

January 2025 | Euronext: ALBPS – OTC: BPTSY


LIVE HEALTHIER LONGER



Forward Looking Statements



This presentation contains forward-looking statements. Forward-looking statements include all statements that are not historical facts. In some cases, you can identify these forward-looking statements by the use of words such as «outlook », «believes», «expects», «potential», «continues», «may», «will», «should», «could», «seeks», «predicts», «intends», «trends», «plans», «estimates», «anticipates» or the negative version of these words or other comparable words. These forward-looking statements include statements regarding Biophytis' anticipated timing for its various BIO101 (20-hydroxyecdysone) clinical trials and expectations regarding commercialization. Such forward-looking statements are based on assumptions that Biophytis considers to be reasonable.

However, there can be no assurance that the statements contained in such forward-looking statements will be verified, which are subject to various risks and uncertainties including, without limitation, delays in patient recruitment or retention, interruptions in sourcing or supply chain, its ability to obtain the necessary regulatory authorizations, COVID-19-related delays, and the impact of the current pandemic on the Company's clinical trials. The forward-looking statements contained in this presentation are also subject to risks not yet known to Biophytis or not currently considered material by Biophytis.

Accordingly, there are or will be important factors that could cause actual outcomes or results to differ materially from those indicated in these statements. Please refer to the «Risk Factors» section of the Company's 2023 Full Year Financial Report available on BIOPHYTIS website (www.biophytis.com) and to the risks discussed in the Company's registration statement on Form F-1 and other reports filed with the Securities and Exchange Commission (the "SEC"). We undertake no obligation to publicly update or review any forward-looking statement, whether as a result of new information, future developments or otherwise, except as required by law.



A clinical-stage biotechnology company specialized in the development of therapeutics for **muscular and metabolic diseases**



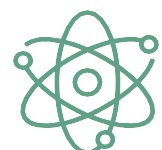
HQ location: Paris, France
Other locations in Sao Paulo, BR and Cambridge, MA US



Founded: 2006



Euronext growth Paris (ALBPS)
OTC market (BPTSY)



Drug discovery: biology of aging for developing drugs for age-related diseases



Multiple partnerships

Academical partnerships



Industrial partnerships



Pharmaceutical partnership





BIOPHYTIS' People: Expertise & Passion



Stanislas Veillet
CEO, cofounder



Rob van MAANEN
Chief Medical Officer



Pierre DILDA
Chief Scientific Officer



Edouard BIETH
Chief Business Officer



Waly DIOH
Chief Operations Officer



Chiara BACCELLI
Chief Pharmaceutical Operation,
Officer & Quality Assurance Director



Our Clinical Pipeline



Candidate	Indication	Program	Preclinical	Phase 1	Phase 2	Phase 3	Regulatory	Market
BIO 101 20-hydroxyecdysone 	Sarcopenia 		[Red bar]				Partnering	
	Obesity 		[Red bar]					
	Covid-19 		[Blue bar]				Partnering	
	DMD 		[Blue bar]					
BIO 203	Dry AMD 		[Green bar]					
	Stargardt 		[Green bar]					

xxx orphan diseases

BIO101 (20-hydroxyecdysone): First-in-class drug candidate

New molecular target

- Activation of MAS receptor¹ (renin-angiotensin system)
- Regulation of smooth, cardiac and skeletal muscle metabolism
- Stimulation of muscular and respiratory functions

POC & safety in clinical studies

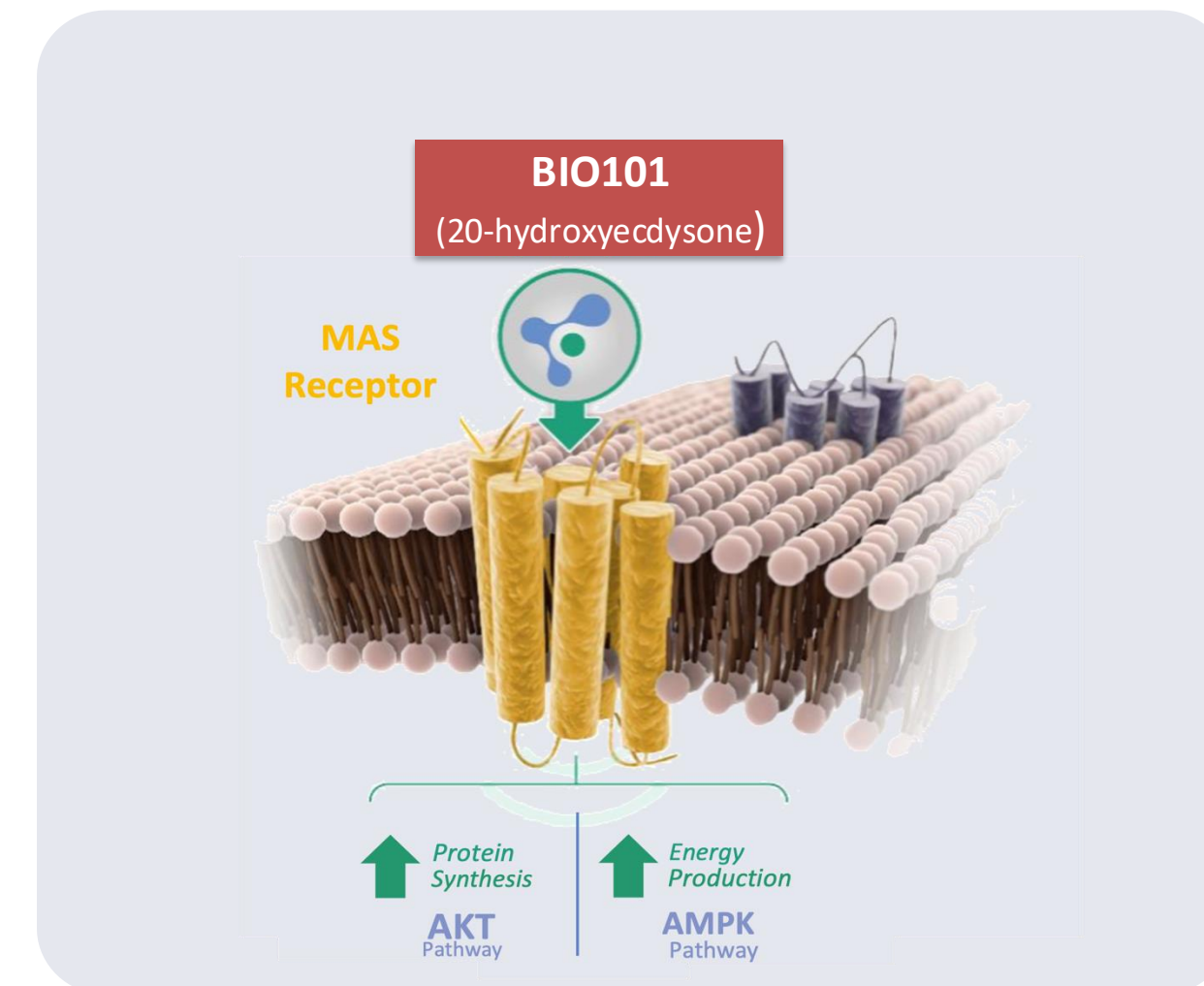
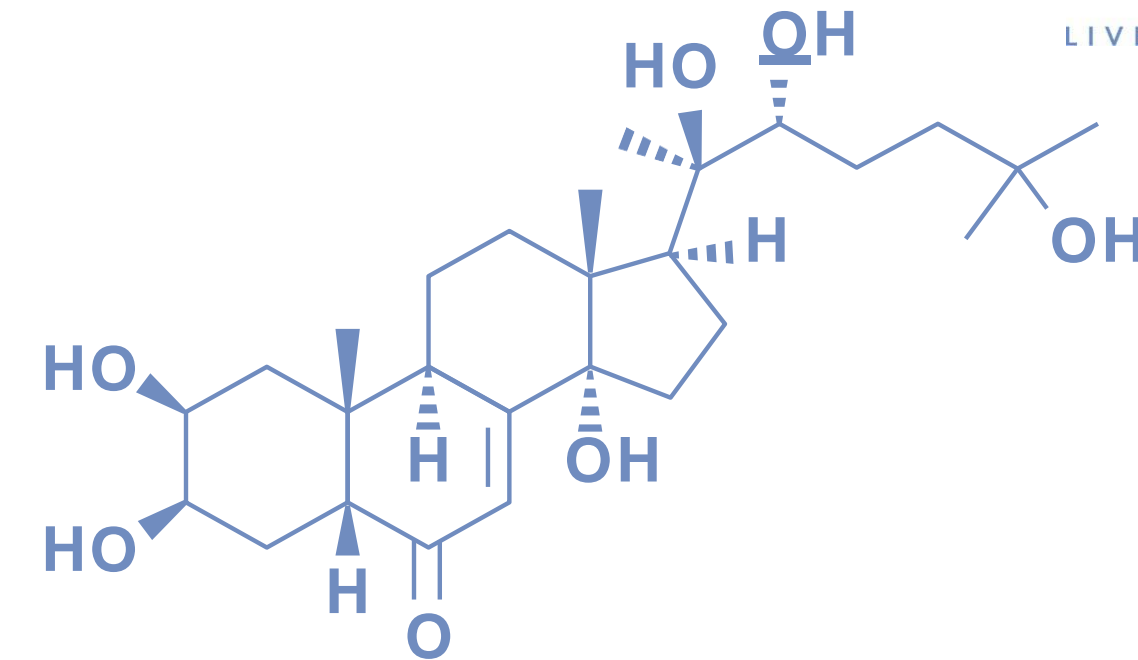
- Clinical studies in healthy elderly and obese adults (Phase 1)
- Clinical study in sarcopenic & obese sarcopenic elderly (Phase 2)
- Clinical study in severe Covid-19 (Phase 2-3)

Convenient administration & affordable cost

- API manufactured at industrial scale
- Oral with adult and pediatric formulations

Rock-solid IP

- 14 patent families, 44 granted in key countries



BIO101 (20-hydroxyecdysone) activates MAS receptor and triggers downstream two signaling-pathways in myocytes: AKT & AMP

Licencing-out BIO101 in sarcopenia (SARA) and Covid-19 (COVA) to regional pharma partners



Executing the partnering strategy

- Objective: license-out BIO101 (20-hydroxyecdysone) to regional pharmaceutical companies for its co-development and future commercialization
- Scope: regional deals with leading regional pharma companies, with focus on phase 3 ready programs (SARA, COVA)
- Revenue: license deal with upfront, milestones and royalties on future sales



LATAM: Exclusive license agreement with Blanver for BIO101

NORTH AMERICA: Active research for partnerships



EUROPE: Active research for partnerships





ASIA: On going discussions with regional pharma partners



€108 M licensing deal with Blanver for LATAM region (June 2024)

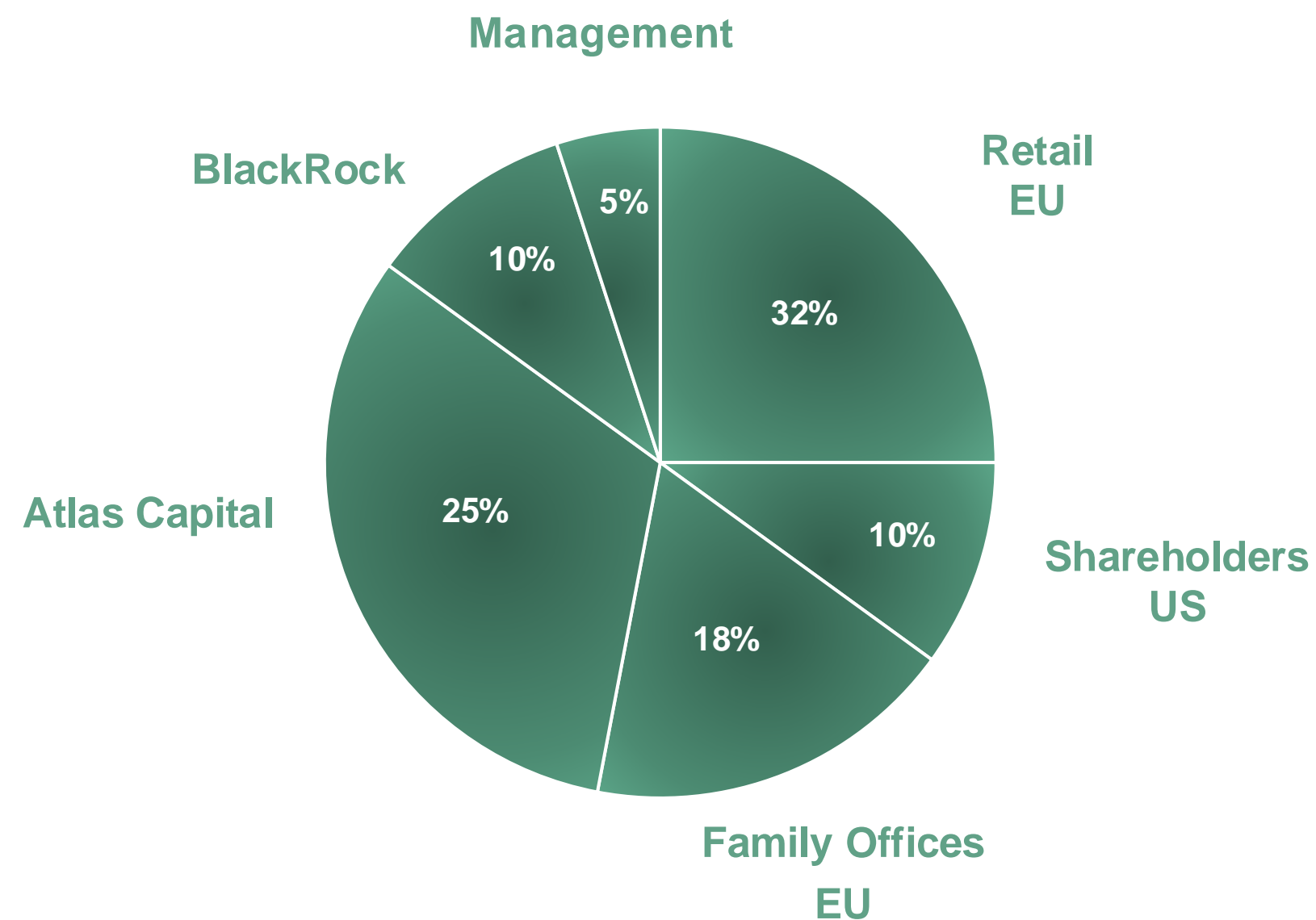
- Blanver is a Brazilian pharmaceutical company with 30+ history specialized in infectious and musculo-skeletal diseases.
- Blanver has been granted an exclusive license agreement covering the 4 indications under development for BIO101 in Latin America
- Biophytis will receive an upfront payment and additional payments based on the achievement of targets, for a total amount of up to €108 million, as well as double-digit royalties on future product sales.
- Sergio Frangioni, CEO of Blanver, said: *“Innovation is part of Blanver’s DNA, and we are delighted to partner with Biophytis to introduce this unique medicine to Latin America”*

Key milestones in the development of BIO101 (20-hydroxyecdysone)

	Achieved in the last 12 months	Anticipated in the next 12 months
	Approval to start the OBA Phase 2 study in the US.	Start of OBA phase 2 study
	€108M licensing partnership with brazilian pharma Blanver for LATAM	Partnering in Asia and/or Europe Start of second phase 3 study depending on the pandemic evolution.

Financial data

Shareholding structure



Number of shares: 15.855.846 (January 10, 2025)



Key financial figures

Listing Euronext (ALBPS) and US market (OTC)

- Cash position: €2.2m (June 30th, 2024)
- New cash of €2.5m in January 2025 through a capital increase
- Commitment to convert €4m of financial debt (BlackRock and Atlas) at €0.30 per share, corresponding to 13.4 million shares.



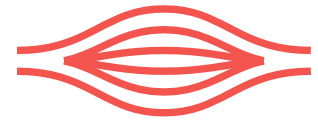
Analyst Coverage

- H.C. Wainwright – Joe Pantginis, Ph.D.
- Kepler Cheuvreux – Nicolas Pauillac
- Invest Securities – Jamila El Bougrini, Ph.D.

BIO101 (20-hydroxyecdysone)

in Obesity





Muscle wasting associated with pharmacology treatment of obesity: an unmet medical need



Obesity is a serious chronic disease

1bn

Adults and children are currently living with obesity globally.

3x

The global prevalence of obesity has more than tripled since 1975.

\$4tn

The global cost of treating obesity-related complications is expected to rise by over \$4 trillion by 2035.

Up to 40%

Total weight loss that comes from muscle when obese patients are treated with GLP-1RA.

“

**nature
biotechnology**

After obesity drugs' success, companies rush to preserve skeletal muscle
Nature Biotechnology. 2024 42(3):351-353





« [There is a need to] counter the side effects of dramatic weight loss [induced by GLP-1s]. [Biotechs] are searching whether it is possible for people to lose weight on these GLP-1 RA agonists without losing muscle. »

”





Competition : BIO101 is the only muscle agent in development focusing on muscle strength

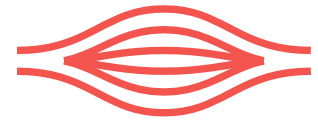
Drug	Company	Mode of action	Main Effect	Safety & Side effects	Administration route	Status
BIO101		MAS Receptor activator	Muscle strength (<i>knee extension determined by dynamometry</i>)	BIO101 has been very well tolerated in 277 individuals across multiple clinical studies	Oral	Phase 2
Azelaprag		APJ agonist	% change in overall weight loss	Hepatotoxicity (Liver transaminitis) (5)	Oral	Phase 2 halted
Bimagrumab		Activin type II receptor blocker	Changes in body weight, waist circumference, and body composition	Muscle spasms and diarrhea (2)	Intravenous	Phase 2
Enobosarm		Selective Androgen Receptor modulator	Total lean body mass	Increased hepatic transaminases, fatigue, hypercalcaemia (1)	Oral	Phase 2



There is no drug registered for muscle preservation in obesity

Source: (1) Lancet Oncol 2024; 25: 317–25 (2) JAMA Network Open. 2020;3(10):e2020836. doi:10.1001/jamanetworkopen.2020.20836 (4) DrugBank entry on Trevogrumab (5) <https://bioagelabs.com/azelafrag>





Potential attributes of BIO101 (20-hydroxyecdysone) in obese patients treated with GLP-1RA



Effects on muscle wasting:

- Preservation of muscle strength
- Reduction of muscle mass loss
- Improvement of mobility



Effects on fat tissues:

- Increase of fat mass loss



Convenient and safe administration :

- Oral route
- Adequate safety demonstrated in adults from trials in other indications





Supportive preclinical data in obesity



Metabolic effects in obese mice :

- Protective effect of BIO101 (20-hydroxyecdysone) in mice fed an obesity-inducing high-fat diet, preventing adipose tissue development
 - Anti-obesity effect by increase in energy expenditure
-

Muscle function in mice fed high fat :

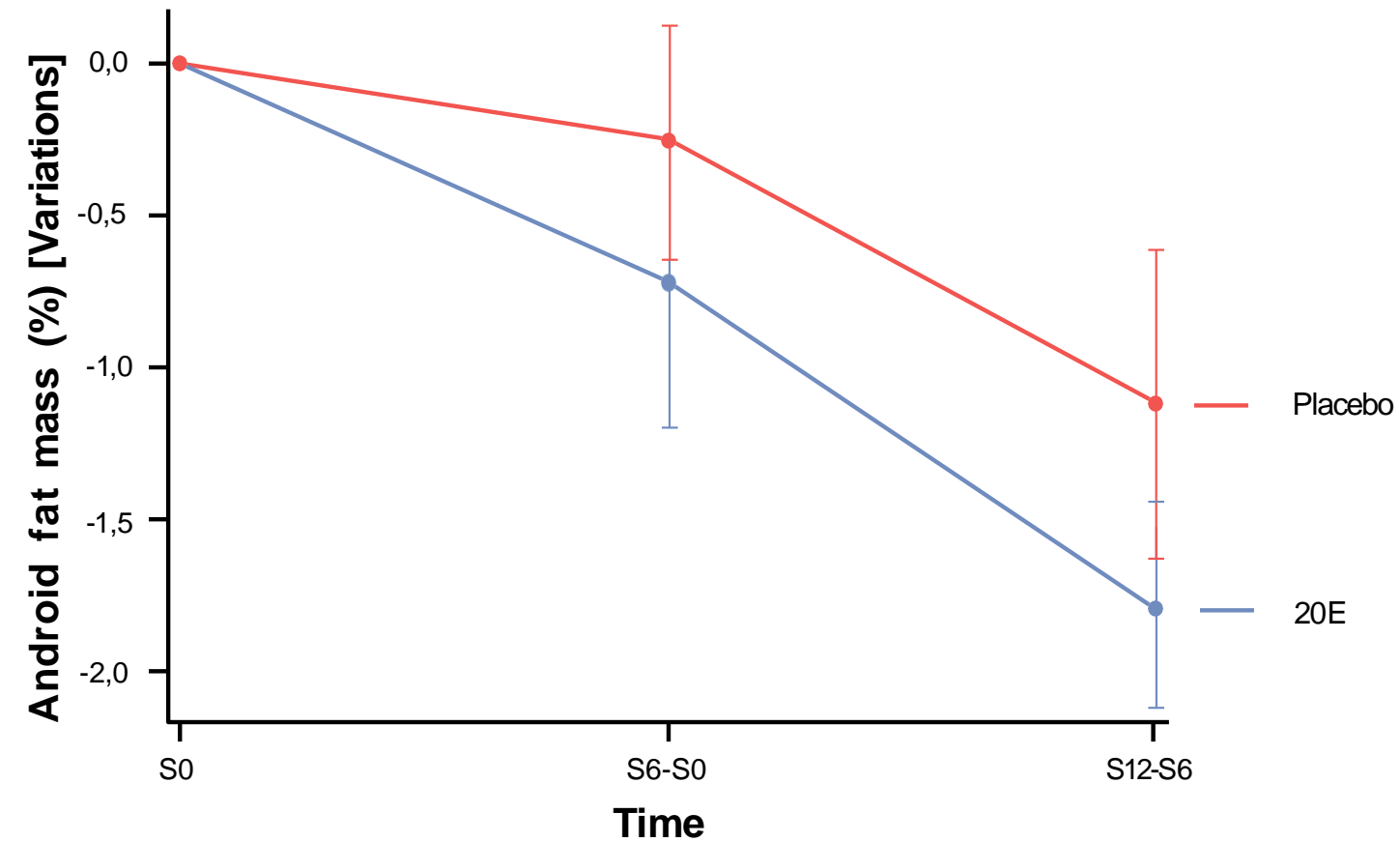
- Improved physical performances in adult and old animals orally treated with BIO101 (20-hydroxyecdysone)



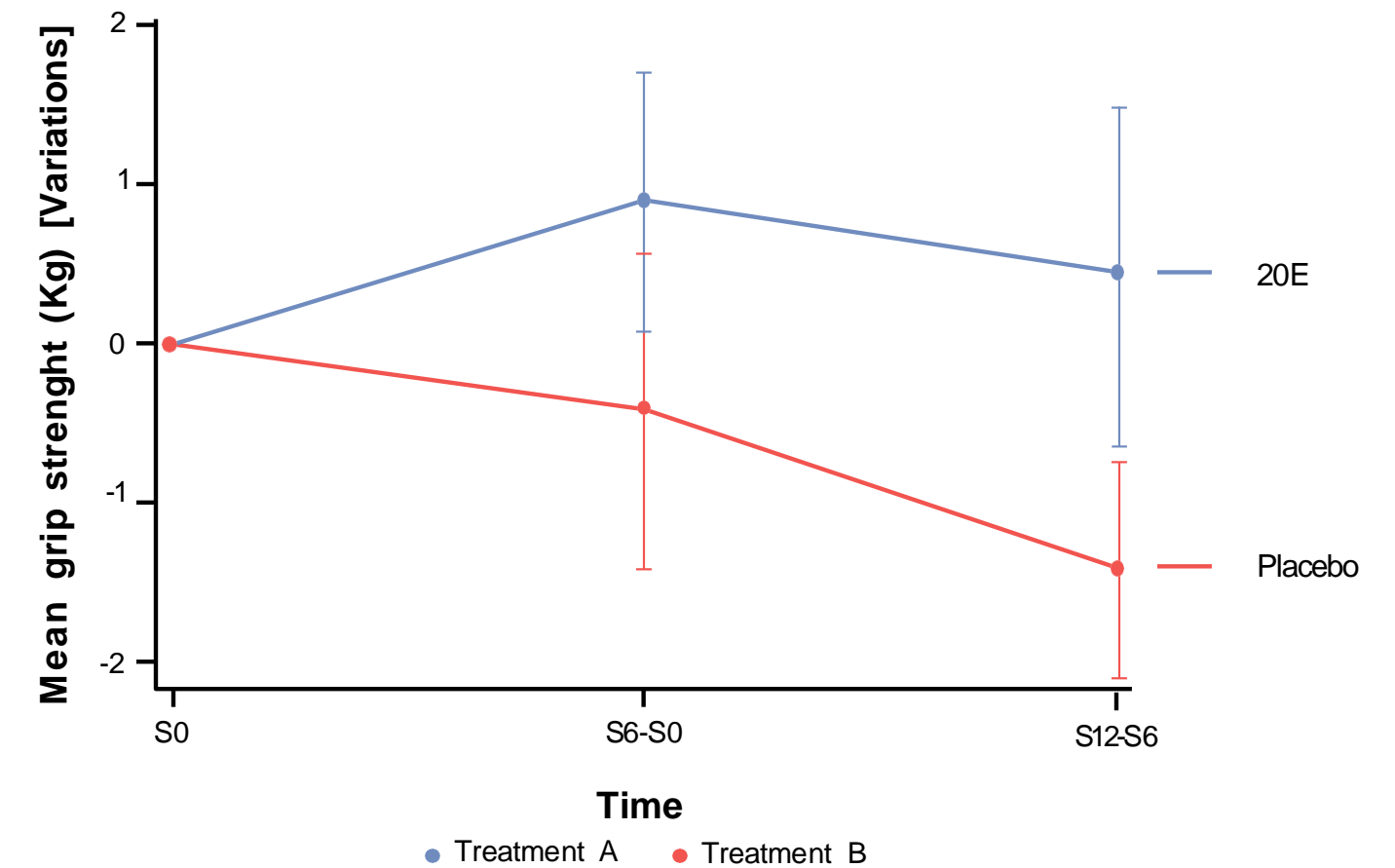
Promising clinical data in obese patients on hypocaloric diets for weight-loss



Android fat mass ($p=0.0386$)



Handgrip strength patients with weight loss $>5\%$ ($p=0.0974$)



20-hydroxyecdysone (20E) daily dose of 37.5 mg (given in the form of a dietary supplement) compared to placebo (n=58)
12 weeks study, with weight loss on hypocaloric diet for six weeks (S0-S6) followed by a normocaloric diet for six weeks (S6-S12)

Source: Foucault 2012. AgroParisTech, 2012. NNT : 2012AGPT0041. pastel-00998299





OBA – Phase 2 development plan



Design

- Randomized, double-blind, placebo-controlled phase 2 trial
- Assess efficacy and safety of BIO101 (20-hydroxyecdysone) 350 mg BID administered orally over 21 weeks

Endpoints

- Primary
 - Muscle strength (knee extension)
- Secondary
 - Walking speed (6-minute walking test)
 - Muscle strength normalized in relation to muscle mass
 - Weight, muscle mass and fat mass
 - Symptoms reported by patients (PROs)

Patient Population

- 164 obese patients treated with GLP-1 RAs, together with hypocaloric dieting
- Obese patients (BMI ≥ 30) or overweight (BMI ≥ 27 with one or more sequelae e.g. hypertension and sleep apnoea)

Product

350 mg b.i.d of BIO101 (20-hydroxyecdysone)

2024

2025

IND in the USA

First patient enrolled

Last patient

Report of the results



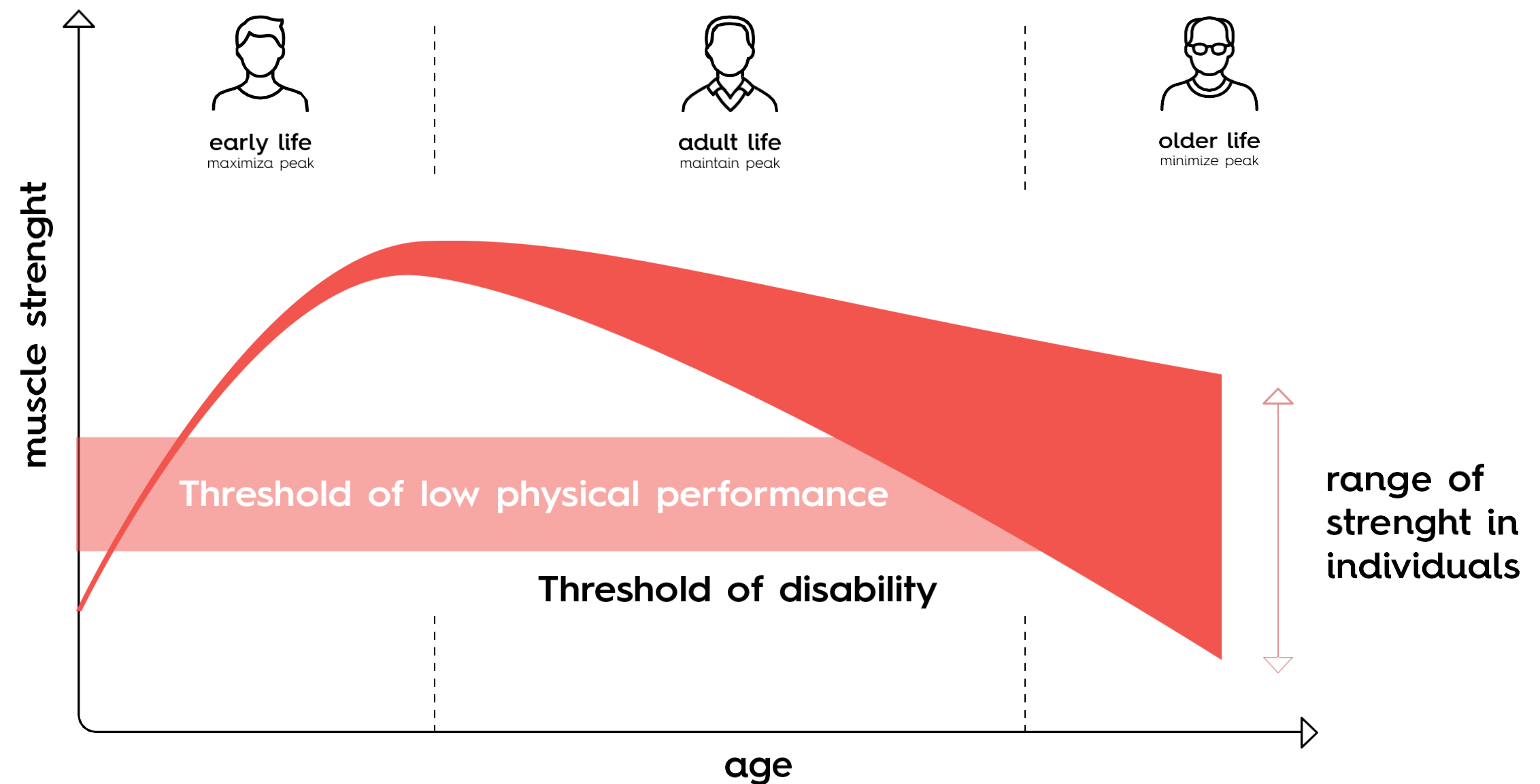
BIO101 (20-hydroxyecdysone)
in SARCOPENIA





Sarcopenia is an aged related disease, with no approved drug

Sarcopenia is a syndrome defined by many consortia including the EWGSOP (The European Working Group on Sarcopenia in Older People), characterized by **progressive and generalized loss of skeletal muscle mass, strength and function** associated with an increased risk of adverse events such as disability, poor quality of life and death.





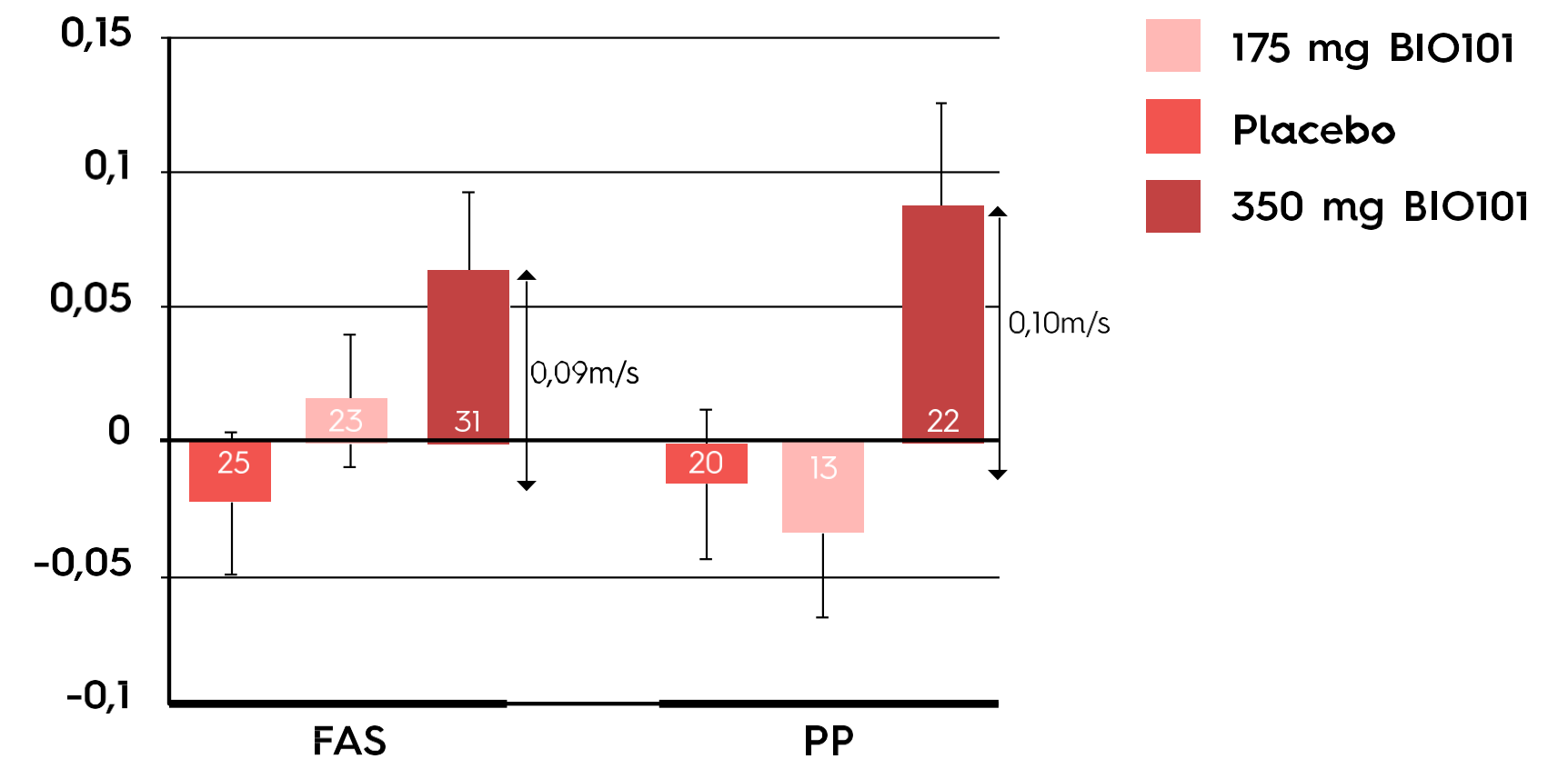
Promising results obtained in SARA-INT phase 2 trial



BIO101 (20-hydroxyecdysone) significantly improves the 400 MWT gait speed, the primary endpoint, in the PP population after 6 months of treatment

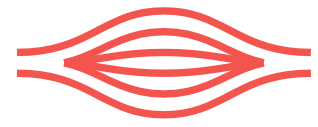
- Global, double-blind, randomized, placebo-controlled trial in patients with aged-related sarcopenia at risk of mobility disability to evaluate safety and efficacy of BIO101 (20-hydroxyecdysone)
- At the highest dose of 350 mg bid: clinically meaningful improvement of 0.10 m/s in the PP population (significant, $p=0.008$) compared to placebo for the 400MWT gait speed after 6 months of treatment
- This gait speed level of 0.10 m/s is known to be associated with a reduction in mobility disability and mortality in the elderly
- BIO101 (20-hydroxyecdysone) demonstrated the same effects on mobility in the **sarcopenic obese subpopulation**.

Change from baseline at M6 Gait speed



Treatment effect is nominally significant in PP population at M6 ($p = 0.008$)





SARA-31 – Phase 3 development plan



Design

- Global, double-blind, randomized, phase 3 placebo-controlled trial
- Assess safety and efficacy of BIO101 (20-hydroxyecdysone) 350 mg BID administered orally over at least 52 weeks, as compared to placebo
- Treatment effect based on estimation of the risk of mobility disability

Endpoints

- Primary
 - Major Mobility Disability (MMD) assessed by the inability to complete the 400-meter walk test (400MWT) within 15 min
- Secondary
 - Gait speed 4-meter from Short Physical Performance Battery (SPPB)
 - Handgrip Strength (HGS)
 - Patient Reported Outcomes (PRO)

Patient Population

- Age: 65 years old or over
- Low mobility measured by Short Performance Physical Battery: $SPPB\ 3 \leq SPPB \leq 7$
- Low Handgrip Strength (HGS < 20 and < 35 kg in female and male)
- Slow walkers (gait speed < 0.8 m/s)
- Reporting a loss of motor function over the last year

Product

2023

2024

2025

2026

350 mg b.i.d of BIO101 (20-hydroxyecdysone)

CTA in Europe/US

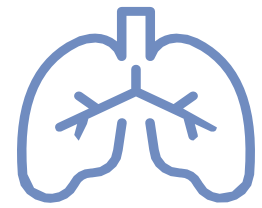
SARA-31 Phase 3 (depending on partnership)



BIO101 (20-hydroxyecdysone)

in Duchenne Muscular Dystrophy

MyODA



The life of patients with Duchenne Muscular Dystrophy (DMD)



Orphan genetic disease affecting 1/5,000 boys at birth (220,000 patients worldwide)

"Duchenne is every child and parents' worst nightmare come true"
(Victoria, Mother of Dougie)



0 to 2 yo
No symptoms



5 yo
Motor delay



Around 12 yo
Wheel chair



Around 19 yo
Trachestomy
& Assisted ventilation



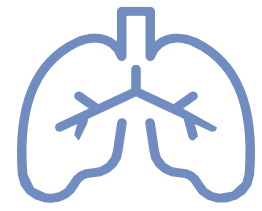
Around 25 yo
Heart failure



Around 30 yo
Death
(cardio-respiratory causes)

- Degenerative : **every muscle is slowly and inexorably damaged** (dystrophin deficiency)
- **It can affect anyone** : 1/3 arise from random spontaneous genetic mutations, which may occur during any pregnancy

MyODA



There is no effective treatment

Despite research progress, no treatment is able to cure or effectively control the progression of the disease



Corticosteroids (Prednisone, deflazacort, vamorolone)

They are the standard of care, but their use is controversial and not uniformly recommended.

- **Mobility loss delayed by 2 years**
- **Serious side effects** (weight gain, behaviour disorders, muscle wasting, osteoporosis, cataracts, high blood pressure...)
- **Long-term use associated with more serious sequelae** (69% of complications reported in non-ambulatory patients)

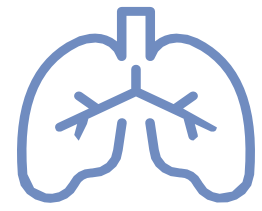
Source : Orphanet J. Rare Dis. doi.org/10.1186/s13023-021-01758-9

Gene-based Therapies (exon skipping, microdystrophin, etc...)

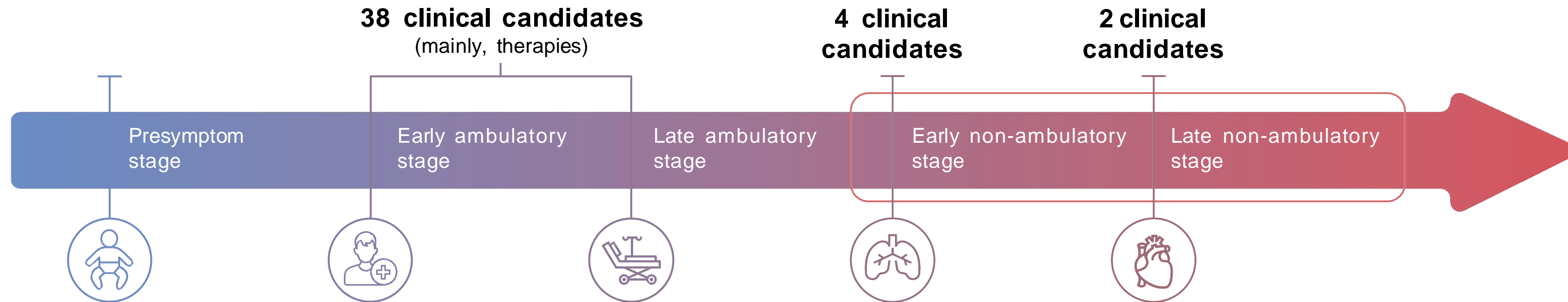
Gene therapies have been considered to be a revolution for the past 35 years, but what are the concrete results today?

- **Limited effectiveness¹**
- **Highly restricted number of addressed patients** (e.g. 13% for eteplirsen², and limited to young patients)
- **Toxicity issues** (several deaths suspected³)
- **Outrageously expensive** (\$3,2M/patient for Elevidys⁴)



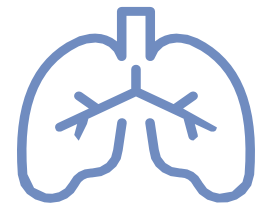


Late-stage patients are neglected



- No approved drug for their **specific respiratory problems**
- Excluded from current clinicals trials
- Average age for tracheostomy is **19 years old**¹

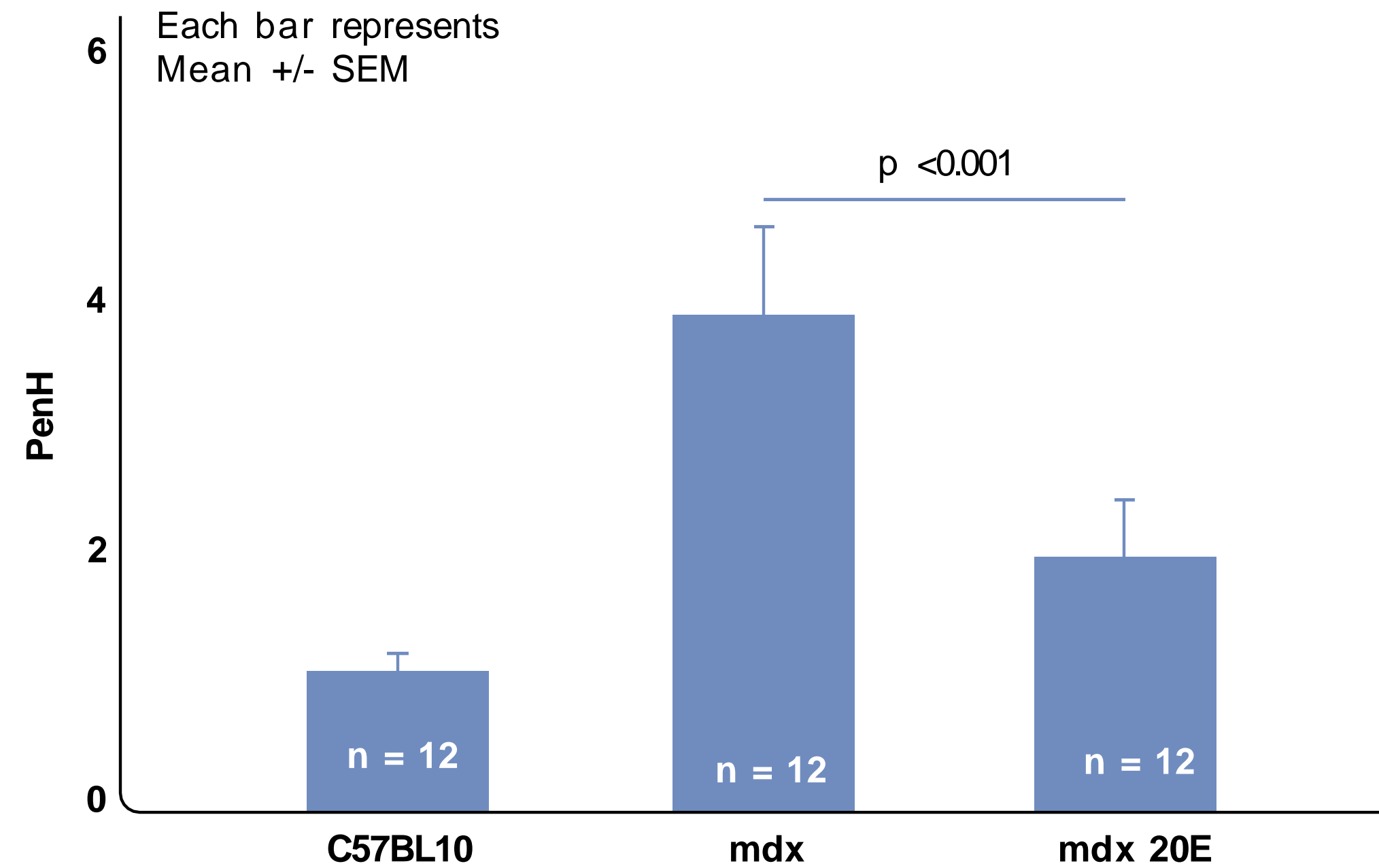
- How do we treat these children who cannot breathe properly as we speak ?
- Ventilatory assistance, mostly invasive, has extended life expectancy by 10 years, **BUT with drastic impact on quality of life**

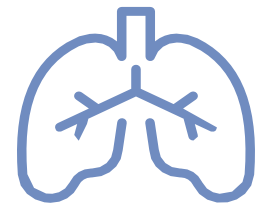


Our solution: A first-in-class medication

BIO101 (20-hydroxyecdysone) aims to improve breathing capacity

Improvement in airway responsiveness (PenH) in C57BL10-mdx mice.





Preparing to start phase 1-2 clinical study in DMD



Design

- A Randomized, Double-Blind, multi-center Phase 1-2 Study
- Evaluate the Safety, Efficacy, Pharmacokinetics, and Pharmacodynamics of BIO101 (20-hydroxyecdysone) in Non-Ambulatory DMD Patients with Respiratory Deterioration.
- Pediatric oral formulation (powder) of BIO101 (20-hydroxyecdysone)

Endpoints

- Primary
 - change from baseline in Forced vital capacity (FVC)
- Secondary : The Peak Expiratory Flow (PEF), Performance of Upper Limbs (PUL) scale, Grip strength (MyoGrip)
- Part 1 (N=15): Safety, tolerability & PK - 7 days of escalating dose)
- Part 2 (N=45): Safety and efficacy on respiratory function (FVC, PEF) of one dose for 48 weeks

Patient Population

- Age: ≥ 12 years old
- Non-ambulatory DMD patients
- Patients at risk of respiratory failure

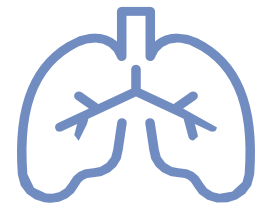


in SARIs

(Severe Acute Respiratory Infections)

Ruvembri™ is the marketing name for BIO101 (20-hydroxyecdysone) in Covid-19 indication





Targeting Hospitalized Patients with severe respiratory symptoms due to COVID-19



Patients **aged 45 and above**, with proven COVID-19, and severe respiratory symptoms:

- With evidence of respiratory decompensation ≤ 7 days before start of study medication, meeting one of the following :
 - Tachypnea: ≥ 25 breaths per minute
 - Arterial oxygen saturation 92% or less

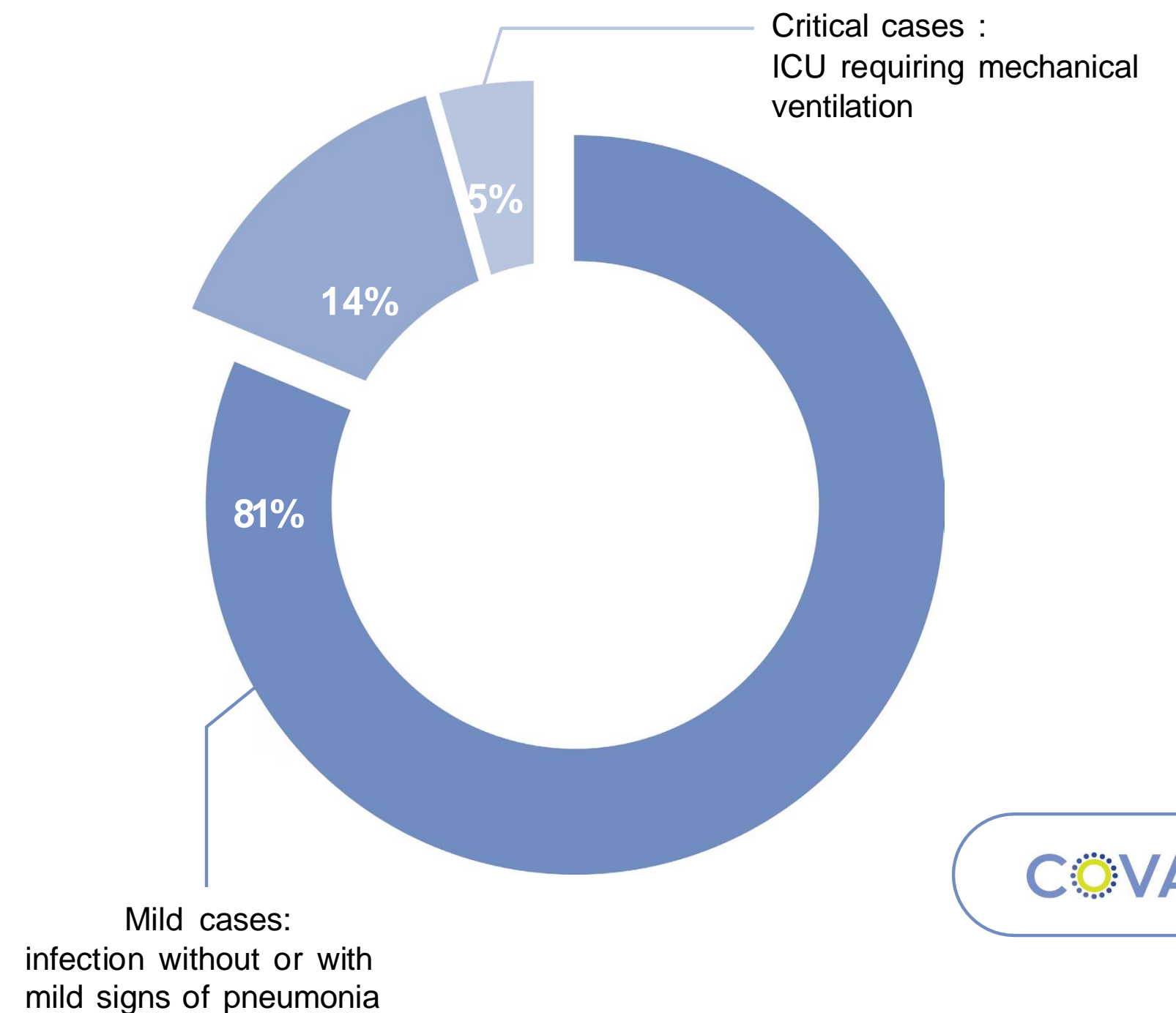
Hospitalized patients with respiratory failure estimated to 15-18% of hospitalized patients: ca **500 new patients per day or 180,000 patients/year in the USA** (CDC data, October 27, 2022)

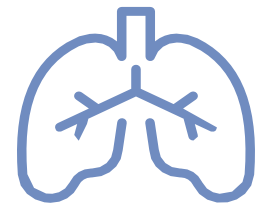


Allowed medications :

- Antiviral agents such as remdesivir, Paxlovid™
- Anti-inflammatory agents such as dexamethasone, tocilizumab™

Severe cases: hospitalized with hypoxemia, tachypnea or pneumonia





Phase 2-3 COVA clinical study to evaluate of Ruvembri™ in the treatment of severe forms of COVID-19



THE LANCET

EClinicalMedicine

Published by THE LANCET

Design

- Administration of 350 mg b.i.d of BIO101
- Global, multi-center, double-blind, placebo-controlled group Phase 2-3 sequential (2 parts) adaptive design
- International study including 37 clinical centers in US, Brazil, France & Belgium

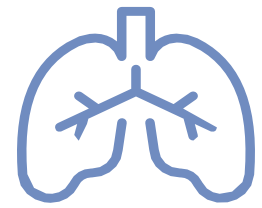
Endpoints & Study Follow-Up

- Primary endpoint : proportion of patients with respiratory failure or early death within 28 days
- Secondary endpoints : mortality at 28 and 90 days; discharge at 28 days
- End of study: Q2 2022 (N=237) after early study termination

Inclusion criteria

- Age : 45 years old or over
- Hospitalized for severe respiratory symptoms and with proven Covid-19 infections
- Patients with hypoxemia (<92%) or tachypnea (> 25 breaths/min)
- All authorized Covid-19 drugs (anti-viral or anti-inflammatory)

COVA



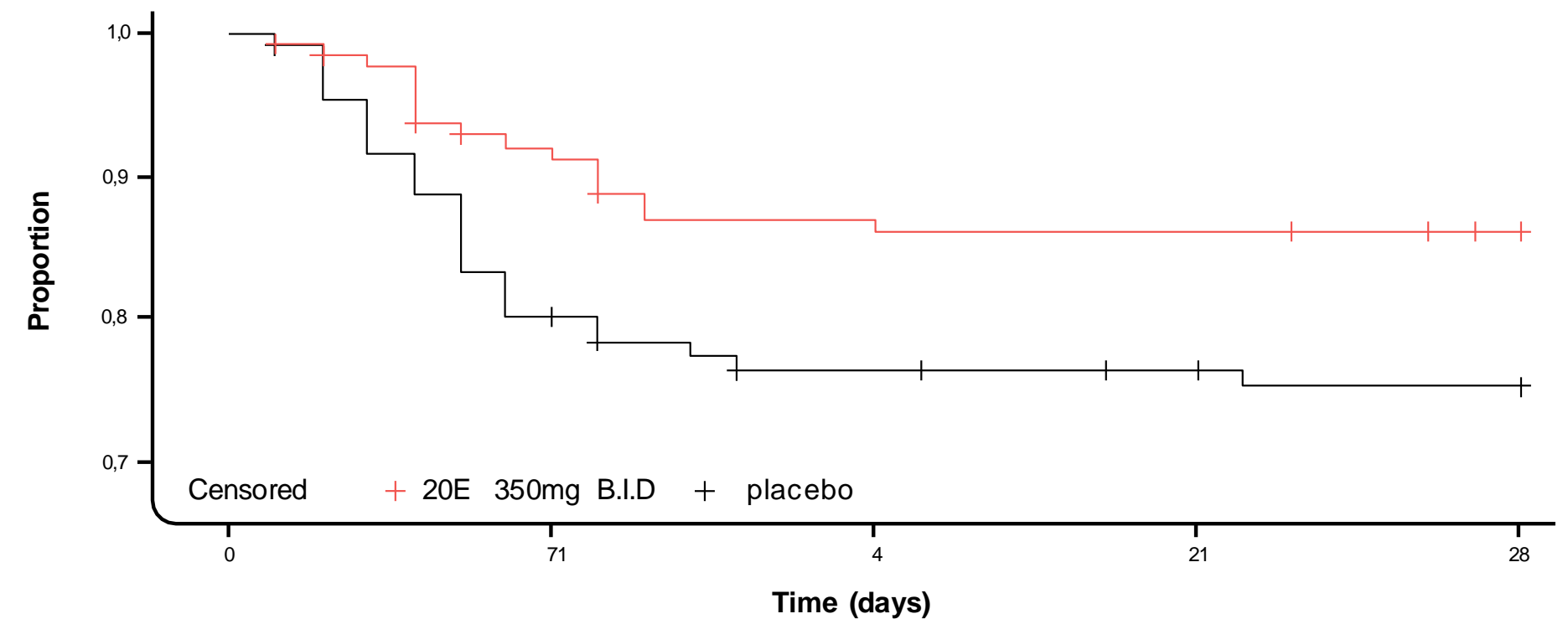
Positive results strongly supporting therapeutic potential of Ruvembri™ in severe COVID-19 : respiratory failure or early death



Respiratory Failure or early death : The study met primary endpoint

- Reduction in the risk of early death or respiratory failure at day 28 by 44% (p=0.043, CMH test)
- Time to early death or respiratory failure over 28 days was lower (p=0.022, Kaplan Meier analysis)
- Post hoc analysis confirmed the reduction in **the risk of early death or respiratory failure** in the ITT population and in the PP population

Proportion without respiratory failure or early death, Kaplan-Meier Analysis, ITT population



THE LANCET

EClinicalMedicine
Published by THE LANCET



Source: Lobo et al., eClinicalMedicine 2023 : 102383.
Published Online : <https://doi.org/10.1016/j.eclinm.2023.102383>

Source: Lobo et al., 2024. www.thelancet.com Vol 68 February, 2024

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Professor, Director of the International Institute of Medicine and Science

Has led clinical trials for new treatments for HIV/AIDS



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THANK YOU

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