



ORYZON GENOMICS, S.A.

Pursuant to the provisions of article 228 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

SIGNIFICANT FACT

ORYZON announces that it has received approval from the UK Medicines Agency (MHRA) to conduct a Phase IIa clinical trial with ORY-2001 in patients of Alzheimer's Disease (AD).

The pressrelease that will be distributed today is attached.

Madrid, 29 May 2018

ORYZON receives approval in the United Kingdom to start ETHERAL: a Phase IIa clinical trial in Alzheimer's Disease with ORY-2001

This trial was approved in the previous weeks in Spain and France

This approval concludes the regulatory start of this trial in Europe

This is the second Phase II study in CNS with the drug

MADRID, SPAIN and CAMBRIDGE, MA, May 29, 2018 – Oryzon Genomics (ISIN Code: ES0167733015, ORY), a public clinical-stage biopharmaceutical company leveraging epigenetics to develop therapies in diseases with strong unmet medical need, has announced today that it has received approval of a Clinical Trial Application (CTA), the European IND equivalent, from the Medicines & Healthcare products Regulatory Agency (MHRA) and also from the Ethics Committee to conduct a Phase IIa clinical study with ORY-2001 in patients of Alzheimer's disease (AD) in the UK. The trial had been previously approved in Spain and France. With this approval, the trial is now fully operationally deployed upon receipt from the corresponding regulatory authorities of all initially scheduled approvals.

The study, named ETHERAL (**E**pigenetic **T**HERapy in **AL**zheimer's Disease), is being conducted in different European hospitals across Spain, France and UK, and is designed as a randomised, double-blind, placebo-controlled, 3-arm, 26 weeks parallel-group study to evaluate the safety and tolerability of ORY-2001 in patients with mild and moderate Alzheimer's disease. The study will involve 90 patients and incorporates measurements in the different domains of the disease as secondary endpoints, including memory and behavior alterations. It will also monitor the variations of diverse, yet significant CSF biomarkers. The company is aiming to launch a twin study in the US soon.

ORY-2001 is an oral and brain penetrant drug that selectively inhibits LSD1 and MAOB. The molecule acts on several levels, reduces cognitive impairment, memory loss and neuroinflammation, and at the same time has neuroprotective effects. The company has recently reported in several scientific conferences that ORY-2001 exerts in preclinical models a holistic action on different types of alterations also seen in patients with AD and other neurodegenerative disorders. ORY-2001 may act as a disease modifying drug. In AD patients and other neurodegenerative disorders, cognitive deterioration is often accompanied by episodes of agitation, aggression, psychosis, apathy and depression. In preclinical studies, ORY-2001 not only restores memory but reduces the exacerbated aggressiveness of SAMP8 mice, a model for accelerated aging and Alzheimer's disease, to normal levels and also reduces social avoidance and enhances sociability in various murine models. In addition, ORY-2001 exhibits fast, strong and durable efficacy in several preclinical models of multiple sclerosis (MS). The company has already started a Phase IIa clinical study with ORY-2001 in patients with MS.

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. The company has one of the strongest portfolios in the field. Oryzon's LSD1 program has rendered two compounds in clinical trials. In addition, Oryzon has ongoing programs for developing inhibitors against other epigenetic targets. The company has a strong technological platform for biomarker identification and performs biomarker and target validation for a variety of malignant and neurodegenerative diseases. The company has offices in Spain and USA. For more information, visit www.oryzon.com.

FORWARD-LOOKING STATEMENTS

This communication contains forward-looking information and statements about Oryzon Genomics, S.A., 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 Genomics, S.A. believes that the expectations reflected in such forward-looking statements are reasonable, investors and holders of Oryzon Genomics, S.A. 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 Genomics, S.A., 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 Genomics, S.A. to the *Comisión Nacional del Mercado de Valores*, which are accessible to the public. Forward-looking statements are not guarantees of future performance. The auditors of Oryzon Genomics, S.A. have not reviewed them. 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 Genomics, S.A. 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 Genomics, S.A. on the date hereof. Except as required by applicable law, Oryzon Genomics, S.A. 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. The Company's securities may not be offered or sold in the United States absent registration or an exemption from registration. Any public offering of the Company's securities to be made in the United States will be made by means of a prospectus that may be obtained from the Company or the selling security holder, as applicable, that will contain detailed information about the Company and management, as well as financial statements.

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