**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

# MATERIAL FACT

ORYZON announces that preliminary efficacy data from the first patient in its iadademstat's Phase IIa CLEPSIDRA clinical trial in small cell lung cancer will be presented today at the IASLC 20th World Conference on Lung Cancer (WCLC).

These data are summarized in the attached pressrelease that will be distributed today.

Madrid, 9 September 2019

# ORYZON's iadademstat shows efficacy signs in relapsed SCLC

- **Preliminary data from CLEPSIDRA Phase II trial presented at WCLC 2019**
- **Case study showing a 86% tumor reduction**
- **Additional data will be presented late September at ESMO-2019**

**MADRID, SPAIN and CAMBRIDGE, MA, UNITED STATES, September 9th, 2019 -** 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 needs, today presents the first preliminary data from CLEPSIDRA, a Phase II trial investigating iadademstat in combination with standard-of-care in relapsing small cell lung cancer (SCLC) patients.

The company will present a poster, at the IASLC 20th World Conference on Lung Cancer (WCLC), being held in Barcelona, Spain, entitled "*Iadademstat shows preliminary efficacy signals in relapsed ED-SCLC patients: A case report within CLEPSIDRA, a Phase II trial of iadademstat in combination with platinum-etoposide in biomarker-positive patients*".

The preliminary data, corresponding to the patient with the longest treatment period in the study, show that:

- Administration of the combination of iadademstat with standard-of-care during the first 6 cycles produced a tumor reduction of 78.7%, as defined by RECIST criteria
- The only toxicity of the combination was hematological
- Subsequent administration of iadademstat alone for four consecutive cycles proved to be safe and well tolerated without producing any hematological toxicity
- Reduction of main lesions and metastasis continues with iadademstat in monotherapy, reaching an overall tumor reduction of 86.3% by RECIST criteria according to CT scan evaluation in Cycle 8.

Dr. Roger Bullock, Oryzon's Chief Medical Officer stated "Although this Phase II trial is just starting and we have only a limited number of patients in the study, the impressive tumor reduction observed in this first patient is very promising. These data show that iadademstat has the potential to make a difference to patients with SCLC, and we are looking forward to presenting more data from the trial with this exciting drug candidate in the coming weeks."

The company plans to present additional data from the first 10 patients of the CLEPSIDRA study during the ESMO-2019 International Conference, to be held from September 27 to October 1 in Barcelona.

CLEPSIDRA ("*A **C**ombination trial of **L**SD1 and **E**top-**P**latinum in **S**mall Cell Lung Cancer in Biomarker-**ID** **R**elapsed p**A**tients)* is a Phase IIa trial of iadademstat, being conducted in several hospitals in Spain. CLEPSIDRA is enrolling relapsed ED-SCLC patients and is designed as a single-arm, open-label study of iadademstat in combination with the standard of care treatment with platinum/etoposide, in order to evaluate the safety and tolerability as well as the clinical effect (including time to response, duration of response, objective response and overall survival) of the combination. Patients are stratified by certain proprietary biomarkers characterized by their ability to identify LSD1i-sensitive SCLC tumors. The study is divided into two parts, the first one to optimize the dose of the combination, and the second one to evaluate the combination's efficacy. The study is designed to recruit up to 36 patients.

WCLC 2019 is held from September 7 to 10. For more info about the congress, please visit WCLC 2019's website

A copy of the poster is available here

**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 neurodegenerative diseases. Oryzon has offices in Spain and the United States. For more information, visit www.oryzon.com

**About Iadademstat**

Iadademstat (ORY-1001) is a small oral molecule, which acts as a highly selective inhibitor of the epigenetic enzyme LSD1 and has a powerful differentiating effect in hematologic cancers (See Maes et al., Cancer Cell 2018 Mar 12; 33 (3): 495-511.e12.doi: 10.1016 / j.ccell.2018.02.002.). A first Phase I/IIa clinical trial with iadademstat in refractory and relapsed acute leukemia patients demonstrated the safety and good tolerability of the drug and preliminary signs of antileukemic activity, including a CRi. Beyond hematological cancers, the inhibition of LSD1 has been proposed as a valid therapeutic approach in some solid tumors such as small cell lung cancer (SCLC), medulloblastoma and others. Oryzon is conducting two Phase IIa clinical trials of iadademstat in combination; the first one in combination with azacitidine in elderly AML patients (ALICE study) and the second one in combination with platinum/etoposide in second line SCLC patients (CLEPSIDRA study).

**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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