



MaaT Pharma Microbiota as a Therapy

Company Presentation
June 2022

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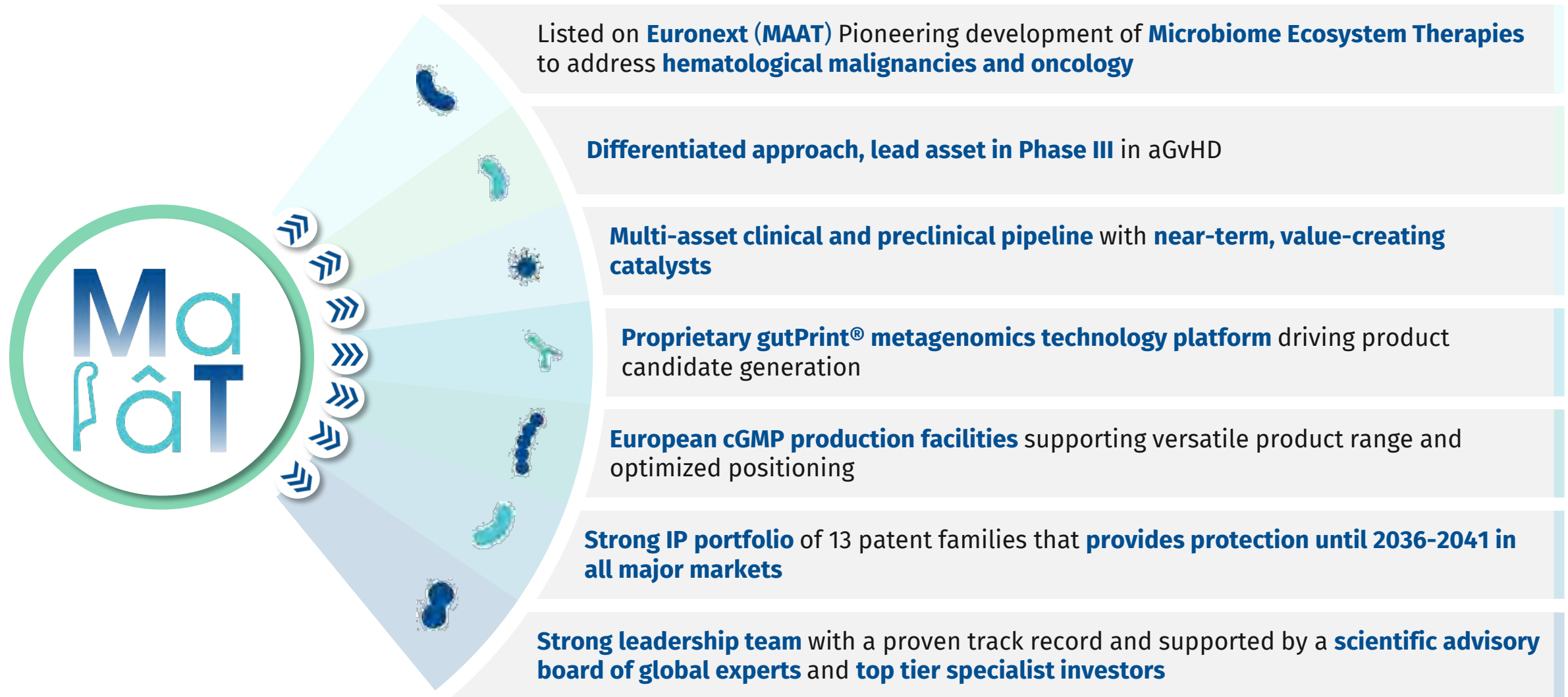
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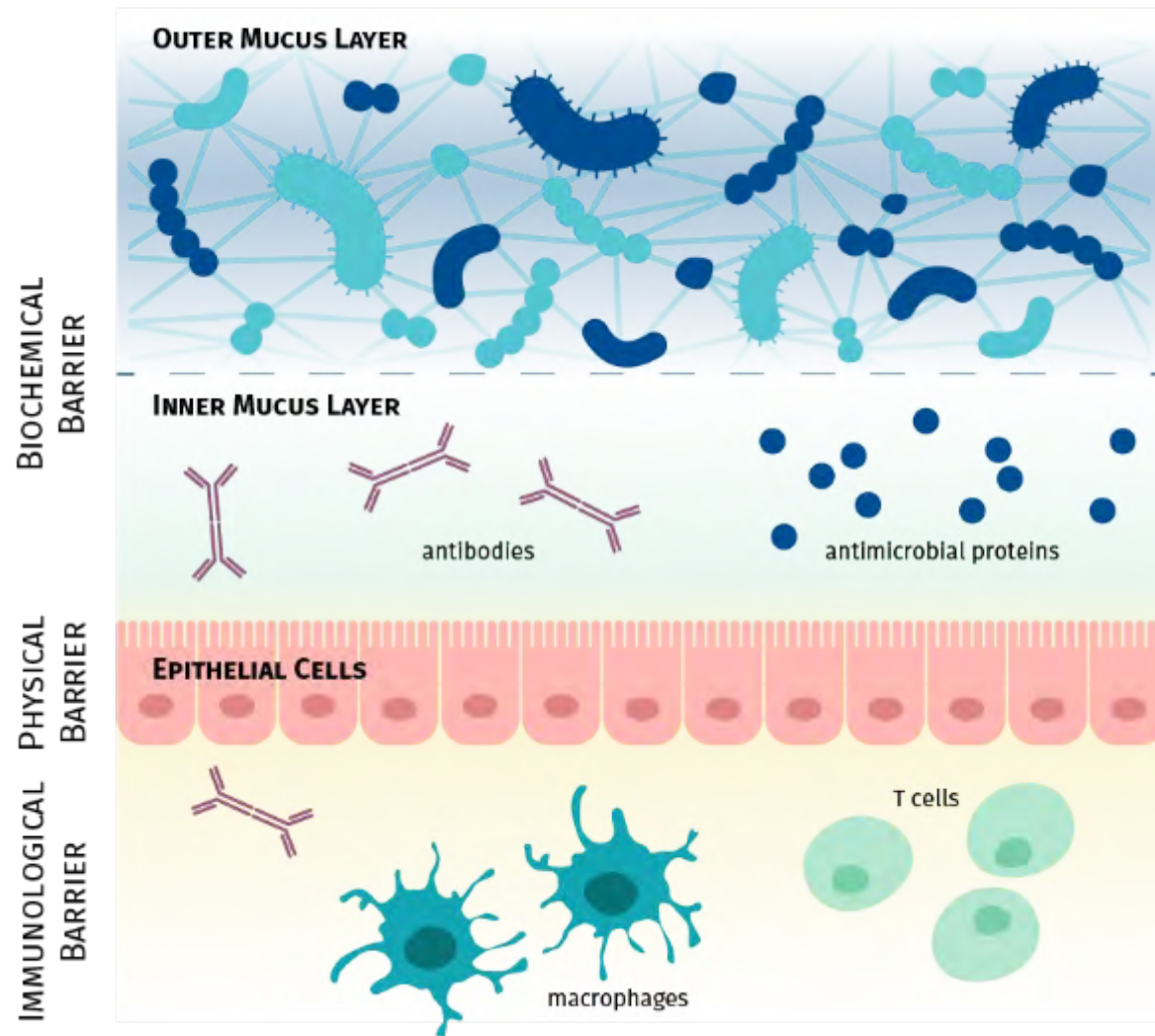
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A Uniquely-Positioned Microbiome Company



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

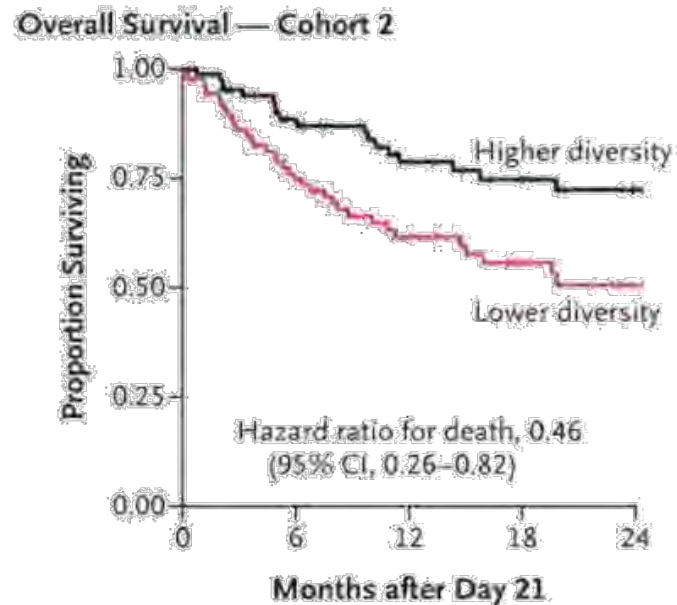
- A diversified microbiome contributes to the education and modulation of our immune system throughout life
- Bacterial richness and mucus layer prevent colonization by pathogens and improve gut barrier
- 80% of cellular host defense are localized in the gut (including innate and adaptive systems)

Diversity matters!

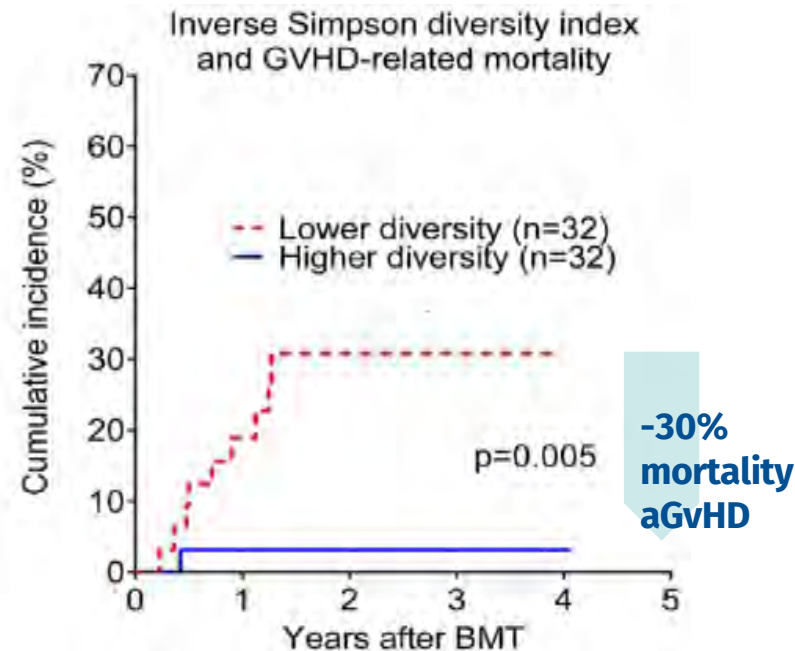
Higher gut microbiome diversity is associated with ...

Liquid Tumors

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



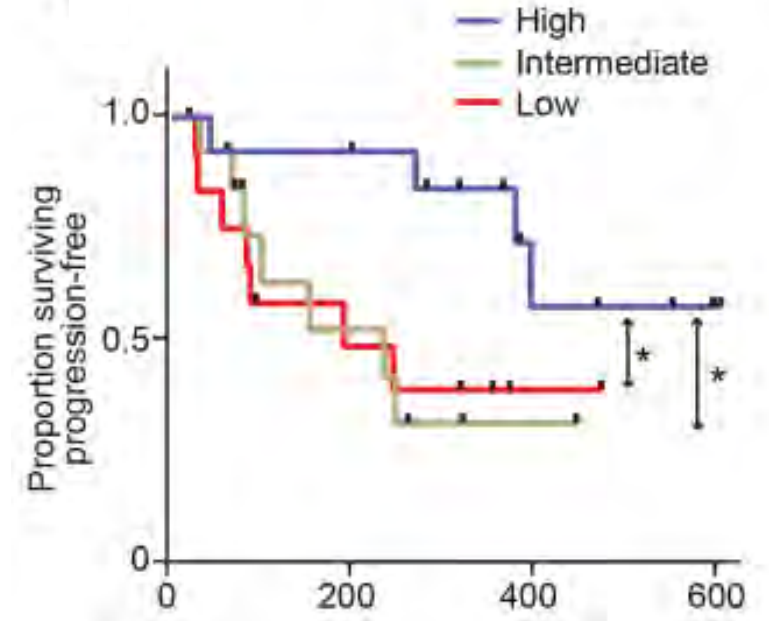
Lower incidence and lower mortality from aGvHD^{*,2}



MaaT Pharma MET Inverse Simpson (mean): 24

Solid Tumors

Higher response rate to ICI* in patients with metastatic melanoma³

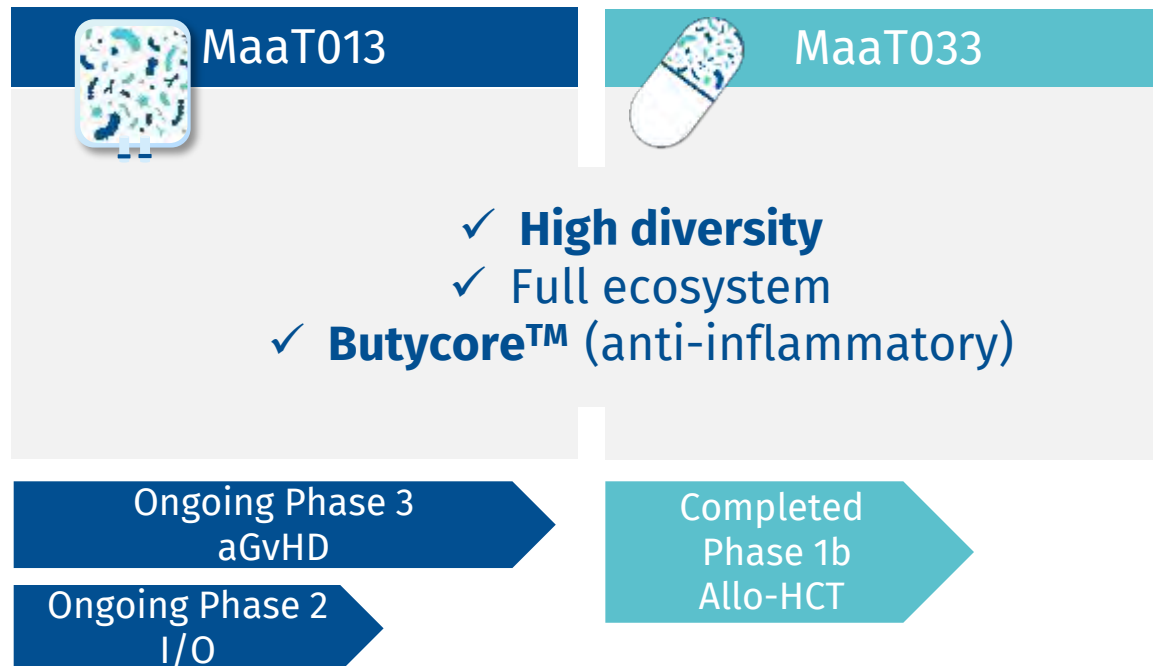


*allo-HSCT: allogeneic hematopoietic stem cell transplantation; aGvHD: acute Graft-vs-host-Disease; ICI: Immune Checkpoint Inhibitors
¹Peled, J.U. & al N Engl J Med 2020;382:822-34; ²Ghani, 2021; ³Jenq RR. et al, Biol Blood Marrow Transplant 21 (2015) 1373e1383; Pamer, Blood, 2014 ; Gopalakrishnan et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

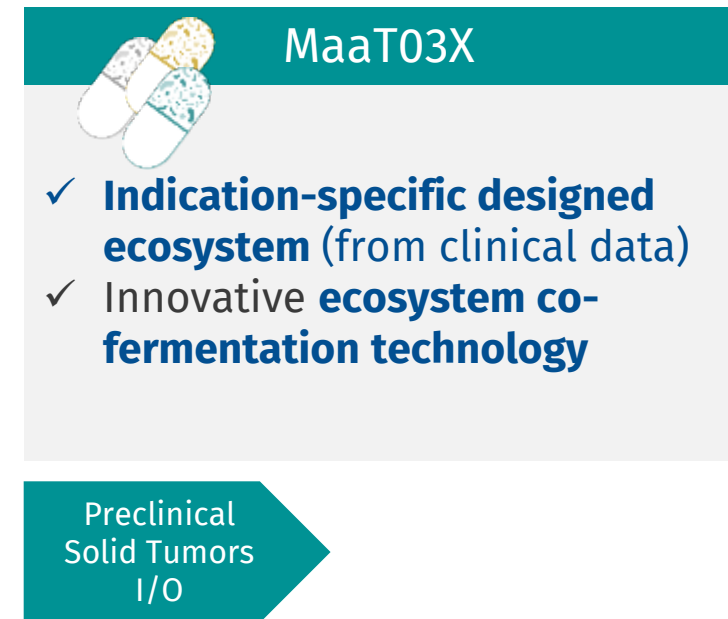
MaaT Pharma's Microbiome Ecosystem Therapy (MET) platform has generated a diverse line of product candidates



Native

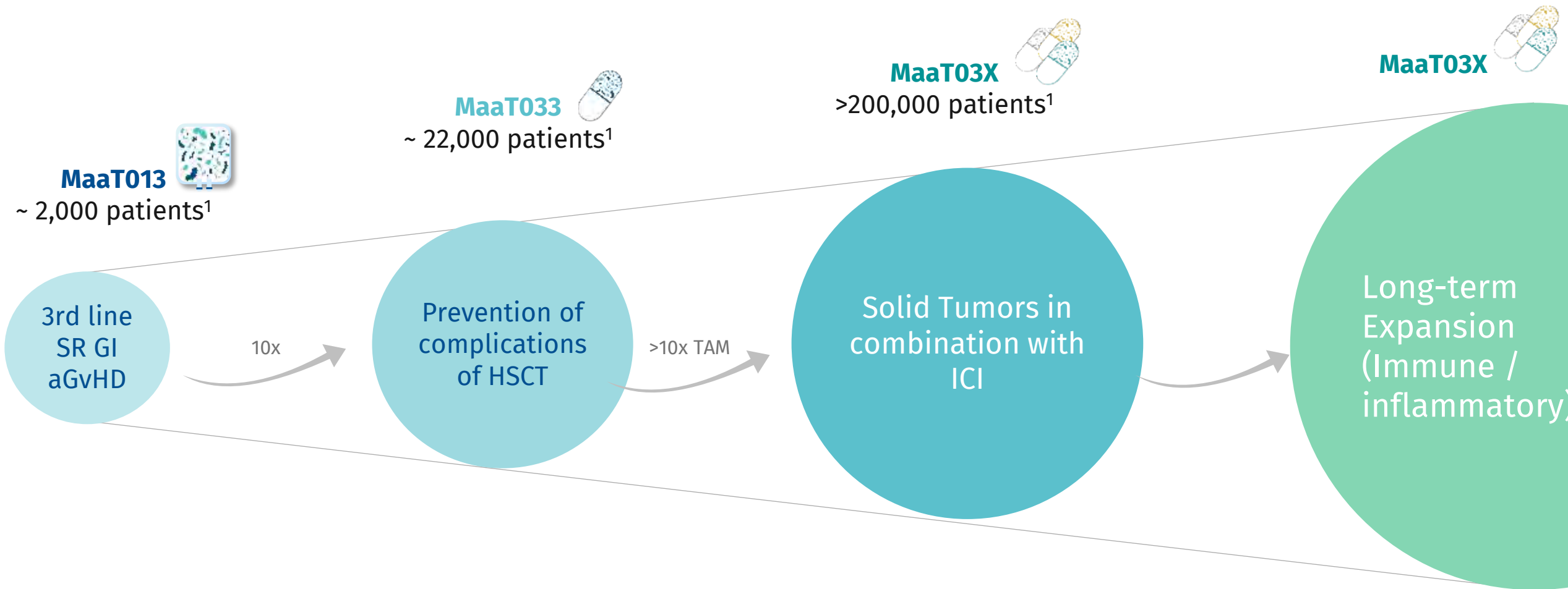


Co-fermented



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

Looking ahead: addressing growing market opportunities with severe medical need



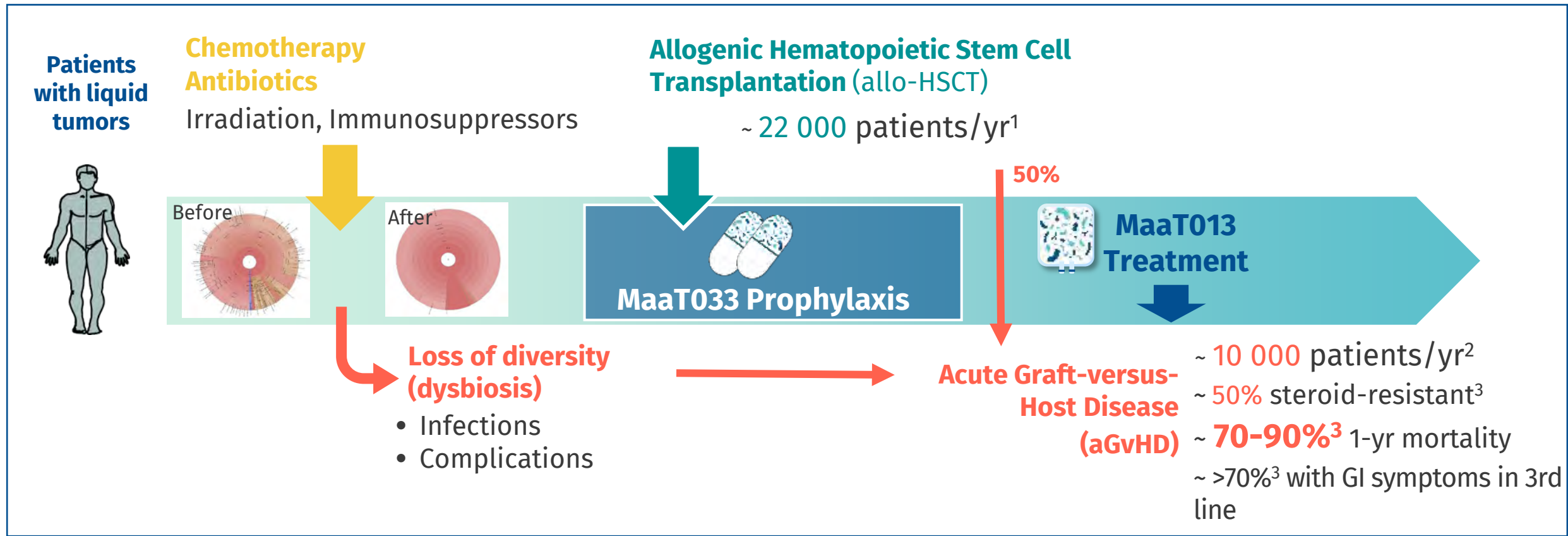


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Hemato-Oncology

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



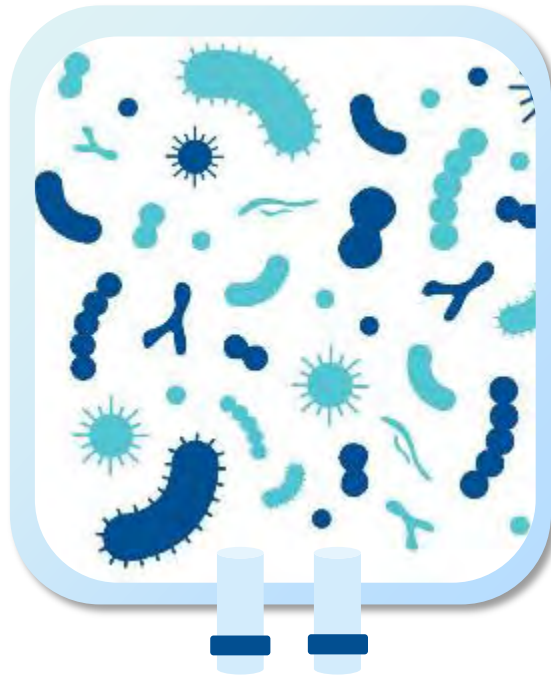
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Hemato-Oncology

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



MaaT013: restore the microbiome to *cure* acute Gastro-Intestinal graft vs. Host disease.



Characteristics

Pooled microbiota: a high-richness, high-diversity, full ecosystem, Microbiome Ecosystem Therapy containing Butycore™



Administration

3 doses (150 mL enema bag)



Available Clinical Data

- ✓ HERACLES Phase 2 Clinical Trial, N=24, 2L
- ✓ Early Access Program, data from N=52, 3L-6L, program still ongoing
- >110 patients treated to date



Efficacy evaluation (GI ORR at Day28)

Complete response (CR), Very Good Partial Response (VGPR), Partial Response (PR)



Current indication

Gastrointestinal acute Graft-versus-Host Disease

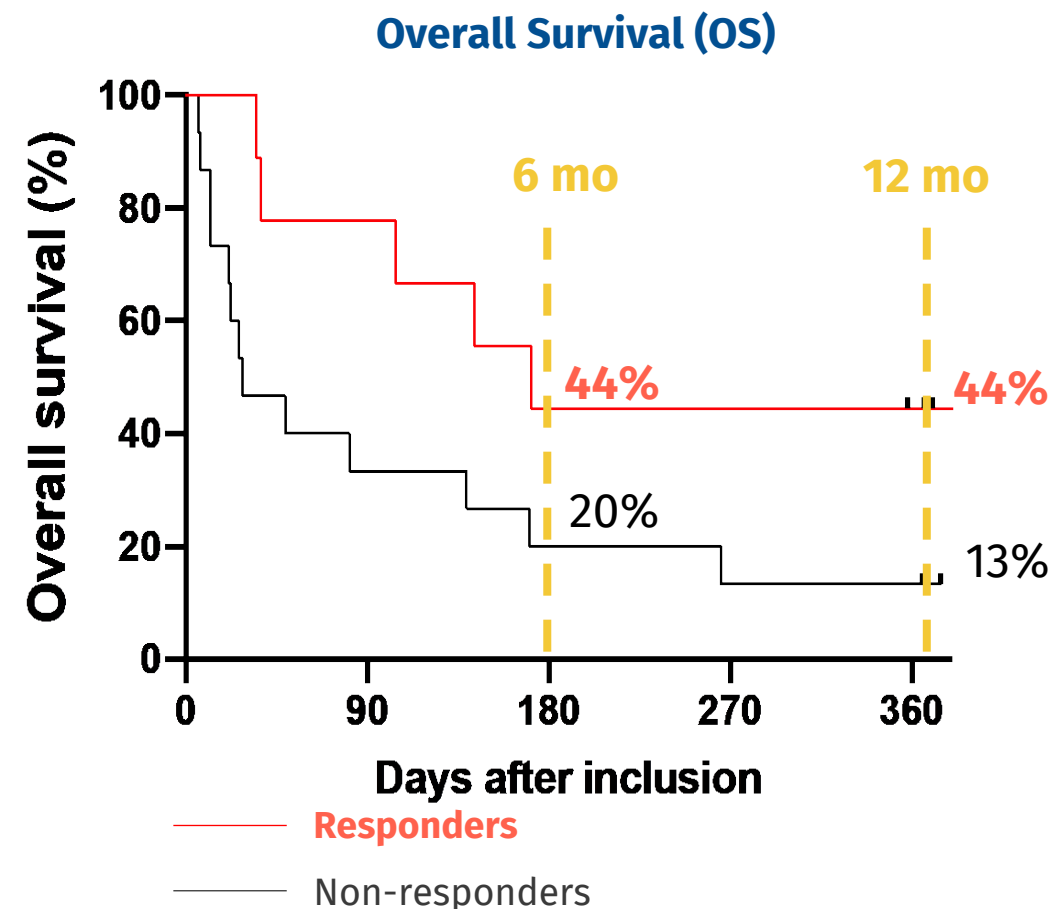
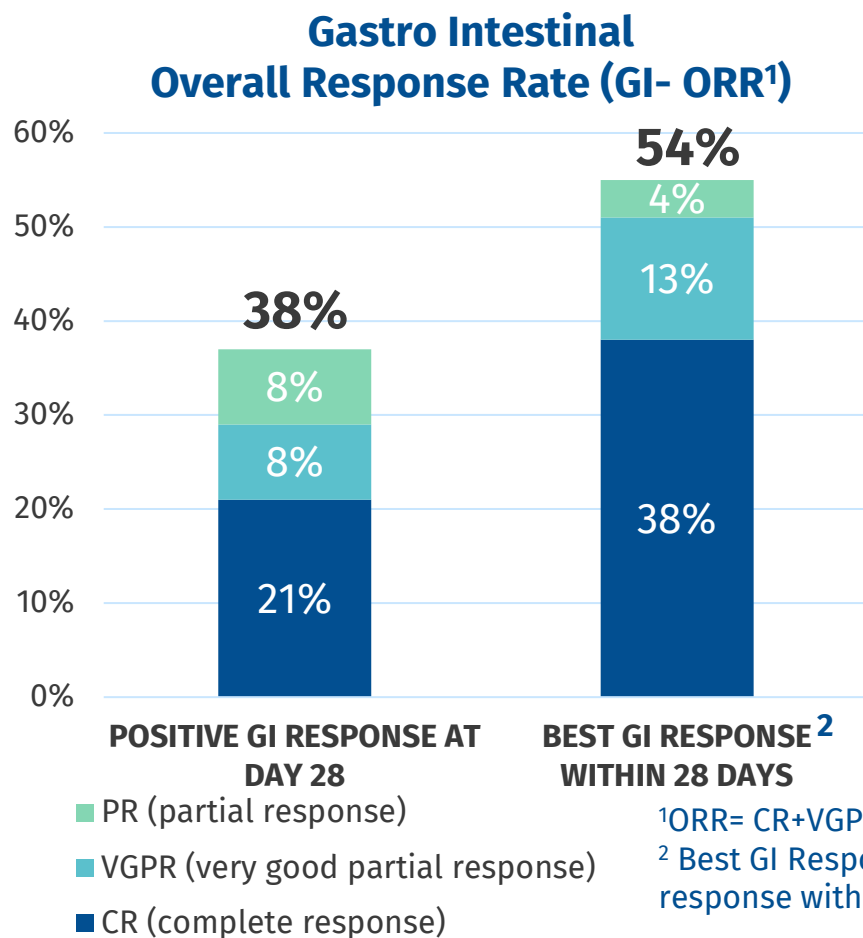
MaaT013 has received Orphan Drug Designation from FDA and EMA



HERACLES Phase 2 Clinical Trial

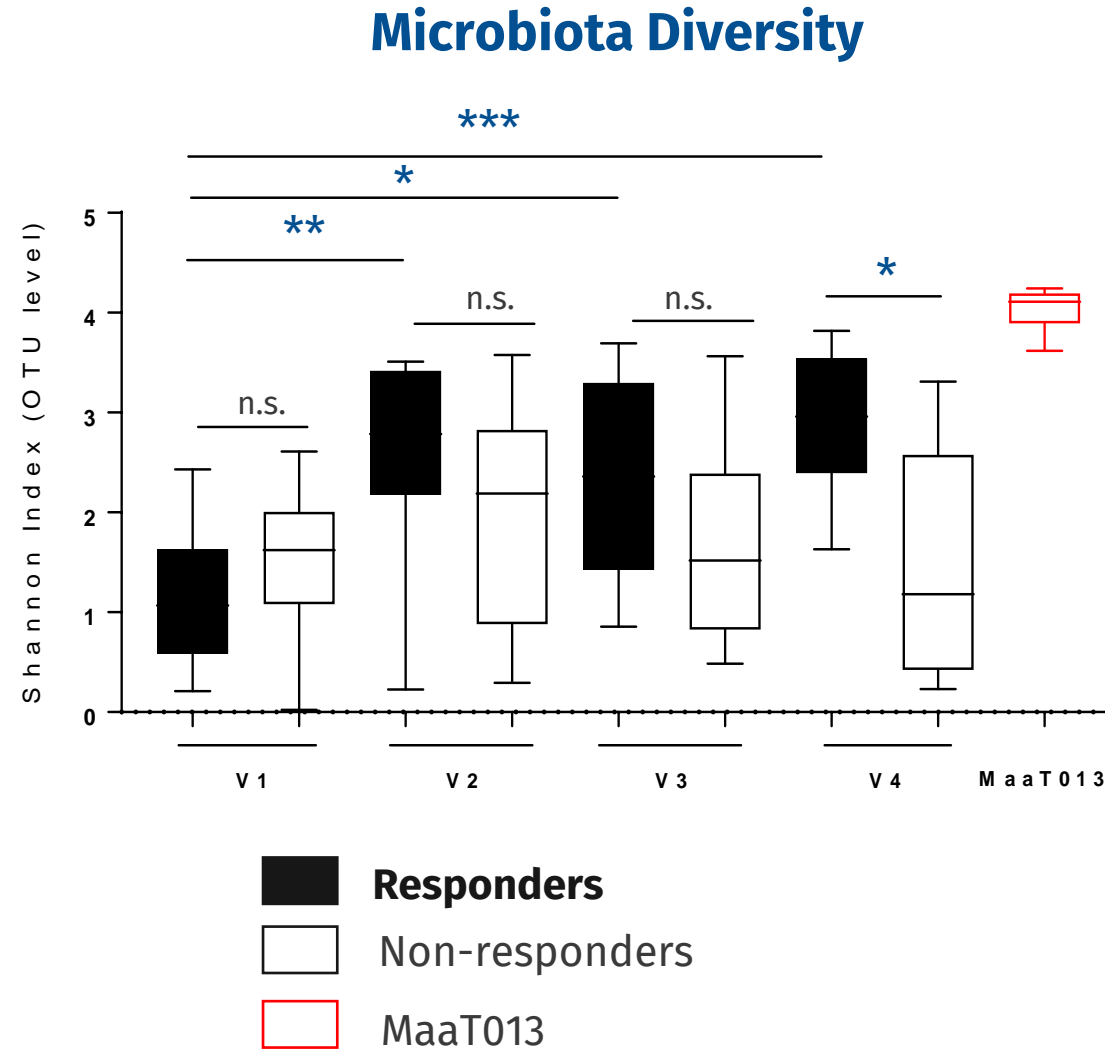
Promising results in a very severe (III-IV) GI aGvHD population

- N=24 patients, 96% grade III (4% grade IV), 2nd line (Steroid-resistant)
- Very good safety and tolerability profile





HERACLES: MaaT013 increases Responders' gut microbiome diversity

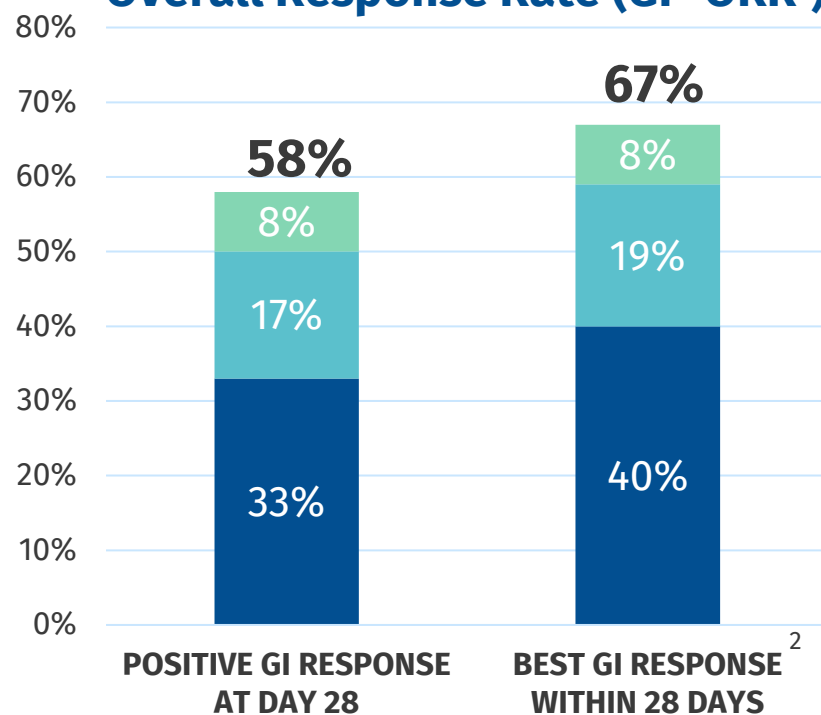




Early Access Program (EAP): Promising confirmation in an advanced, severe and more diverse GI aGvHD population

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

Gastro Intestinal Overall Response Rate (GI- ORR¹)

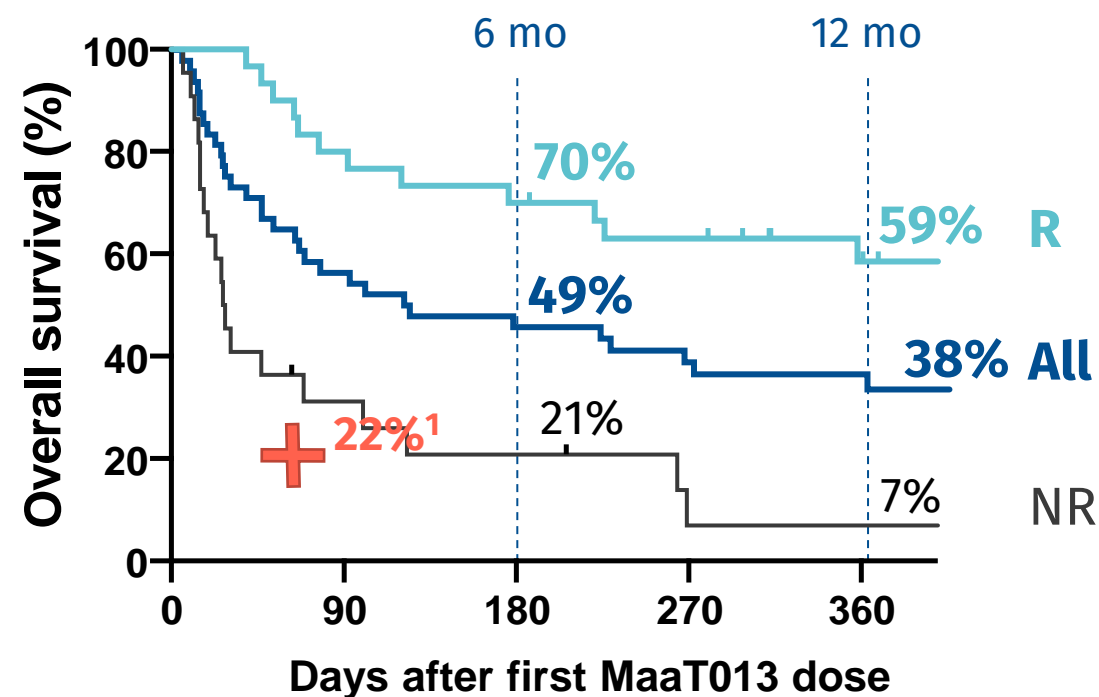


■ PR (partial response)
■ VGPR (very good partial response)
■ CR (complete response)

¹ORR= CR+VGPR+PR

² Best GI Response: Any response within 28 days

Overall Survival Rate Responders vs. Non responders



¹OS expected in ruxolitinib-resistant patients at 2 months (REACH1 study)



The ARES Phase 3 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



MaaT033: An optimized oral capsule to restore and maintain a healthy gut microbiome in patients with severe dysbiosis



Characteristics

Pooled microbiota: high-richness, high-diversity, full ecosystem, Microbiome Ecosystem Therapy containing Butycore™



Administration

Oral (a lyophilized capsule)



Clinical program

- ✓ CIMON Ph1b: Dose-finding study (completed)
- Planning OR-ALLO Phase 2/3 trial: Prevention of allo-HSCT complications



Indication

OS improvement through prevention of allo-HSCT complications



Phase Ib CIMON study aimed to determine MaaT033 dose for further clinical development



After induction therapy
with intensive chemotherapy

Cohort 5 (n= 6)

Not performed, due to sufficient data from Cohort 1-4

Cohort 4 (n= 6)

_____ / day, 2 weeks

Cohort 3 (n= 6)

_____ / day, 1 week

Cohort 2 (n= 6)

_____ / day, 1 week

Cohort 1 (n = 3)

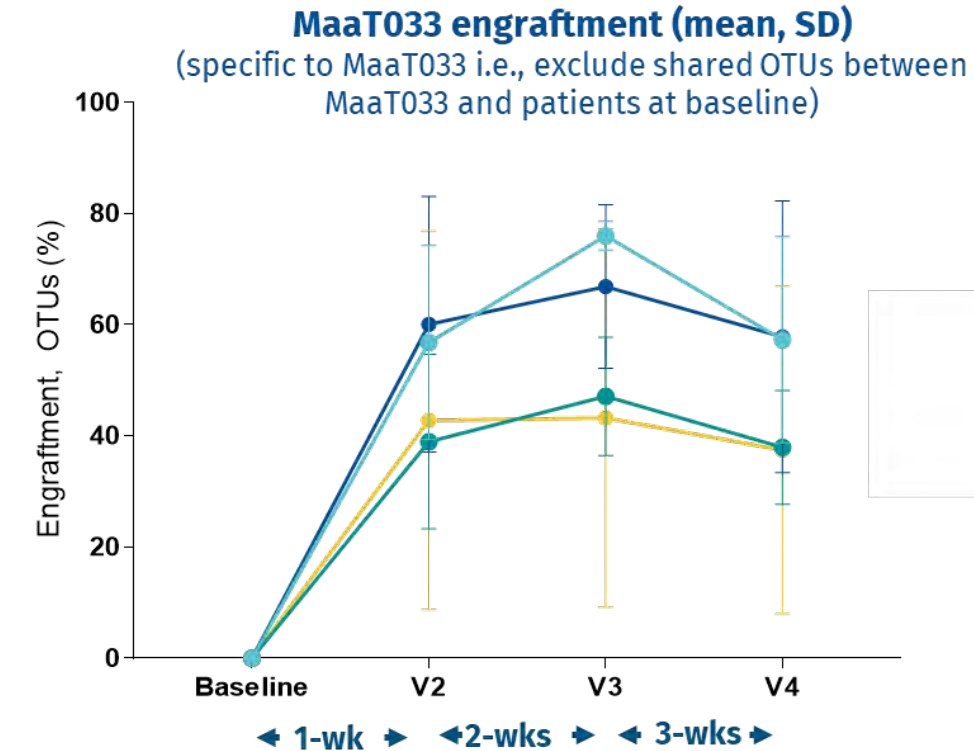
● ● / week, twice



Dose escalation



Phase Ib CIMON study : Positive topline engraftment and safety data



C1 (n=3): 1/w
C2 (n=6): 1/d
C3 (n=6): 3/d
C4 (n=5): 3/d

- First clinical POC of MaaT033 oral formulation
 - ✓ Robust and persistent engraftment
 - ✓ Good safety profile:
 - 21 patients exposed, 20 completed.
 - 100% drug compliance.
 - 4/4 positive DSMB meetings

→ **Dose selected for planned Phase II-III pivotal OR-ALLO study**
(342 patients, RCT, double-blind, placebo-controlled, evaluating overall survival after allo-HSCT)



CIMON results open an attractive market opportunity: Improving survival in patients receiving allo-HSCT



United States

c. 7,800
primary
procedures



EU 5

c. 9,600
primary
procedures



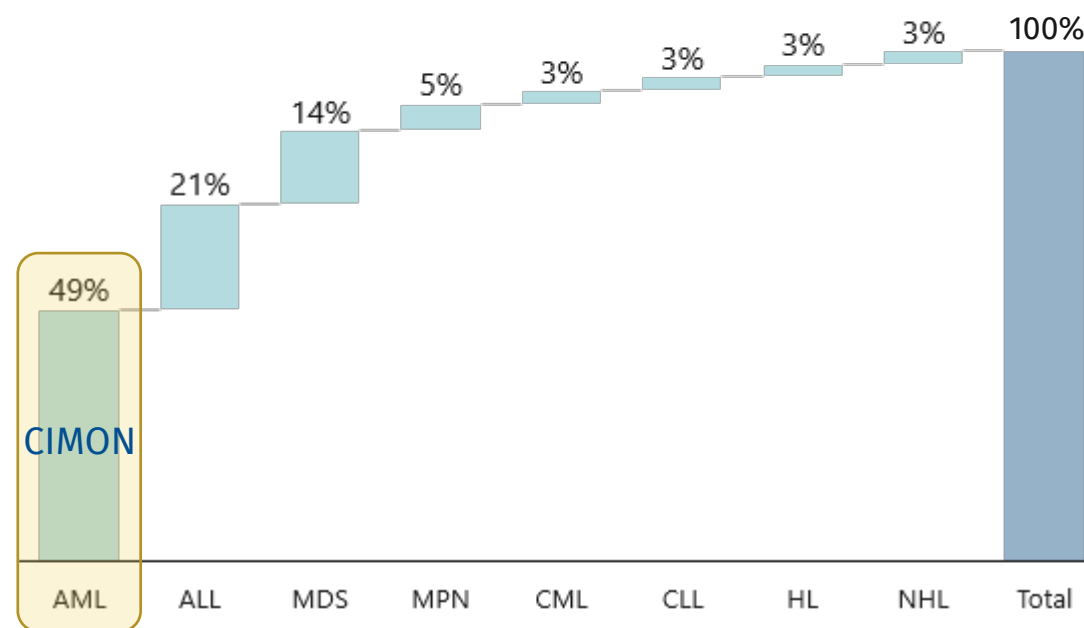
Japan

c. 3,000
primary
procedures

Additional 7%-10% recurrent procedures

Approximately 22,500 procedures/year

Hematological Malignancy Patients Receiving Allo-HSCT¹



AML : acute myeloid leukemia; ALL : acute lymphoblastic leukemia ; MDS : myelodysplastic syndrome; MPN : myeloproliferative neoplasms ; CML: chronic myeloid leukemia ; CLL : chronic lymphocytic leukemia ; HL: Hodgkin's Lymphoma ; NHL: Non Hodgkin Lymphoma



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Immuno-Oncology
Solid Tumors

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

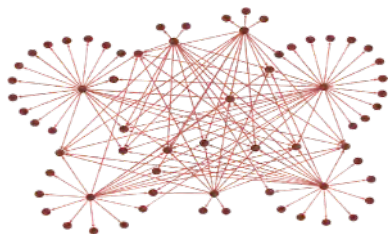
FMT from ICI responders to ICI non-responding patients with metastatic melanoma

✓ **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, Diversity and composition of gut microbiome drive survival and ICI toxicity in patients receiving ICI^{1,2,3,4}
- FMT from ICI responders (R) could induce response in metastatic melanoma non-responders (NR)^{5,6}

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

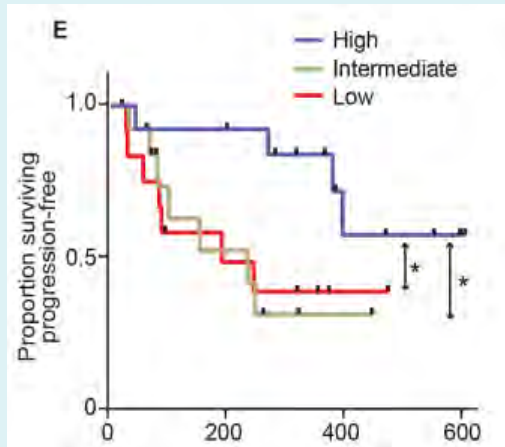
¹. Gopalakrishnan et al, Science 2018; ². Matson, et al Science 2018; ³. Routy et al, Science 2017;

⁴. Mc Culloch et al, Nat Med 2022; ⁵. Baruch et al, Science 2021; ⁶. Davar et al, Science 2021

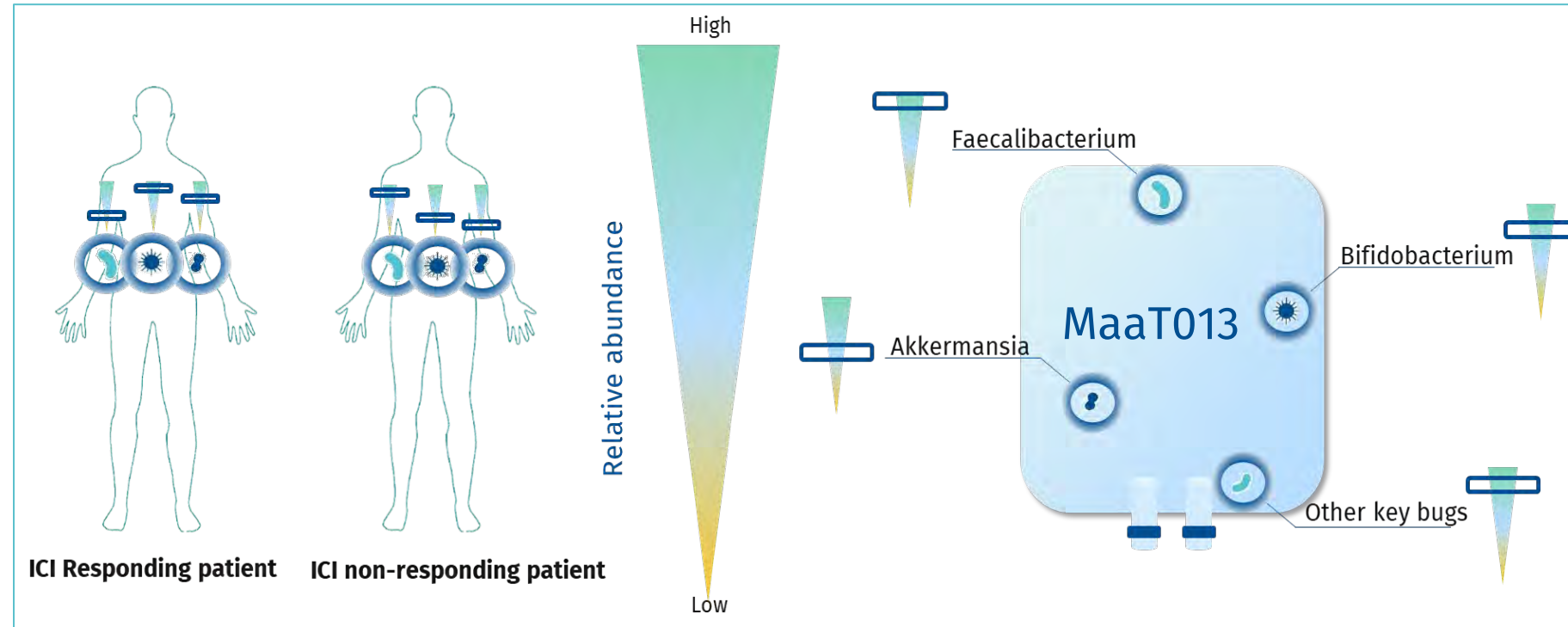


MaaT013 ensures high diversity and contains specific bacterial strains that have been identified to improve ICI response

Higher microbiome richness → better response rate to ICI in patients with metastatic melanoma



Gopalakrishnan et al, Science 2018



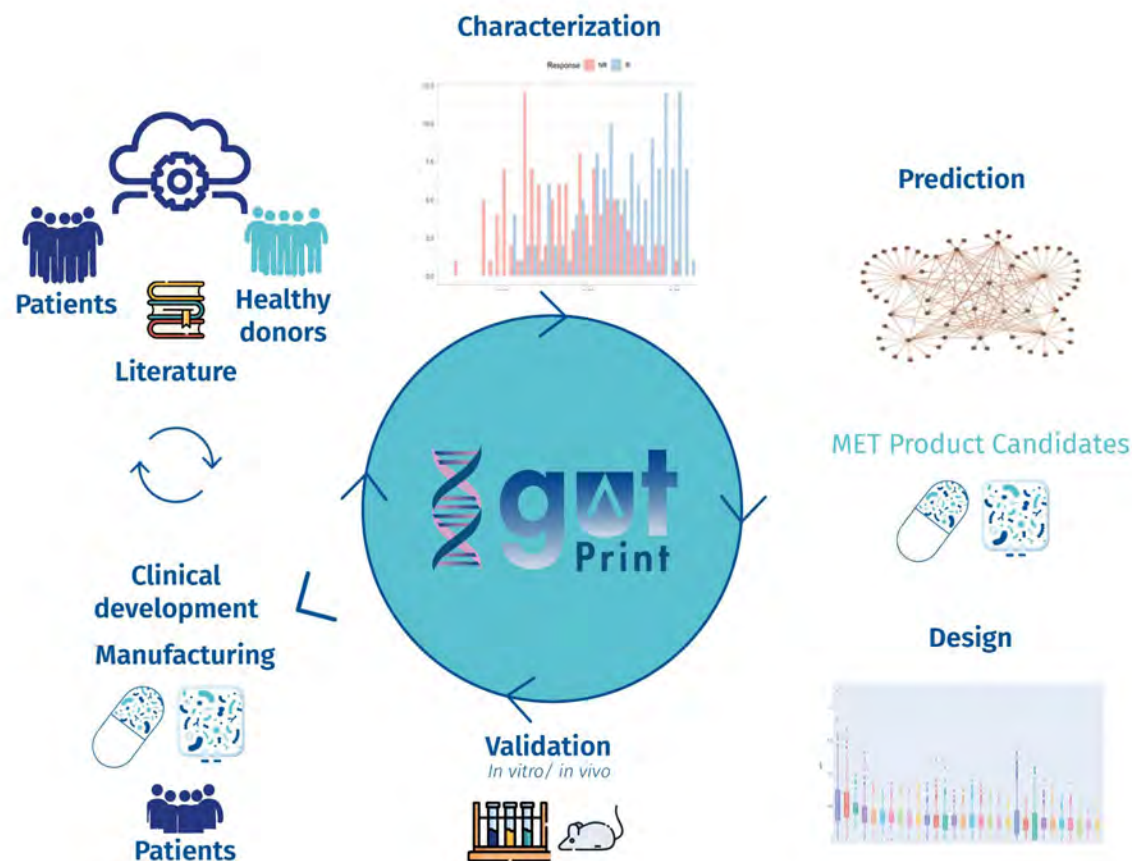
Ongoing Phase IIa PICASSO trial¹, in collaboration with **Assistance Publique - Hôpitaux de Paris** (sponsor).

- ✓ **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

¹ Registered trial #NCT04988841

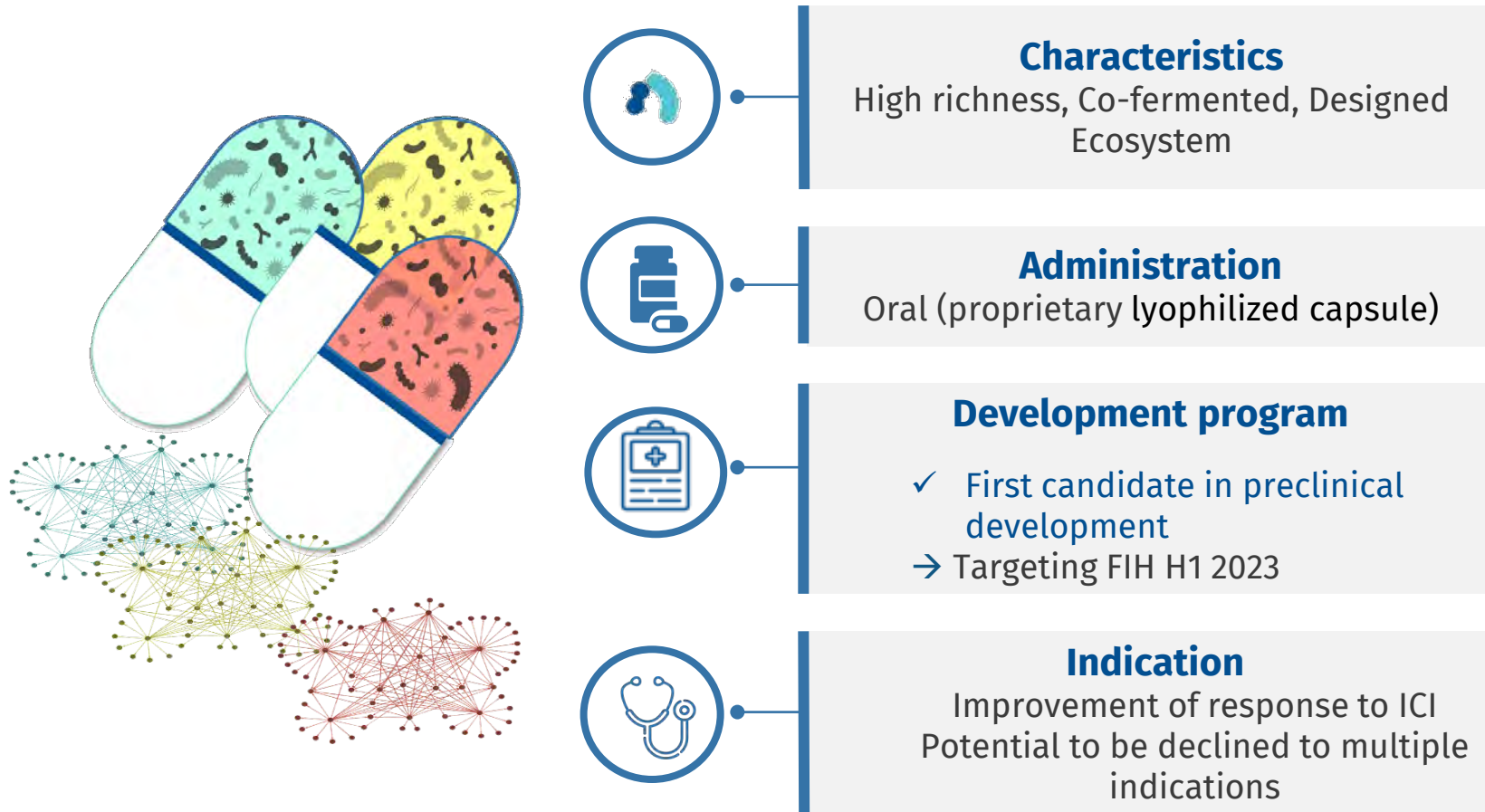


Proprietary gutPrint® platform synergizes multi-source data to generate innovative and indication-specific microbiome ecosystem therapies

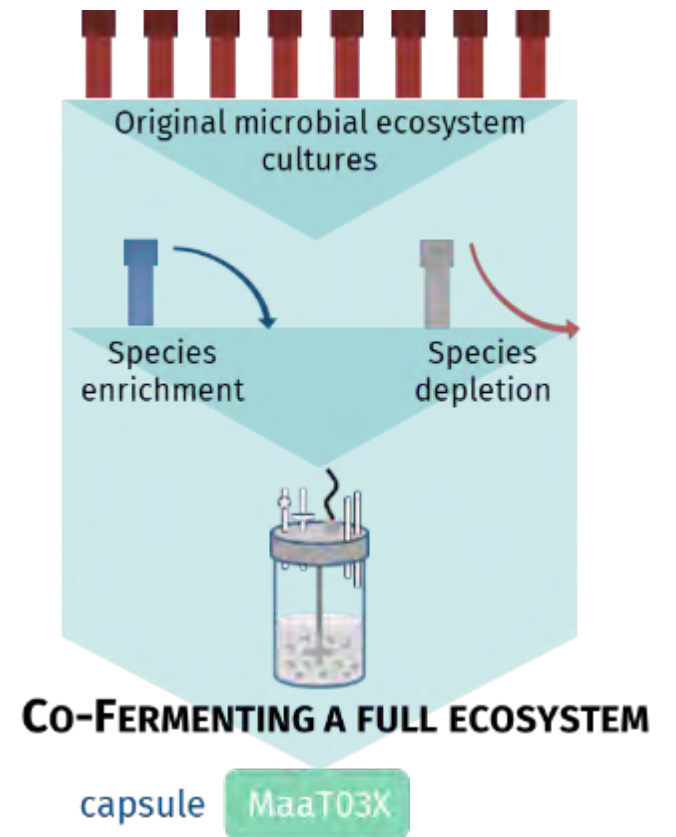


gutPrint® is the engine that drives MaaT Pharma's MET product candidate generation capabilities to broaden and strengthen the pipeline

MaaT03X: Modulate the gut microbiome to improve response to Immune Checkpoint Inhibitors treatment in solid tumors



Customizable, donor-independent, scalable process



Building Europe's largest specialized cGMP manufacturing facility for Microbiome Ecosystem Therapies



Partnership with  Skyepharma



Building a dedicated 1,500 square meter site (which could be doubled).



Designed to support commercial manufacturing of MaaT013 and MaaT033 and clinical manufacturing of MaaT03X products



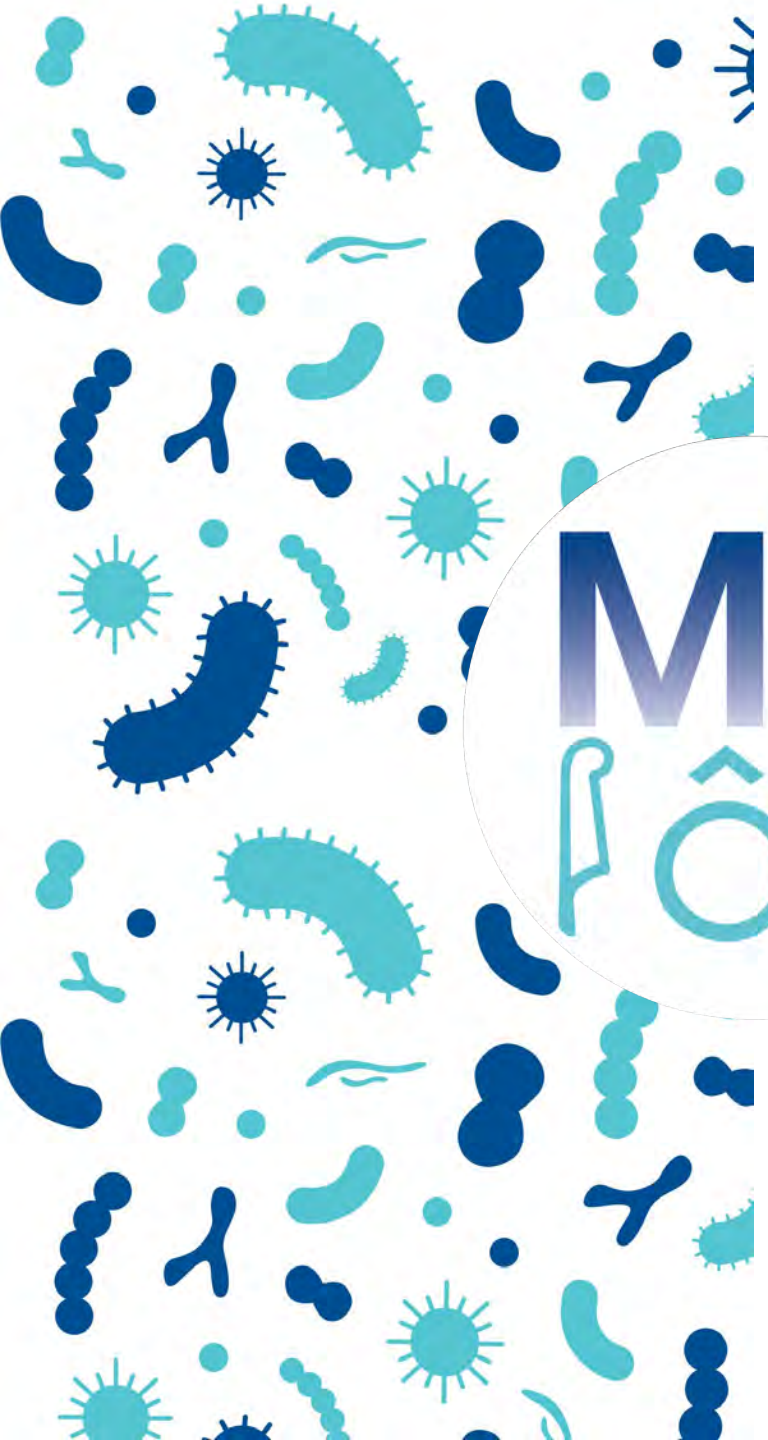
Skyepharma already manufactures approved drugs for the USA and Europe



Building will host manufacturing and R&D activities






Artist's representation of future plant



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Key Upcoming Milestones

Delivering on our objectives

	Clinical program	Milestones announced at IPO (Nov 2021)	Status
Onco-hematology	 MaaT013 (pooled enema) <i>FDA & EMA Orphan Drug Designation</i>	Launch of the first Phase 3 trial in oncology in the world	✓
	 MaaT033 (pooled capsule) <i>Post allo HSCT</i>	Completion of Phase 1b trial and positive preliminary safety and engraftment data	✓
Immuno-oncology	 MaaT013 (pooled enema) <i>Improving ICI responses in metastatic melanoma</i>	Launch of Phase 2 trial* - POC * Sponsored by AP-HP	✓
	 MaaT03X (fermented capsule) <i>Undisclosed indications</i>	Preclinical activities to enter clinical development in H1 2023	✓
cGMP production	 Increasing cGMP production capacities 	Partnership with Skyepharma to build the first and largest exclusive Microbiome Ecosystem Therapies facility in Europe	✓

Looking ahead

	Clinical program	Next Step	Expected timeline
Onco-hematology	 MaaT013 (pooled enema) <i>FDA & EMA Orphan Drug Designation</i>	Intermediate review	H1 2023
		ORR	H2 2023
	 MaaT033 (pooled capsule) <i>Post allo HSCT</i>	Launch of Phase 2/3 OR-ALLO (pivotal)	Q4 2022
Immuno-oncology	 MaaT013 (pooled enema) <i>Improving ICI responses in metastatic melanoma</i>	Interim partial data review	H1 2023
	 MaaT03X (fermented capsule) <i>Undisclosed indications</i>	Start of Phase 1/2	H1 2023
cGMP production	 Increasing cGMP production capacities 	Opening of the first and largest exclusive Microbiome Ecosystem Therapies facility in Europe	2023

* Sponsored by AP-HP
June 2022

Key differentiators of MaaT Pharma from other microbiome competitors

Leveraging the complexity of the microbiome

Pioneering a **full ecosystem approach** to restore host/microbiome **immune symbiosis**, based on proprietary **AI** and manufacturing capacities

Manufacturing versatility

cGMP manufacturing scalability for both native and co-fermented products and building of a new plant

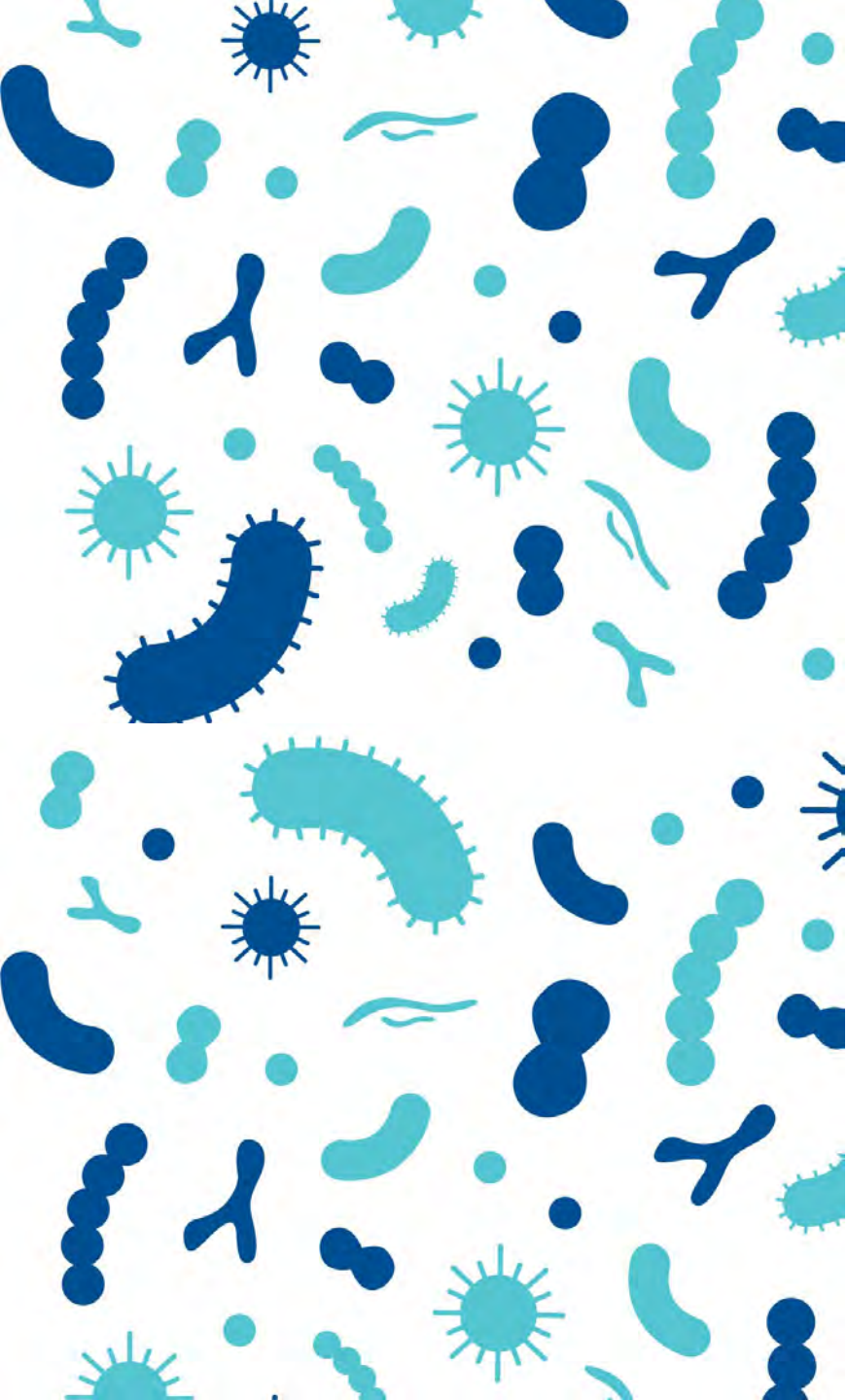
MaaT

Oncology focus

Addressing **high unmet needs** in the hemato-oncology and immuno-oncology therapeutic areas

Established proof of concept

First company to reach Phase 3 testing for a microbiome product in oncology globally



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