and statements. These risks and uncertainties include those discussed or identified in the documents sent by Oryzon to the Spanish Comisión Nacional del Mercado de Valores (CNMV), which are accessible to the public. Forward-looking statements are not guarantees of future performance and have not been reviewed by the auditors of Oryzon. You are cautioned not to place undue reliance on the forward-looking statements, which speak only as of the date they were made. All subsequent oral or written forward-looking statements attributable to Oryzon or any of its members, directors, officers, employees or any persons acting on its behalf are expressly qualified in their entirety by the cautionary statement above. All forward-looking statements included herein are based on information available to Oryzon on the date hereof. Except as required by applicable law, Oryzon does not undertake any obligation to publicly update or revise any forward-looking statements, whether as a result of new information, future events or otherwise. This press release is not an offer of securities for sale in the United States or any other jurisdiction. Oryzon’s securities may not be offered or sold in the United States absent registration or an exemption from registration. Any public offering of Oryzon’s securities to be made in the United States will be made by means of a prospectus that may be obtained from Oryzon or the selling security holder, as applicable, that will contain detailed information about Oryzon and management, as well as financial statements.

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