

24 April 2025 • Press Release

ORYZON Raises €30 Million in Capital Increase

- Issued 12,765,958 new shares
- Priced at €2.35 per share, representing a 15.44% discount on the 5-day
 VWAP
- Strong investor demand, with the offering significantly oversubscribed
- €15 million anchored by a leading US institutional investor
- Proceeds to fund clinical development and corporate initiatives

MADRID, SPAIN and CAMBRIDGE, MA, UNITED STATES, April 24th, 2025 - Oryzon Genomics, S.A. (ISIN Code: ES0167733015, ORY), a clinical-stage biopharmaceutical company and a European leader in epigenetics, announced today the successful completion of a \leq 30 million capital increase through the issuance of 12,765,958 new shares, priced at \leq 2.35 per share. This pricing reflects a 15.44% discount to the 5-day volume-weighted average price (VWAP) of \leq 2.779 per share prior to April 22, 2025, and a 19.38% discount to the April 22 closing price of \leq 2.915. The offering, which was structured as straight equity with no warrants attached, attracted strong demand and was significantly oversubscribed. A US-based institutional investor anchored the round with a \leq 15 million order, with the remaining demand filled by investors across the US, Europe, and Spain. The capital increase was capped at \leq 30 million by the company's Board of Directors.

Maxim (U.S.), All-Invest (EU/UK), and Singular Bank (Spain) acted as bookrunners. Banco Sabadell, S.A. acted as the Settlement and Technical Pre-financing Agent for the Capital Increase, while Gómez-Acebo & Pombo Abogados, S.L.P. (Spanish law) and Gibson, Dunn & Crutcher LLP (U.S. law) served as legal advisors to the Company, and Loeb & Loeb LLP acted as U.S. legal counsel to Maxim Group LLC.

The funds raised will be used to:

- Advance the clinical development of vafidemstat for the treatment of aggression in CNS disorders such as Borderline Personality Disorder (BPD) and Autism Spectrum Disorder (ASD)
- Expand ongoing clinical programs in oncology and hematology
- Support CMC activities and preparations for Phase IIb-III trials, as well as other regulatory initiatives
- Strengthen the company's balance sheet to support future partnership or M&A negotiations
- Cover general administrative expenses and financial obligations
- Enhance legal and audit readiness for a potential dual listing on Nasdaq

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Dr. Carlos Buesa, CEO of Oryzon, CEO of Oryzon, commented: "This financing marks a key milestone that enables us to continue progressing our clinical programs with vafidemstat in aggression-related CNS disorders and to further develop our oncology and hematology pipeline. It also reinforces our financial position, supporting strategic conversations with investors and potential partners.

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 ladademstat

ladademstat (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). ladademstat 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). ladademstat is currently being evaluated in combination with gilteritinib in the ongoing Phase Ib FRIDA trial in patients with relapsed/refractory AML with FLT3 mutations, and in combination with azacitidine and venetoclax in 1L AML in an investigator-initiated study led by OHSU and in a trial sponsored by the U.S. National Cancer Institute (NCI) under the Cooperative Research and Development Agreement (CRADA) signed between Oryzon and the NCI to collaborate on further clinical development of iadademstat in different types of hematologic and solid cancers. 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 collaborative Phase II trial with the Fox Chase Cancer Center (FCCC) in combination with paclitaxel in R/R neuroendocrine carcinomas, and 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. Oryzon is further expanding the clinical development of iadademstat through additional investigator-initiated studies. Iadademstat has orphan drug designation for SCLC in the US and for AML in the US and EU.

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 two Phase IIa clinical trials in aggressiveness in patients with different psychiatric disorders (REIMAGINE, see Ferrer et al, Psychiatry & Clin Neurosci, 2025, doi.org/10.1111/pcn.13800) and in aggressive/agitated patients with moderate or severe AD (REIMAGINE-AD), with positive clinical results reported in both. Additional finalized Phase IIa clinical trials with vafidemstat include the ETHERAL trial in patients with Mild to Moderate AD, where a significant reduction of the inflammatory biomarker YKL40 was observed after 6 and 12 months of treatment, and the pilot, small-scale SATEEN trial in Relapse-Remitting and Secondary Progressive MS, where anti-inflammatory activity was also observed. Vafidemstat has also been tested in a Phase II in severe Covid-19 patients (ESCAPE) assessing the capability of the drug to prevent ARDS, one of the most severe complications of the viral infection, where it showed significant anti-inflammatory effects in severe Covid-19 patients. Vafidemstat is currently advancing as a Phase III-ready asset in Borderline Personality disorder (BPD) following completion of the global, randomized, double blind Phase IIb PORTICO trial (final data presented at ECNP-2024). Following receipt of the minutes from the End-of-Phase II meeting with the FDA to discuss PORTICO's results, the company announced plans to move forward with a Phase III PORTICO-2 trial in agitation/aggression in BPD (FDA submission planned in 1H2025). Vafidemstat is also being investigated in a doubleblind, randomized, placebo-controlled Phase IIb trial in negative symptoms of schizophrenia (EVOLUTION trial, recruitment ongoing). The company is also deploying a CNS precision medicine approach with vafidemstat in genetically-defined patient

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subpopulations of certain CNS disorders and is evaluating a clinical trial in Kabuki Syndrome patients. The company is also exploring the clinical development of vafidemstat in other neurodevelopmental syndromes.

FORWARD-LOOKING STATEMENTS

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