



MaaT
RÔLE

MaaT Pharma Microbiota as a Therapy

Company Presentation
December 2021

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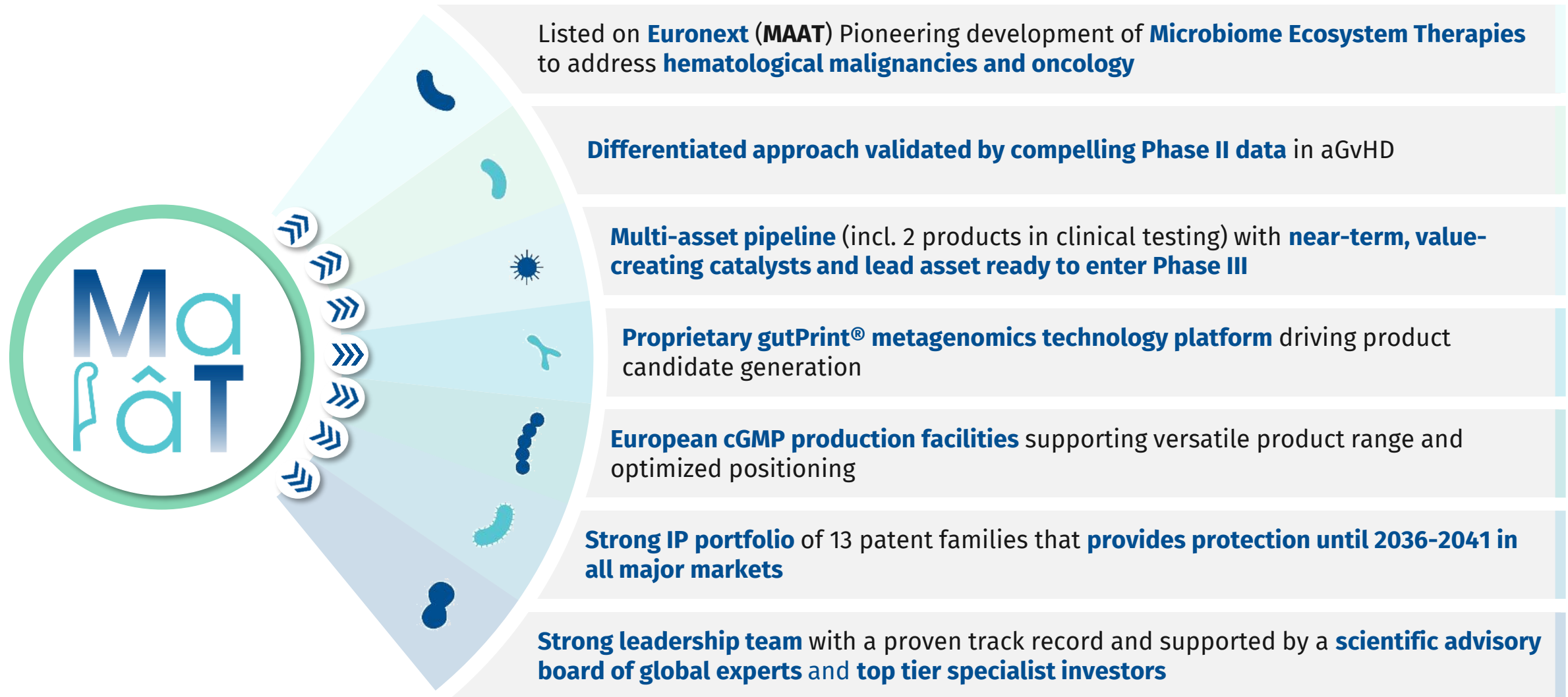
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A Uniquely-Positioned Microbiome Company



Management Team



Siân Crouzet
Chief Operating Officer



Hervé Affagard
Founder & CEO



Dr. Carole Schwintner
Chief Technology Officer



Dr. Savita Bernal
Chief Business Officer



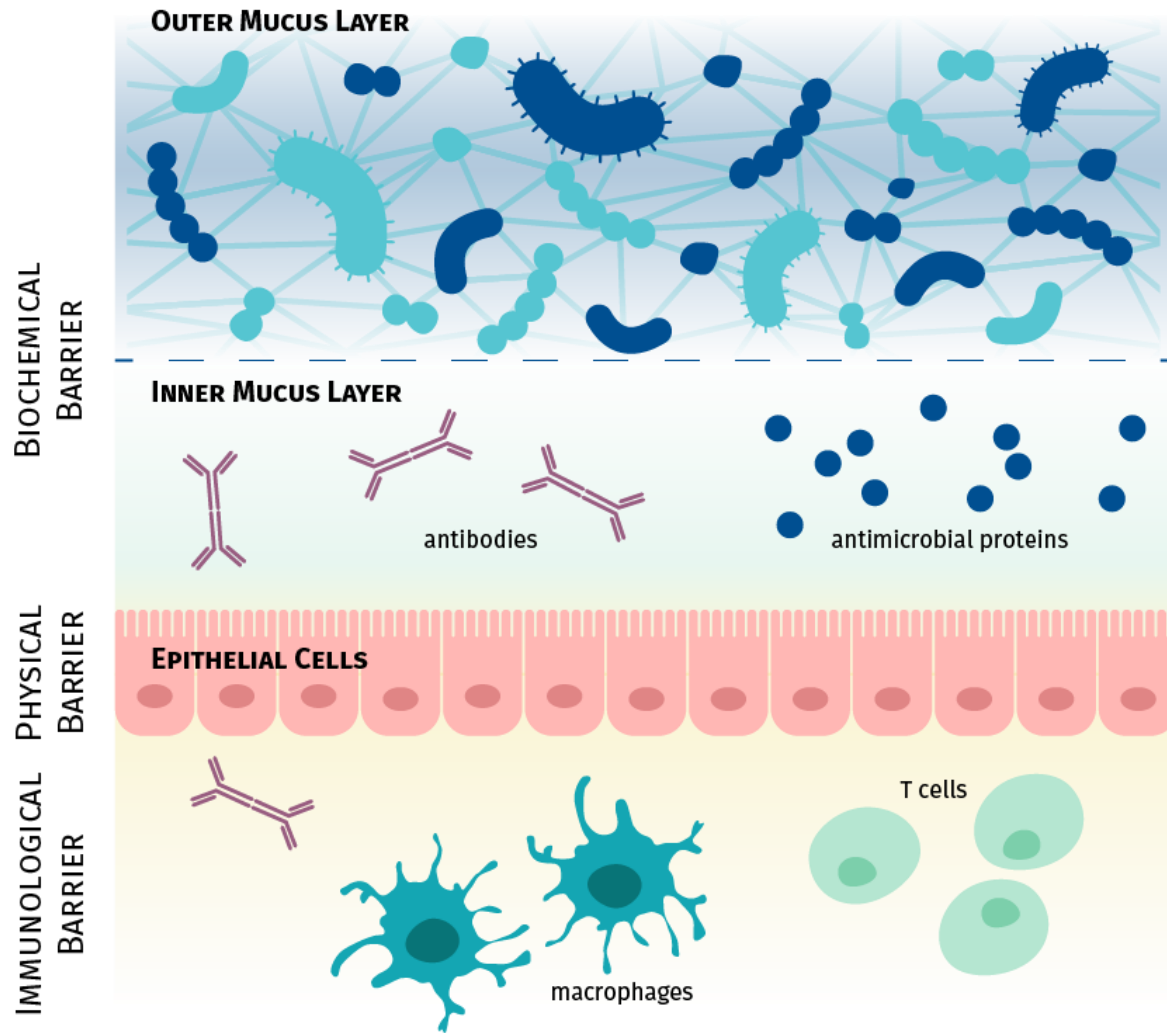
Dr. John Weinberg
Chief Medical Officer



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

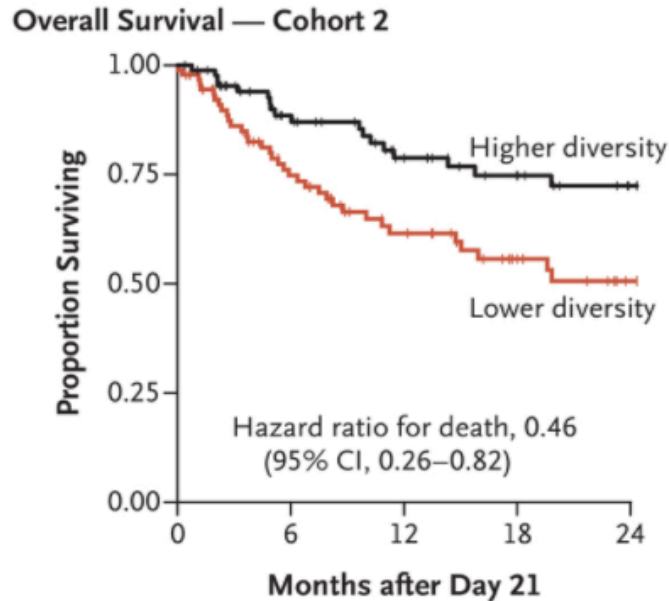
- A diversified microbiome contributes to the education and modulation of our immune system throughout life
- Bacterial richness and mucus layer prevent colonization by pathogens and improve gut barrier
- 80% of cellular host defense are localized in the gut (including innate and adaptive systems)

Diversity matters!

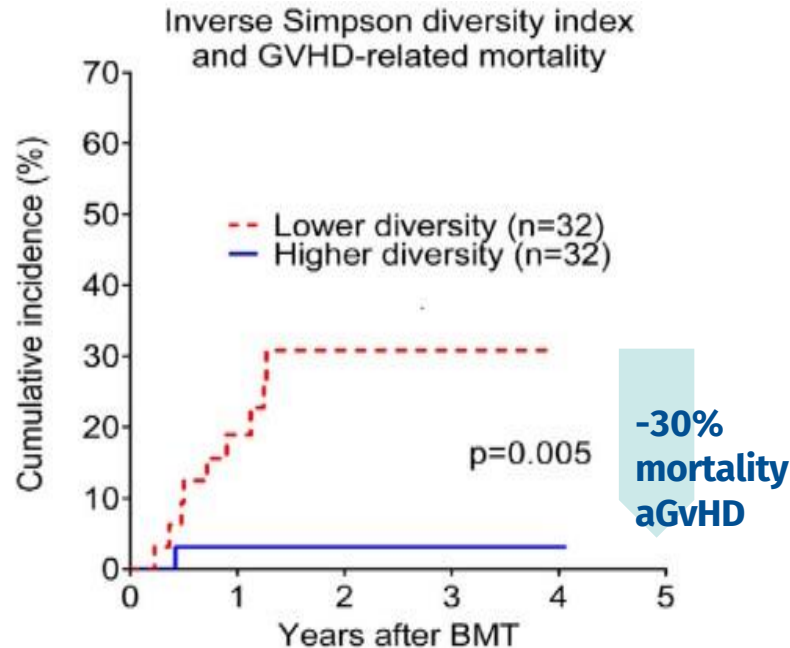
Higher gut microbiome diversity is associated with ...

Liquid Tumors

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



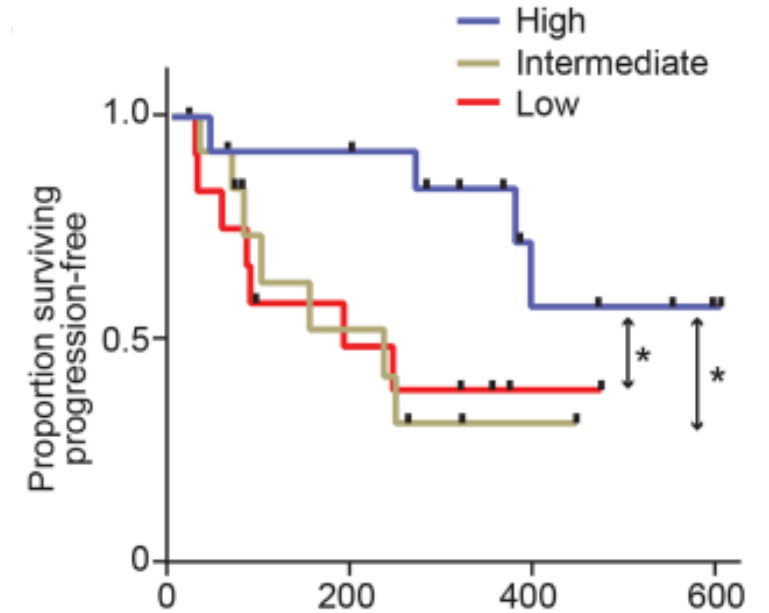
Lower incidence and lower mortality from aGvHD^{*,2}



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

MaaT013



MaaT033



- ✓ High diversity
- ✓ Full ecosystem
- ✓ **Butycore™** (anti-inflammatory)

Co-fermented

MaaT03X



- ✓ Indication-specific designed ecosystem (from clinical data)
- ✓ Innovative ecosystem co-fermentation technology

Entering Phase 3
aGvHD

Entering Phase 2
I/O

Phase 1
Allo-HCT

Preclinical
Solid Tumors
I/O

¹ **Butycore**: Group of 15 different genera known to produce short-chain fatty acids with anti-inflammatory properties
Corporate Presentation December 2021



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

MaaT013
~ 2,000 patients¹

MaaT033
~ 22,000 patients¹

MaaT03X
>200,000 patients¹

3rd line SR
GI
aGvHD

10x TAM

Prevention of complications of HSCT

>10x TAM

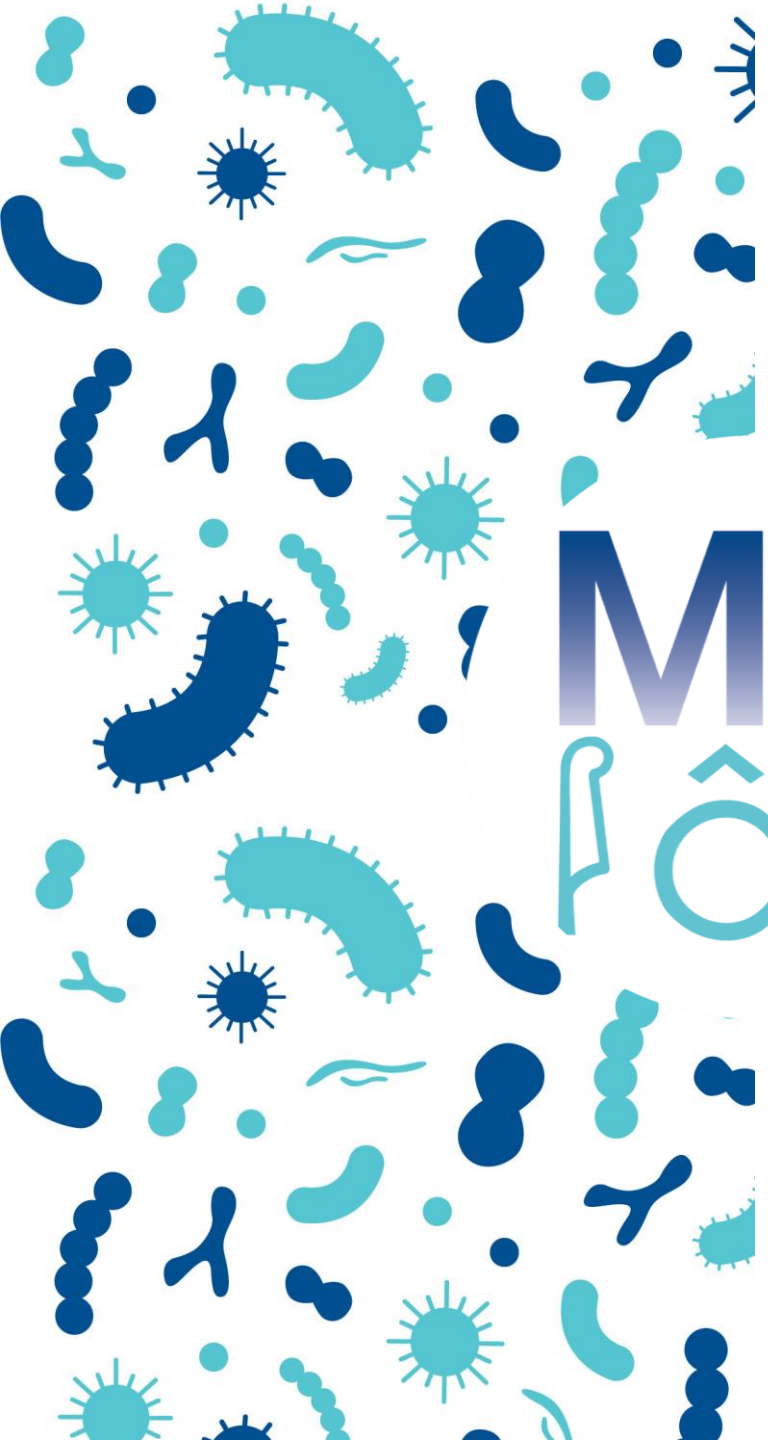
Melanoma, NSCLC, and other solid tumors



Long-term Expansion (Immune / inflammatory)

MaaT Pharma has a clear roadmap to **expand its total addressable market (TAM)** with **frequent market approval (MA) milestones** and **continuous product candidate generation**

¹ EU5, US, and Japan

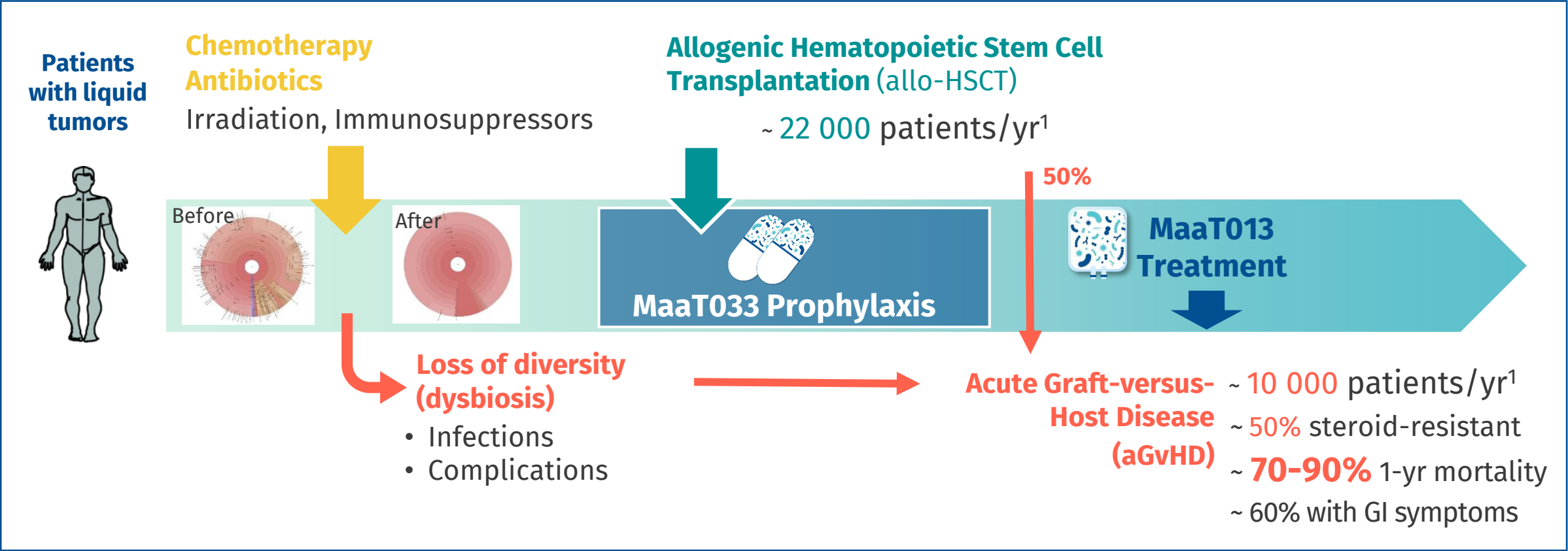


Mã Rất

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 procedues with an additional 7%-10% recurring), 2. EU5 + US



Ma
Rôl

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)
 - 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)

MaaT013 has received Orphan Drug Designation from the **FDA and EMA** for aGvHD

¹ Program is ongoing – 63 patients treated as of Oct 2021

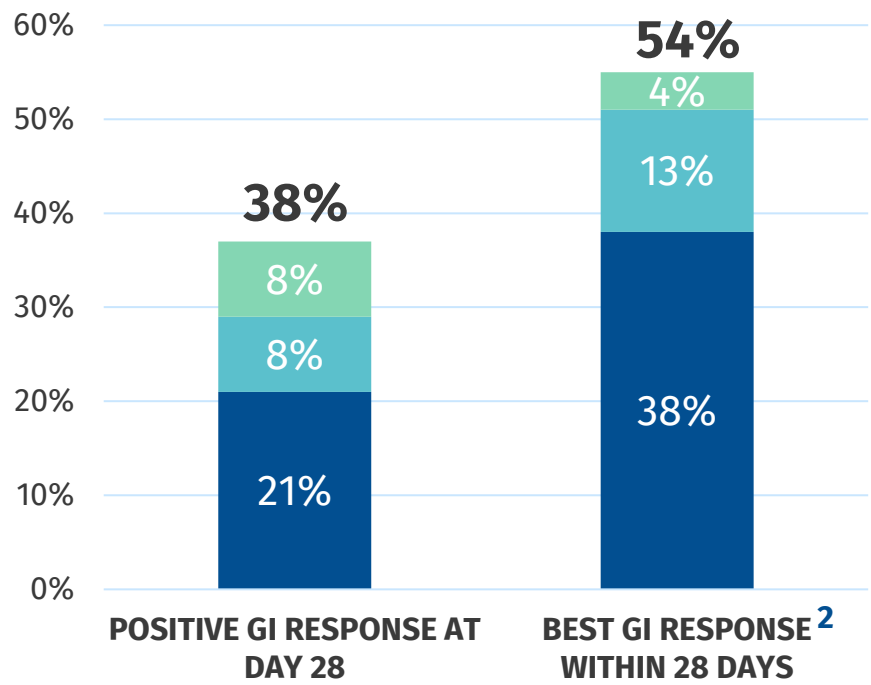


HERACLES Phase 2 Clinical Trial

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

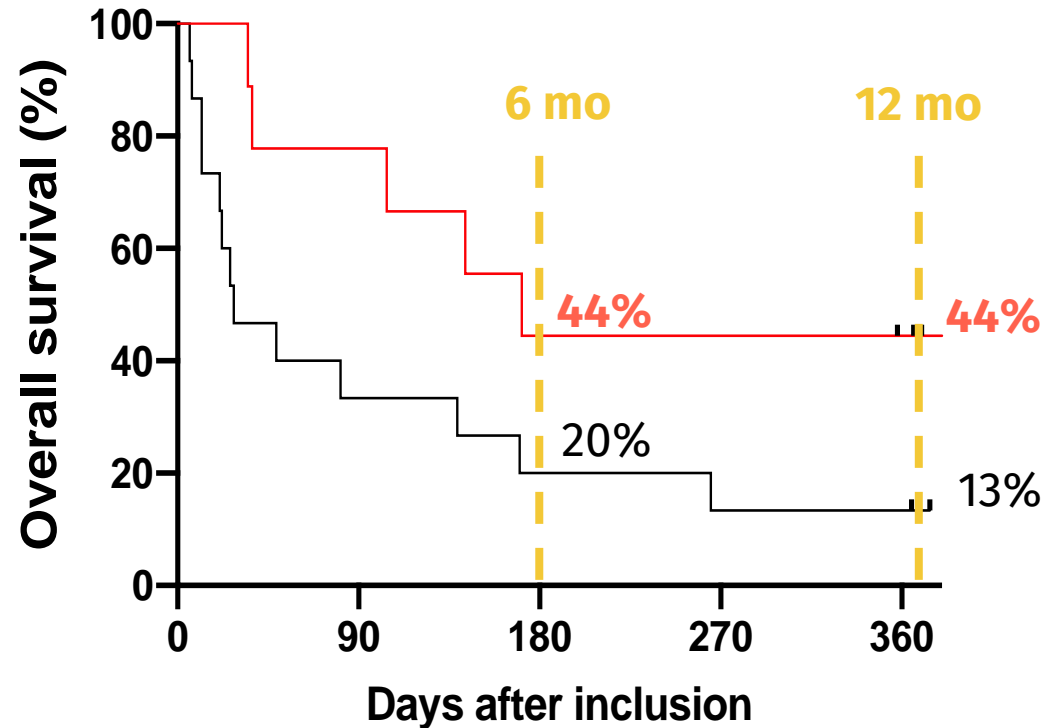
Gastro Intestinal Overall Response Rate (GI- ORR¹)



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

¹ORR= CR+VGPR+PR
² Best GI Response: Any response within 28 days

Overall Survival (OS)

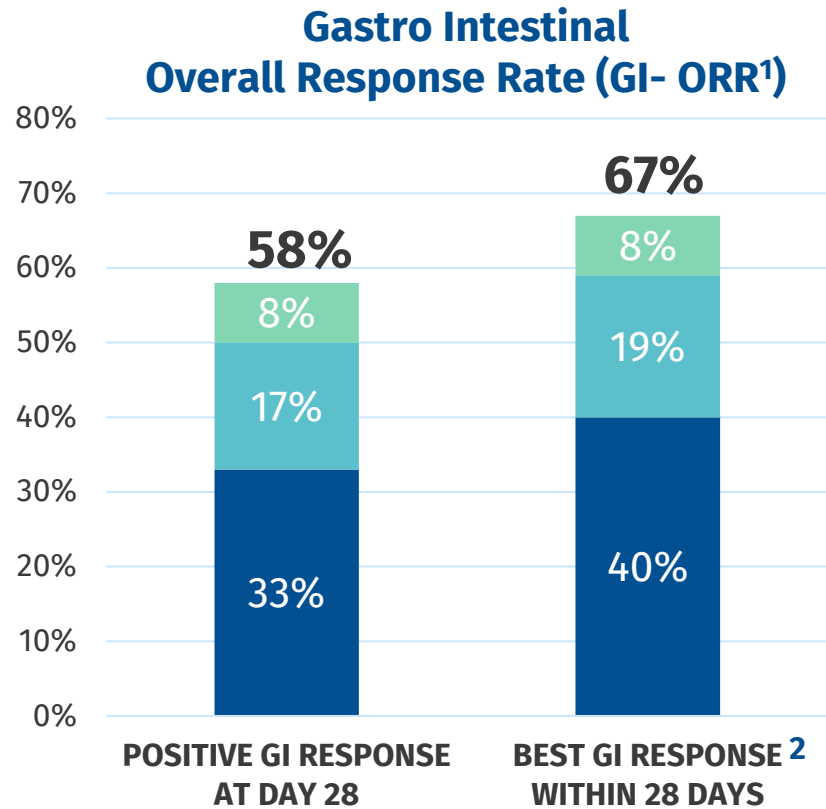


— Responders
 — Non-responders

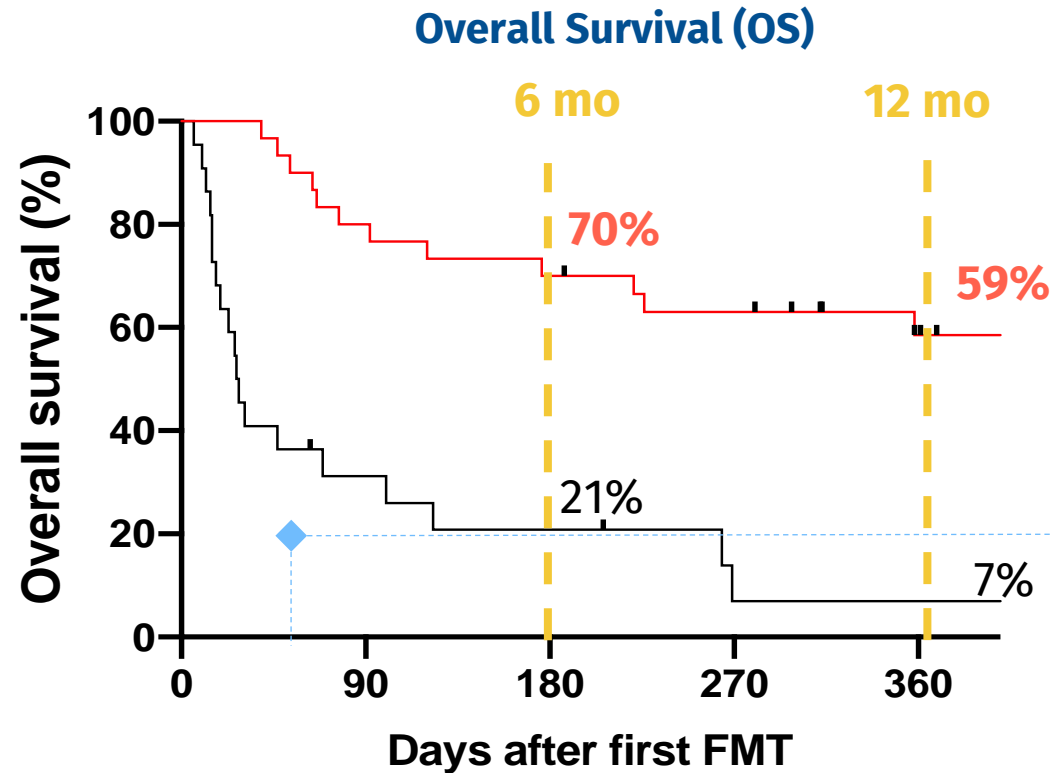


Early Access Program (EAP): Promising confirmation in an advanced, severe and more diverse GI aGvHD population

- 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) ¹ORR= CR+VGPR+PR
 ■ VGPR (very good partial response) ² Best GI Response: Any response within 28 days
 ■ CR (complete response)



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

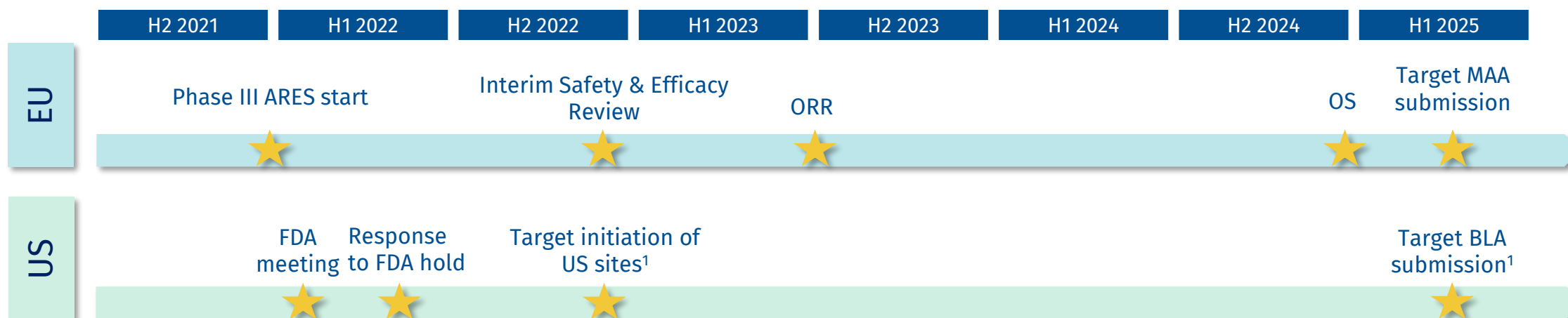
EUROPE :

- ✓ Study design reviewed by EMA through Scientific Advice procedure in Q1 2021
- ✓ CTA approved in 2 European countries and submitted to a third. Expected to expand to additional EU countries.

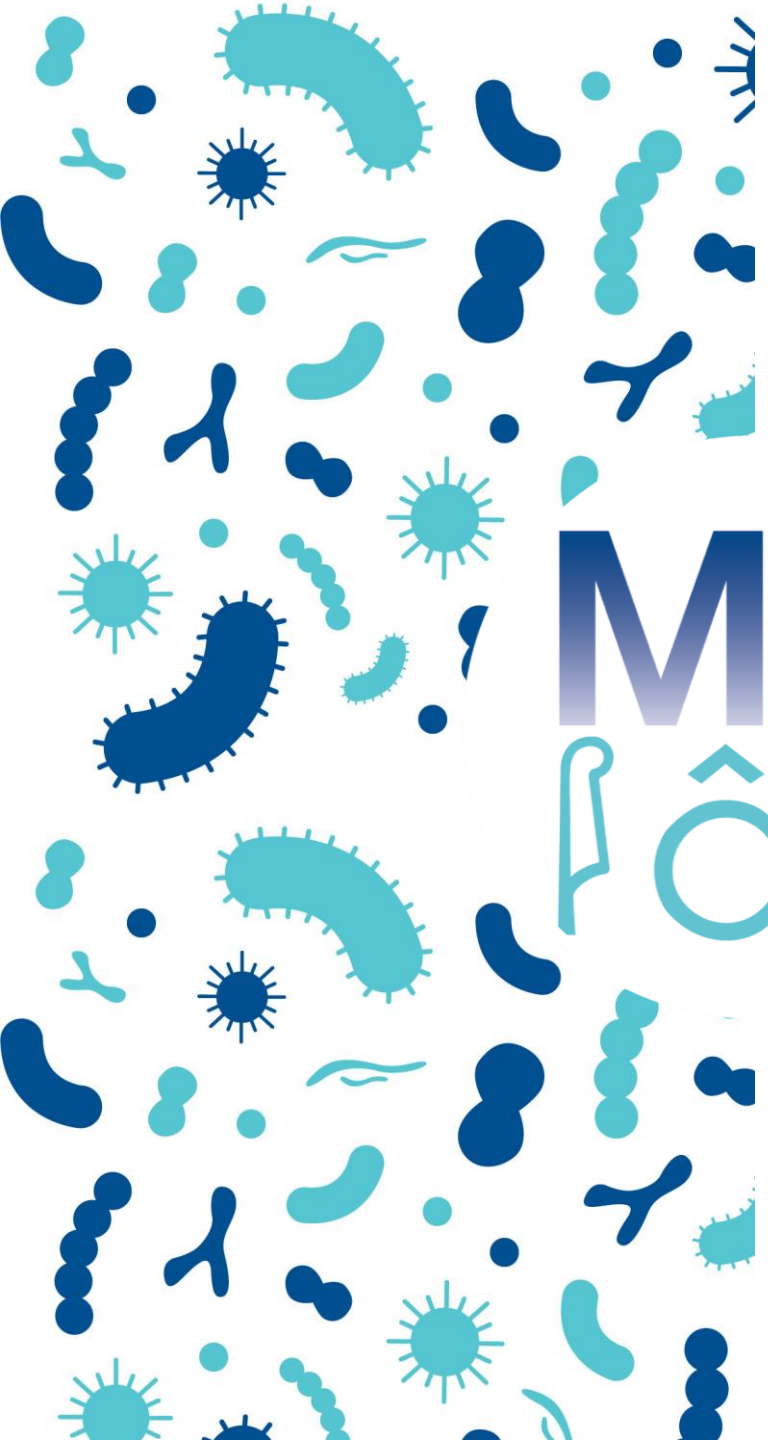
USA:

- FDA requested further information – on clinical hold.
- Will submit 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

Targeted Timelines ARES Phase III Trial



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



Mã Rất

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

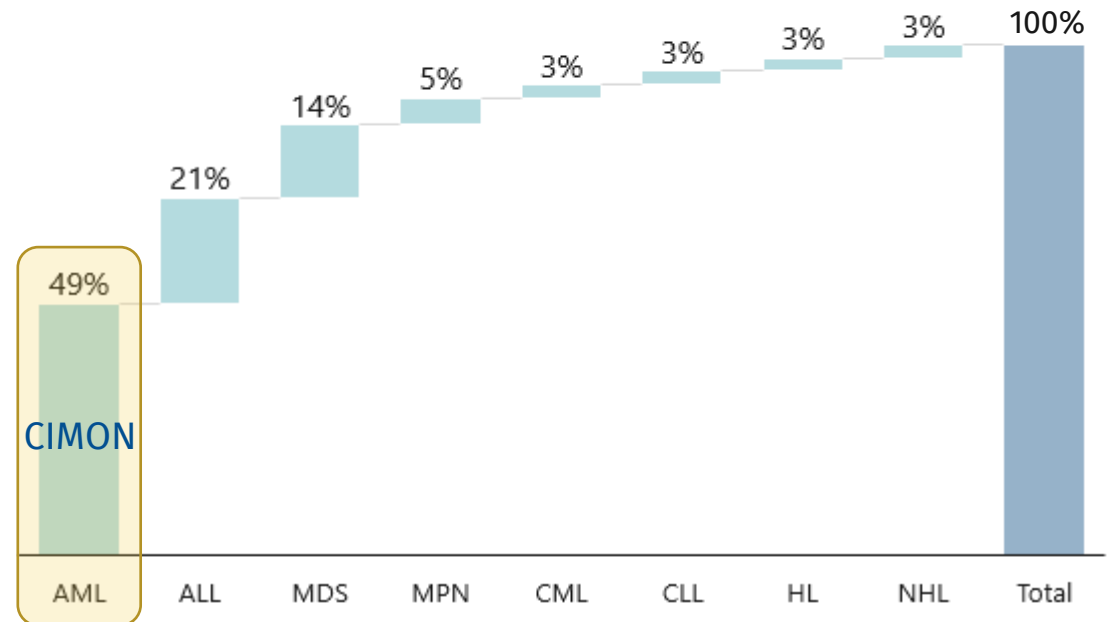


Japan

c. 3,000
primary
procedures

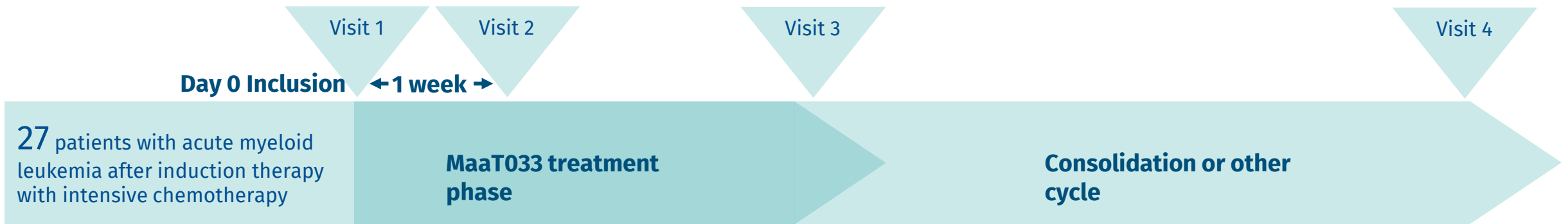
Additional 7%-10% recurrent procedures

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

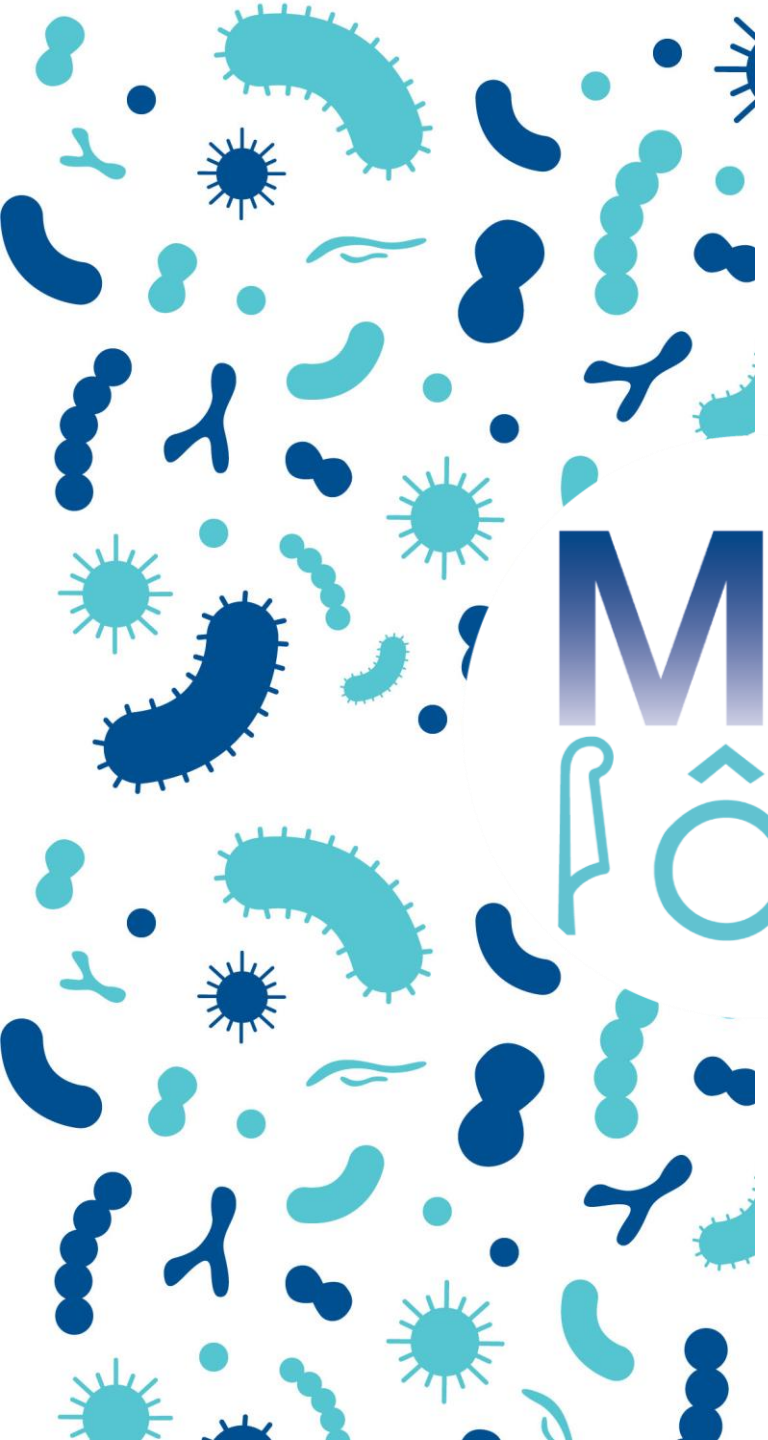
The ongoing Phase Ib CIMON study will determine MaaT033 dose for a Phase III study in post-allogeneic HSCT complication prevention



- Explores safety and the recommended dose of **orally administered MaaT033** in AML patients post induction chemotherapy
- **Primary Endpoint:** Dose limiting toxicity-related treatment emergent (serious) adverse events
- 5 Dose cohorts dosed daily for one to two weeks
- Trial is being conducted in six separate hospitals in France

Cohort 4 of 5 fully enrolled as of December 2021 with dose recommendation on track for H1 2022

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



Mã Rất

Immuno-Oncology
Solid Tumors

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

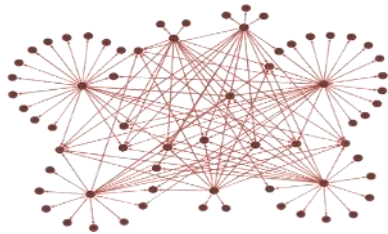
FMT from ICI responders to ICI non-responding patients with metastatic melanoma

✓ **6/15**

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

✓ **3/10**

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

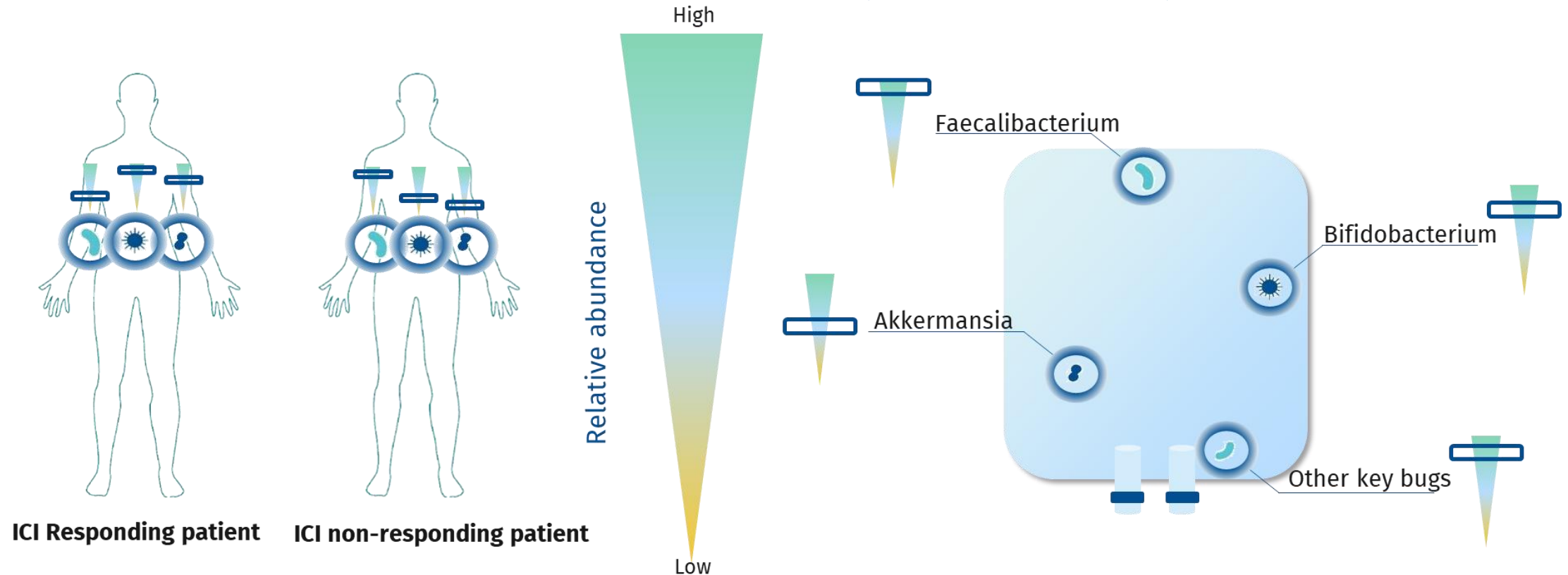


- Immune check-point inhibitors (ICI) therapies have established themselves as key therapeutic options in solid tumors, but ORR may be as low as 20% in some indications.
- Richness and Diversity of gut microbiome drive survival in patients receiving ICI^{1,2,3}
- FMT from ICI responders (R) could induce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

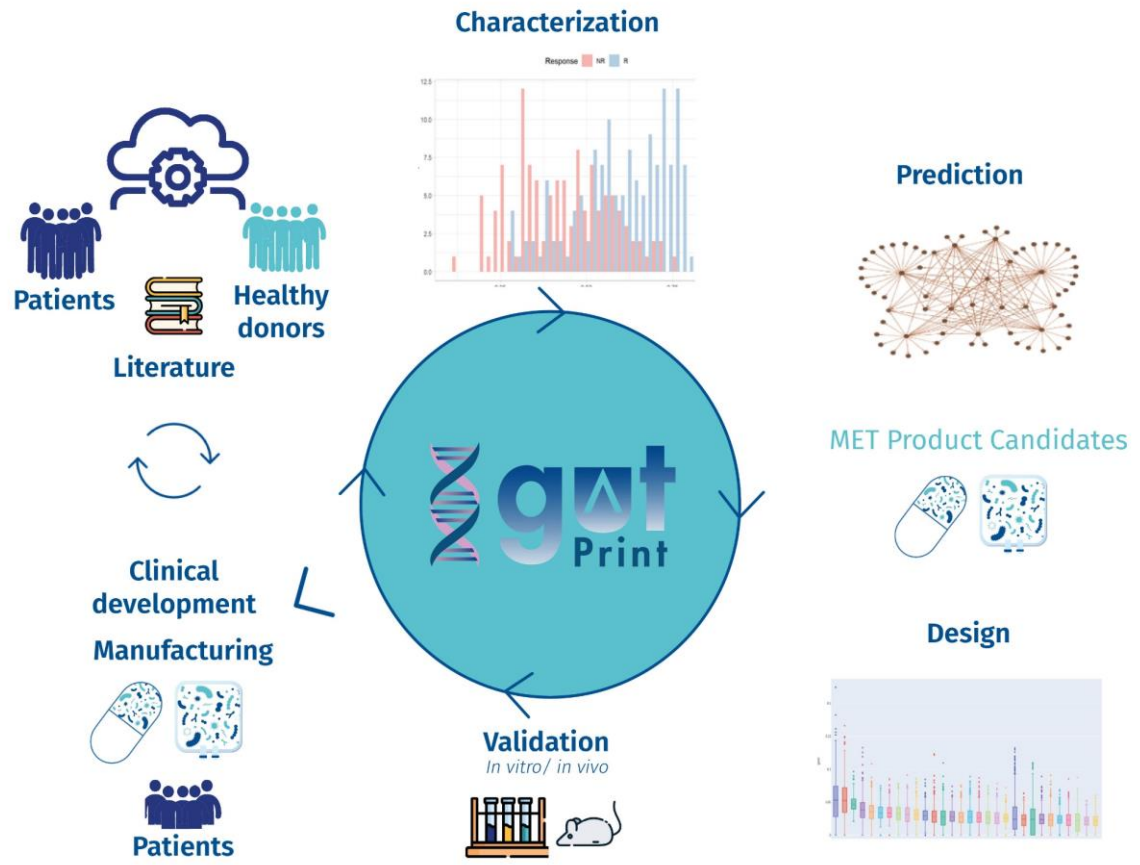


MaaT013 ensures high diversity and contains specific bacterial strains that have been identified to improve ICI response



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

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



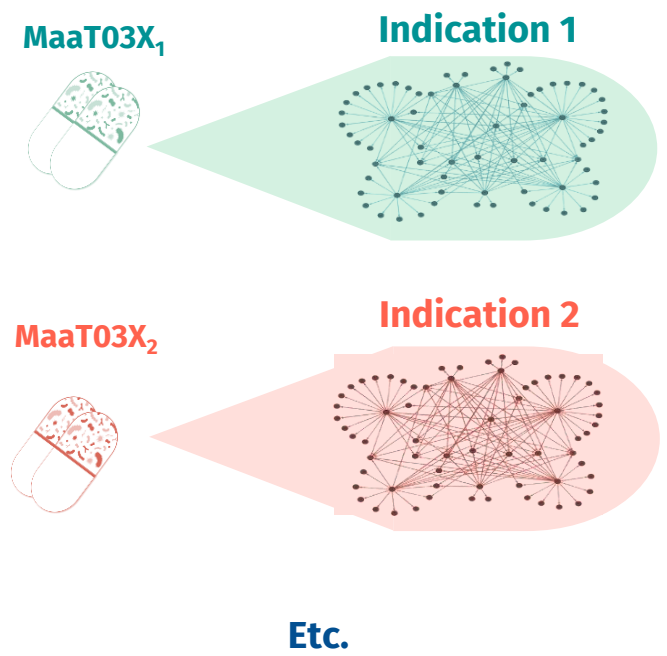
gutPrint® is the engine that drives MaaT Pharma's MET product candidate generation capabilities that broaden and strengthen the pipeline



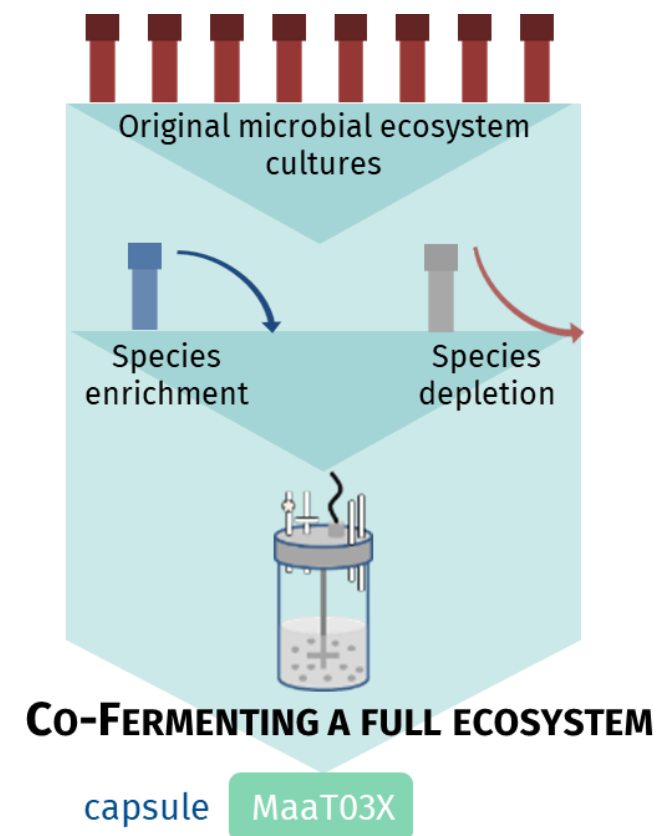
The customizability and scalability of the MaaT03X line allows it to potentially address several solid tumor indications

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

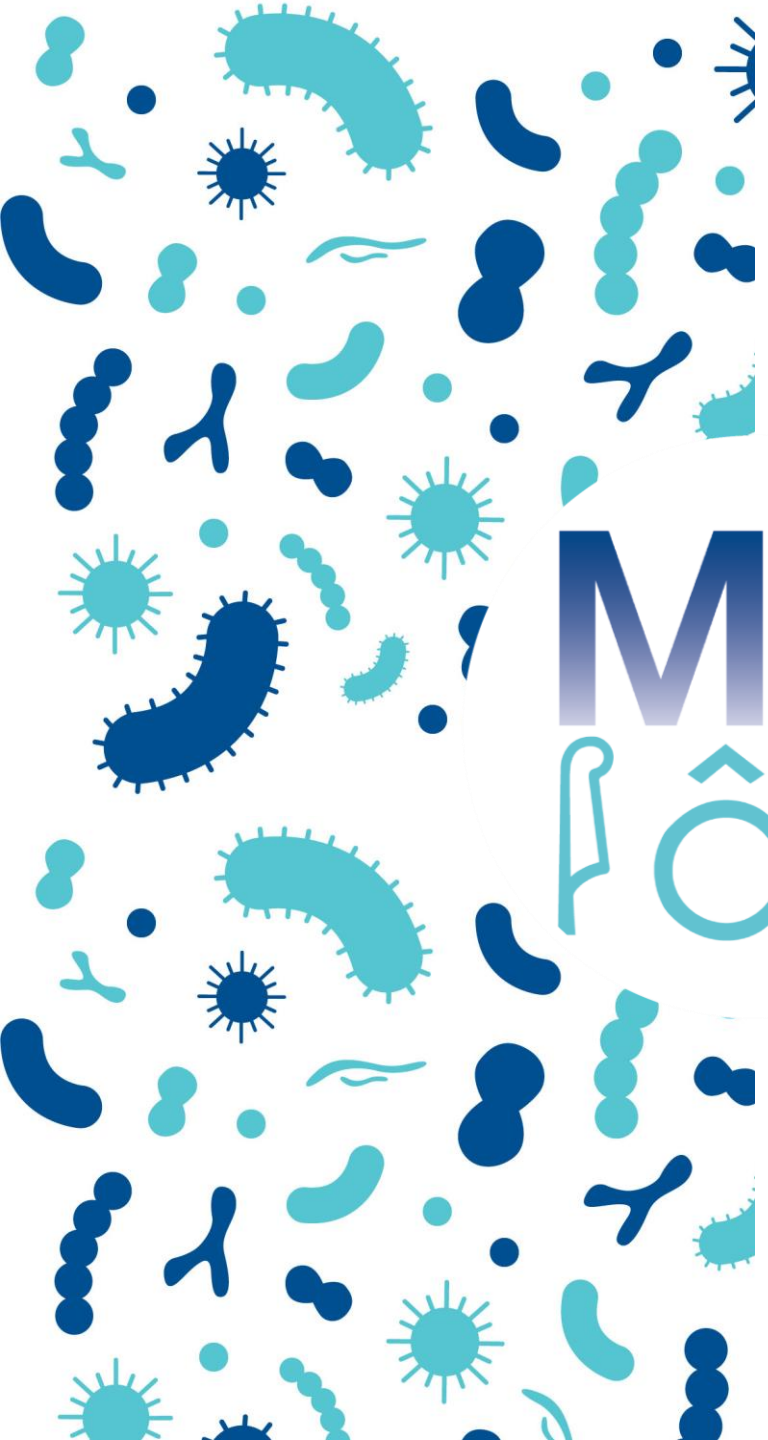
Designable according to indication-specific microbial signatures



Customizable, donor-independent, scalable process



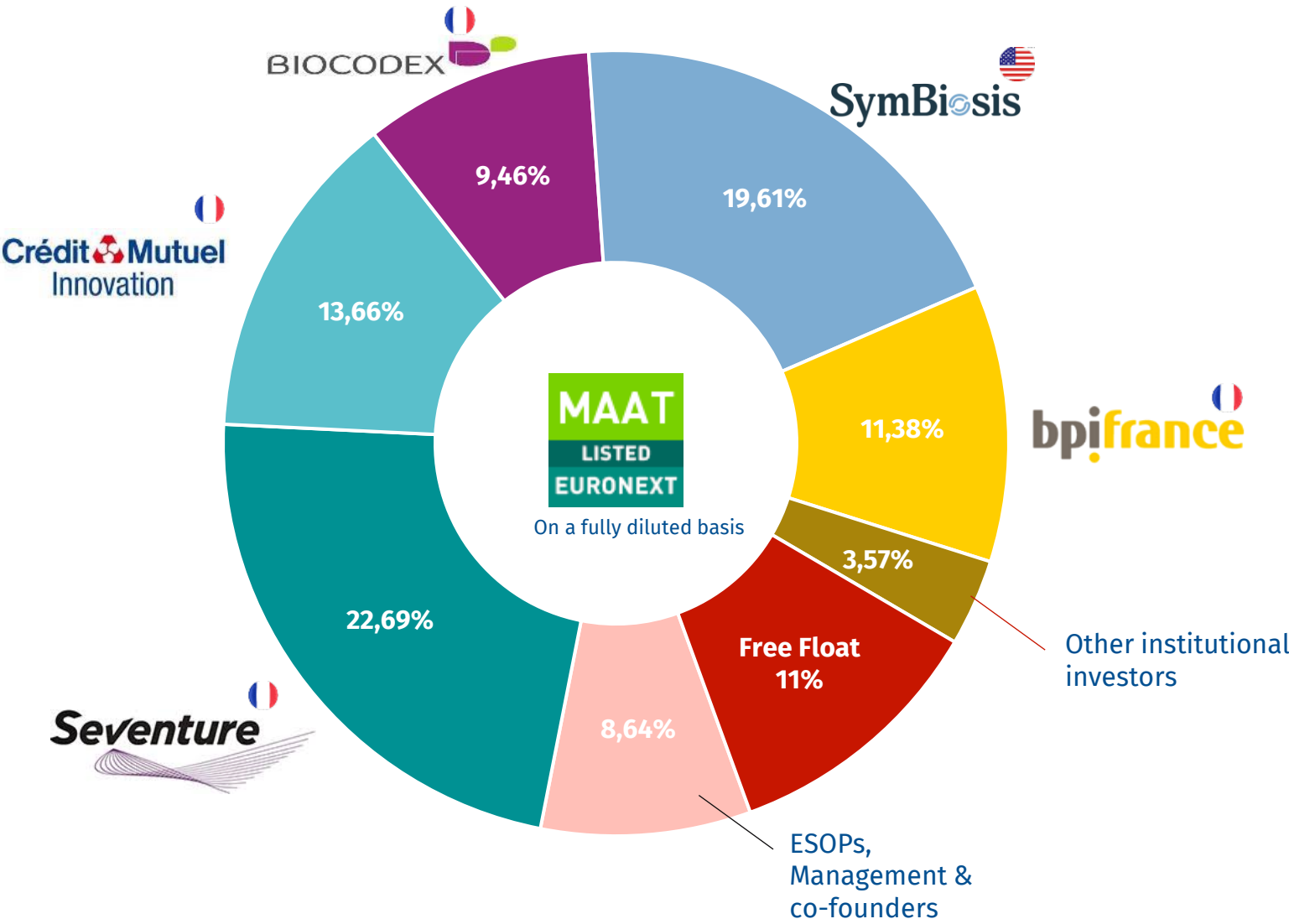
First candidate in preclinical testing – Expected to enter clinical testing in H1 2023




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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
- 

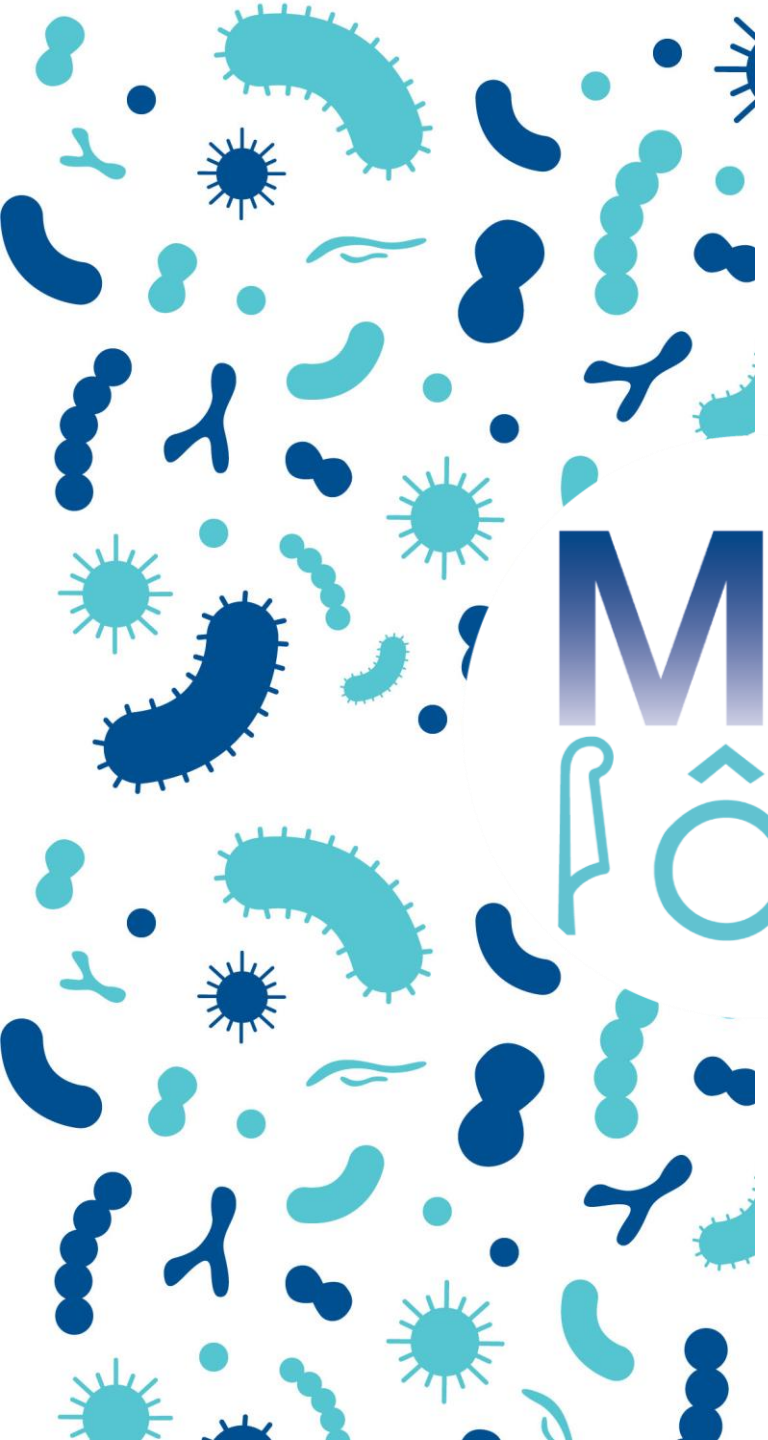
Martine George
Non-Executive Director
Oncologist
- 

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

 Intermediate review
 Results  Launch

2021	2022		2023		2024		2025	
H2	H1	H2	H1	H2	H1	H2	H1	H2

Graft-vs-Host-Disease
MaaT013 (pooled enema)
FDA & EMA Orphan Drug Designation

Phase III ARES in EU (pivotal)¹



ORR



Overall Survival



Post-allogeneic HSCT complications
MaaT033 (pooled capsule)

Phase Ib



Phase II/III OR-ALLO (pivotal)

Immuno-Oncology potentiation - Melanoma
MaaT013 (pooled enema)

Phase IIa PICASSO²



Undisclosed
MaaT03X (fermented capsule)

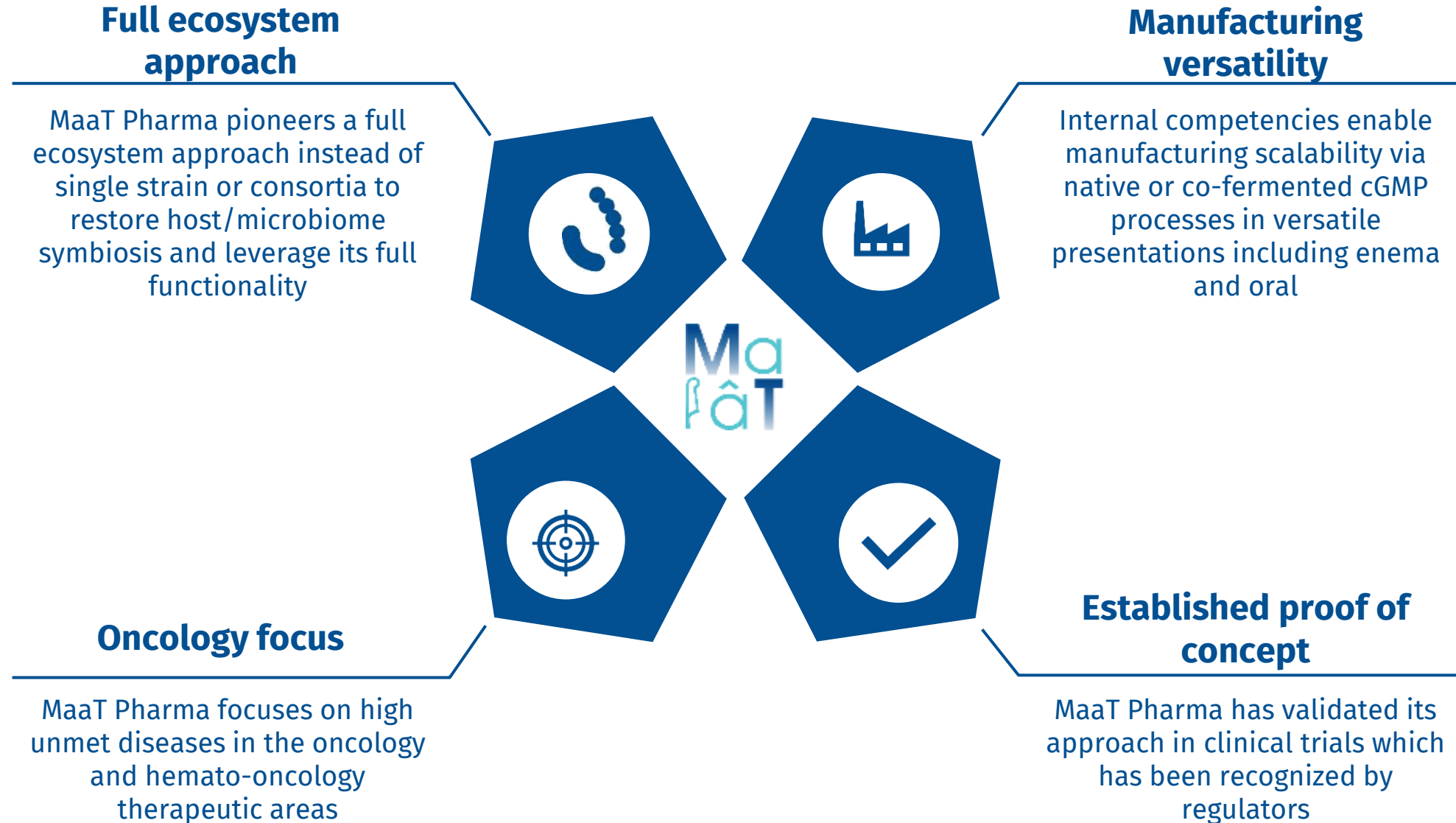
Phase I/II

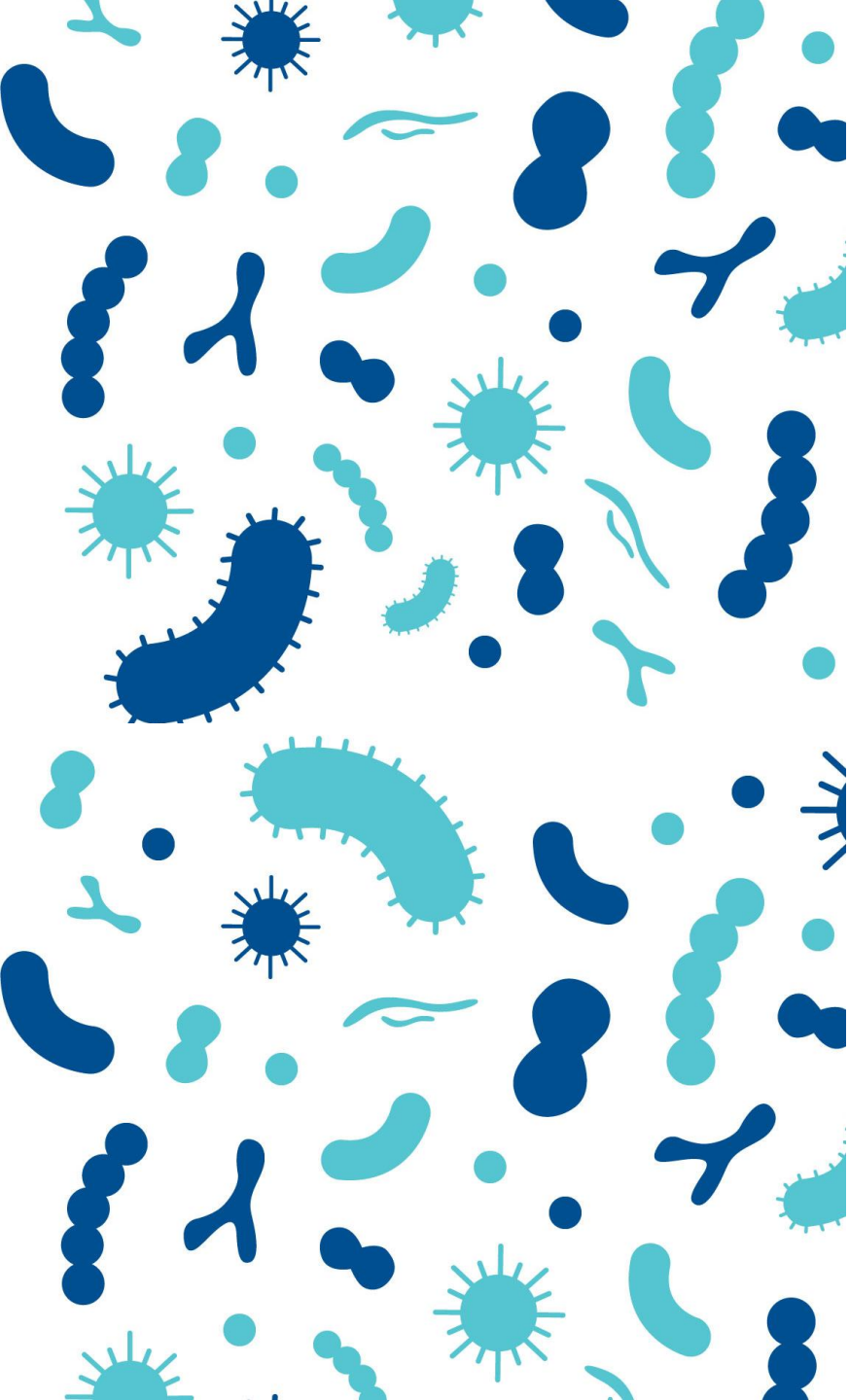


¹expansion to US sites in H2 2022 subject to IND approval in the US;

²Investigator sponsored trial where MaaT Pharma supplies the drugs and performs the microbiome profiling using its gutPrint® platform

Key differentiators of MaaT Pharma from other microbiome competitors





THANK
YOU