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### An experienced and complementary management team



Guy-Charles Fanneau de La Horie, **CEO | MBA, Veterinary Doctor** 

Over 25 years of experience in the international pharmaceutical and biotech industry













Philippe Rousseau, MBA **Chief Operating Officer** 

- Over 20 years of experience in senior positions in the life sciences industry
- Ex-Deputy CEO ABCDx











**Cindy Fevre R&D Director, PhD** 

- International R&D experience
- Microbiologist, expert in bacteriology
- PhD at the Pasteur Institute on Antimicrobial Resistance





#### Frédérique Vieville **Quality Director, PharmD**

- 20 years of experience as Quality Director
- Doctor in Pharmacy and Biotechnology Engineering (Universities Blaise Pascal Clermont II, Montpellier I and Polytech Clermont-Ferrand)











#### **Brigitte Palestro Medical Director, MD**

- 24 years of experience in the pharmaceutical industry
- Graduate of the Lyon I and Paris XII Faculties of Medicine
- Medical Director in large pharmaceutical groups

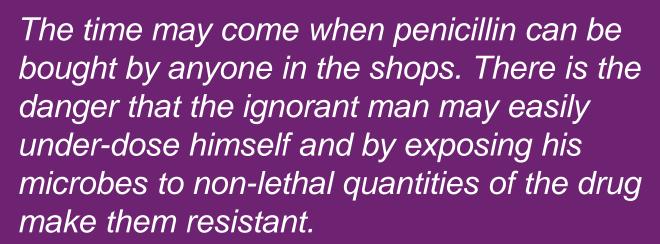










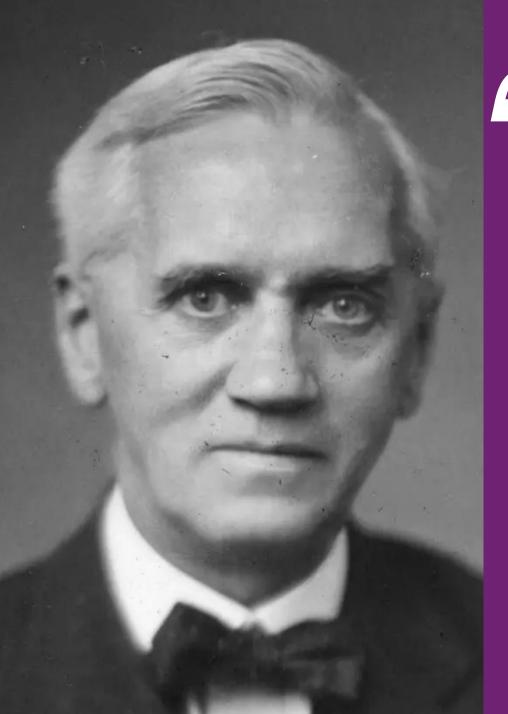


Sir Alexander Fleming, Inventor of penicillin Nobel Prize speech





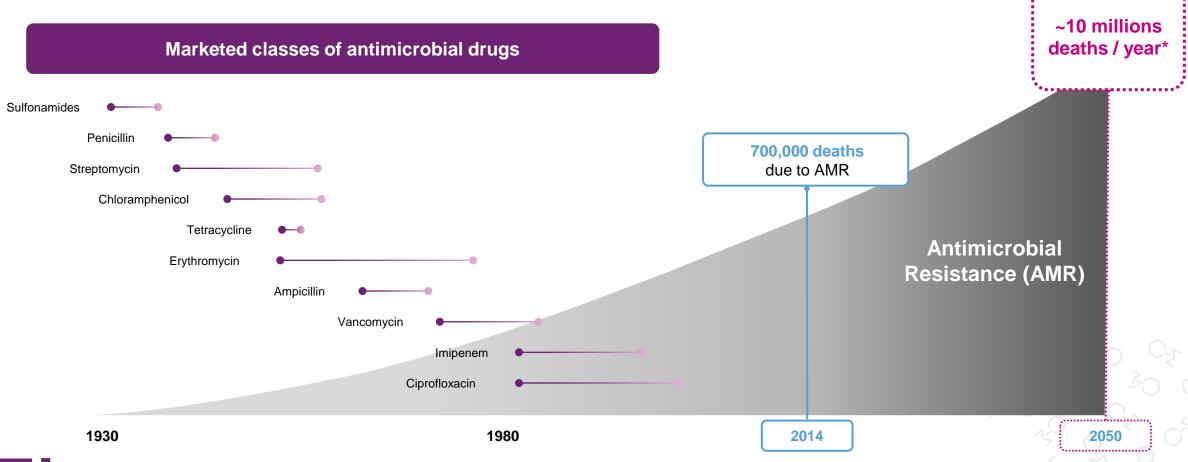








## Antimicrobial resistance, a critical global public health issue

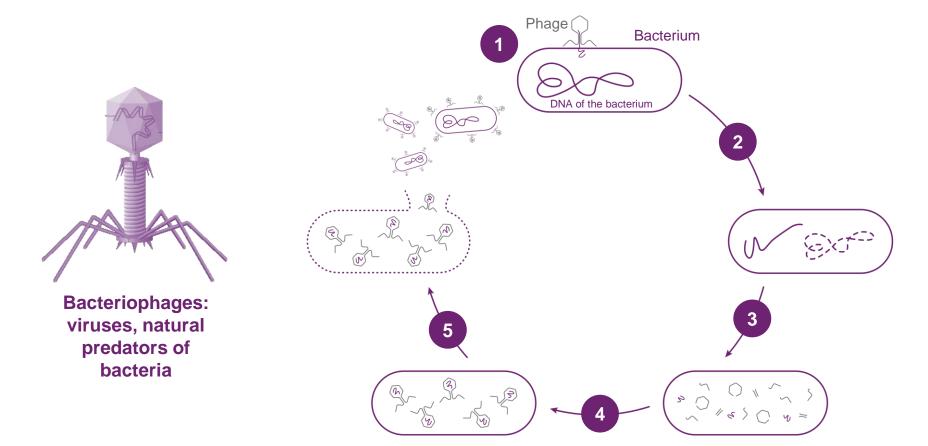




Absolute need to address antimicrobial resistance with an effective response

Sources: Jim O'Neill's Report, 2016 - Stephen R. Palumbi, "Humans as the world's greatest evolutionary force", Science, vol. 293, 2001, p. 1786-1790 (PMID 11546863)

## Phage therapy: a natural solution to treat resistant infections



#### Unique mode of action

- Specificity
- Speed (less than 45 min)
- Self-replication down to the last bacterium

C¥

Phage therapy allows simple, effective and well-tolerated treatments

## Pherecydes' innovative approach: precision phage therapy

- SELECT phages targeting a maximum number of bacterial strains to better "cover" the targeted bacterial species
  - → Pherecydes technology makes it possible to screen and characterize large quantities of phages in order to select only the most efficient ones
- PRODUCE on a large scale the retained phages in conditions of optimal purity
  - → Pherecydes has industrial partnerships to produce its phages in accordance with GMP standards
- RETAIN for each treatment, the active phage(s) on the bacterial strain in question
  - → Pherecydes identifies in vitro the most effective target phages in order to propose a customized treatment

Compassionate treatment in 22 patients in France



Individualized treatments adapted to each case

## Favorable regulatory opinion

Design of the next phase I/II study in line with the recommendations of the European Medicines Agency (EMA)

### Unique R&D skills combined with an unquestionable IP policy

R&D IP 0 Patents

#### **Recovery of natural** phages

The phages are recovered in their natural environment (sewers,...)

#### Screening

R&D teams select phages of interest

#### Characterization

The phages are isolated and characterized by Next-Generation Sequencing (NGS)

#### **Protection**

Filing of patents on individual phages, their variants and their associations

4 patents already granted in high-potential areas

















## **Tangible fundamentals for controlled development**

## Pherecydes Pharma, at the forefront of phage therapy against the most resistant bacterial infections

1 priority

To address 3 of the most critical bacterial infections according to the WHO

families of phages

- anti-Staphylococcus aureus
- anti-Pseudomonas aeruginosa
- anti-Escherichia coli

22 patients

treated as part of compassionate treatments in prestigious hospital centers









4 patents

including some issued in the United States, Europe, Japan, Hong Kong and Australia

21 experienced staff members

including 1 MD, 1 Pharm D, 1 DVM & 5 PhD





Developments within a regulatory framework validated by key health authorities



Network of prestigious scientific partners











01

Antimicrobial resistance: a fast growing market with strong medical needs

## Antimicrobial resistance: a global health issue declared a priority by the WHO





700,000 deaths worldwide in 2014<sup>1</sup>

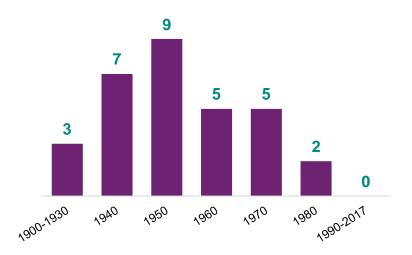
35,000 deaths in the US in 2017<sup>2</sup> 33,000 deaths in Europe in 2019<sup>3</sup>

10,000,000 deaths expected/year by 2050<sup>4</sup>



#### Very few new weapons

No. of new classes of antibiotics approved<sup>5</sup>

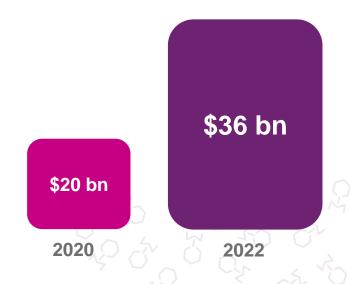






#### **Pressure on health systems**

Annual cost of antimicrobial resistant infections in the United States<sup>6</sup>





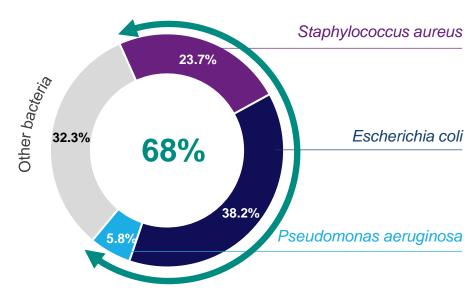
#### A growing and extremely heavy societal cost for health systems

Sources: (1) https://www.who.int/news/item/29-04-2019-new-report-calls-for-urgent-action-to-avert-antimicrobial-resistance-crisis, (2) https://www.edc.gov/drugresistance/biggest-threats.html, (3) https://www.edc.europa.eu/sites/default/files/documents/surveillance-antimicrobial-resistance
Europe-2019.pdf; (4) Report by Jim O'Neill, 2016, requested by the British PM, (5) Carb-X https://carb-x.org/about/global-threat, (6) https://www.marketresearchfuture.com/reports/hospital-acquired-infections-market-2576

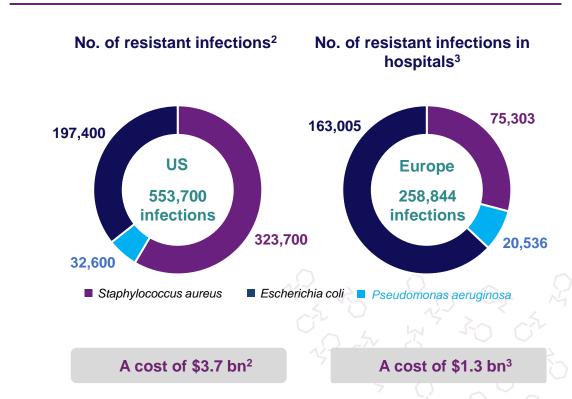
## 3 bacteria targeted by Pherecydes representing 2/3 of resistant hospitalacquired infections

Distribution of resistant infections by type of bacteria

68% of resistant infections in hospitals are caused by the 3 Pherecydes targets<sup>1</sup>



Addressable market corresponding to resistant infections caused by these 3 bacteria





## A cost of \$5 billion for healthcare systems in Europe and the United States

Sources: (1) Opatowski M - Hospitalisations with infections related to antimicrobial-resistant bacteria from the French nationwide hospital discharge database, 2016; (2) Hôpital et ville - CDC (Centers for disease Control and Prevention) – Antibiotic Resistance Threats in the US; (3) https://www.ecdc.europa.eu/sites/default/files/documents/surveillance-antimicrobial-resistance-Europe-2019.pdf

## The global market for resistant infections represents a unique opportunity

#### Considerable number of resistant infections

> 800,000 resistant bacterial infections / year<sup>1</sup>

> 670,000 resistant bacterial infections / year<sup>2</sup>





Cost of antimicrobial resistance

Between \$20,000 and \$80,000/patient<sup>3</sup>

#### Mortality and hospitalizations rising sharply<sup>4</sup>

**Mortality x2.4** 

Hospitalizations extended from 9.3 to 13 days

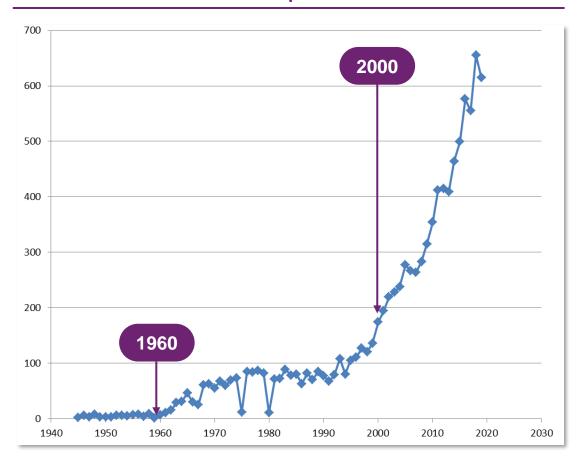
#### Price of new generation of antibiotics

\$8,000 - \$10,000/patient<sup>5</sup>

Source: (1) CDC (Centers for disease Control and Prevention) – Antibiotic Resistance Threats in the US; (2) <a href="https://www.ecdc.europa.eu/sites/default/files/documents/surveillance-antimicrobial-resistance-Europe-2019.pdf">https://www.ecdc.europa.eu/sites/default/files/documents/surveillance-antimicrobial-resistance-Europe-2019.pdf</a>, (3) <a href="https://www.ecdc.europa.eu/sites/default/files/documents/surveillance-antimicrobial-resistance-Europe-2019.pdf</a>, (3) <a href="https://www.ecdc.europa.eu/sites/default/files/documents/surveillance-antimicrobial-resistance-Europe-2019.pdf</a>, (4) <a href="https://www.ecdc.europa.eu/sites/default/files/documents/surveillance-antimicrobial-resistance-Europe-2019.pdf</a>, (5) <a href="https://www.ecdc.europa.eu/sites/default/files/antimicrobial-resistance-Europe-2019.pdf</a>, (6) <a href="https://www.ecdc.europa.eu/sites/default/files/antimicrobial-resistance-Europe-2019.pdf</a>, (7) <a href="https://www.ecdc.europa.eu/sites/default/files/antimicrobial-resistance-Europe-2019.pdf</a>, (8) <a href="https://www.ecdc.europa.eu/sites/antimicrobial-resistance-Europe-2019.pdf">https://www.ecdc.europa.eu/sites/antimicrobial-resistance-Europe-2019.pdf</a>, (9) <a href="https://www.ecdc.europa.eu/sites/antimicrobial-resistance-Europe-2019.pdf">https://www.ecdc.europa.eu/sites/antimicrobial-resistance-Europe-2019.pdf</a>, (9) <a href="https://www.ecdc.europa.eu/sites/antimicrobial-resistance-Europe-2019.pdf">https://www.ecdc.europa.eu/sites/antimicrobial-resistance-Europe-2019.pdf</a>, (9) <a href="https://www.ecdc.europa.eu/sites/antimicrobial-resistance-Europe-2019.pdf">https://www.ecdc.europa.eu/sites/antimicrobial-resistance-Europe-2019.pdf</a>, (9) <a href="https://www.ecdc.europe-antimicrobial-resistance-Europe-2019.pdf">https://www.ecdc.europe-antimicrobial-resistance-Europe-2019.pdf</a>, (9) <a href="https://www.ecdc.europe-antimicrobial-resistance-Europe-antimicrobial-resistance-Europe-antimicrobial-resistance-Europe-antimicrobial-resistance-Europe-antimicrobial-

### **Growing scientific and media interest in phage therapy**

## Exponential growth in the number of scientific publications

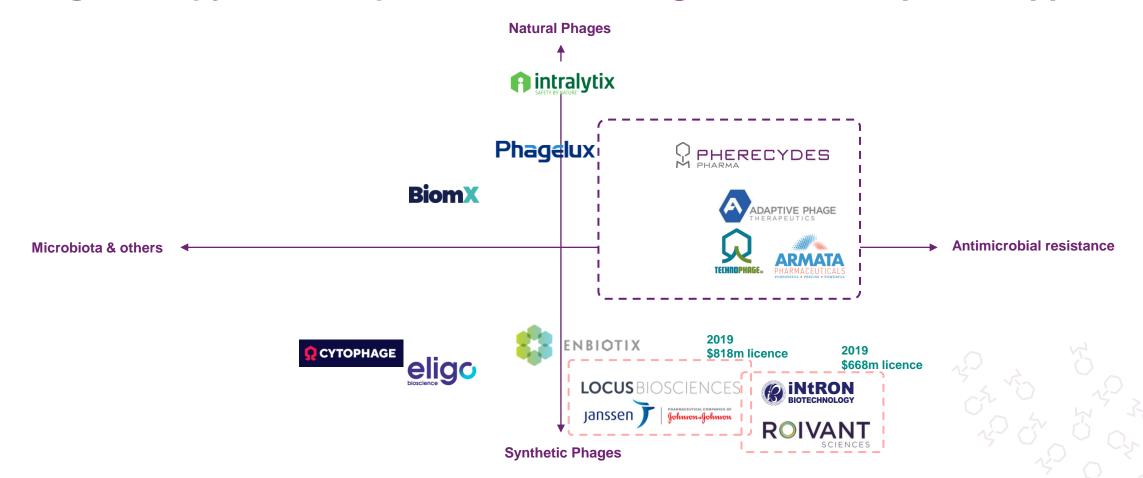


## Therapeutic interest of phages widely publicized in the media



Source: Company on the basis of PubMed publications

## Phage therapy is developed for different targets and with specific approaches



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Pherecydes, at the forefront of research with a rich and diversified portfolio in human health



02

# Pherecydes' precision phage therapy: an innovative approach

## Precision phage therapy: a unique concept developed by Pherecydes

#### **Physicians**

Sending of the patient's bacterial strain with the agreement of the ANSM



## PHERECYDES

- Analysis by Phagogram (<48h)</li>
  - Only active phages are selected



#### **Pharmacists**

Phages are assembled in sterile conditions in a physiological solution



#### **Patients**

Phages are administrated according to a procedure defined by the physician and the pharmacist



## PHERECYDES

Phagogram - selection of "custom-made" phages



In vitro measurement of phage activity on the collected bacterial strain



Selection of the most active phages on the pathogenic strain



Assembly of the selected phages by the hospital pharmacy

Currently internalized and in the future in partnership



An approach that can be replicated for an unlimited number of bacterial targets

## Pherecydes Platform: selection of phages combining therapeutic and industrial efficacy

Selected Coverage of the reference **Activity on clinical strains Phages** panel\* **Targets Selection of phages** Therapeutic benefits In compassionate treatment meeting the criteria: 100% **78%** of efficiency Staphylococcus aureus of purity of production capacity 98% 80% Pseudomonas aeruginosa 91% Escherichia coli n.a.



## Competitive advantage constituting a major barrier to entry

Note: \* Panels used: Staphylococcus aureus: CNR Staphylococci; Pseudomonas: De Soyza; E. Coli: Donamur – national reference center for enterobacteria

## Pherecydes' 1st target: Staphylococcus aureus





- 2<sup>nd</sup> bacterium most responsible for hospital-acquired infections in Europe
- High global priority according to the WHO¹
- >25% of Staphylococcus aureus infections are resistant

Initial targeted indications

- Bone and joint infections on prosthesis: up to 2% of hip and 3% of knee prostheses
- Diabetic foot ulcer: 15% of patients with an infection rate of 40-80%, leading cause of lower limb amputations in developed countries

#### Results to date

- 2 toxicological studies revealed no signs of toxicity:
  - an acute toxicity study was performed on mice
  - a 21-day local tolerance and toxicity study was conducted on mini-pigs
- Lytic activity demonstrated in a biofilm



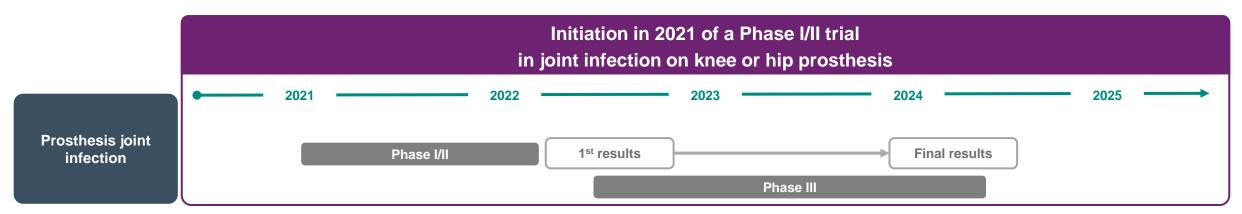
#### **Intellectual Property**

- Filing of patents on individual phages and their variants in 2018
- Protection of all associations including at least one phage

Sources: (1) https://www.who.int/fr/news/item/27-02-2017-who-publishes-list-of-bacteria-for-which-new-antibiotics-are-urgently-needed; (2) CDC - U.S. Department of Health and Human Services - Centers for Disease Control and Prevention

#### Pherecydes' 1st target: Staphylococcus aureus





#### Design of the study

- Comparative Phase I/II study on 60 to 80 patients\*
- Collaboration with attached Reference Centers for the Treatment of Complex Bone and Joint Infections (CRIOAc)
- Treatment with anti-Staphylococcus aureus phages active on each strain (inclusion condition)
- Primary endpoint: the efficacy and tolerance of phages in association with the DAIR procedure (Debridment, Antimicrobials, Implant Retention) 3 months post-operation



#### Main recommendations of the EMA:

Randomized and controlled study

Standardization of reference antimicrobial treatment

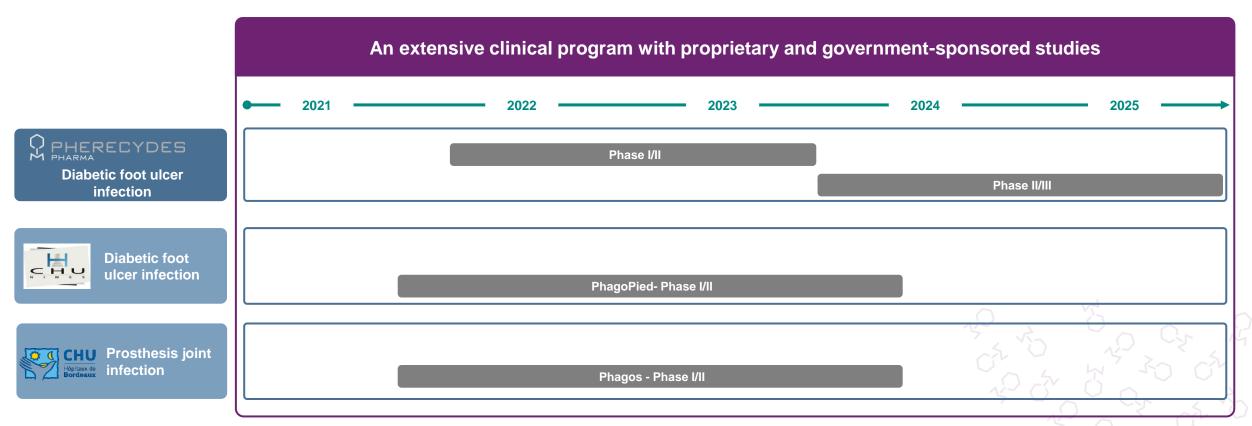


#### Provide proof of concept of the clinical value of precision phage therapy

Note: \* Number of patients and design refined following the results of a retrospective study currently being conducted with CRIOAc

## Pherecydes' 1st target: Staphylococcus aureus



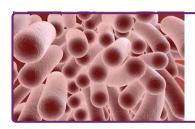




A variety of clinical programs to de-risk the development of this key asset

## Pherecydes' 2<sup>nd</sup> target: *Pseudomonas aeruginosa*





- One of the most difficult bacteria to treat
- Global critical priority according to the WHO¹
- A very strong impact on patient mortality

Initial targeted indications

- Ventilation-associated pneumonia
- Pneumonia associated with cystic fibrosis

Results to date

- Significant decrease in the pulmonary bacterial load
- Regulatory preclinical development (toxicity studies) completed
- Excellent results with nebulization administration in an animal model of infection
- Very good tolerance, including with intravenous injections, as part of 1st compassionate treatments
- Production on a bacterial strain genetically modified to ensure the purity of the phages

PHERECYDES

- Next steps Provision of phages under Early Access Program
  - Evaluation of a clinical protocol in a cystic fibrosis affected population

Intellectual Property

- Patents on individual phages and their variants in the USA, Australia, Japan and Europe
- Protection of all associations including at least one phage

Sources: (1) https://www.who.int/fr/news/item/27-02-2017-who-publishes-list-of-bacteria-for-which-new-antibiotics-are-urgently-needed (Company)

## Compassionate treatments: 22 patients have benefited from Pherecydes' phage therapy in France

- Several routes of administration tested, including intravenous administration
- +7 different indications treated with a majority of bone and joint infections

Indication / Infection	Targeted bacteria
Knee prosthesis	<ul><li>Staphylococcus aureus</li><li>Pseudomonas aeruginosa</li><li>Staphylococcus lugdunensis (?)</li></ul>
Hip prosthesis	<ul><li>Staphylococcus aureus</li><li>Pseudomonas aeruginosa</li></ul>
Bone Minerals	<ul> <li>Pseudomonas aeruginosa</li> </ul>
Others (Complicated pelvic fracture, cranioplasty, diabetic foot, endocarditis on valve, bacteremia on deep thoracic fistula.)	<ul><li>Staphylococcus aureus</li><li>Pseudomonas aeruginosa</li></ul>

Systematic support of the ANSM and health centers











#### **Tangible results**

- **Excellent results observed in 10 cases of infections:** 
  - phages well tolerated
  - beneficial clinical and/or microbiological effect
- Scientific publication of the results on 3 patients









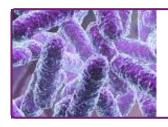




**Encouraging results validating Pherecydes' approach** 

## Pherecydes' 3<sup>rd</sup> target: *Escherichia coli*





- Global critical priority according to the WHO¹
- Main cause of resistant hospital-acquired infections

## Initial targeted indications

Complicated urinary tract infections

## Results to date

- In vitro results: coverage of more than 91% of the representative panel
- First results *in vivo* (mice): demonstration that phages penetrate the kidneys and the bladder after intravenous administration validating the 1<sup>st</sup> selected indication
- Continued preclinical development (including regulatory toxicity studies)
- Development of GMO production strains
- 2 clinical studies planned:
  - phase I/II study: demonstrate antimicrobial activity in the intravenous urinary tract and define the optimal treatment design
  - phase III study: prove the efficacy of the treatment when added to the standard treatment

## PHERECYDES

Intellectual Property

**Next steps** 

- Patents on individual phages and their variants in the USA, Australia, Japan, Europe and Israel
- Protection of all associations including at least one phage

Sources: (1) https://www.who.int/fr/news/item/27-02-2017-who-publishes-list-of-bacteria-for-which-new-antibiotics-are-urgently-needed (Company)

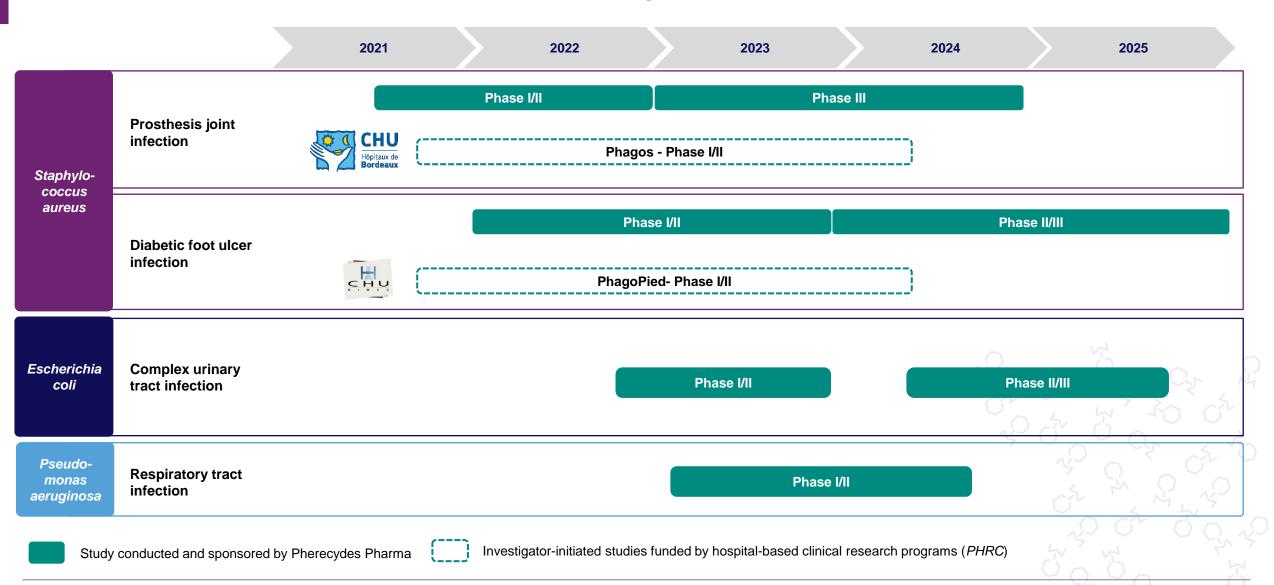


03

Goal: to further establish the leadership position in phage therapy

A de-risked development based on the choice of the priority indications, the compassionate treatments and the Phagogram

## Sustained and diversified clinical development in critical indications



## A process that has already begun, with initial results expected by the end of 2022 in prosthesis joint indications



- Dedicated internal team led by Dr. Brigitte Palestro, Medical Director
- 25 years of experience in clinical research and regulatory issues
- Regular communication on the study's progress









A tier-one CRO with personalized services and advice





#### Notices from regulatory authorities

- Study design developed with the EMA since 2019
- Regular consultations with the ANSM

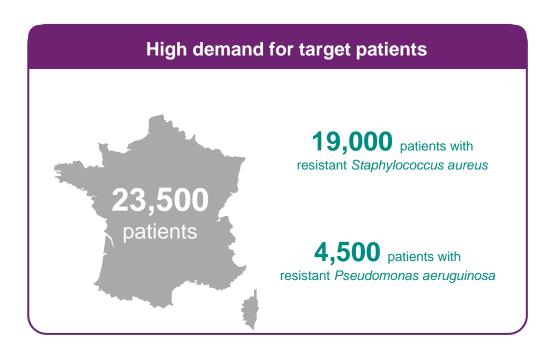
#### **Network of reference centers**

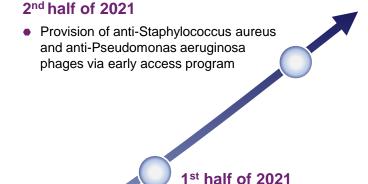
- Principal Investigator: Prof. Tristan Ferry, Infectiologist and Coordinator of CRIOAc Lyon
- Network of 24 French and European reference centers
- Strong recruiting capacity



A robust organization for the completion of this structuring clinical study

### Early access program expected to start H2 2021





Availability of GMP phage batches

#### November 2020

 Production of the first GMP batches

#### August 2020

 GMP production in partnership

#### Early access program System

**Early access program** allows certain categories of patients in France to use drugs that have not yet received full marketing authorization (MA).



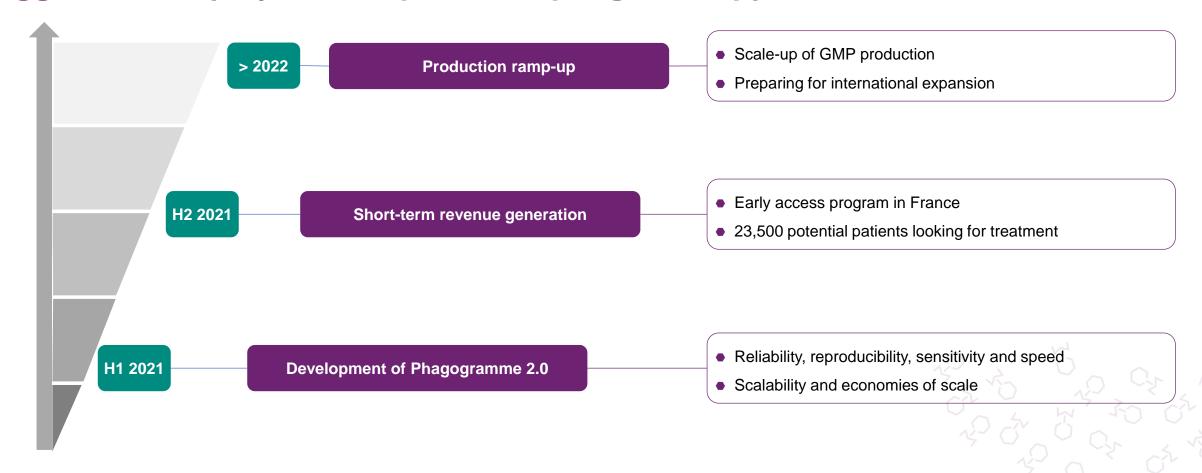
#### March 2019

- Temporary specialized scientific committees with a favorable opinion from the ANSM
- Need for GMP production

## Meeting the most urgent medical needs in the very short term

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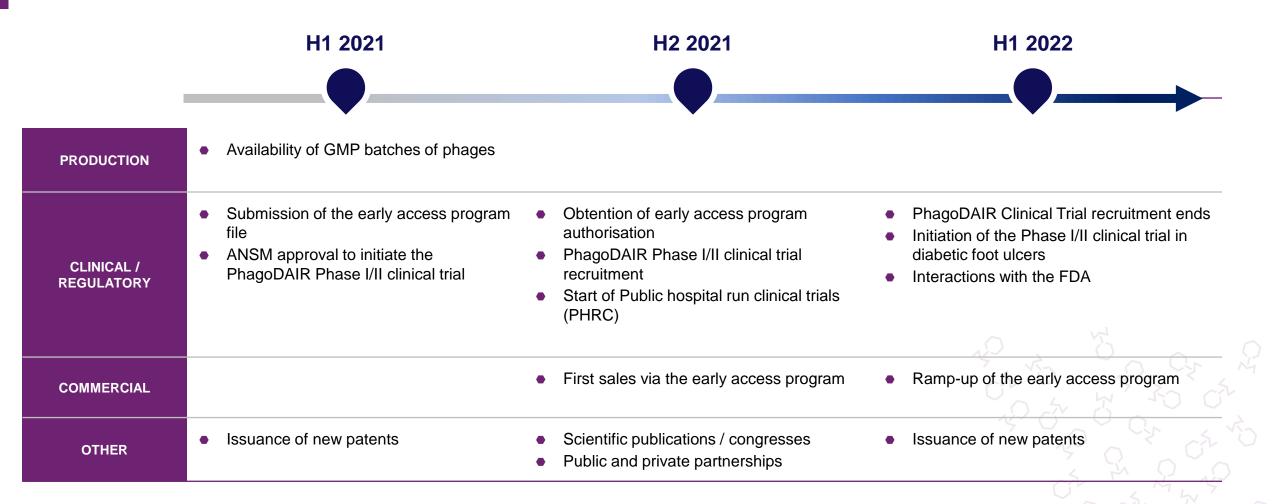
### **Aggressive deployment of precision phage therapy**



2

Leveraging unique competitive advantages to make Pherecydes the leader in precision phage therapy

#### Intense newsflow over the next 18 months



## Pherecydes: a de-risked development based on IP, choice of priority indications and success of compassionate treatments

Filling a critical unmet need for treatment of resistant bacterial infections with no satisfactory therapeutic solution to date: a fast-growing market estimated at \$5 billion

A diversified product portfolio, protected by international intellectual property rights, targeting the 3 of the most resistant and dangerous bacteria according to the WHO

Priority development of the treatment of prosthesis joint infections due to Staphylococcus aureus, where Pherecydes has already obtained promising results in compassionate treatment

Strong relationship with French and European health authorities and reference centers for complex bone and joint infections, enabling the development of phage therapy

A ramp-up supported by potential revenues thanks to the availability of phages, as of H1 2021, via the French early access program

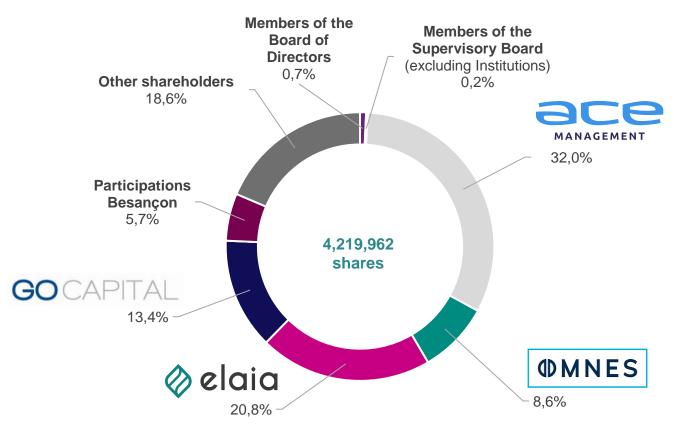
A team of experts strongly involved in the Company's success



## Appendix

## **Key financials**





#### €20 million in financing since inception

€13 million raised from current investors

**€7 million** in non-dilutive financing (grants and repayable advances from regions), Bpifrance, the European Commission and the French army)

## **Experienced Supervisory Board**



Didier Hoch Chairman

- Other assignments:
- Board Director of DBV Technologies, Genticel, Goliver Therapeutics,
   Pevion Biotech, OSE Immunotherapeutics, etc.
- 2015-2018: Co-founder and Chairman, Big Booster
- 2000-2010: Chairman, Sanofi Pasteur MSD
- **1998-2000:** VP, Rhône-Poulenc Rorer



Maryvonne Hiance Vice-Chairwoman

- Other assignments:
  - Vice-Chairwoman and Strategy Director of OSE Immunotherapeutics
  - 2016-2020: Chairwoman, France Biotech
  - 2008-2016: Chairwoman, Effimune



Franck Lescure
Permanent representative
of Elaia Partners



**Leila Nicolas**Permanent representative of Go Capital



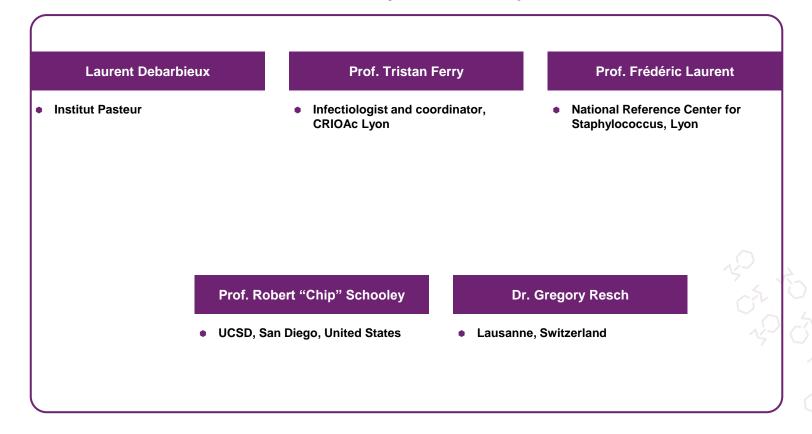
**Guy Rigaud** Member



ACE Management
Non-voting Board member

## The support of prestigious experts

## Pherecydes Pharma also benefits from the support of internationally renowned experts:



#### The terms and conditions of Pherecydes Pharma's IPO

**SHARE CODES** 

Label: PHERECYDES PHARMA

Mnemonic: ALPHE

ISIN Code: FR0011651694

INDICATIVE SCHEDULE OF THE OPERATION

January 4, 2021

Release of the Document d'Information (Part 1)

**January 14, 2021** 

Release of the *Document d'Information* (Part 2)

Opening of the Closed Price Offer (CPO) and the Global Offering (GO)

**January 28, 2021** 

Closing of the CPO at 5 p.m. (Paris time) at the counters and at 8 p.m. (Paris time) on the Internet Closing of the GO at 5 p.m. (Paris time)

January 29, 2021

Result of the Offer

**February 2, 2021** 

Payment and delivery of the CPO and the GO

**February 3, 2021** 

First trading of Pherecydes Pharma shares on Euronext Growth

Copies of the *Document d'Information* are available free of charge from Pherecydes Pharma (20 boulevard Benoni Goullin, 44200 Nantes) and on the websites of Pherecydes Pharma (www. pherecydes-pharma.com) and Euronext (www.euronext.com). Investors are invited to draw their attention to the risks relating to the business described in chapter 4 "Risk Factors" of the *Document d'Information* and in chapter 20 "Risk Factors Related to the Offer" of the second part of the *Document d'Information*. These risks should be taken into consideration by investors before making any investment decision. This document does not constitute an offer of securities of Pherecydes Pharma in any jurisdiction in which such an offer would violate applicable laws and regulations, or an offer to sell Pherecydes Pharma shares in the United States. The shares of Pherecydes Pharma may not be offered or sold in the United States absent registration or an exemption from registration under the U.S. Securities Act of 1933, as amended. Pherecydes Pharma does not intend to register the offer mentioned in this document in the United States. This document may not be published, transmitted or distributed, directly or indirectly, in the United States, Canada, Australia or Japan.

\* Income tax reduction, see section 22.11 of the Document d'Information

## At the forefront of the fight against bacterial infections

#### Structure of the offer

- Offer to the public in France
- Global Institutional Offering in France and outside France (excluding the United States)

Indicative Price Range applicable to the Closed Price Offer and the Global Offering

€XX per share

#### Nature of the offered shares

New and existing actions

#### Initial size of the offer

Issuance of 1 097 092 new shares representing approximately
 €20.0 based on the mid-point of the range

#### **Extension Clause**

A maximum of 82 282 new shares, i.e. 7.5% of the initial size of the offer

#### **Over-allotment Option**

N/A

#### Total gross amount of the operation

 €21.5 million in case of full exercise of the Extension Clause on the basis of the mid-point of the price range

#### Commitments to subscribe

- Elaia has committed to subscribe for €990,250 in cash,
- Go Capital has committed to subscribe for €713,000 in cash,
- Fa Diese has committed to subscribe for €200,000 in cash,

#### Forbearance and retention commitments

- Company: 180 days
- Shareholders: 18 months