



**ORYZON**

**A GLOBAL LEADER IN EPIGENETICS**

INVESTOR PRESENTATION

MADX: ORY

September 2019

# LEGAL NOTICE

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## Company Highlights

- ✓ A **clinical stage** biopharmaceutical company developing innovative therapies in the field of **Epigenetics**
- ✓ **Two molecules** already with **positive data in humans**
- ✓ Large IP portfolio with technology fully developed in-house
- ✓ MADX: ORY A **publicly traded** company on the **Spanish Stock Exchange**
- ✓ Integrated in the **IBEX Small Cap Index**

- ✓ **Raised an aggregate of circa €85M** (in 2015-2019)
- ✓ **Cash runway** expected till **2H2021\***
- ✓ One of the most **LIQUID** companies in the MicroCap group in the Spanish Stock Market
  - ✓ 45.7 M Shares outstanding. Fully diluted
  - ✓ 350,000 daily volume (Avg Traded Volume in 2018)
  - ✓ +88M shares negotiated in 2018 / ≈5 months for share full turnover

\* On July 26<sup>th</sup>, the company completed a Private Placement with International Investors raising gross proceeds of €20M (circa \$22.2M at the exchange rate on that day)



BOLSA DE MADRID



ORYZON GENOMICS SA  
BALANCE SHEET DATA (UNAUDITED)<sup>1</sup>  
(Amounts in thousands US \$)

	June 30th, 2019	June 30th, 2018
Cash and cash equivalents	27,868	30,986
Marketable securities	669	165
Total Assets	<u>73,125</u>	<u>68,352</u>
Deferred revenue	0	0
Total Stockholders' equity	<u>50,888</u>	<u>40,697</u>

<sup>1</sup> Spanish GAAPs

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# Epigenetic Modifications : New Targets for Drug Development

- ✓ Epigenetic dysfunctions are associated with aberrant gene expression and disease
- ✓ Epigenetic drugs can restore these transcriptional imbalances
- ✓ Lysine specific histone demethylase 1 (LSD1), aka KDM1A, removes methyl marks at mono- and dimethyl-H3K4 (histone H3 lysine 4) and H3K9 (histone H3 lysine 9)
- ✓ LSD1 is the most abundant histone demethylase in the prefrontal cortex

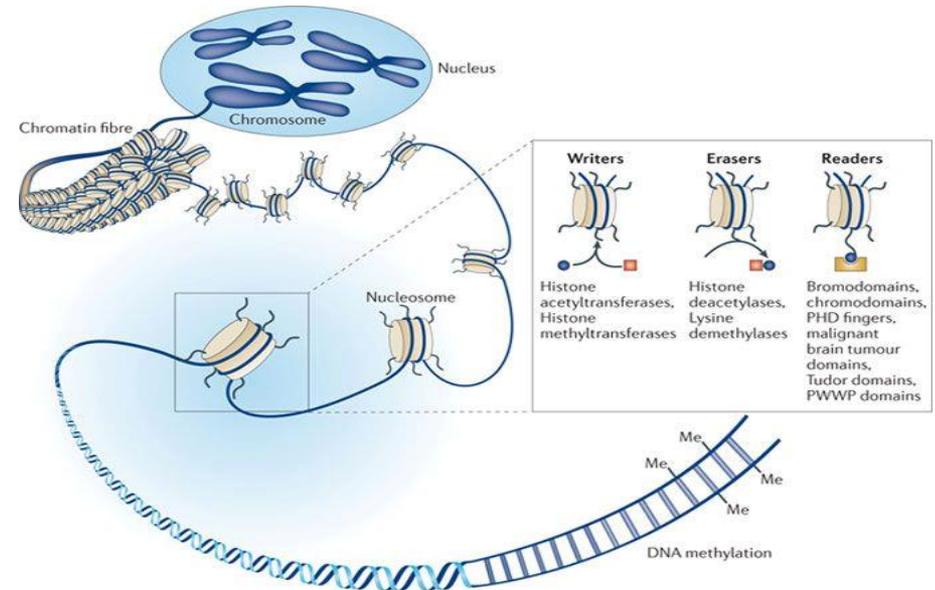
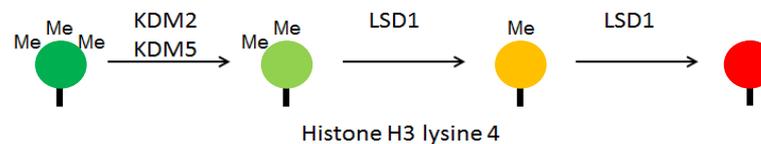


Figure from Arrowsmith et al. *Nature Reviews Drug Discovery* volume 11 (2012)



## Lysine specific histone demethylase 1 (LSD1): an epigenetic “eraser” that removes methyl groups from histones



LSD1 expression and activity can both block and promote gene expression



LSD1 plays an important role in cancer, CNS, inflammatory and viral diseases



# Oryzon is pioneering epigenetics in CNS and active in oncology

A broad pipeline to address unmet medical needs with an attractive market opportunity

INDICATION	STUDY*	RESEARCH	PRECLINICAL	PHASE I	PHASE IIA	PHASE IIB	PHASE III
<b>VAFIDEMSTAT (ORY-2001) - dual LSD1-MAO B inhibitor</b>							
Aggression in BPD	REIMAGINE / PORTICO (*)	[Progress bar]					
Aggression in ADHD	REIMAGINE / ENTRANCE (*)	[Progress bar]					
Aggression in ASD	REIMAGINE / COLONNADE (*)	[Progress bar]					
Aggression in AD	REIMAGINE-AD / GATEWAY (**)	[Progress bar]					
Alzheimer's disease (Mild Moderate)	ETHERAL monotherapy	[Progress bar]					
Multiple Sclerosis (RR & SP)	SATEEN monotherapy	[Progress bar]					
<b>IADADEMSTAT (ORY-1001) - selective LSD1 inhibitor</b>							
AML (Elderly Unfit)	ALICE Combo w Aza	[Progress bar]					
SCLC (First Line Relapsed)	CLEPSIDRA Combo w Platinum/Etoposide	[Progress bar]					
<b>ORY-3001 - selective LSD1 inhibitor</b>							
Non Oncological	Preclinical finished	[Progress bar]					
<b>OTHER PROGRAMS</b>							
Undisclosed		[Progress bar]					

\* IN BLUE, NEW PHASE IIB STUDIES UNDER PREPARATION OR EVALUATION

\*\* Contingent to + results in REIMAGINE-AD

## CNS Market Need

45 million people with AD worldwide; 20% shows aggressiveness

Aggression is a common feature in many psychiatric diseases. +50% in ADHD (\*\*\*)

Global BPD market expected to grow to \$2.6B in 2027

AD main disruptions: memory loss, aggression and apathy. AD global costs per annum of \$605B

## Oncology Market Need

Global AML market of \$990m in 2019. Room for new Combos according to KoLs

SCLC is a serious unmet medical need, with a MOS of 8–12 months and 5% 2-year OSR

Global SCLC market +300,000 patients/y. FDA approved label extension of Pembro but only 19% of ORR (\*\*\*\*)

Projections of Rova-T when in Phase III were +5B peak sales/y

(\*\*\*) J Child Adolesc Psychopharmacol. 2016 Feb 1; 26(1): 19–25.

(\*\*\*\*) Keynote study in 83 patients





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IN CNS EPIGENETICS

**VAFIDEMSTAT a Phase II Clinical Stage Compound with a broad  
developability in CNS diseases**

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## Vafidemstat (ORY-2001): a “Neuron-fixer”

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- ✓ Vafidemstat is a **small molecule** that selectively inhibits LSD1 and MAO-B
- ✓ LSD1 inhibition is the major driver of the pharmacological action
- ✓ **Excellent Pharmacology**. High **oral** bioavailability
- ✓ **Positive** results in **7 different animal models** and in *in-vitro* models
  - ✓ Cognition
  - ✓ Neuroprotection
  - ✓ Neuroinflammation
  - ✓ Social Withdrawal / Apathy
  - ✓ Aggression / Agitation
  - ✓ Others
- ✓ Epigenetic **MoA** that modulates **neuroinflammation** and expression of key **plasticity neuronal genes**
- ✓ Biomarkers identified
- ✓ **Good Safety in humans in Phase I+II trials with +220 participants** so far
- ✓ **BBB penetrance** and (indirect) human brain target engagement established
- ✓ Pharmacologically active in humans

**In Phase IIa in multiple clinical studies**

**Phase IIb studies under preparation**

# Vafidemstat, and LSD1 inhibition, improves cognition

In Alzheimer's SAMP8 model vafidemstat restores memory by the NORT model

In Huntington disease R6/1 model vafidemstat improves memory by the NORT model

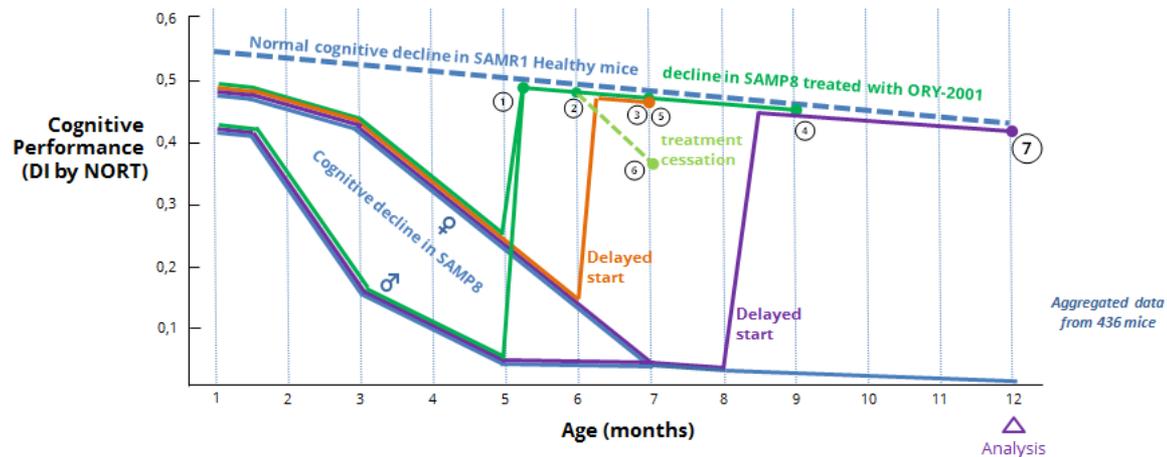
In Schizophrenia SETD1a +/- model iadademstat (ORY-1001) improves working memory

In Psychosis & Schizophrenia NMDA receptor-hypofunction mice model T-448 (TAKEDA) LSD1 inhibitor improves memory

## Vafidemstat Fully Restores Memory Measured by NORT in SAMP8 AD Model

MILD  
MODERATE  
SEVERE

Treatment from month 5 during 1 week (1), 1 month (2), 2 months (3), 4 months (4)  
Treated from month 6 during 1 month (Delayed start-1) (5)  
Treatment from month 5 during 1 month, tested at month 7 (1 month after treatment cessation) (6)  
Treatment from month 8 during 4 months (Delayed start-2) (7)



Cognition and memory impairments are found in AD and dementias but also in Autism, Schizophrenia, Depression, Bipolar disorder and other psychiatric conditions



# Vafidemstat Produces Significant Behavioural Changes

## Vafidemstat Reduces Aggression in the Resident Intruder Test in the SAMP8 AD Mice model



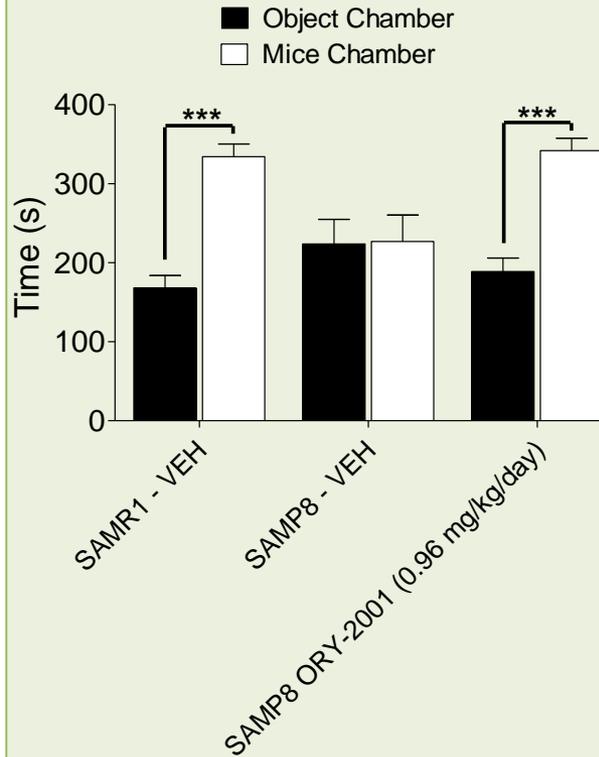
SAMP8 MICE treated with Vehicle Resident Intruder Test



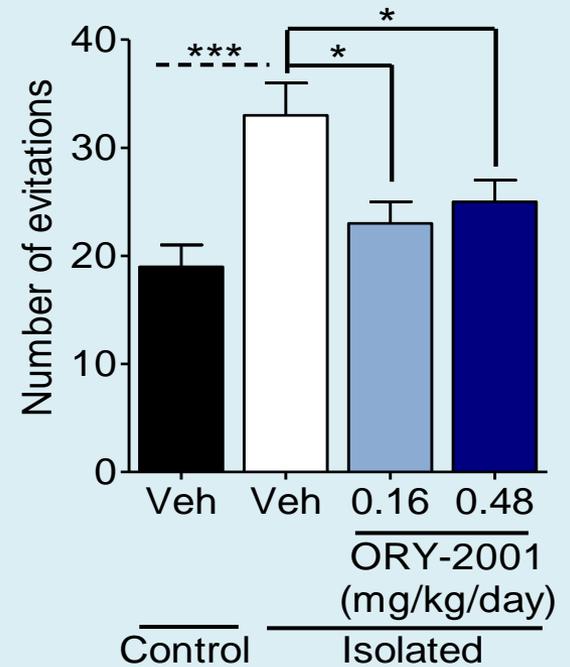
SAMP8 MICE treated with ORY-2001 0.32 mg Resident Intruder Test

## Vafidemstat Enhances Sociability in the Three-Chamber Test in SAMP8 AD Mice

TCT - Females 12M age  
4 months treatment  
Chamber Preference

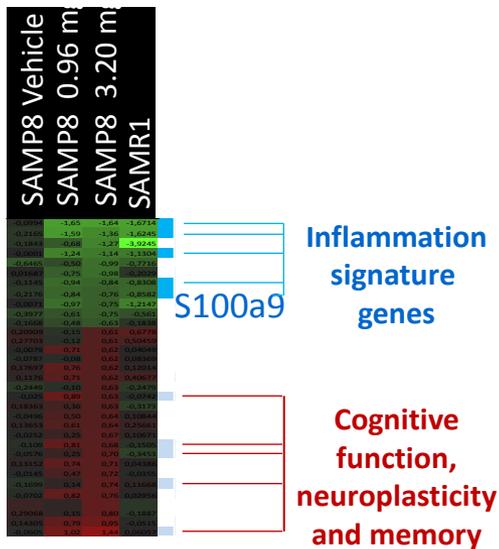


## Vafidemstat Reduces Social Withdrawal in the Rat Isolation Model



LSD1 localizes *in vivo* to enhancers and promoters of confirmed CNS disease risk genes  
 LSD1 binds to TFs that control IEG expression and stress in the PFC-amygdala axis, including SRF

- ✓ vafidemstat **up-regulates** genes associated with:
  - ✓ **Cognition**, notably memory and **executive functioning**
  - ✓ **Neuroplasticity**
- ✓ vafidemstat potentiates the response capacity of IEGs to stress
- ✓ vafidemstat **reduces** the expression of **inflammatory** genes including S100A9 and others



Vafidemstat potentially **down-regulated** the expression of a subset of genes related to immune reaction and **inflammation** as **S100A9** involved in OPC defective remyelination

Genes **up-regulated** in SAMP8 mice by vafidemstat included

**Baiap3**: involved in retrograde trafficking

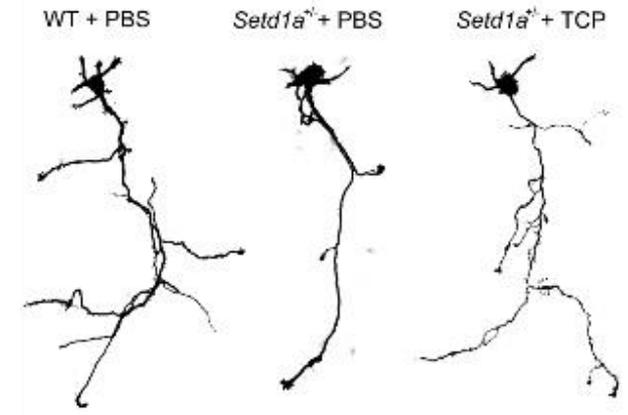
**Prph**: mutated in Amyotrophic Lateral Sclerosis (ALS),

**Fabp7**: upregulation in drosophila favors long term memory consolidation

**Doc2a**: activity-dependent modulator of excitatory synaptic transmission, relevant to memory formation

**Kremen2** and **Rspo1**, regulators of the WNT pathway

LSD1 inhibition rescues the axon branching deficits in the *Setd1a*<sup>+/-</sup> mice



In vitro axon branching rescue assays ORY-1001 was 1000-fold more potent than TCP

Mukai et al 2019 <http://dx.doi.org/10.1101/529701>  
 Recapitulation and reversal of schizophrenia-related phenotypes in *Setd1a*-deficient mice



# Vafidemstat : Safety demonstrated in a Phase I study

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## Safe and well tolerated in a +100 healthy volunteers (young and elderly) Phase I (MAD+SAD) study

- ❖ No hematological impact at planned doses
- ❖ Efficiently crossed the BBB (70-90%)
- ❖ Oral PK - Half Life of 22h allowing once daily oral
- ❖ PK/PD data allowed definition of recommended Phase II doses

## Safe and well tolerated so far in diverse Phase II studies

- ❖ Vafidemstat has been already administered to +220 volunteers and patients
- ❖ Phase IIs (MS, AD, ADHD, BPD and ASD patients) with no safety signals to date
- ❖ Longest exposure to date: 15 months



## Duration

**8 Weeks treatment + 4 weeks of follow up**

## Cohorts to be recruited

### REIMAGINE:

Borderline Personality Disorder	6 patients	Done – Data reported in April 2019	} upscaled later to an aggregated of 30 patients; recruitment finished
Attention Deficit and Hyperactivity Disorder	6 patients	Done – Data reported in April 2019	
Autism Spectrum Disorder	6 patients	Done – Data reported in Sept 2019	

### REIMAGINE-AD:

Alzheimer's Disease	12 patients	Recruitment Finished (1 <sup>st</sup> report expected in Dec 2019)
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## Endpoints

Safety

Efficacy:

Aggression / Agitation measured by CGI-S

Aggression / Agitation measured by CGI-I

Aggression / Agitation measured by NPI A/A 4 items

Psychiatric status measured by NPI Global assessment (12 items)

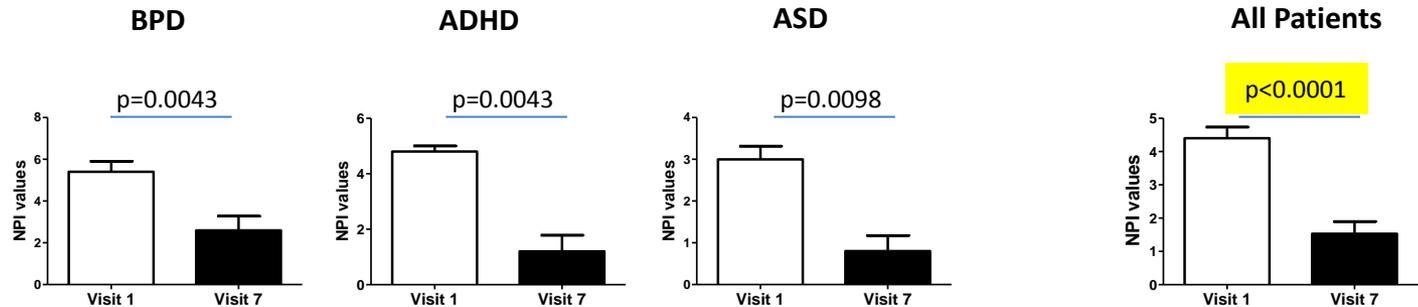
Change in specific disease scales

**Patients of the three psychiatric indications: Borderline Personality Disorder (BPD) Attention Deficit and Hyperactivity Disorder (ADHD) and Autistic Spectrum Disorder (ASD) Treated With Vafidemstat Showed a Reduced Aggressivity**

Secondary Endpoints: Efficacy

Significant improvements in aggression evaluated using the Neuropsychiatric Inventory 4-item agitation/aggression score (**NPI-A/A**)

**NPI-A/A Agitation/Aggression (4 items)**



Also significant improvements in aggression evaluated using the Clinical Global Impression of Severity (**CGI-S**) and Improvement (**CGI-I**) scales

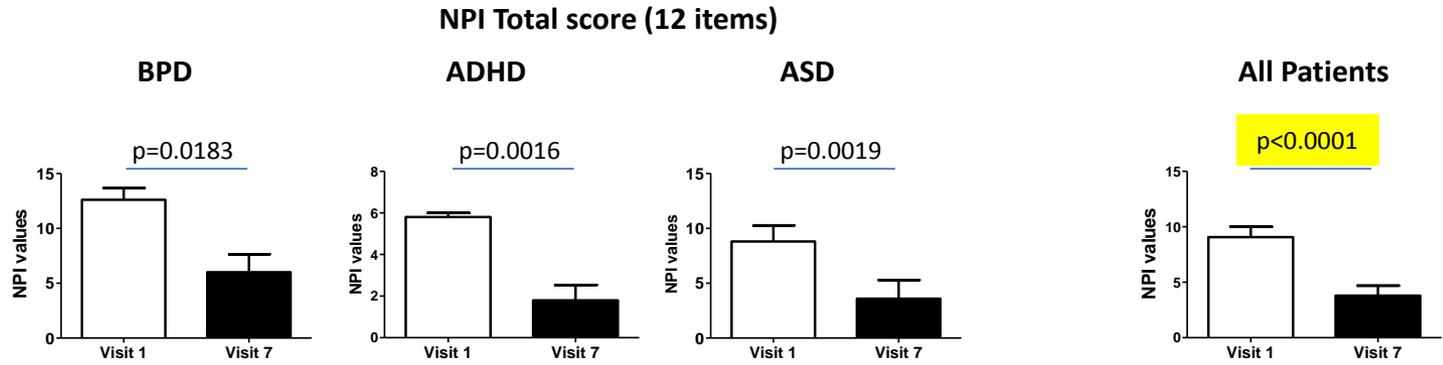
Data presented at



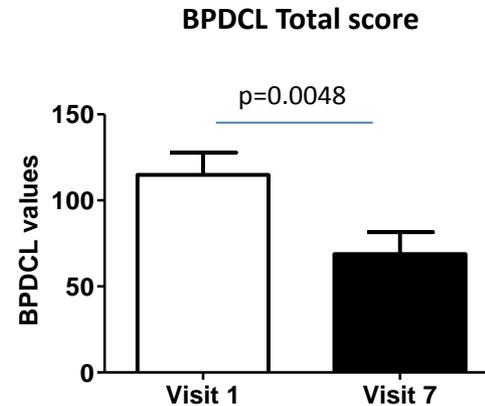
# REIMAGINE the first proof of concept for vafidemstat in human patients

The significant improvements in the NPI global score and overall specific scales for BPD and ADHD suggest that vafidemstat has a **broader psychiatric effect beyond aggression**

Vafidemstat also produced significant improvements on the global NPI score (12 items) in BPD, ADHD and ASD patients



Remarkably, vafidemstat not only improved aggression but also produced significant improvement on the GLOBAL Borderline Personality Disease Checklist (BPDCL) scale



*"I find myself able now to control my negative emotions and my frustration"*

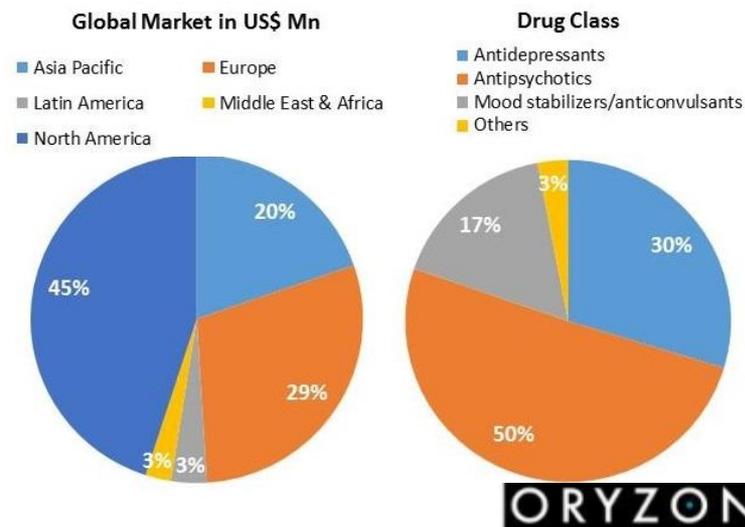
Patient with severe borderline personality disorder during treatment with vafidemstat. Testimony to her psychiatrist Dr. Marc Ferrer



### The company recognizes a significant development potential for vafidemstat in psychiatric indications

- ✓ Vafidemstat may be a **disease modifying therapeutic option for BPD**: reduces aggression and produces an overall improvement of the core features of the disease, with **no sedation and no weight gain**
- ✓ BPD prevalence ranges between 0.5%-1.4% of the total population ( $\leq 9.1\text{M}$  in US+EU5)
- ✓ The treatment of BPD is now based on psychotherapeutic interventions. No drugs currently approved for this condition
- ✓ A **significant unmet medical need**
- ✓ **Global BPD Market, 2018-2027 (US\$), \$2.6B expected in 2027**

- ❖ A new Phase IIb in BPD (*Portico*) under preparation
- ❖ Additional Phase IIb in adult aggressive ADHD (*Entrance*) and ASD (*Colonnade*) under evaluation
- ❖ And if +data in Reimagine-AD, a Phase IIb in AD aggressive patients (*Gateway*) will be performed



## Alzheimer's, the huge need



- ✓ **45 million** people affected worldwide
- ✓ The Global cost of AD is **\$605 billion/year**. No therapeutic options so far
- ✓ 20% of the outpatients and 40% of the inpatient patients display aggressiveness

## Vafidemstat first proposition in AD: a symptomatic drug

- ✓ Vafidemstat is **safe and highly brain-penetrant** in humans
- ✓ Brain target engagement in humans established (indirectly)
- ✓ Positive **effects** in different preclinical models on **memory, aggression, sociability and apathy**, all core features in Mild and Moderate AD patients
- ✓ Biomarkers identified that may be surrogate pharmacological biomarkers
- ✓ Vafidemstat is **pharmacologically active in BPD, ADHD and ASD patients**
- ✓ **Data on aggressiveness in AD to come in December (CTAD-2019 San Diego)**

Vafidemstat may also provide clinical benefit in AD either as a single or multi-symptomatic drug or as a disease modifier

# ETHERAL: Epigenetic **THE**Rapy in **AL**zheimer's Disease

## Besides aggression, vafidemstat may provide also further benefits to AD patients

An ambitious Phase IIa study to provide useful information to design future Phase II/III studies

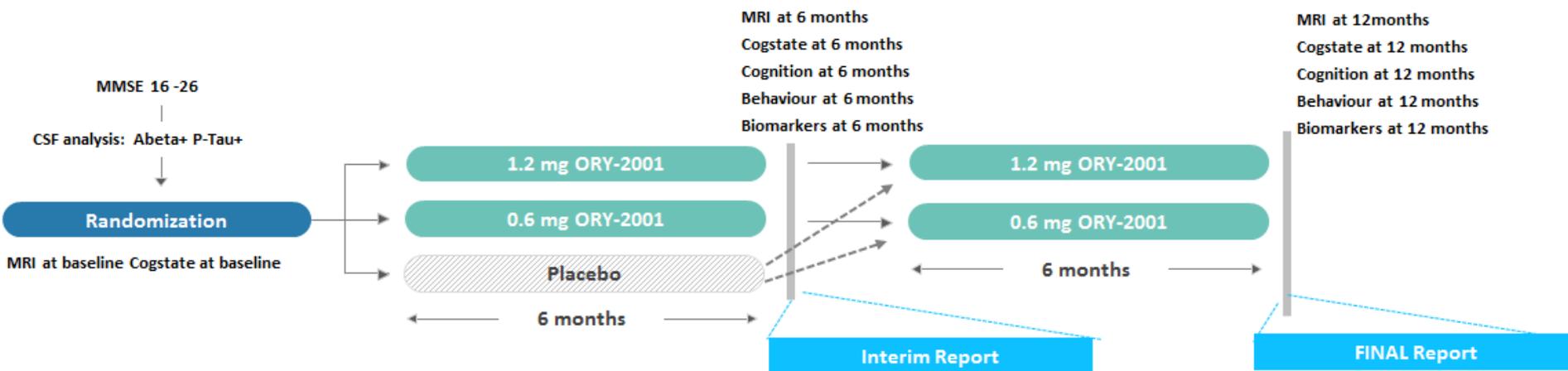
- ✓ **150 Mild to Moderate AD patients** (6+6 months)
- ✓ Primary Objective: Safety & Tolerability
- ✓ Secondary Objectives :
  - ✓ Cognition/Agitation/Apathy/Depression/QoL
  - ✓ Volumetric MRI
- ✓ **Biomarker guided study** (with 8 CSF Biomarkers)



- ✓ 117 patients in EU. 17 sites
- ✓ Spain, France & UK



- ✓ A Twin study in US: around 25-28 patients
- ✓ IND approved mid March
- ✓ US Sites (3) opened
- ✓ FPI recruited in May



One of the main goals in ETHERAL is to establish safety of vafidemstat in long-term administration, at therapeutically relevant doses in a frail and elder AD population



## Vafidemstat treatment appears so far to be safe and well tolerated:

- ❖ Four SAEs were reported in three subjects suspected to be unlikely related to the treatment
- ❖ Platelet, neutrophils and other hematological parameters do not show clinically relevant variations due to vafidemstat treatment
- ❖ No abnormal and clinically significant liver enzymes levels or other laboratory findings have been reported-to-date

## Biomarker and other functional evolution in ETHERAL (blind analysis) is compatible with an informative study

Variations on the S100A9 CSF levels observed in the first 33 patients completing the first 24 weeks of therapy (blind analysis) might be compatible with the preclinical data that shows that vafidemstat decreases S100A9 levels in the CNS. S100A9 is a proinflammatory protein reported to be highly overexpressed in PFC in AD patients



individual patient CSF S100A9 Change from Baseline (CFB) is shown as fold induction after 24 weeks treatment



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## Iadademstat

**A Phase II stage clinical  
compound in Oncology**

## ladademstat (ORY-1001): the most advanced selective LSD1 inhibitor in clinic

- ✓ LSD1 is involved in different cancers. **High levels of LSD1 often correlate with more aggressive forms of cancer and/or bad prognosis**
- ✓ ladademstat is a small molecule that selectively inhibits LSD1. Preclinical positive *in vivo* results in different xenograft models. Best in Class. Full characterization published in top-rank journal.
- ✓ First LSD1i drug to enter into clinical trials. Encouraging results in a FiM Acute Leukemia Phase I/IIa trial

Cancer Cell  
Article



### ORY-1001, a Potent and Selective Covalent KDM1A Inhibitor, for the Treatment of Acute Leukemia

Tamara Maes,<sup>1,6,\*</sup> Cristina Mascaró,<sup>1</sup> Iñigo Tirapu,<sup>1</sup> Angels Estiarte,<sup>1</sup> Filippo Ciceri,<sup>1</sup> Serena Lunardi,<sup>1</sup> Nathalie Guibourt,<sup>1</sup> Alvaro Perdonés,<sup>1</sup> Michele M.P. Luffino,<sup>1</sup> Tim C.P. Somerville,<sup>2</sup> Dan H. Wiseman,<sup>2</sup> Cihangir Duy,<sup>3</sup> Ari Melnick,<sup>3,4</sup> Christophe Willekens,<sup>5</sup> Alberto Ortega,<sup>1</sup> Marc Martinell,<sup>1</sup> Nuria Valls,<sup>1</sup> Guido Kurz,<sup>1</sup> Matthew Fyfe,<sup>1</sup> Julio Cesar Castro-Palomino,<sup>1</sup> and Carlos Buesa<sup>1</sup>

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<sup>5</sup>Drug Development Department (DITEP) and Hematology Department, Gustave Roussy, Université Paris-Saclay, 94805 Villejuif, France

<sup>6</sup>Lead Contact

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<https://doi.org/10.1016/j.ccell.2018.02.002>

### Phase I/IIa acute leukemia - previous data

- ✓ Safe and very well tolerated and therefore a meaningful candidate for combination with other agents
- ✓ PD Biomarkers identified in different subsets of leukemia
- ✓ Antileukemic activity observed in 29% of patients (12/41), including one CRi as Proof of Biological concept

## POTENTIAL ONCOLOGICAL INDICATIONS:

### Solid Tumors



Small Cell Lung Cancer  
Prostate cancer  
Colorectal cancer  
Bladder cancer  
Some breast cancers  
Merkel Cell Carcinoma

### HemONC



AML  
MDS  
Myelofibrosis  
Non Hodgkin  
Lymphoma

### Brain/rare Tumors



Medulloblastoma  
Glioblastoma

**MoA well characterized in SCLC, AML and Medulloblastoma**

## ALICE: An AML trial with LSD1i in Combination with azacitidine in the Elderly

A Phase IIa study to evaluate the safety, tolerability, dose finding and efficacy of iadademstat (ORY-1001) in combination with azacitidine in older patients with AML in first line therapy

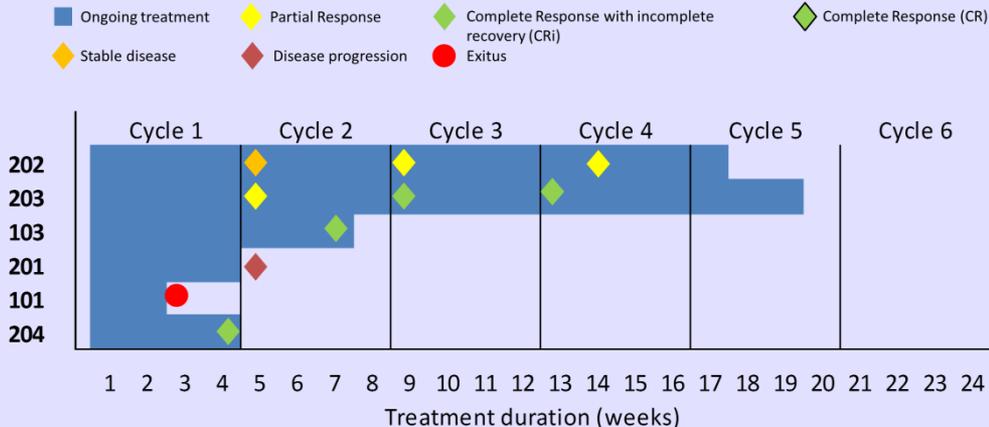
- ✓ Single arm & Open label. Up to 36 patients to be enrolled
- ✓ **Primary endpoint:** Safety and tolerability of the combo with hypomethylating agent Azacitidine
- ✓ **Secondary endpoints:** Responses; time to responses; duration of response; and overall survival

### Preliminary Results

(5 evaluable patients out of 6)



- ✓ Combo **well tolerated**
- ✓ Fast responses (median time to response 1.5 months)
- ✓ **80% OR** (4/5 evaluable patients) : **75% CRi** and 25% PR
- ✓ 1 patient in CRi with decreasing need of transfusions
- ✓ Additional reports to be presented in future Medical Conferences



# Iadademstat opportunity in Small Cell Lung Cancer (SCLC)

- ✓ LSD1 is a **target well characterized in SCLC** and validated in preclinical models. LSD1 inhibitors are effective in several in-vitro and in-vivo models of SCLC
- ✓ Iadademstat produces **complete and durable tumor regression** in different **chemoresistant PDX models**
- ✓ Characterized MoA
- ✓ Identified and patented Biomarkers that are differential in sensitive SCLC cell lines, tumors and plasma from patients
- ✓ Phase II trial ongoing in second line SCLC patients using these **biomarkers to stratify patients and identify super-responders**

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Targeting NOTCH activation in small cell lung cancer through LSD1 inhibition

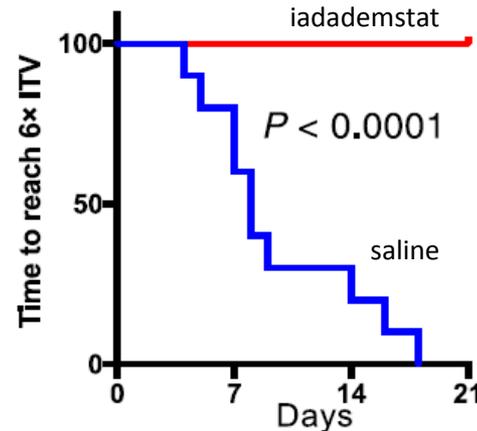
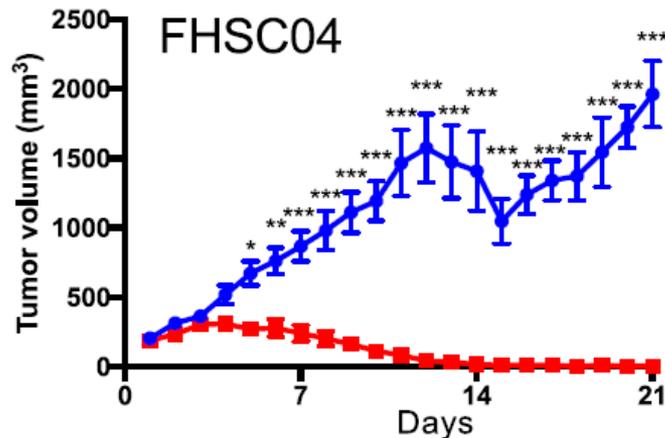
Arnaud Augert<sup>1,\*</sup>, Emily Eastwood<sup>1,\*</sup>, Ali H. Ibrahim<sup>1</sup>, Nan Wu<sup>1</sup>, Eli Grunblatt<sup>1</sup>, Ryan Basom<sup>2</sup>, Denny Liggitt<sup>1</sup>, Keith D. Eaton<sup>1</sup>, ...

Sci. Signal. 05 Feb 2019; Vol. 12, Issue 567, eaau2922; DOI: 10.1126/scisignal.aau2922

Article Figures & Data Info & Metrics eLetters PDF

**Targeted epigenetic therapy for SCLC**

Small cell lung cancer (SCLC) is an aggressive neuroendocrine tumor with no targeted therapeutic options and in which chemotherapy is only partially effective. Previous studies indicated that SCLC growth is suppressed by drugs that inhibit the histone lysine demethylase LSD1. Augert *et al.* found that LSD1 epigenetically suppressed the expression of the gene encoding NOTCH1, enabling the activity of the neuroendocrine cell lineage-associated transcription factor ASCL1. Blocking LSD1 with iadademstat (ORY-1001), a drug that has just been approved for phase 2 clinical trials in leukemia, reactivated NOTCH signaling and



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**CLEPSIDRA:** A Combination trial of LSD1 and Etop-Platinum in Small Cell Lung Cancer in **biomarker-ID** Relapsed pAtients

**A Phase IIa study to assess the safety, tolerability, dose finding and efficacy of iadademstat (ORY-1001) in combination with platinum-etoposide chemotherapy in patients with relapsed, extensive-stage disease small cell lung cancer who are positive to candidate predictive biomarkers**

- ✓ Single arm
- ✓ Open label; 4 sites in Spain
- ✓ Up to 36 patients to be enrolled
- ✓ **Primary end point:** Safety and tolerability of the combo with platinum-etoposide therapy
- ✓ **Secondary endpoints:** RECIST responses; time to responses; duration of response; and overall survival

### **Current status:**

- ✓ 10 Patients enrolled. One patient at cycle 12
- ✓ Satisfactory and dynamic recruitment pace

## Preliminary Results

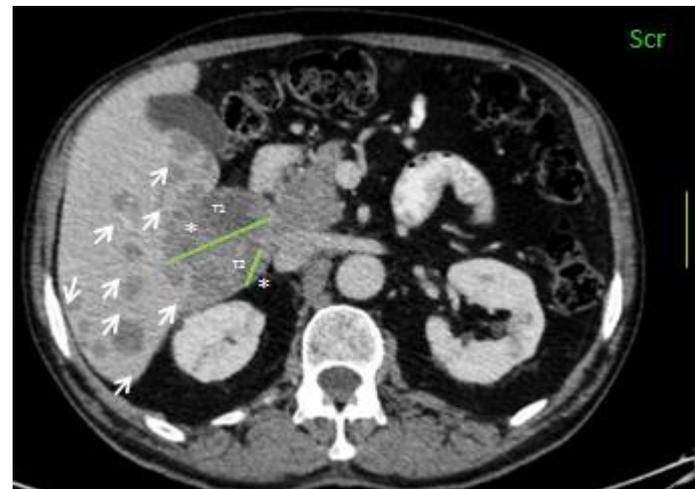


**Case report** from a patient with the longest treatment period in the study presented at IASLC 2019 WCLC:

- ❖ Administration of the combination during the first 6 cycles produced a tumor reduction of 78.7% by RECIST criteria
- ❖ Subsequent administration of iadademstat alone for four consecutive cycles proved to be safe and well tolerated, without producing any hematological toxicity
- ❖ Remarkably, reduction of main lesions and metastasis continues with iadademstat in monotherapy, reaching an overall tumor reduction of 86.3% by RECIST criteria (CT scan evaluation in Cycle 8)



Preliminary results for first 10 patients enrolled to be presented at ESMO-2019



# Anticipating a rich flow of catalysts / clinical data

## Iadademstat Phase IIs in oncology

**CLEPSIDRA**

**ALICE**

**2019**

## Vafidemstat Phase IIs in CNS

**REIMAGINE**

**ETHERAL**

**SATEEN**



**2020**

**ASCO ANNUAL MEETING**

**Chicago; June SCLC & AML data**

**April; Vienna AD Global EU 6m data**

**AD/PD 2020**  
The 15<sup>th</sup> International Conference on Alzheimer's & Parkinson's Diseases

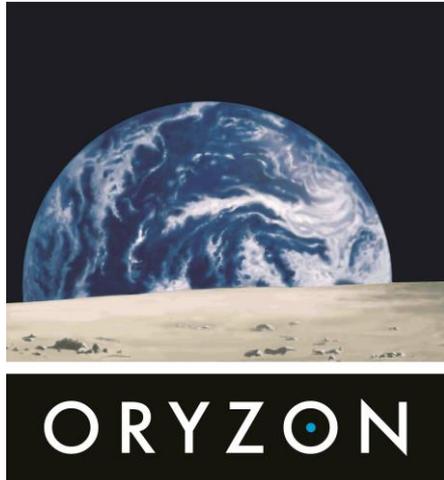
**July; Amsterdam AD Global US 6m data**

alzheimer's association  
**AAIC >20**

**ORYZON**

## ORYZON – a unique investment opportunity in an epigenetic platform

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- ✓ A differential proposition in **EPIGENETICS** drugs in **CNS and ONCOLOGY** around one of the most interesting targets in the field: **LSD1**
- ✓ **2 molecules** in **Phase II** with promising clinical signals in human patients
- ✓ **Pioneers in CNS epigenetics**
  - ✓ Vafidemstat shows efficacy in psychiatric disorders (BPD, ADHD, ASD)
  - ✓ **Phase IIb in Borderline personality disorder under preparation.** Additional options in ADHD or ASD under evaluation
  - ✓ Vafidemstat may be also clinically relevant in neurodegenerative disorders (Phase IIs in MS and AD ongoing)
- ✓ **Most advanced LSD1i (iadademstat) in Oncology**
  - ✓ 2 Phase II trials ongoing in combo with respective SoC in AML and SCLC
  - ✓ **Positive preliminary efficacy results** reported from AML and SCLC trials
  - ✓ **SCLC trial is a biomarker-guided** study to stratify responsive patients
- ✓ **Rich pipeline** of clinical **news** expected in the next 2-4 Qs
- ✓ Clinical Operations in US started and under expansion
- ✓ A **cash efficient** company with a seasoned international management team
- ✓ **€135M market cap.** One of the most liquid stocks in the microcap group in MadridSEXC
- ✓ Perseverant **presence in the US market in the last 4 years.** Two successful PIPEs executed in 2018-19 led by US Investment Banks and with participation of US investors
- ✓ A public company in Europe with **plans to** get dual listed in **NASDAQ**

# EXPERIENCED MANAGEMENT TEAM

▪ CEO



**CARLOS BUESA: CEO & President. Spain/US**

PhD in Biochemistry and Molecular Biology. Founder and CEO since inception. Advanced programs on finance, business development, negotiation skills and human resources. He is also PADE at the IESE Business School. He is Board Member of the VC Fund Inveready and Deputy President of the Spanish BioIndustry Association.

▪ CSO



**TAMARA MAES: CSO & VicePresident. Spain**

PhD in Biotechnology. Founder and Chief Scientific Officer since inception. Responsible of the creation of the whole pipeline of the company and the biological target validation programs. She is SAB member on several public institutions as CSIC and private companies. Since 2016 Scientific Advisor of the ADDF

▪ Medical Director



**ROGER BULLOCK: UK /PT/ Spain**

**Chief Medical Officer**

Graduated in Physiological Sciences at Keble College in Oxford University and got his MB.BS at London University

Extensive experience as clinical researcher, having participated in more than 70 clinical trials in Alzheimer's disease and other CNS conditions

30-year research career, +than 100 peer-reviewed publications and book chap

He has worked as a consultant for companies active in the CNS space, including Lilly and Merck

▪ VP Clinical Development



**MICHAEL T. ROPACKI: US**

**Vice President of Clinical Development**

PhD in Clinical Neuropsychology. Dr. Ropacki has held roles of increasing responsibility for + 10y at Johnson & Johnson, his last as Director of Clinical Development, Neuroscience, Research and Development, for Janssen R&D serving as the Clinical Lead responsible for developing and leading the Cognitive Health in Aging Registry. Prior to that role he served as Global Medical Affairs Leader, Head of Late-Stage Development at Janssen AD Immunotherapy, LLC.

▪ Clin Ops Director



**SONIA GUTIERREZ: Spain**

**Chief of Clinical Operations**

BSc. Pharm. & MSc. & PDD in IESE Business School. More than 20 years of experience in the clinical research and operations area at different Intnal. Pharma & Biotech companies. CNS: +13y in Lundbeck involved in + 40 Clinical Trials in CNS. Experience in oncology and other indications in Regeneron and other companies.

▪ IP Director



**NEUS VIRGILI: Spain**

**Chief IP Officer**

B.Sc. in Organic Chemistry from the University of Barcelona

Qualified European Patent Attorney

She has over 20 years experience in pharmaceutical IP

Since 2011 IP Officer at Oryzon

▪ CFO



**ENRIC RELLO: Spain**

**Chief Financial Officer**

J.D.; PhD in Economics & Business Administration.

PLD - Program for Leadership Development, Harvard Business School.

BSc & MSc in Business Administrations & Laws, HBS Finance Excell. Prog. Harvard Business School.

From 1997 till 2007 CFO of SANDOZ (NOVARTIS), Spanish Arm.

CFO at Oryzon since 2011

▪ BDO



**EMILI TORRELL: Spain**

**Chief BD Officer**

B.Sc. in Sciences, Autonomous University of Barcelona

MBA at ESADE and PDG at IESE Business School

In the business development area from 1990 in the most relevant Spanish companies Prodesfarma, Almirall and Laboratorios Esteve

From 2007 BD Director at Oryzon

- An experienced and respected managerial team in the Biopharmaceutical industry
- Team members have a track record in product discovery & in advancing successfully through product development phases
- Demonstrated ability to close world class deals and to lead, and participate in international consortia

**ORYZON**  
**A GLOBAL LEADER**  
**IN EPIGENETICS**



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