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ORYZON announces first patient dosed in an Investigator-initiated Phase Ib study of iadademstat in extensive stage small cell lung cancer

- **Exploring the combination with atezolizumab and radiation therapy**
- **Study led by Yale University**

MADRID, SPAIN and CAMBRIDGE, MA, UNITED STATES, February 9, 2026 - Oryzon Genomics, S.A. (ISIN Code: ES0167733015, ORY), a clinical-stage biopharmaceutical company and a global leader in epigenetics, announced today that the first patient has been dosed in an investigator-initiated Phase Ib dose-finding trial evaluating iadademstat, Oryzon's potent and selective LSD1 inhibitor, in combination with radiotherapy and an immune checkpoint inhibitor in patients with residual, progressive or recurrent extensive stage small cell lung cancer (ES-SCLC). The study is sponsored and conducted by Yale University.

The trial ([NCT07113691](#)), titled "*Iadademstat and Radiation Therapy With Atezolizumab in Extensive Stage Small-cell Lung Cancer (ES-SCLC) Patients With Persistent, Recurrent or Progressive Disease After First Line Systemic Therapy*", is an open-label, non-randomized Phase Ib study that will evaluate the safety, tolerability, and efficacy of iadademstat combined with atezolizumab and stereotactic body radiation therapy (SBRT) followed by maintenance therapy with atezolizumab and iadademstat. The study will enroll patients with residual, progressive or recurrent ES-SCLC who previously received platinum-based chemotherapy with or without immune checkpoint inhibitor therapy. Dr. Anne Chiang at Yale University is serving as Principal Investigator.

Dr. Carlos Buesa, Oryzon's CEO, stated: "The initiation of this investigator-led study at Yale University represents an important milestone in our efforts to expand the clinical development of iadademstat in small cell lung cancer. Combining LSD1 inhibition with immunotherapy and radiotherapy is a compelling strategy, and we look forward to the insights this trial may provide into the potential role of iadademstat in addressing this aggressive disease with high unmet medical need."

Dr. Chiang, Principal Investigator of the study, added: "Some SCLC patients have had long-term benefit with immunotherapy, and we hope to learn how to extend benefit to more people. Through this study, we aim to boost the immune response by adding radiation and iadademstat to immunotherapy in SCLC patients. The use of paired biopsies will help us understand key events that are occurring in the tumor and its microenvironment as well."

About Oryzon

Founded in 2000 in Barcelona, Spain, Oryzon (ISIN Code: ES0167733015) is a clinical stage biopharmaceutical company and the European leader in epigenetics, with a strong focus on personalized medicine in CNS disorders and oncology. Oryzon's team is composed of highly qualified professionals from the pharma industry located in Barcelona, Boston, and San Diego. Oryzon has an



advanced clinical portfolio with two LSD1 inhibitors, vafidemstat in CNS (Phase III-ready) and iadademstat in oncology (Phase II). The company has other pipeline assets directed against other epigenetic targets like HDAC-6 where a clinical candidate ORY-4001, has been nominated for its possible development in CMT and ALS. In addition, Oryzon has a strong platform for biomarker identification and target validation for a variety of malignant and neurological diseases. 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 FIM Phase I/IIa clinical trial with iadademstat in R/R AML patients demonstrated the safety and good tolerability of the drug and preliminary signs of antileukemic activity, including a CRi (see Salamero et al, J Clin Oncol, 2020, 38(36): 4260-4273. doi: 10.1200/JCO.19.03250). Iadademstat has shown encouraging safety and strong clinical activity in combination with azacitidine in a Phase IIa trial in elder 1L AML patients (ALICE trial) (see Salamero et al., ASH 2022 oral presentation & The Lancet Haematology, 2024, 11(7):e487-e498). Iadademstat is currently being evaluated in combination with azacitidine and venetoclax in 1L AML in an investigator-initiated study (IIS) led by OHSU and in combination with gilteritinib in the company-sponsored Phase Ib FRIDA trial in relapsed/refractory FLT3-mutant AML, with highly encouraging preliminary safety and efficacy data recently reported at ASH-2025 for both trials: 100% ORR and 90% strict CR in 1L AML, and 67% CCR (at the dose under expansion) in R/R AML. Additional studies in hemato-oncology include an IIS in MDS, and trials in myeloproliferative neoplasms and 1L AML both sponsored and conducted by the U.S. National Cancer Institute (NCI) under a Cooperative Research and Development Agreement (CRADA) signed between Oryzon and the NCI. 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), neuroendocrine tumors (NET), medulloblastoma and others. In a Phase IIa trial in combination with platinum/etoposide in second line ED-SCLC patients (CLEPSIDRA trial), preliminary activity and safety results have been reported (see Navarro et al., ESMO 2018 poster). Iadademstat is in a Phase I/II randomized trial in 1L ED-SCLC in combination with ICI sponsored by NCI and led by the Memorial Sloan Kettering Cancer Center. In addition, Oryzon is expanding iadademstat's clinical development into non-oncological hematology indications, with trials in sickle cell disease (approved by EMA, enrolling) and essential thrombocythemia (submitted to EMA). Iadademstat has orphan drug designation for SCLC in the US and for AML in the US and EU.

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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