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These factors should not be construed as exhaustive and should be read in conjunction with the other cautionary statements that are included in the registration statement and in the listing prospectus. We undertake no obligation to publicly update or review any forward-looking statement made in this presentation, whether as a result of new information, future developments or otherwise, except as required by law.



A clinical-stage biotechnology company specialized in age-related diseases



Our goal

Prevent disabilities (muscular, respiratory and vision) and increase health span for patients suffering from age-related diseases Our small molecules are aimed at stimulating biological resilience and are developed through a drug discovery platform based on a reverse pharmacology approach



COVID-19 & Neuromuscular diseases

Drug candidate **Sarconeos (BIO101)** in clinical development for :

Respiratory failure resulting from SARS-Co-V2 infection

Sarcopenia: Phase 2 An age-related degeneration of skeletal muscle

Duchenne's Muscular Dystrophy (DMD): IND granted A rare pediatric genetic neuromuscular disease



Retinal diseases

Pre-clinical drug candidate **Macuneos (BIO201)** for diseases of the retina, such as dry **Age-related Macular Degeneration** (AMD) and Stargardt disease

Modern drug discovery process, inspired by traditional medicine

Our technology

Reverse pharmacology for drug candidates in Age Related diseases

Build a proprietary collection of natural molecules & analogs from medicinal plant, produced under biotic or abiotic stress

Screen in cellular models of age related diseases and identification of targets & pathways Selection of best drug candidates based on animal models of aging or genetic diseases







- Sarconeos (BIO 101)
- Macuneos (BIO201)

- Small molecules: natural and/or NCE (new chemical entity)
- New key target against aging
- Preclinical proof-ofconcept & safety
- IP on use, process and composition of matter



Executive team







Shire





Stanislas Veillet - Founder & CEO

- PhD in genetics, AgroParisTech
- 25+ years in biotech; Pharmacia-Monsanto, Danone Group



Samuel Agus - CMO

- MD, PhD, Board-certified Neurologist
- 15+ years pharma/biotech experience including Abbott, Shire and Teva Pharmaceuticals



Pierre Dilda - CSO

- PhD in pharmacology (Paris V)
- 25 years experience in pharmaceutical research, in both academic and industrial settings



Waly Dioh - COO

- PhD in phytopathology (Paris XI) and MBA
- 21+ years biotech experience in France and the U.S. and R&D at Monsanto



Evelyne Nguyen- CFO

- 30+ years of experience in Corporate Finance for International Pharma & Biotech companies (BMS, LFB)
- Expertise in cross-borders transactions between Europe, US and Asia



Our clinical pipeline

Candidate	Indication	Program	Preclinical	Phase 1	Phase 2	Phase 3
Sarconeos (BIO101) Macuneos (BIO201)	Covid-19	COVA				
	Sarcopenia	SARA				
	DMD	MYODA				
	Dry AMD	MACA				
	Stargardt					

• Second generation drug candidates, BIO103 and BIO203, are life-cycle extension candidates in the preclinical Phase



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Key milestones

- COVA started in the Belgium, Brazil, France and US
- COVA completion of patient enrollment (Part 1) in January 2021
- COVA interim analysis expected in Q1 2021
- COVA completion of patient enrollment (Part 2) expected in Q1 2021
- COVA topline results and expected in Q2 2021
- SARA-INT (Phase 2) patient enrollment completed in March 2020 SARA-INT last patient out (LPO) completed in Dec 2020
- □ SARA-INT topline trial results expected in Q2 2021

MYODA (Phase 1 to 3) IND approved in Belgium and US

□ MYODA FPI expected in H1 2021 depending on COVID-19 pandemic evolution



Sarconeos (BIO101) activates MAS receptor, a key factor for muscle and respiratory functions

- MAS receptor: a key component of the Renin-Angiotensin System (RAS)
- Triggers two important downstream signaling-pathways in myocytes:



PI3K/AKT/mTOR: Increases **protein synthesis**, preserving muscle mass and increasing **muscle strength**

AMPK/ACC Stimulates energy production, increasing muscle strength and mobility



Sarconeos (BIO101) for respiratory failure in COVID-19

- SARS-CoV-2 uses ACE2 to penetrate the lungs destabilizing RAS system and causing respiratory failures
- Sarconeos (BIO101) activates MAS receptor, a key component of the protective arm of the RAS system





We believe Sarconeos (BIO101) improves muscle strength and mobility in animal model



Administration of 50 mg/kg/day of Sarconeos (BIO101) demonstrated a statistically significant (p<0.01) improvement in maximum running velocity (Vmax) compared to "old" control mice, compensating almost completely for the loss of mobility due to aging



Administration of 50 mg/kg/day of Sarconeos (BIO101) demonstrated a preservation of muscle strength while immobilized (d0-d14) compared to vehicle control in hind limbimmobilized mice

1. Results were presented in a poster at the SCWD conference in December 2016 in Berlin, Germany.



Early data suggests Sarconeos (BIO101) improves respiratory functions in animal model



C57BL10-mdx mice treated with 50 mg/kg/day of Sarconeos (BIO101) over 8 weeks showed a-improvement in airway responsiveness (PenH) as compared to untreated control C57BL10-mdx mice



1. Results were presented in October 2019 at the WMS conference in Copenhagen, Denmark in a poster presentation;

2. Results were presented in March 2019 at the annual international congress of Myology in Bordeaux, France



Phase 1 study (SARA-PK) results in elderly healthy volunteers with Sarconeos (BIO101)



Sarconeos (BIO101) showed a dose dependent effect on muscle growth and repair (PIIINP) and a dose dependent negative e effect on muscle wasting (myoglobin) for Sarconeos BIO101

- Single and multiple ascending doses tested in 54 healthy young and elderly (over 65 years) volunteers
- Safety profile : No Severe Adverse Events
- Two active doses (175 & 350 mg b.i.d.) have been selected for the upcoming Phase 2 studies



Sarcopenia: a large unmet medical need with no currently **#SARA** approved drugs

- Age-related degeneration of skeletal muscle characterized by a loss of muscle mass, strength and functional issues such as the ability to stand and/or walk
- A major cause of mobility disability, resulting in a loss of independence and increased risk of adverse events (for example falls), which can shorten life expectancy

Sarconeos (BIO101):

- ✓ Only drug candidate in Phase 2 currently being tested for sarcopenia
- ✓ Myostatin inhibitors halted for lack of effectiveness in neuromuscular diseases



SARA-INT: Phase 2 clinical trial in sarcopenia

#SARA

- Global, double-blind, randomized, placebo-controlled trial: NCT03452488
- Recruitment completed March 2020 for 233 elderly patients with sarcopenia at risk of mobility disability over 22 centers in the US and Belgium

Objectives

- Assess safety and efficacy of two doses of Sarconeos (BIO101) administered orally with a meal over 26 weeks, compared to placebo
- Treatment effect on improvement of physical function and on decrease of risk of mobility disability

Key Endpoints

Primary

 400-meter walk test (400MWT)
 - 0.05 m/s is considered the minimal meaningful change

Key secondary

- Changes in time to rise from a chair.
- 400MWT responder analysis
- Patient reported outcomes (PRO)

- Age (≥65 or over)
- Low mobility measured by Short Performance Physical Battery (SPPB) ≤8 out of 12
- DEXA body composition as measured by ALM/BMI (appendicular lean mass / body mass index)
- Able to exercise for 30 minutes per day 5 days per week





SARA-INT: topline results expected in Q2 2021



Dr. Roger Fielding, PhD, Director of the Nutrition, Exercise Physiology & Sarcopenia Laboratory at **Tufts University** in Boston and Principal Investigator of SARA-INT trial

Product	2019	2020	2021
175 & 350 mg b.i.d of Sarconeos (BIO101)		SARA-INT Phase 2	

- No safety issue to date, with multiple DSMB/DMC Meetings with the conclusion that the benefit risk ratio is positive.
- Last patient out in December 2020 with 196 patients having completed the study.
- Top line trial results expected in Q2 2021.



COVA Study: targeting hospitalized patients with respiratory failure, not intubated



- Patients aged 45 and above, with proven COVID-19, and severe respiratory symptoms:
 - With evidence of respiratory decompensation ≤4 days before start of study medication, meeting one of the following:
 - Tachypnea: ≥ 25 breaths per minute
 - Arterial oxygen saturation ≤92%
- Allowed medications:
 - antiviral agents such as Remdesivir, Bamlanivimab,
 - anti- inflammatory agents such as Dexamethasone





COVA : Sarconeos (BI0101) evaluating prevention of further respiratory deterioration linked to COVID-19



•	This is a Phase 2/3	Part		Goal	Analysis by the DSMB/DMC	Number of participants
	 seamless study design Multinational, multi-centric Double-blind, placebo controlled Group sequential (2 parts), adaptive 		1. 2.	Obtain safety and tolerability data on (BIO101) Obtain an indication of activity for BIO101	IA1: 1 st interim analysis Decide on the beginning of part 2 recruitment Assess indication of activity of BIO101	50 1:1 randomization
			Re-a step	issess the sample size for 2	IA2: 2 nd interim analysis to confirm sample size for Part 2	155 (an addition of 105 participants) 1:1 randomization
	 design Sarconeos (BIO101) 350mg BID vs. placebo 			firmation of the effect of 101 in preventing further iratory deterioration	-	310, potentially increased by 50%(up to 465, based on interim analysis 2)1:1 randomization
	Product			2020		2021
	350 mg b.i.d of Sarconeos (BIO101)				COVA Study	



DMD: No cure and limited treatment options



Proportion of ambulatory class in DMD¹



Ventilation support

Late nonambulatory (age 16 or older)



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■ Late ambulatory (age 8–11)
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Early ambulatory (age 5–7)

- Rare, genetic neuromuscular disease in male children characterized by accelerated degeneration of muscles, responsible for loss of mobility, respiratory failure and cardiomyopathy, leading to premature death.
- No known cure and limited treatment options, including corticosteroids and targeted therapies (exon-skipping in U.S. & stop codon in EU) that treat approximately 13% of DMD patients with specific genetic mutations.
- We received **orphan drug designation** (ODD) in 2018 from the FDA and EMA for Sarconeos (BIO101) in DMD.



MYODA: IND granted by FDA to start Phase 2/3 clinical study



Product	2020 2021		2022	2023
Sarconeos (BIO101)			MYODA (Phase 2/3)	

Design	Patients	Regulatory Status
 Global, multicenter, double-blind, placebo-controlled, seamless, Phase 1-2-3 clinical trial 	 Non-ambulatory DMD patients: Part 1: 18 participants Part 2: an addition of 30 	 Orphan drug designation in US and Europe granted in 2018
 Part 1: Safety, tolerability & PK (initial 7 days of dosing of escalating dose of Sarconeos BIO(101) 	 participants Part 3: up to 200 participants Interim analysis at the end of 	• FDA IND and CTA in Belgium granted in 2020
 Part 2: Efficacy of Sarconeos (Respiratory function after dosing for 52 weeks) 	 parts 1 and 2 Enrollment in the U.S. and EU Patient advocacy group support 	
 Part 3: Efficacy of Sarconeos (Respiratory function after dosing for 52 weeks) 	AFM Téléthon in France	

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MACA

Dry AMD is an unmet medical need with no approved drugs

- AMD is a common eye disorder among people over 50 that affects the central part of the retina, known as the macula
- Can impair functions such as reading, driving, and facial recognition, and has a major impact on QoL and the ability to live independently
- Multifactorial disease that we believe is mainly caused by accumulation of A2E (a byproduct of the visual pigment cycle) that leads to retinal degeneration





Projection of AMD prevalence in Europe (in M, mean projection)¹



- 85 90% of AMD patients have dry AMD in some form; either early, intermediate or late stage, known as geographic atrophy (GA)
- No approved treatments for any stage of dry AMD, including GA
- We are developing Macuneos to treat patients with intermediate dry AMD to prevent the development to advanced stages (wet AMD + GA), which lead to severe vision loss



Macuneos (BIO201) mechanism of action: Noncanonical activation of PPARs

- We believe Macuneos potentially counteracts the phototoxic effects of A2E by selective noncanonical activation of the transrepressive activity of PPAR α and PPAR β/δ in the retina
- Most other PPAR ligands mainly exhibit canonical activity and are associated with side effects



Anti-inflammatory activity (promotes the expression of anti-inflammatory genes)

Anti-oxidant activity (promotes the expression of anti-oxidant genes)

Anti-apoptotic activity (enables pathways that prevent cell death)

MACA

Macuneos (BIO201) protects the retina in rodent models of dry AMD and Stargardt disease



Results were presented in May 2016 at the ARVO conference in Seattle, WA in a poster presentation and published in PLoSONE (Fontaine *et al.;* 2016).

MACA

Appendix



Scientific advisory board









University College London Hospitals NHS Foundation Trust





Pr. Jean Mariani, Chairman

- Professor of neuroscience and biology of aging and Director of Charles Foix Institute of Longevity at Sorbonne University
- Emeritus Professor (PU-PH) at the Sorbonne University's School of Medicine



Dr. Roger Fielding

- Professor of Medicine, Tufts University School of Medicine
- Director and Sr. Scientist Jean Mayer USDA Human Nutrition Research Center on Aging



René Lafont

- Co-Founder & Professor emeritus and former Dean of the life sciences department at Sorbonne University
- 185 scientific articles + 59 reviews and book chapters



Pr. Jose-Alain Sahel

- Chair of the department of ophthalmology at University of Pittsburgh School of Medicine and director of the UPMC eye center
- Founder and director of the Vision Institute in Paris and professor at the Sorbonne's medical school



Dr. Thomas Voit

- Professor, University College London
- Director of the Research Centre of the Great Ormond Street Hospital for Children



Dr. Ivana Kim

- Associate Professor Harvard Medical School, Massachusetts Eye and Ear
- Co-Director of the Harvard Medical School Department of Ophthalmology AMD Center of Excellence; Associate Scientist, Massachusetts Eye and Ear



Board of directors

🕑 Replimune

MONSANTO 🛔











Stanislas Veillet - Founder & CEO

- PhD in genetics, AgroParisTech
- 25+ years in biotech; Pharmacia, Monsanto, Danone & Cargill



Pr. Jean Mariani

- Professor of neuroscience and biology of aging and Director of Charles Foix Institute of Longevity at Sorbonne University
- Emeritus Professor (PU-PH) at the Sorbonne University's School of Medicine



Jean M. Franchi

- Independent Director,
- Replmmune CFO
- 30+ years as finance director, including 15 years at Genzyme



Dimitri Batsis

- Independant Director
- Entrepreneur, Founder of Zeni Corporation, Drone Volt
- 20 years in the High-Tech sector



Nadine Coulm

- Independant Director
- 20 years of experience as CFO in charge of IR & Financing with Korian, FNAC, Darty Danone & Casino

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From a Sorbonne University spin-off to a successful clinical-stage biotechnology company: 90M€ raised to date

 Incorporated of Sept 26, 2006 First patents filed in 2007 Raised €800 K with Seventur Partners 		 IPO on Altern Paris that rais €10.0 M Opened U.S. subsidiary in Cambridge, N Raised €6.0M private placement 	IA	 Initiated SARA INT Phase 2 clinical trial Received orph drug designat for Sarconeos (BIO101) in DI in US and Euro Raised €10M in notes 	ran (ion (VID E ope E n • 3	aunch of COVA project with Garconeos BIO101) in COVID-19, started in US, Europe and Brazil BOM€ raised in new equity
2006 - 2008	2009- 2012	2015	2016-2017	2018	2019	2020
SORBONNE UNIVERSITÉ CREATEURS DE FUTURS DEPUIS 1257	 2009 €2.2 M Round A with VC's 2009 launcheo first human clinical trials for Sarconeos (BIO101) 2012 €3.0 M Round B with VC's 		 2016 initiate and complete SARA-PK Phat clinical trial 2017 raised €18.0 M in private placements Set up €15M bond financia Initiated SAR OBS clinical te 	ng A-	 Set up €24M bond financing Filed and granted IND for Sarconeos (BIO101) in DN 	r

Intellectual Property portfolio – Neuromuscular & respiratory diseases

- We hold exclusive commercial rights through licenses of each of our drug candidates.
- IP is jointly owned with Sorbonne University & sometimes with other academic research institutions^{1.}
- Patent portfolio covers 10 patent families, including a total of <u>24 co-owned issued patents</u> and a total of <u>26 co-owned patent applications</u>.
- Issued patents: 5 European, 2 U.S., and 17 in ROW, including **China, Japan**.
- Pending applications: 2 European, 5 U.S., and 19 in ROW, including **China, Japan, South Korea**



10 families of patents covering production process, second generation compounds and various applications such as sarcopenia, myopathies (DMD), disuse atrophy, spinal muscular atrophy, respiratory function and COVID-19



Intellectual Property portfolio – Retinal Diseases

- We hold exclusive commercial rights through licenses of each of our drug candidates.
- IP is jointly owned with Sorbonne University & sometimes with other academic research institutions.
- Patent portfolio covers 5 patent families, including a total of <u>16 co-owned issued patents</u> and a total of <u>10 co-owned patent applications</u>.
- Issued patents: 4 European, 3 U.S., and 9 in ROW, including **China, Japan**.
- Pending applications: 10 in ROW, including **China, Japan, South Korea.**



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Thank you

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