



MaaT Pharma Microbiota as a Therapy

Company Presentation
29 March 2022

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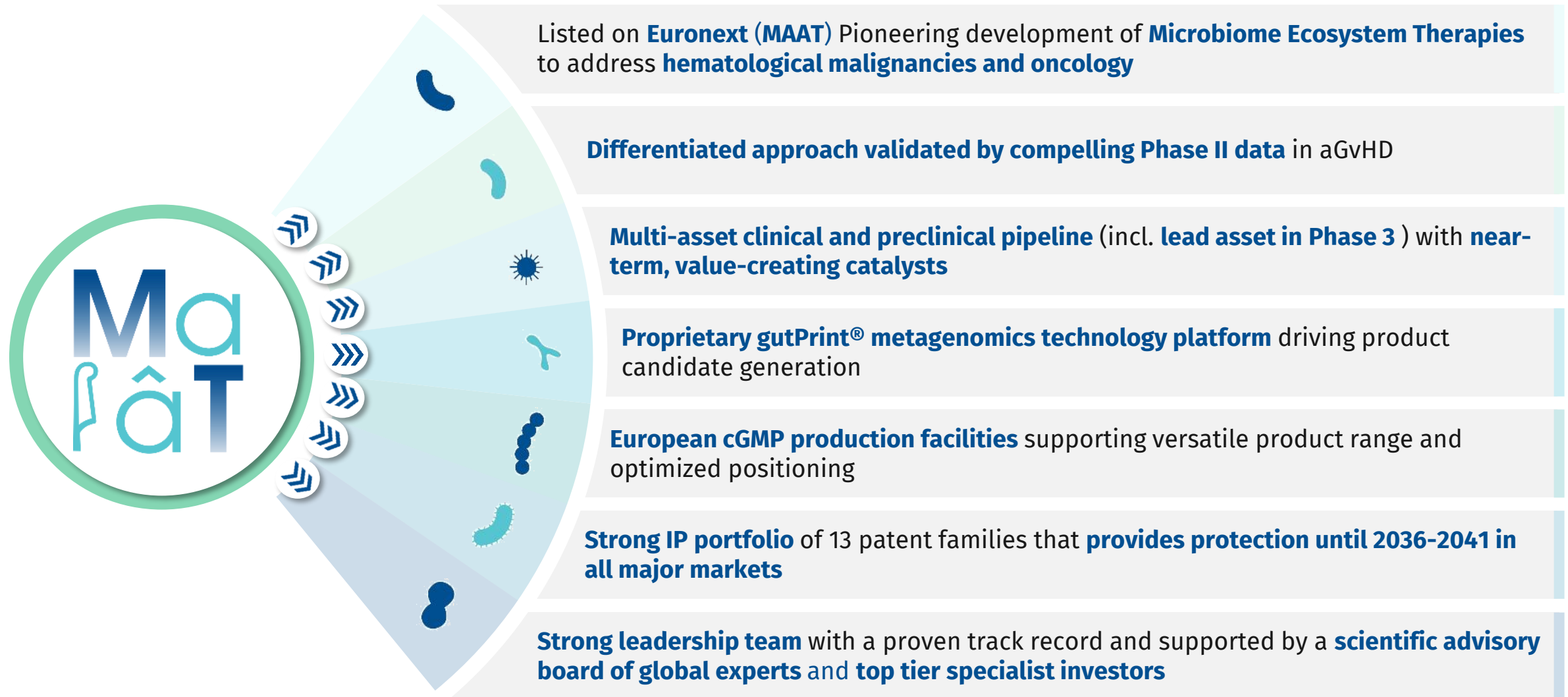
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A Uniquely-Positioned Microbiome Company



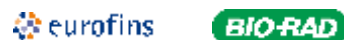
Management Team



Siân Crouzet
Chief Operating Officer



Hervé Affagard
Founder & CEO



Dr. Carole Schwintner
Chief Technology Officer



Dr. Savita Bernal
Chief Business Officer



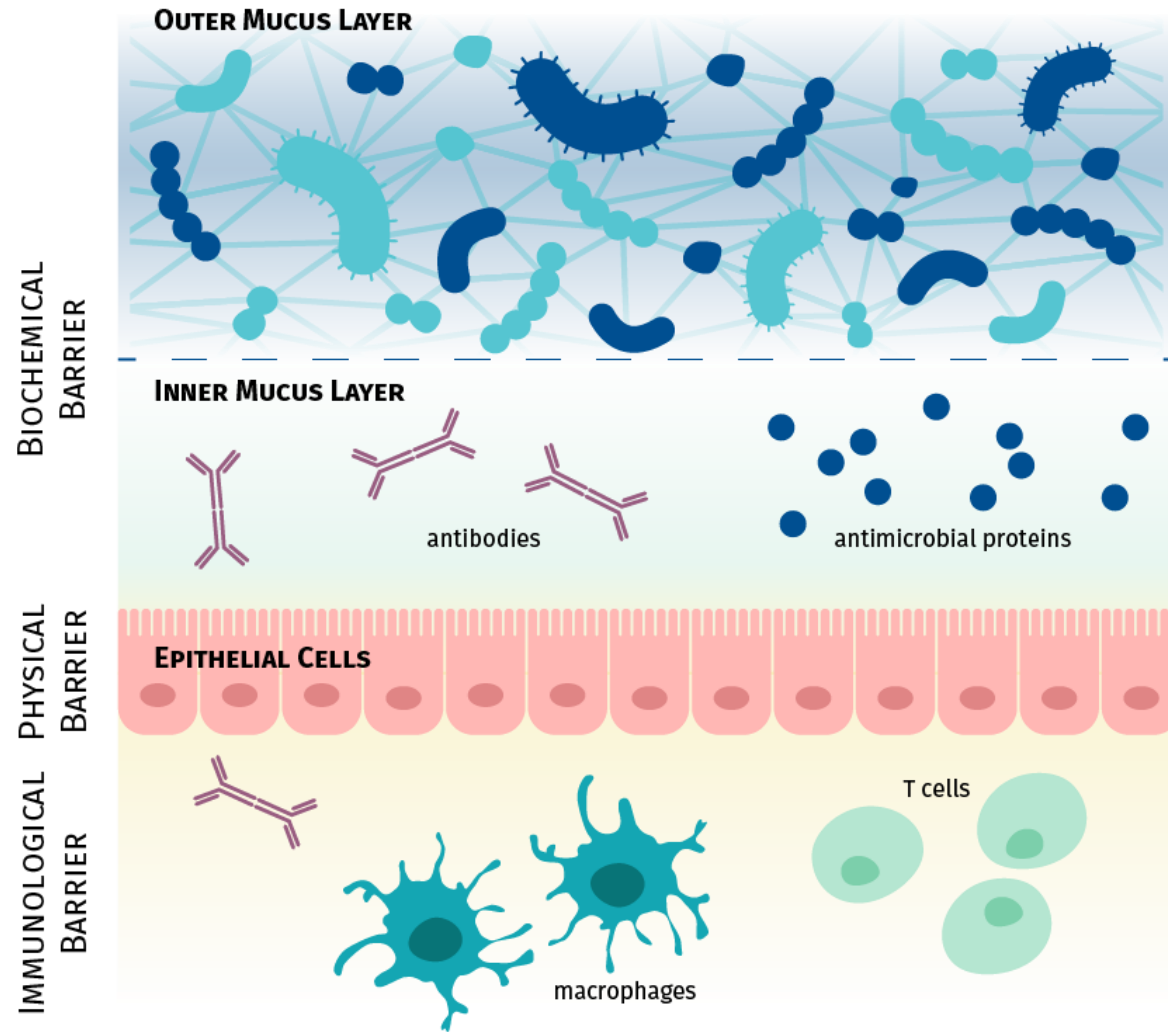
Dr. John Weinberg
Chief Medical Officer



Dr. Isabelle Adeline
Chief of Staff



Host – Microbiota Interactions are Critical for a Functional Immune System



Cross-section of a healthy gut

A rich and diversified gut ecosystem actively modulates the immune system functionality

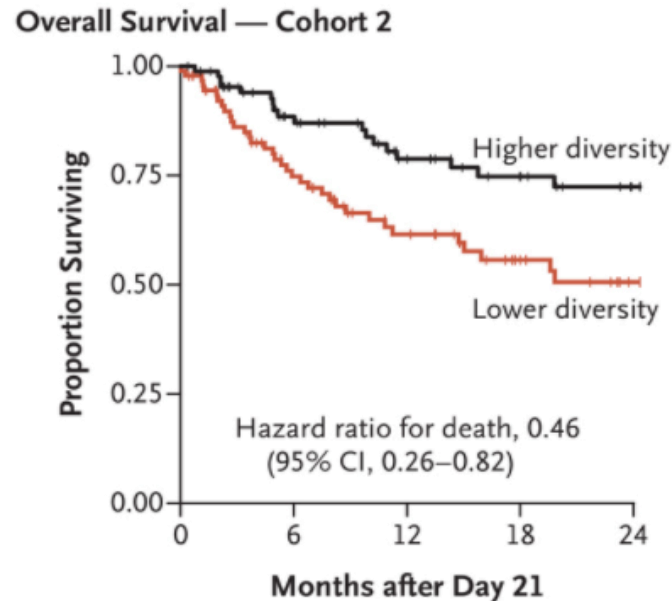
- A diversified microbiome contributes to the education and modulation of our immune system throughout life
- Bacterial richness and mucus layer prevent colonization by pathogens and improve gut barrier
- 80% of cellular host defense are localized in the gut (including innate and adaptive systems)

Diversity matters!

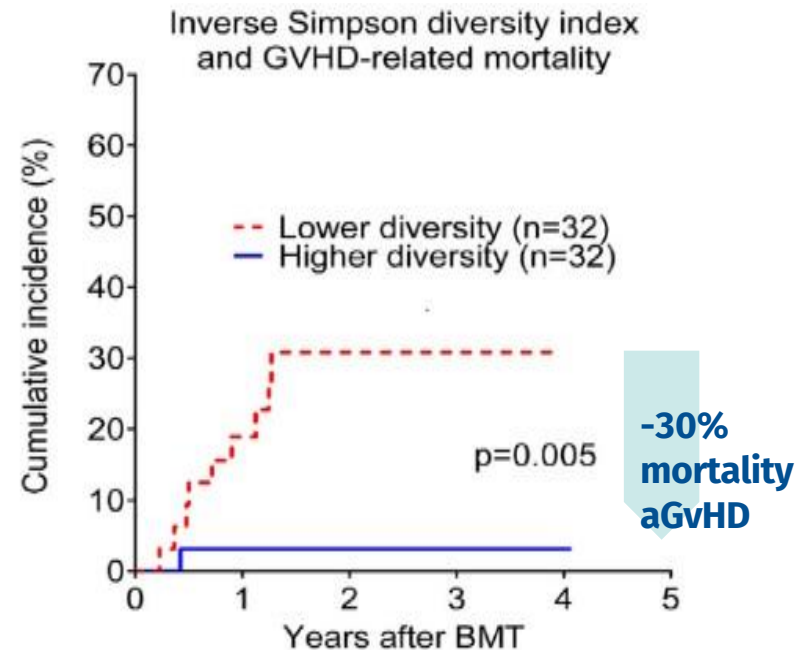
Higher gut microbiome diversity is associated with ...

Liquid Tumors

Higher survival rate in patients receiving allo-HSCT ^{*,1}



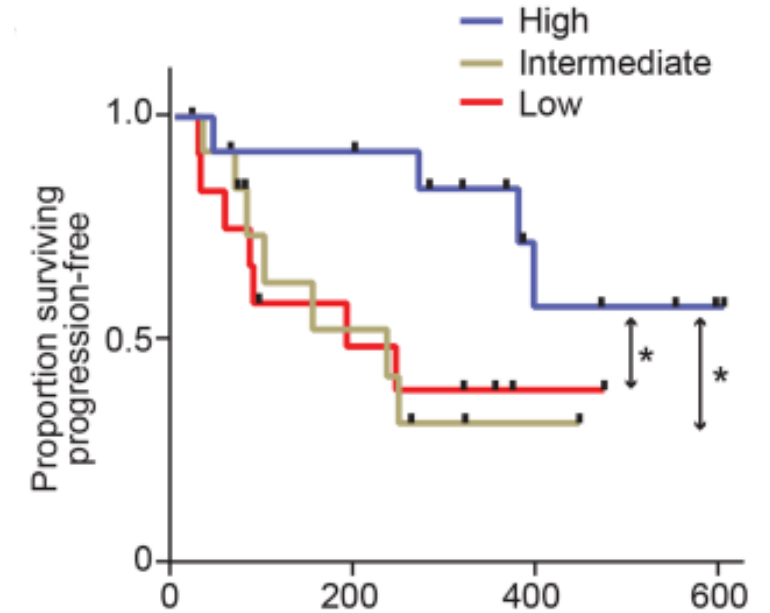
Lower incidence and lower mortality from aGvHD^{*,2}



MaaT Pharma MET Inverse Simpson (mean): 24

Solid Tumors

Higher response rate to ICI* in patients with metastatic melanoma³

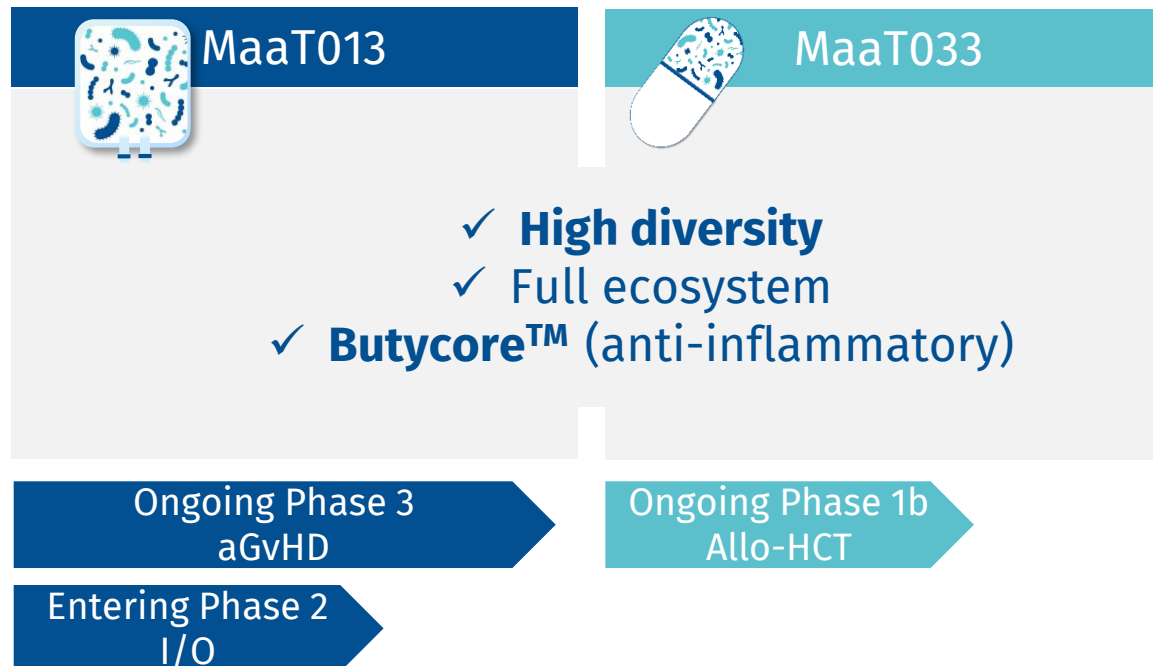


*allo-HSCT: allogeneic hematopoietic stem cell transplantation; aGvHD: acute Graft-vs-host-Disease; ICI: Immune Checkpoint Inhibitors
¹Peled, J.U. & al N Engl J Med 2020;382:822-34; ²Ghani, 2021; ³Jenq RR. et al, Biol Blood Marrow Transplant 21 (2015) 1373e1383; Pamer, Blood, 2014 ; Gopalakrishnan et al., Science, 2017, Routy et al, Science, 2018 ; Vetizou et al Science 2015;

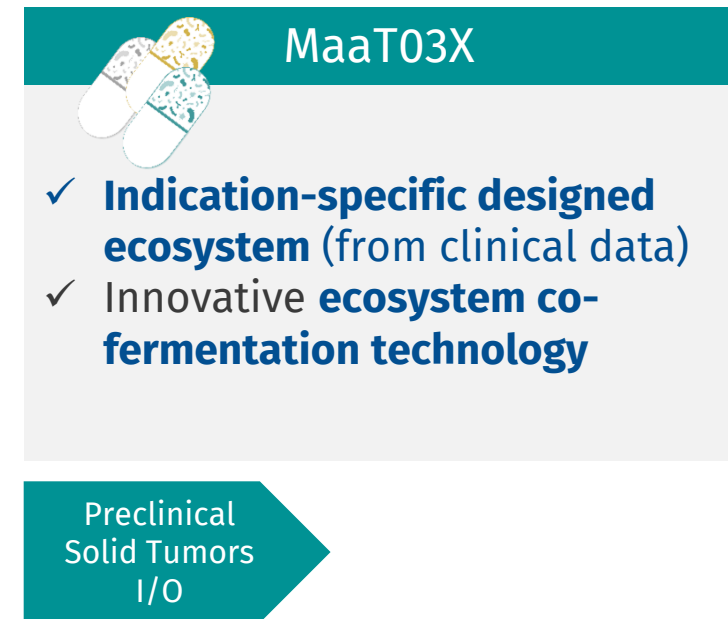
MaaT Pharma's Microbiome Ecosystem Therapy (MET) platform has generated a diverse line of product candidates



Native

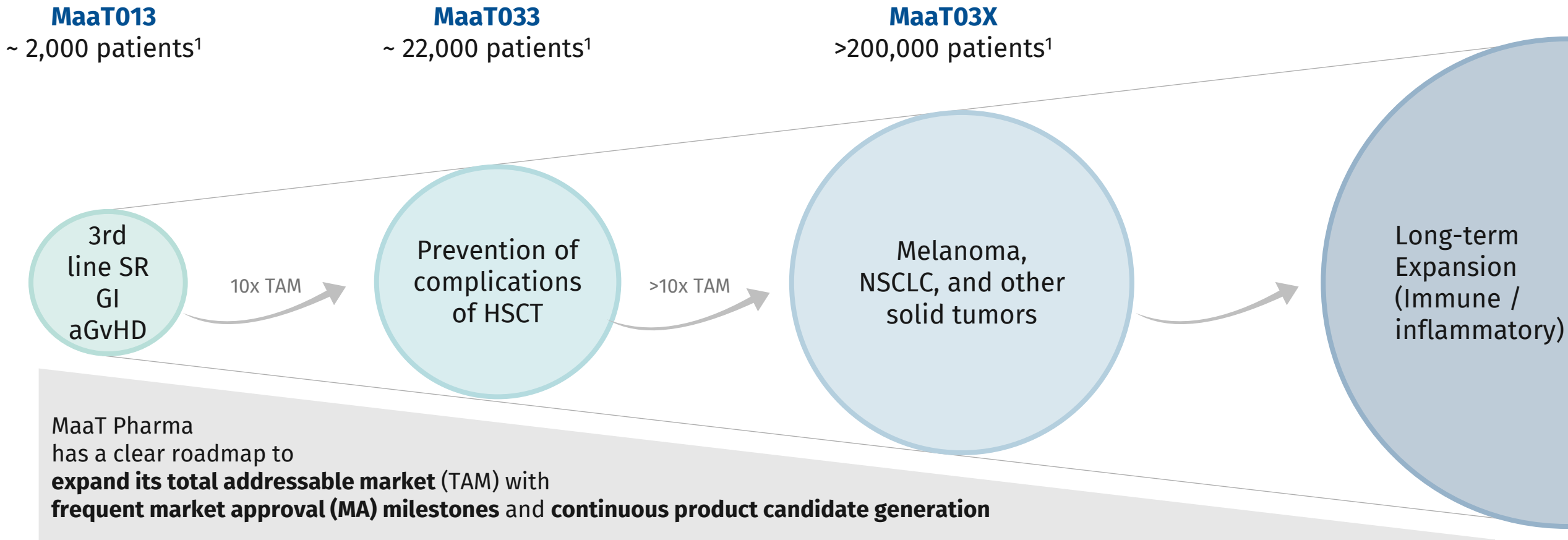


Co-fermented

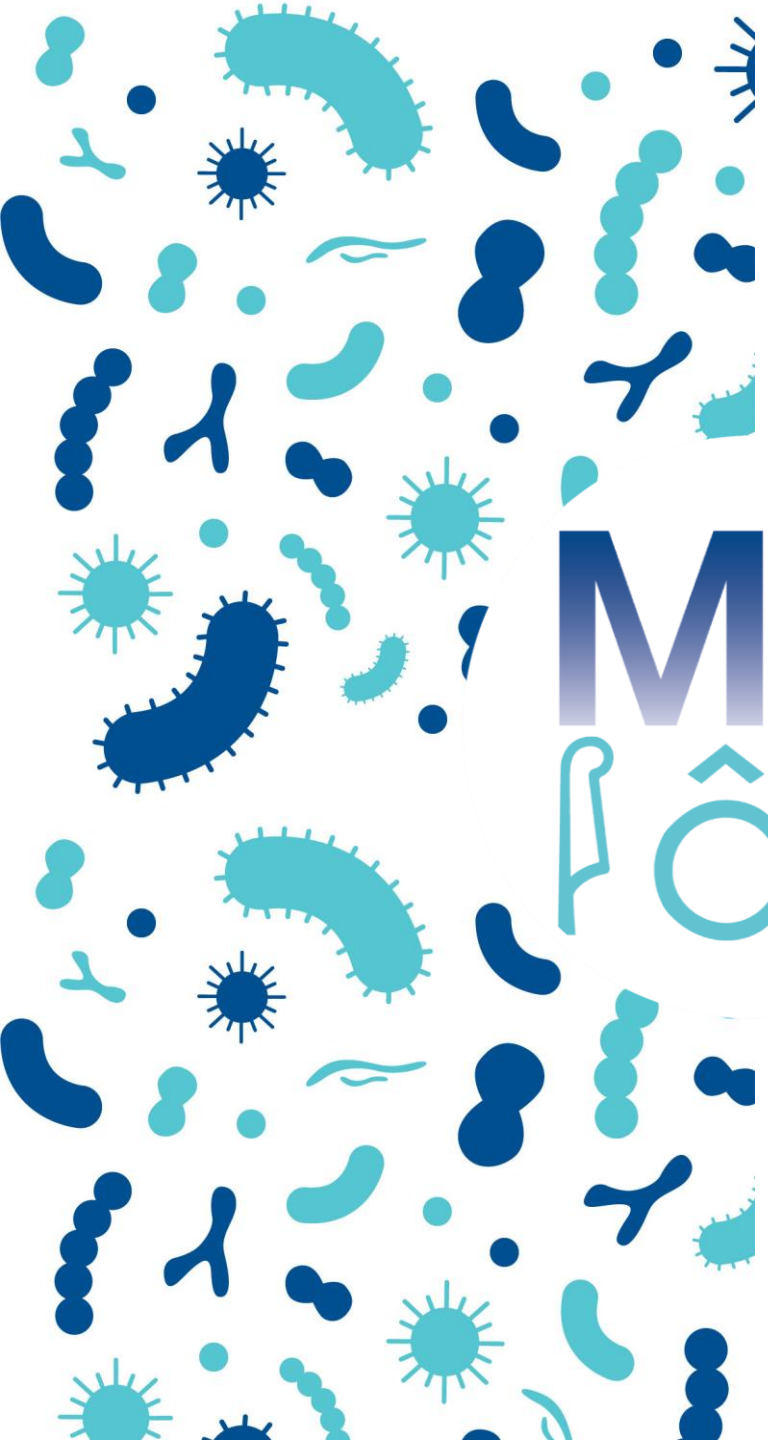


¹ **Butycore**: Group of 15 different genera known to produce short-chain fatty acids with anti-inflammatory properties

MaaT Pharma's approach and platform enable a rapid build-up of the addressable population that can benefit from its therapies



¹ EU5, US, and Japan

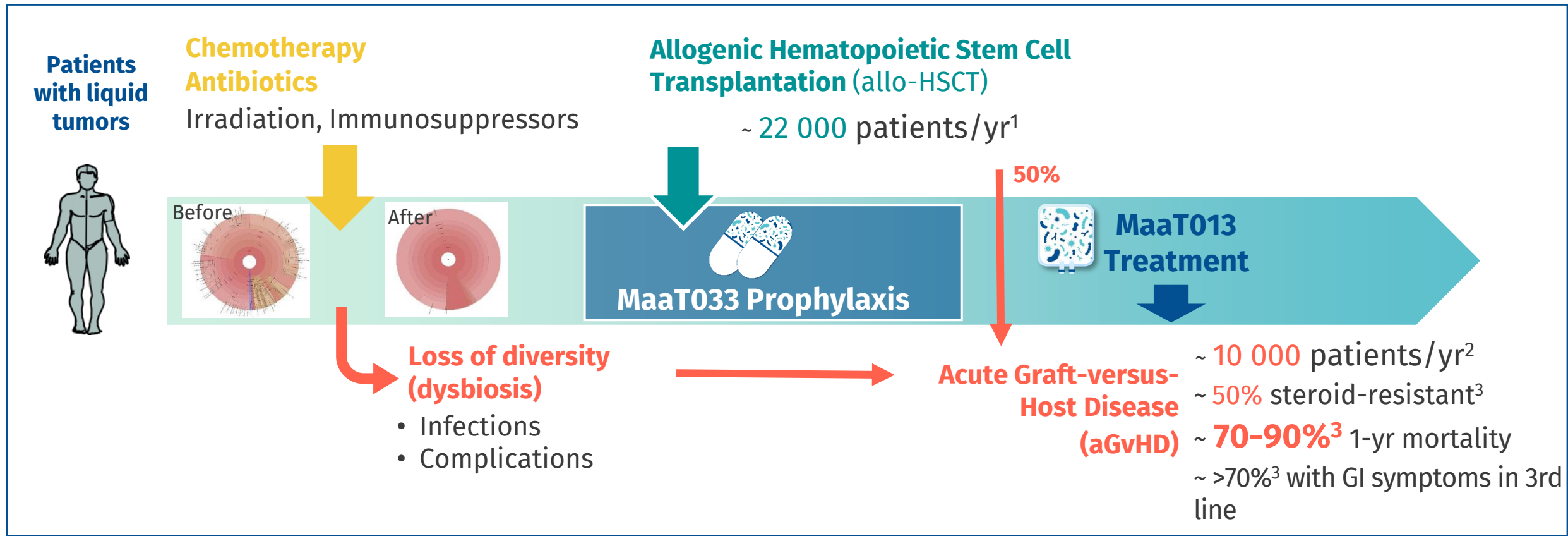


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Hemato-Oncology

MaaT013 and MaaT033 aim to restore the gut microbiota to improve survival in patients with liquid tumors

Intestinal dysbiosis is associated with higher mortality in hemato-oncology



1. EU5 + US : (~ 20 500 primary procedures with an additional 7%-10% recurring), 2. EU5 + US, ³ According to MAGIC database



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Hemato-Oncology

Treatment of acute Graft-vs-host-Disease
(aGvHD)



Two complementary approaches generating data on MaaT013

Phase 2 clinical trial - HERACLES

- Phase 2 clinical trial HERACLES ([NCT03359980](#))
 - N=24 patients
 - 4 countries
- Gastro-intestinal aGvHD grade III-IV (most severe)
- Steroid-refractory
- 3 doses of MaaT013 as a monotherapy over 2 weeks
- As 2nd line of treatment
- Follow-up at 28 days (GI-response) and after 12 months (overall survival)

Early Access Program/Compassionate Use (formerly « ATU »)

- Authorized by the French regulator (ANSM)
 - Results published on N=52¹ patients
 - France
- Gastro-intestinal aGvHD grade II-IV
- Steroid-refractory or steroid-dependent
- 3 doses of MaaT013 as monotherapy or in combination over 2 weeks
- After 1 to 6 lines of treatment
- Follow-up at 28 days (GI-response) and after 12 months (overall survival)

¹ Program is ongoing. >80 patients treated as of March 2022

MaaT013 has received Orphan Drug Designation from the [FDA](#) and [EMA](#) for aGvHD

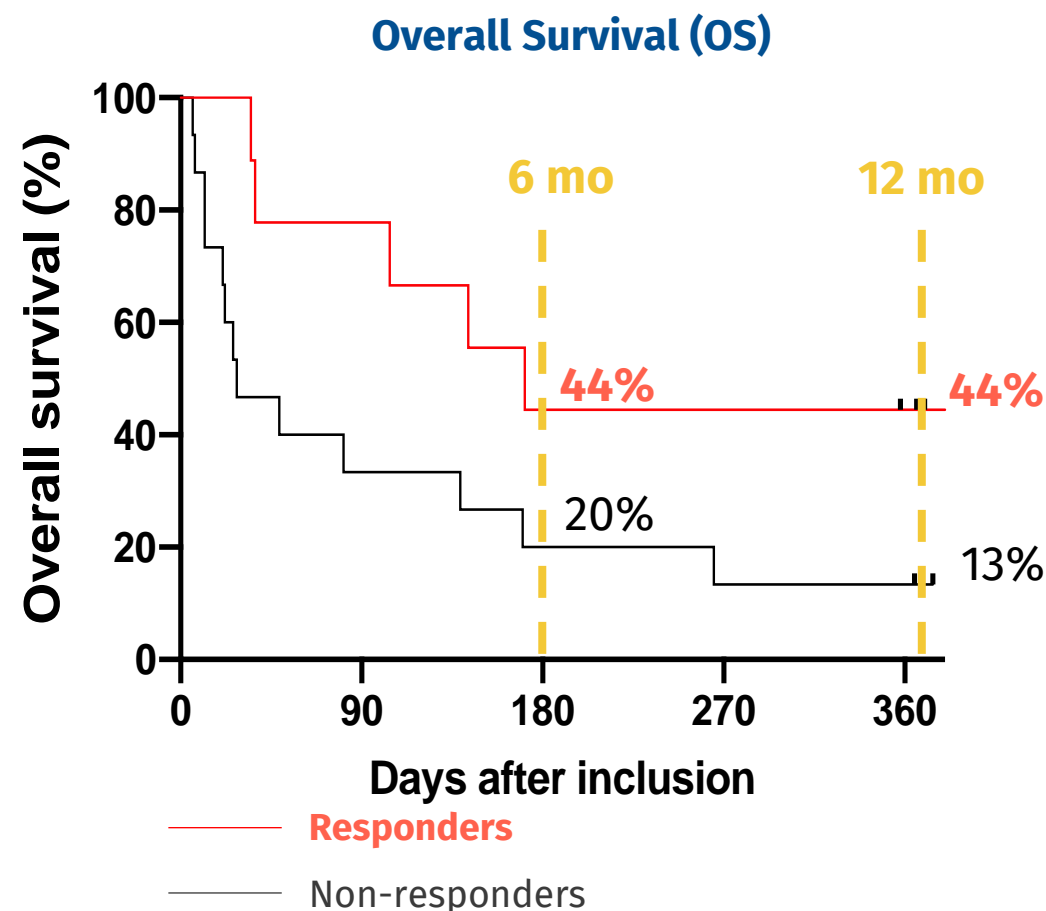
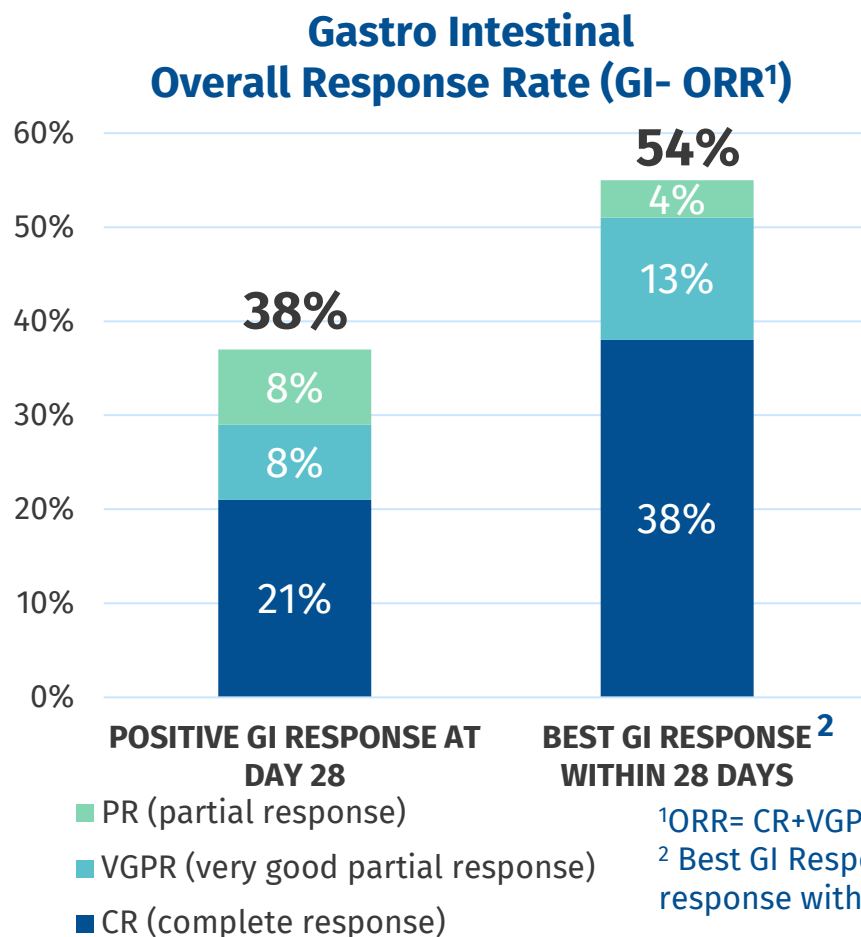




HERACLES Phase 2 Clinical Trial

Promising results in a very severe (III-IV) GI aGvHD population

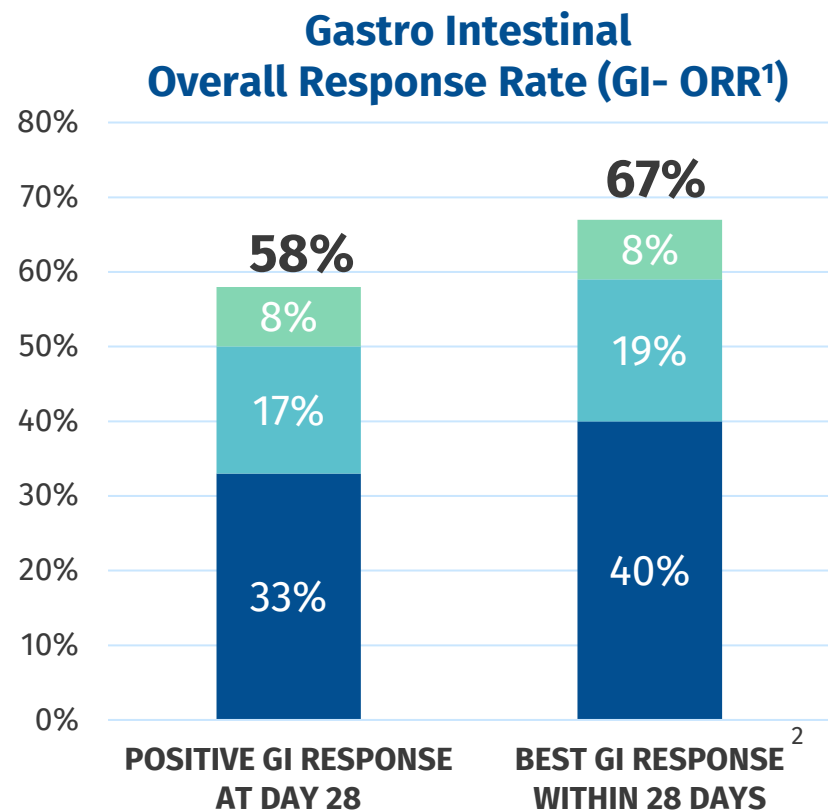
- N=24 patients, 96% grade III (4% grade IV), 3 doses, 2nd line (Steroid-resistant)
- Microbiota analysis shows better engraftment of MaaT013 and higher gut microbiome diversity after treatment in Responders
- Very good safety and tolerability profile





Early Access Program (EAP): Promising confirmation in an advanced, severe and more diverse GI aGvHD population

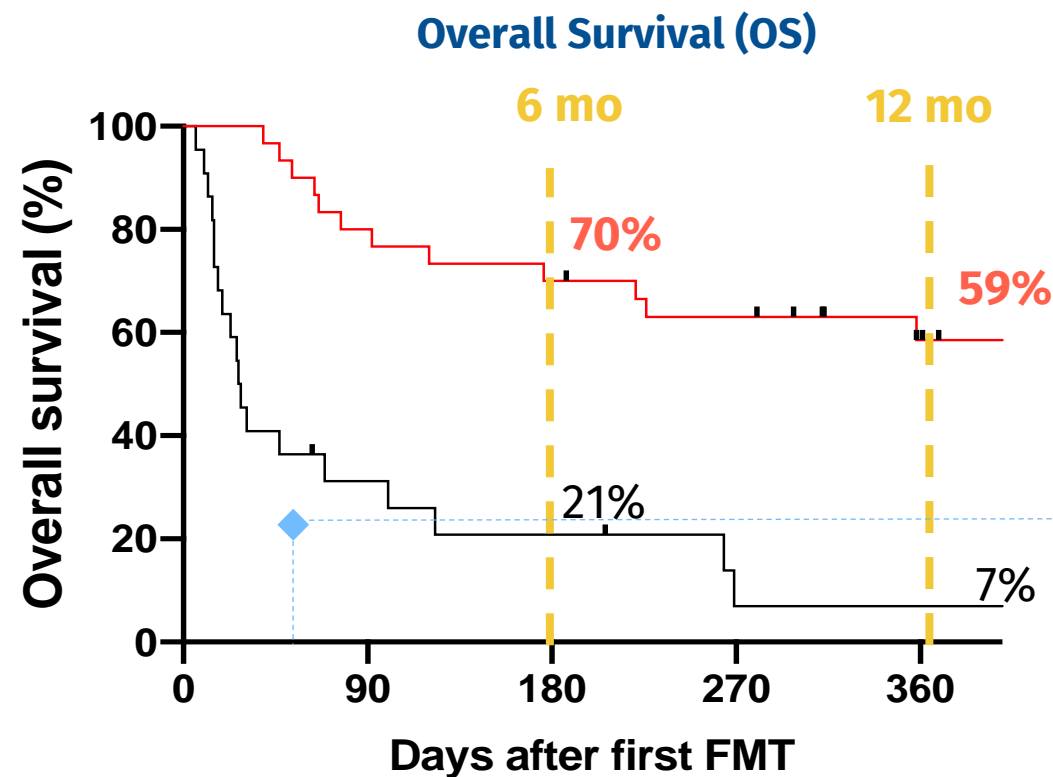
- N=52 patients : 83% steroid-resistant ; 94% grade III, Up to 6 lines of prior treatment (median: 3 ; 77% have received ruxolitinib); 3 doses
- Good tolerability and safety profile in a fragile population



■ PR (partial response)
■ VGPR (very good partial response)
■ CR (complete response)

¹ORR= CR+VGPR+PR

² Best GI Response: Any response within 28 days



22% expected OS at 2 months in ruxolitinib-resistant patients (REACH1 study)

— Responders
— Non-Responders



The ARES Phase III study is designed to establish MaaT013 as the 3rd line agent in GI aGvHD treatment

- Pivotal single arm trial of MaaT013 as 3rd line (steroid-resistant & ruxolitinib-resistant) in n=75 GI-aGvHD patients
- Primary endpoint: GI-ORR at Day28

EUROPE :

- ✓ First patient dosed in Q1 2022
- ✓ CTA approved in 3 European countries. Expected to expand to additional EU countries

USA:

- FDA requested further information – on clinical hold.
- Submitted a request for a “Type A” meeting to the FDA by the end of 2021, with the support of well-respected regulatory consultants, aiming to resolve the clinical hold and expand ARES to US sites. Exchanges ongoing.

Targeted Timelines ARES Phase III Trial



¹ subject to the lifting of the FDA clinical hold ; ORR: overall response rate ; OS: overall survival ; MAA: Market approval application; BLA: Biological License Application



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Hemato-Oncology

Allogeneic-HSCT Complication Prevention

Prevention of complications of allo-HSCT offers an attractive market opportunity for MaaT Pharma to address with MaaT033

- MaaT033, an oral formulation of MaaT013’s drug substance, aims to prevent complications from allo-HSCT in all patients receiving the intervention



United States

c. 7,800
primary
procedures



EU 5

c. 9,600
primary
procedures

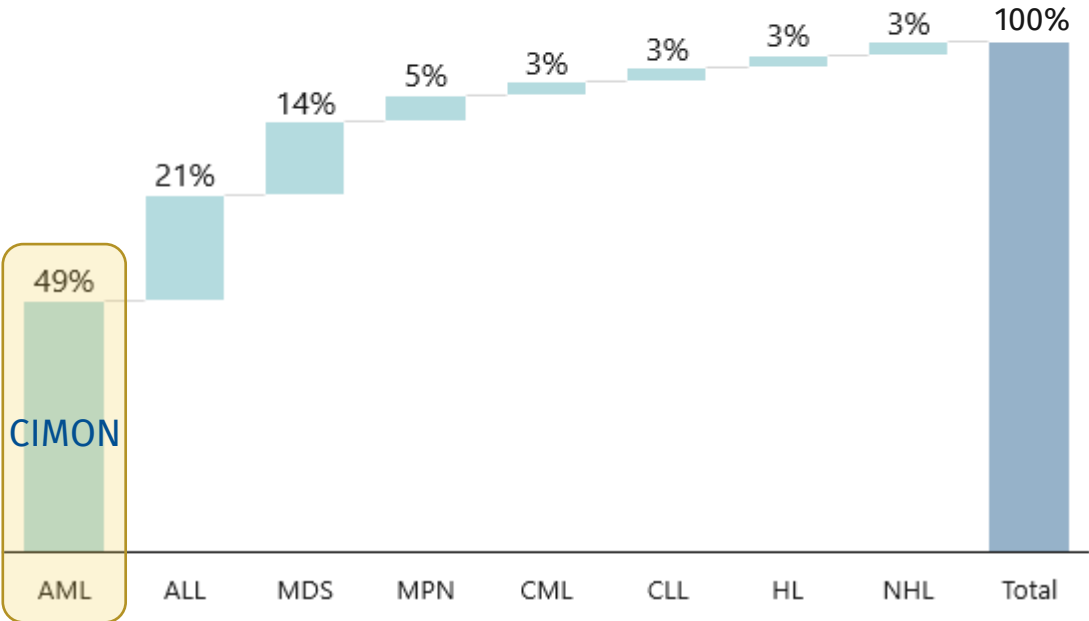


Japan

c. 3,000
primary
procedures

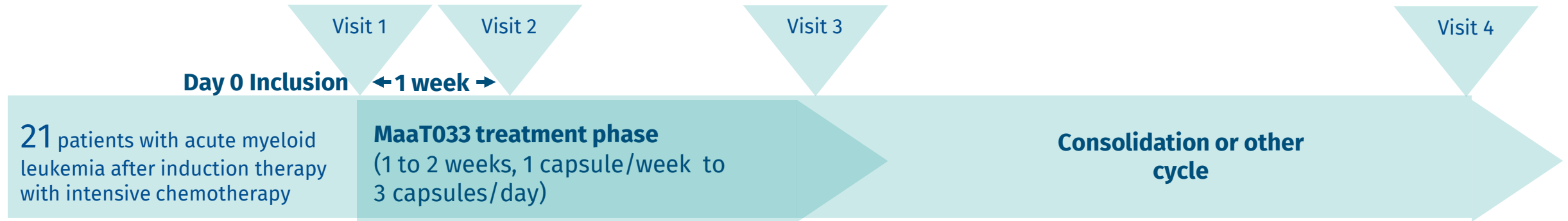
Additional 7%-10% recurrent procedures

Hematological Malignancy Patients Receiving Allo-HSCT¹



LAM (AML) : acute myeloid leukemia; LAL (ALL) : acute lymphoblastic leukemia ; SMD (MFS) : myelodysplastic syndrome; NMP (MPN) : myéloproliférative neoplasms ; LMC (CML) : chronic myeloid leukemia ; LLC (CLL) : chronic lymphocytic leukemia ; LH (HL) : Hodgkin’s Lymphoma ; LNH (NHL) : Non Hodgkin Lymphoma

Phase Ib CIMON study aims to determine MaaT033 dose for a Phase III study in post-allogeneic HSCT complication prevention



- CIMON explored safety and recommended dose of **orally administered MaaT033** in AML patients post induction chemotherapy. **Primary Endpoint:** Dose limiting toxicity-related treatment emergent (serious) adverse events
- MaaT033 achieved
 - ✓ 100% drug compliance
 - ✓ **Good overall safety profile** : 4 positive DSMB (4 cohorts evaluated) , majority of mild adverse events
 - ✓ **Good microbiome engraftment and engraftment persistence**

→ **Final results expected H1 2022**

→ **Next Phase II-III pivotal study (Allo-HSCT, RCT, ~340 patients, OS) planned to start H2 2022**



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Immuno-Oncology
Solid Tumors

A diverse gut microbiome increases survival in patients receiving immune checkpoint inhibitors (ICI)

FMT from ICI responders to ICI non-responding patients with metastatic melanoma

✓ **6/15**

Non-responders
→ Responders
(Davar et al, 2021)

✓ **3/10**

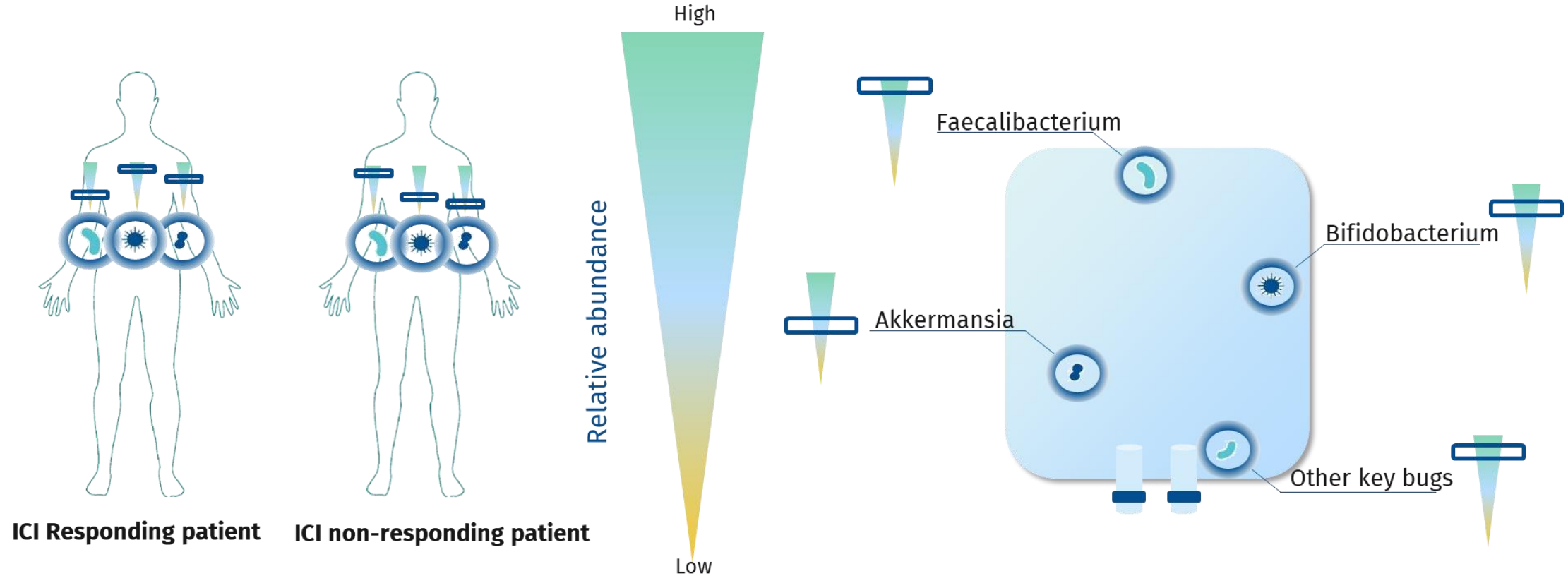
Non-responders
→ Responders
(Baruch et al, 2021)



- Immune check-point inhibitors (ICI) therapies have established themselves as key therapeutic options in solid tumors, but ORR may be as low as 20% in some indications.
- Richness and Diversity of gut microbiome drive survival in patients receiving ICI^{1,2,3}
- FMT from ICI responders (R) could induce response in metastatic melanoma non-responders (NR) (Baruch et al, *Science* 2021, Davar et al, *Science* 2021)

→ Leveraging the gut microbiome richness, diversity and its key functional networks may be a game-changer in immuno-oncology in the coming years

MaaT013 ensures high diversity and contains specific bacterial strains that have been identified to improve ICI response

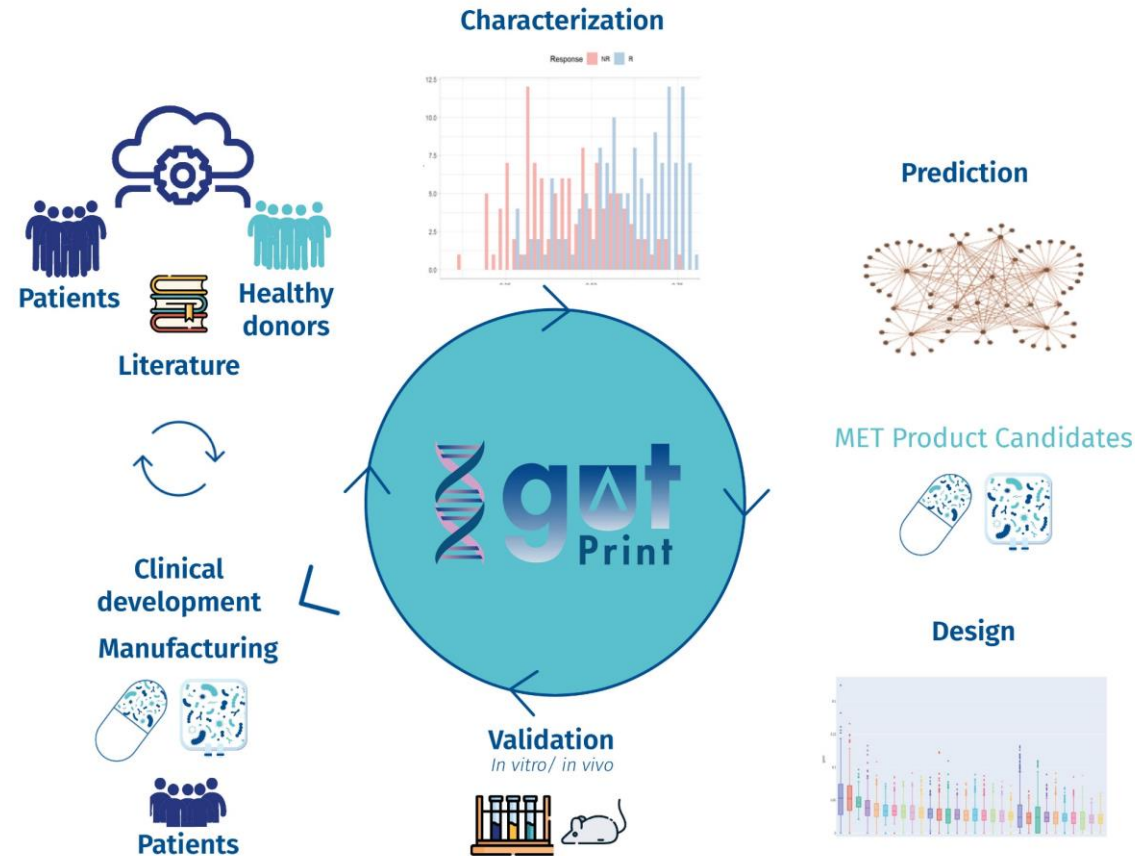


Phase IIa PICASSO trial², in collaboration with **APHP** (sponsor), ready to start (approved by ANSM)

- ✓ **RCT** [MaaT013 + ICI] vs. [Placebo + ICI] in **60** metastatic melanoma patients
- ✓ Assessing **Safety** and **Efficacy** (iRECIST) of MaaT013 vs. placebo after 23 weeks of treatment

¹Gopalakrishnan and al., Science 2018, Routy and al, Science 2018; Matson and al. Science 2018, ²Registered trial #NCT04988841

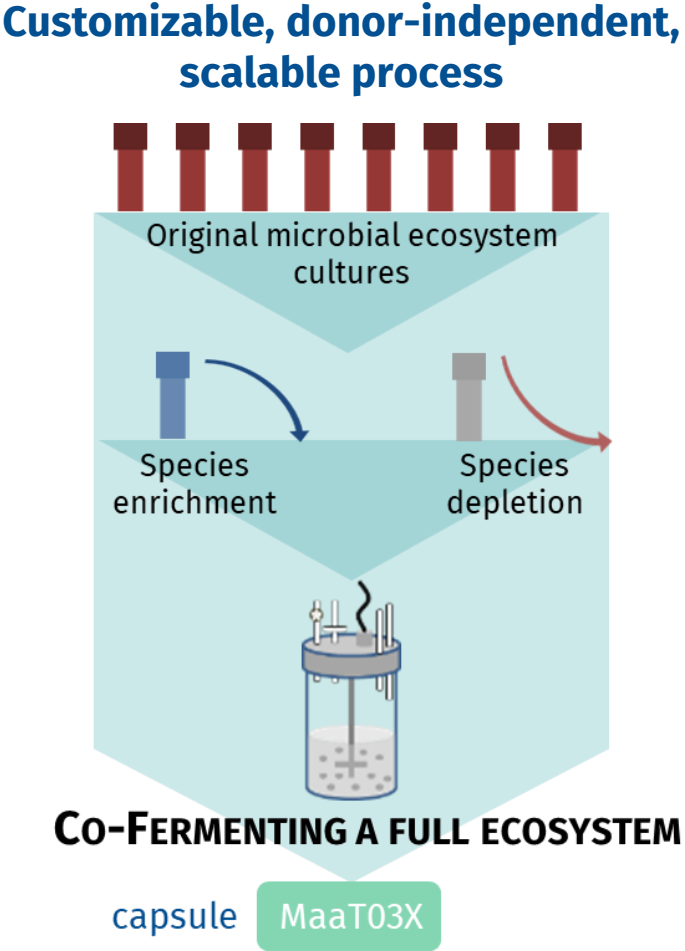
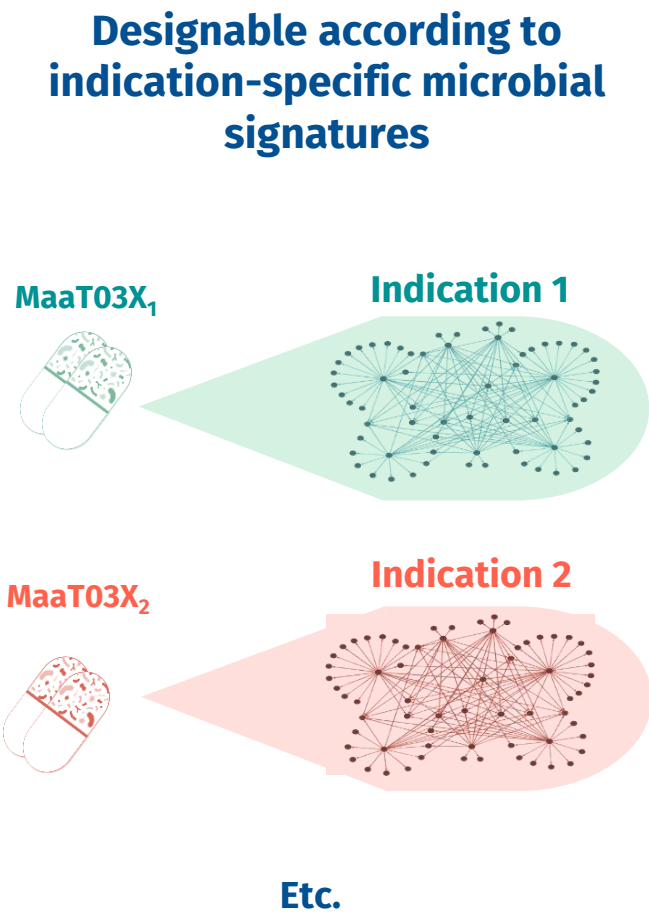
Proprietary gutPrint® platform synergizes multi-source data to generate innovative and indication-specific microbiome ecosystem therapies



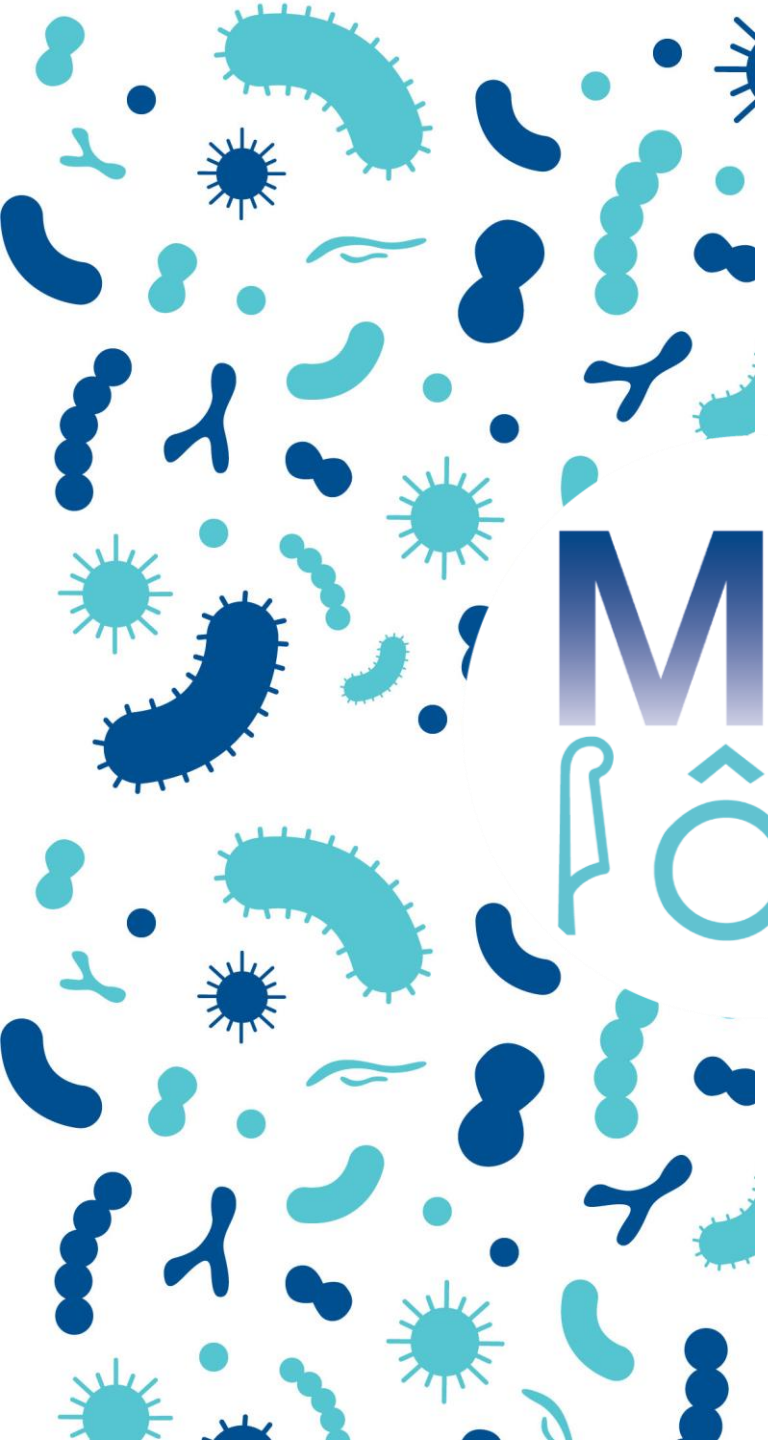
gutPrint® is the engine that drives MaaT Pharma's MET product candidate generation capabilities to broaden and strengthen the pipeline

The customizability and scalability of the MaaT03X line allows it to potentially address several solid tumor indications

	Fermented (MaaT03X)
Ecosystem design	Full
Richness & diversity	High
Scalability	Improved
Administration route	Enema and oral
Customizability	Yes
Tumor Types	Multiple (undisclosed)



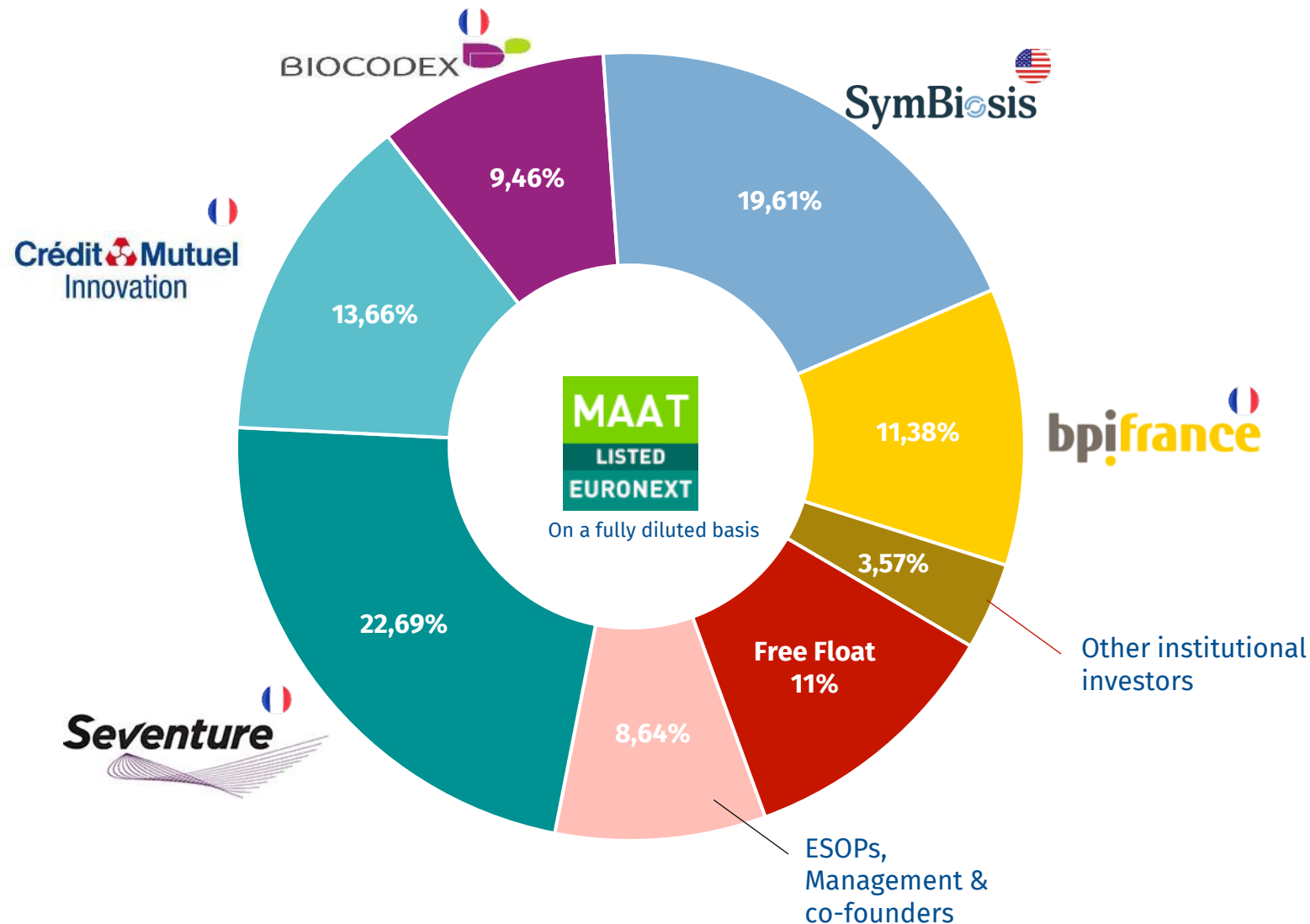
First candidate in preclinical testing – Expected to enter clinical testing in H1 2023



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Shareholding structure

MaaT Pharma is listed on Euronext Paris – 35.7M€ IPO Nov. 2021



BOARD OF DIRECTORS



Jean-Marie Lefèvre

Chairman & Non-Executive Director

President - Biocodex



Isabelle de Crémoux

Non-Executive Director

CEO & Managing Partner - Seventure



Claude Bertrand

Non-Executive Director

General Director R&D - Servier



Martine George

Non-Executive Director

Oncologist



Jean Volatier

Non-Executive Director

CFO - Inventiva



Dorothée Burkel

Non-Executive Director

Chief Corporate and People Operations Officer - PartnerRe



Muriel Prudent

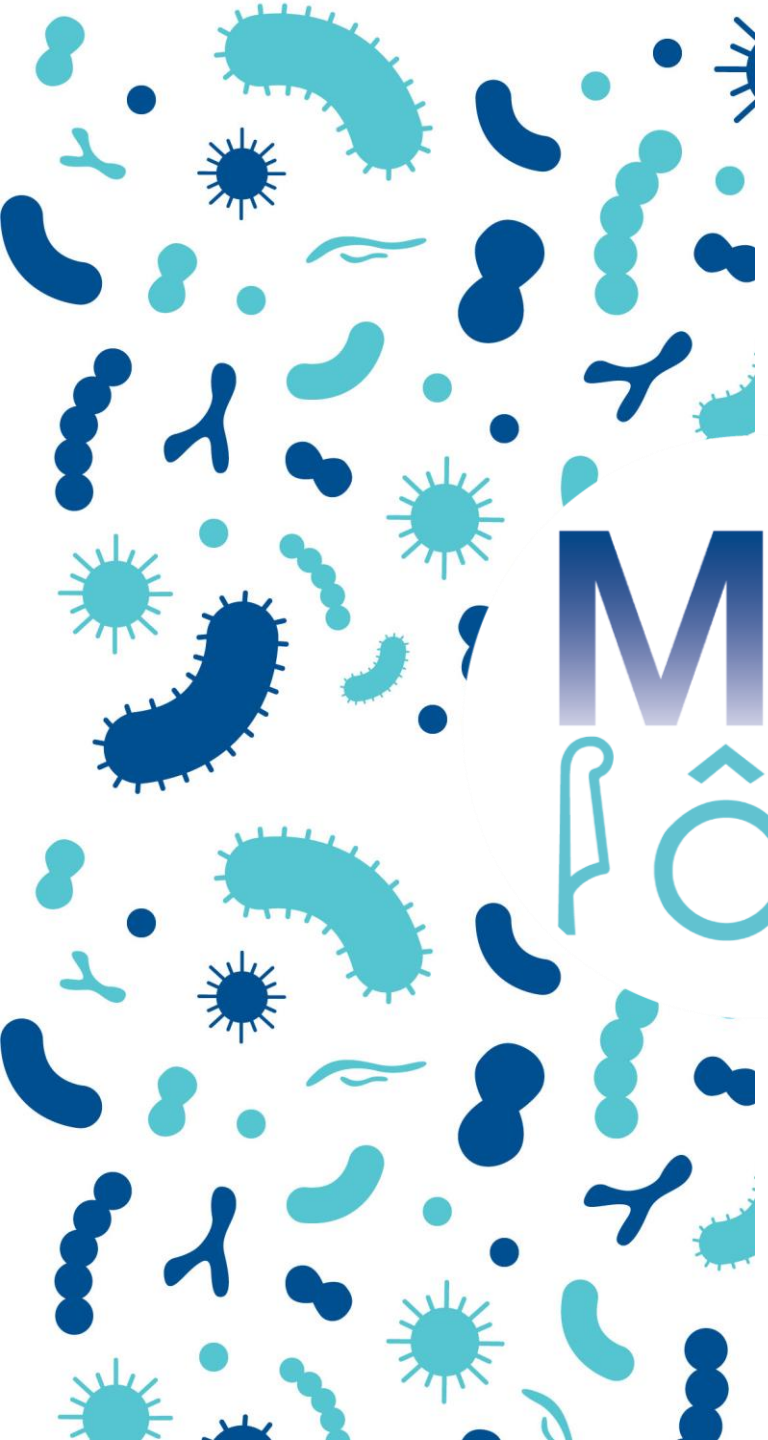
Censor

VC Investment Manager – Fonds PSIM - Bpifrance



Hervé Affagard

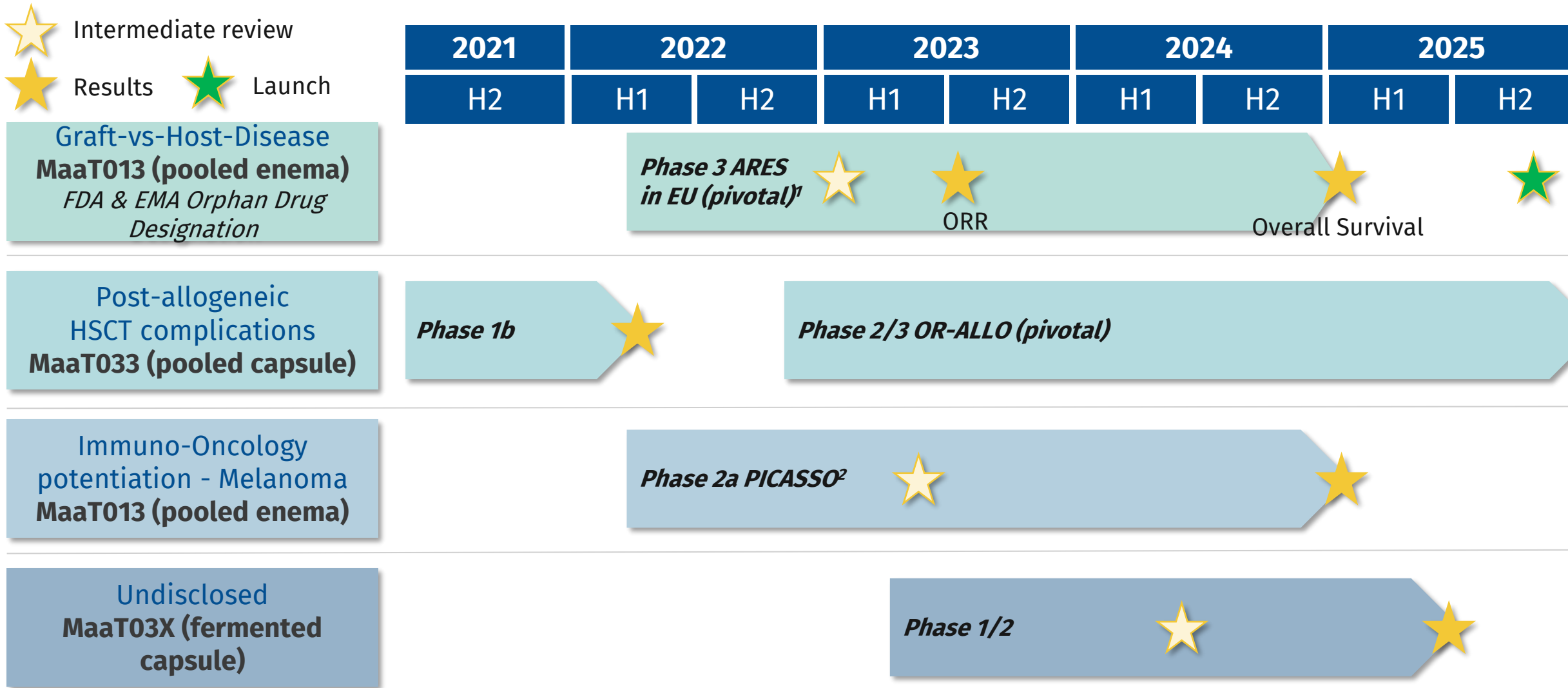
Executive Director



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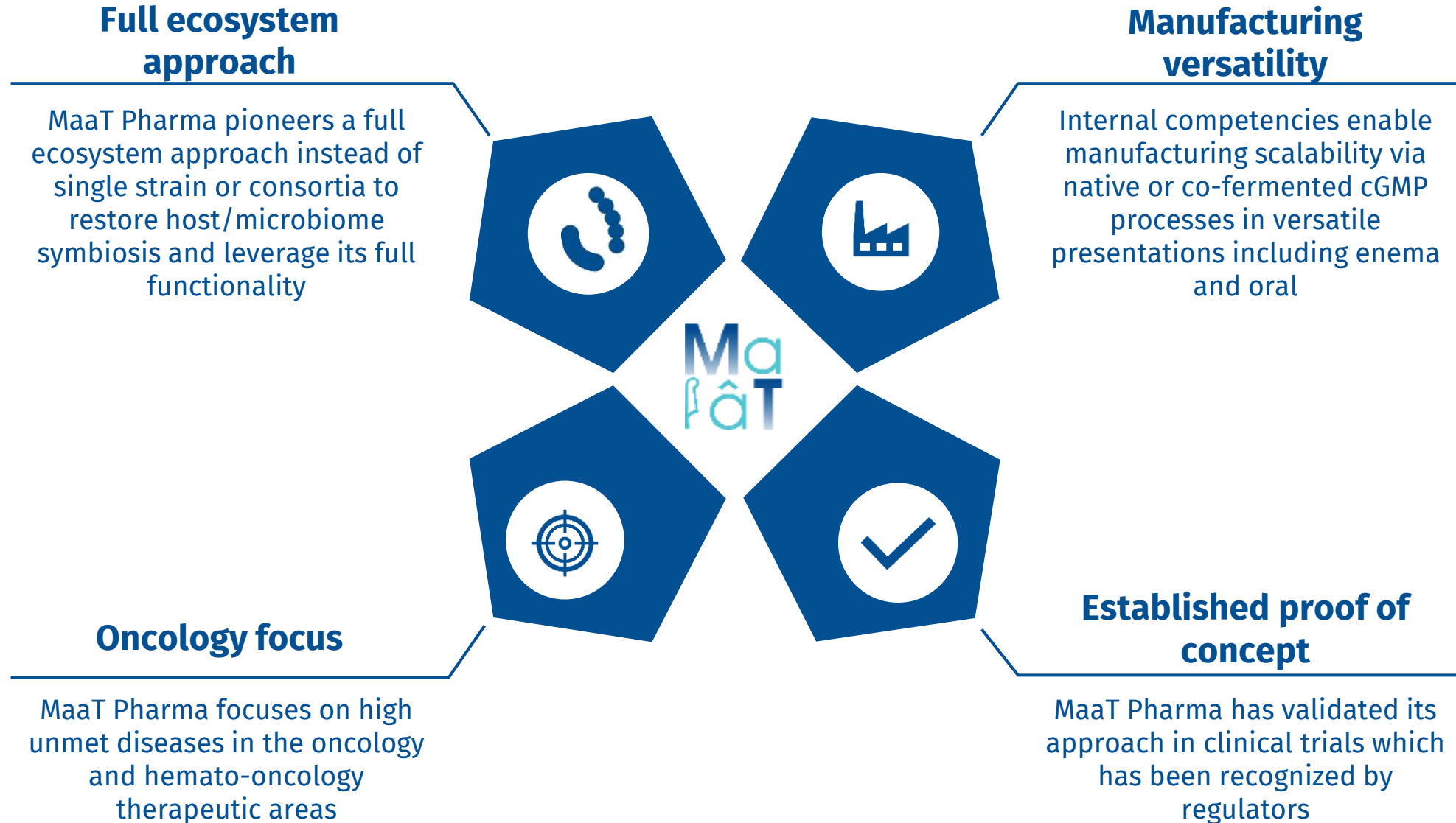
Key Upcoming Milestones

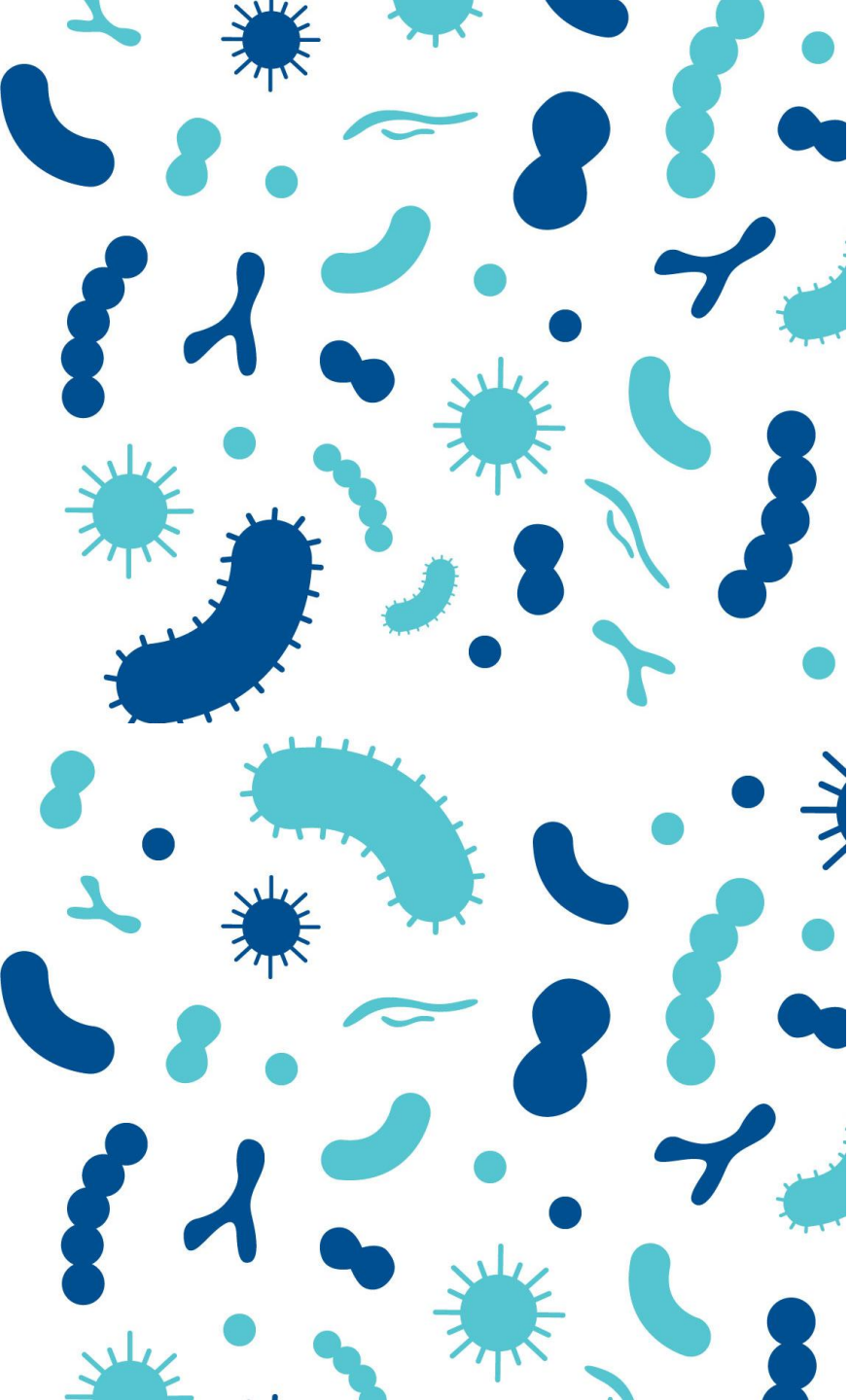
MaaT Pharma's development plan produces a steady flow of meaningful and value-creating news in both the near and long term



¹expansion to US sites in H2 2022 subject to IND approval in the US;
²Investigator sponsored trial (AP-HP) where MaaT Pharma supplies the drugs and performs the microbiome profiling using its gutPrint® platform
March 2022

Key differentiators of MaaT Pharma from other microbiome competitors





THANK
YOU