



SERES
THERAPEUTICS

Seres Therapeutics Investor Presentation
September 12, 2024



Disclaimers

Forward Looking Statements

This communication contains forward-looking statements within the meaning of the Private Securities Litigation Reform Act of 1995. All statements contained in this communication that do not relate to matters of historical fact should be considered forward-looking statements, including statements about the potential benefits of any of our products or product candidates; the ultimate safety and efficacy data of SER-155; study results; plans to seek FDA feedback; clinical data and clinical trials; our intentions related to the development of SER-155; our intention to seek Breakthrough Therapy Designation; the ability of live biotherapeutics to prevent or reduce infections; or the timing of any of the foregoing; the financial terms, timing and completion of the sale of VOWST assets to SPN; the receipt of future payments and the use of proceeds of the transaction; the timing and results of our clinical studies and data readouts; future product candidates, development plans and commercial opportunities; operating plans and our future cash runway; our ability to generate additional capital; our planned strategic focus; anticipated timing of any of the foregoing and other statements which are not historical fact.

These forward-looking statements are based on management's current expectations. These statements are neither promises nor guarantees, but involve known and unknown risks, uncertainties and other important factors that may cause our actual results, performance or achievements to be materially different from any future results, performance or achievements expressed or implied by the forward-looking statements, including, but not limited to, the following: (1) we have incurred significant losses, are not currently profitable and may never become profitable; (2) our need for additional funding; (3) our history of operating losses; (4) the restrictions in our debt agreement; (5) our novel approach to therapeutic intervention; (6) our reliance on third parties to conduct our clinical trials and manufacture our product candidates; (7) the competition we will face; (8) risks associated with our clinical trials; (9) whether the FDA grants Breakthrough Therapy Designation; (10) our ability to protect our intellectual property; (11) our ability to retain key personnel and to manage our growth; and (12) risks related to the proposed transaction under the Purchase Agreement with SPN for the sale of the VOWST business to SPN. These and other important factors discussed under the caption "Risk Factors" in our Quarterly Report on Form 10-Q filed with the SEC, on August 13, 2024, and our other reports filed with the SEC could cause actual results to differ materially from those indicated by the forward-looking statements made in this communication. Any such forward-looking statements represent management's estimates as of the date of this communication. While we may elect to update such forward-looking statements at some point in the future, we disclaim any obligation to do so, even if subsequent events cause our views to change. These forward-looking statements should not be relied upon as representing our views as of any date subsequent to the date of this communication.

Disclaimers

Important Information About the Transaction and Where to Find it

In connection with the proposed transaction involving Seres Therapeutics, Inc. (“Seres”) and Société des Produits Nestlé S.A. (“SPN”), Seres filed a definitive proxy statement with the Securities and Exchange Commission (the “SEC”). Seres may also file other relevant material with the SEC regarding the proposed transaction. Beginning on August 26, 2024, Seres mailed the definitive proxy statement to its stockholders. **INVESTORS AND STOCKHOLDERS OF SERES ARE URGED TO READ THE DEFINITIVE PROXY STATEMENT AND OTHER RELEVANT MATERIALS CAREFULLY AND IN THEIR ENTIRETY WHEN THEY BECOME AVAILABLE BECAUSE THEY CONTAIN OR WILL CONTAIN IMPORTANT INFORMATION ABOUT SERES AND THE PROPOSED TRANSACTION.** Investors may obtain a free copy of these materials (when they are available) and other documents filed by Seres with the SEC at the SEC’s website at www.sec.gov or from Seres at its website at ir.serestherapeutics.com.

Participants in the Solicitation

Seres and certain of its directors, executive officers and other members of management and employees may be deemed to be participants in soliciting proxies from its stockholders in connection with the proposed transaction. Information regarding the persons who may, under the rules of the SEC, be considered to be participants in the solicitation of Seres’ stockholders in connection with the proposed transaction is set forth in Seres’ definitive proxy statement for its stockholder meeting, which was filed with the SEC on August 26, 2024, at which the proposed transaction will be submitted for approval by Seres’ stockholders. You may also find additional information about Seres’ directors and executive officers in Seres’ Annual Report on Form 10-K for the fiscal year ended December 31, 2023, which was filed with the SEC on March 5, 2024, Seres’ Definitive Proxy Statement for its 2024 annual meeting of stockholders, which was filed with the SEC on March 5, 2024, and in subsequently filed Current Reports on Form 8-K and Quarterly Reports on Form 10-Q.

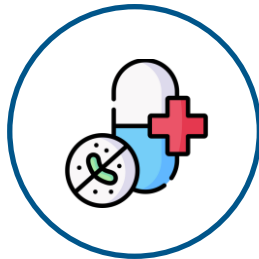
September 2024:

SER-155 Phase 1b placebo-controlled Cohort 2 study results in allo-HSCT

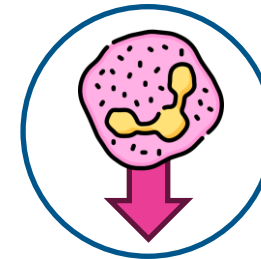
- Phase 1b study in adults undergoing allogeneic hematopoietic stem cell transplant (allo-HSCT) to assess SER-155 safety and pharmacology, with additional endpoints such as incidence of bacterial bloodstream infections and related medical consequences such as antibiotic use and febrile neutropenia
- Demonstrated generally well tolerated safety profile and confirmed drug bacteria strain engraftment
- SER-155 administration associated with following outcomes compared to placebo at day 100 post-HSCT:



Significant reduction
in bacterial
bloodstream infections



Significant reduction
in systemic antibiotic
exposure



Lower incidence of
febrile neutropenia

- Company to seek Breakthrough Therapy designation from the FDA given the high unmet medical need associated with bloodstream infections

Results support Seres' strategy to develop SER-155 and other live biotherapeutics to prevent serious bacterial infections in medically vulnerable patient populations

Transforming patient outcomes using proprietary consortia of live biotherapeutics

Strong foundation

- Validated platform with VOWST® clinical and regulatory success
- Asset sale strengthens balance sheet, expected to extend runway into Q4 '25
- Wholly-owned cultivated pipeline: SER-155, SER-147, beyond

Favorable Phase 1b clinical data in SER-155 allo-HSCT

- Well tolerated safety profile; no treatment-related SAEs
- Drug bacteria engraftment as expected
- Significant reduction in bloodstream infections and systemic antibacterial exposure
- Lower incidence of febrile neutropenia

Blockbuster opportunity

- Accelerate SER-155 development in allo-HSCT
- Potential to initiate multiple clinical studies in the next 12-18 months
- Potential to evaluate SER-155 in additional patient populations at high risk of serious bacterial infections (e.g., autologous HSCT, CAR-T, blood cancers, solid organ transplant)

Expansive potential

- SER-147 designed to prevent infections in chronic liver disease; anticipate IND readiness in H2 '25
- Current focus of preventing life-threatening infections
- Potential to treat immune-related diseases (including IBD)

Validated platform: Seres pioneered the development and FDA approval of VOWST as the first-ever oral live microbiome therapeutic



VOWST™
(fecal microbiota spores, live-brpk)

FDA approved (April 2023)
to prevent the recurrence of
C. difficile infection in adults

DRAMATIC CLINICAL BENEFIT –
Preventing infection recurrence

Approximately

88%

sustained clinical response rate
(*C. diff.* recurrence, at up to 8 weeks)

VOWST asset sale a transformational transaction for Seres



- **VOWST asset purchase agreement provides infusion of capital and supports pipeline development**
- **Asset sale expected to extend operational runway into Q4 2025**
- **Will retire debt and other obligations**

KEY FINANCIAL TERMS

\$100M upfront payment to Seres, less ~\$20M in net obligations due to an affiliate of SPN*

\$15M equity investment by SPN at closing

\$60M prepaid sales-based milestone at closing

\$75M in deferred payments due in 2025 (less ~\$1.5M in employment-related payments)

\$275M in potential future sales-based milestone payments (subject to reductions for interest on prepaid milestone payment)

Shareholder meeting scheduled September 26; transaction expected to close shortly thereafter subject to customary closing conditions

Potential to treat a range of vulnerable patient populations

Target population characteristics



GI microbiome functional disruption



Immune suppression



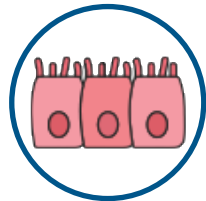
Antibiotic use



Neutropenia



Time in hospital/care settings



Lost epithelial or mucosal barrier integrity

Potential to prevent bacterial infections and immune-related disease

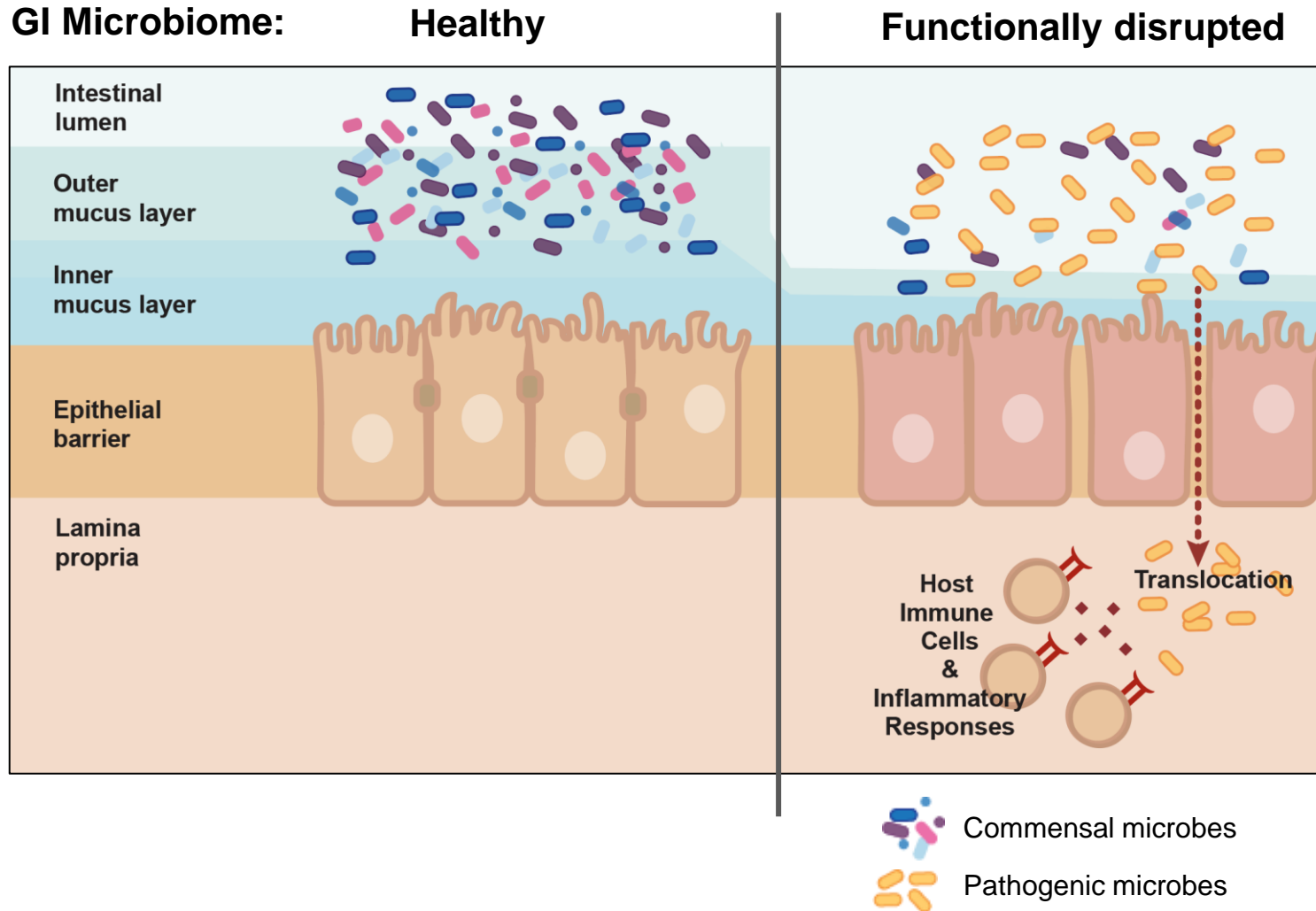
Prevent life-threatening infections (current focus)

- Blood cancers (including HSCT, CAR-T)
- Solid organ transplant
- ICU & long-term care patients
- Chronic liver disease

Treat immune-related diseases

- Inflammatory bowel disease
- Graft vs. host disease (GvHD)
- Checkpoint colitis
- Radiation enteritis

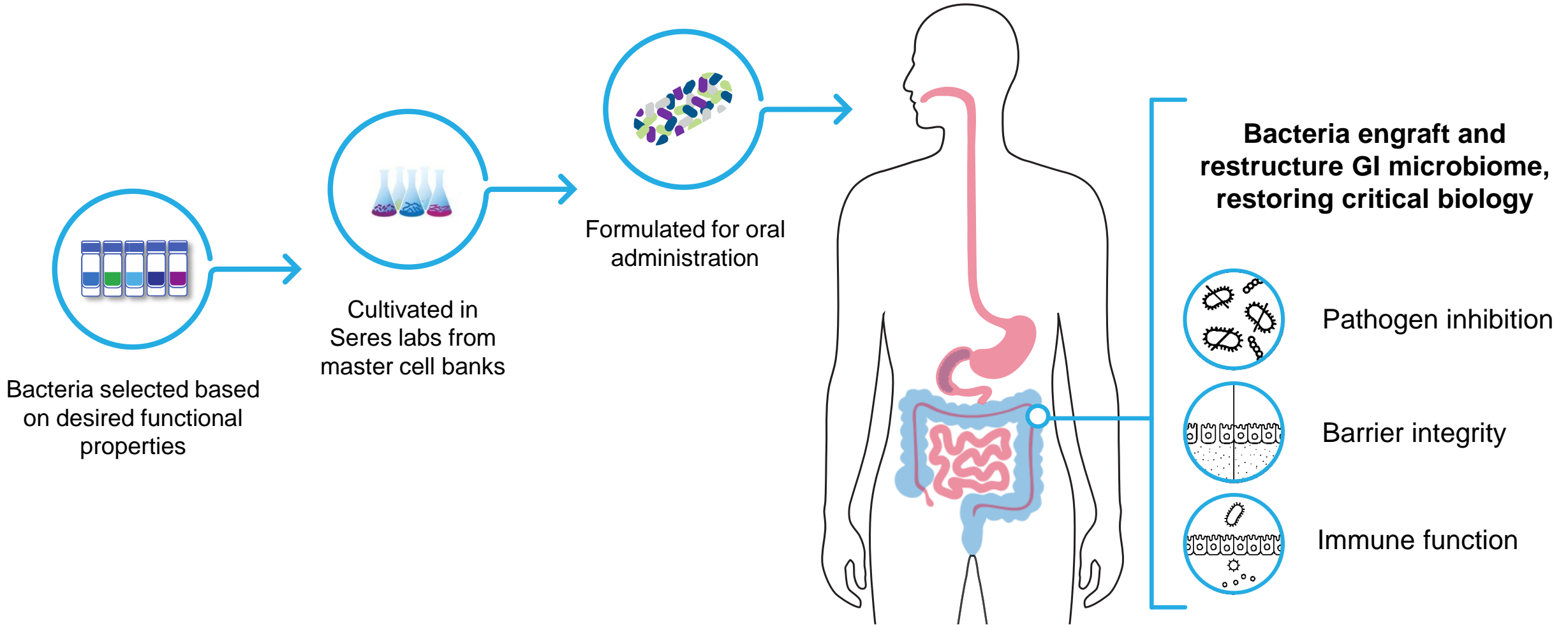
GI microbiome functional disruption leads to disease susceptibility



IMPACTED BIOLOGY

- Pathogen domination
- Lost barrier & mucosal integrity
- Unfavorable immune responses

Seres' biotherapeutics designed to restore functionality and health



Seres' biotherapeutics and pipeline candidates have well tolerated safety profile, reducing development risk

- ✓ Based on GI bacteria naturally **found in healthy humans**, and not associated with disease
- ✓ VOWST product profile includes **well tolerated safety** without drug-related serious adverse events
- ✓ **Well tolerated safety profile in multiple clinical trials** and patient populations, including medically vulnerable allo-HSCT recipients

Safety profile has potential to mitigate a primary cause of drug development failure

Near-term focus on SER-155 and SER-147 with potential to address expansive therapeutic opportunities



- Reduces risk of recurrent *C. diff* infections
- Well tolerated safety profile

Program	Lead Indication & Development Stage	Therapeutic Objectives	Potential Additional Indications
SER-155	<u>Allogeneic HSCT:</u> Phase 1b Cohort 2 (placebo controlled) data announced Sept. '24	Reduce incidence of serious bacterial infections (e.g., BSIs), febrile neutropenia, and GvHD	<ul style="list-style-type: none"> • Autologous HSCT • CAR-T • Blood cancers • Solid organ transplant
SER-147	<u>Chronic liver disease:</u> Anticipate IND ready in H2 '25	Reduce incidence of serious bacterial infections (e.g., SBP, BSIs) and related complications	<ul style="list-style-type: none"> • ICU patients • Long-term care patients

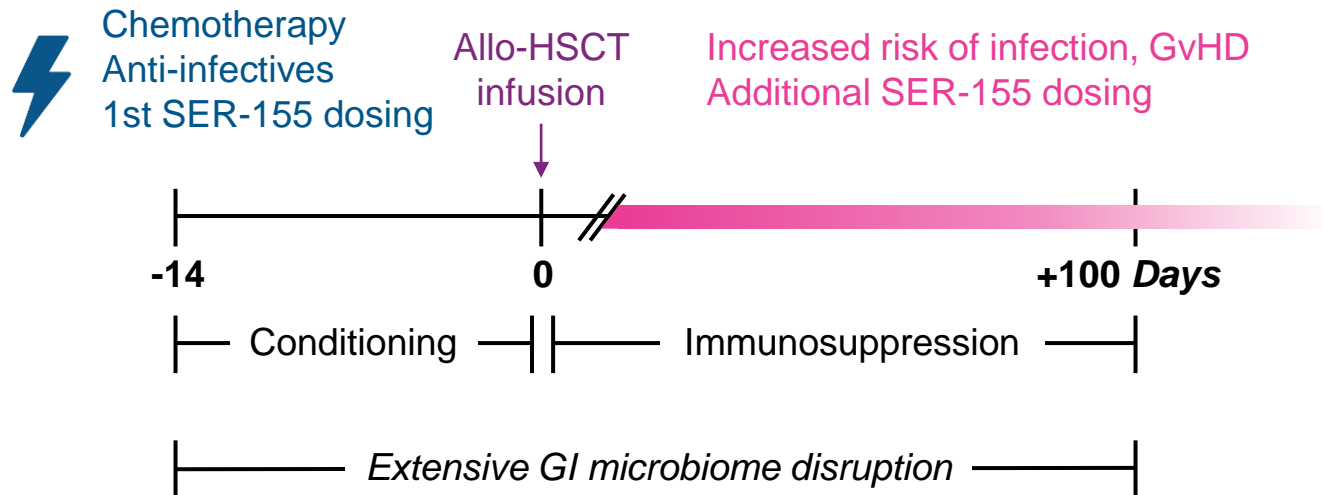
- Immediate therapeutic focus: prevent life-threatening infections
- Future: potential to treat immune-related diseases

SER-155 is designed to reduce life-threatening complications of allo-HSCT

SER-155

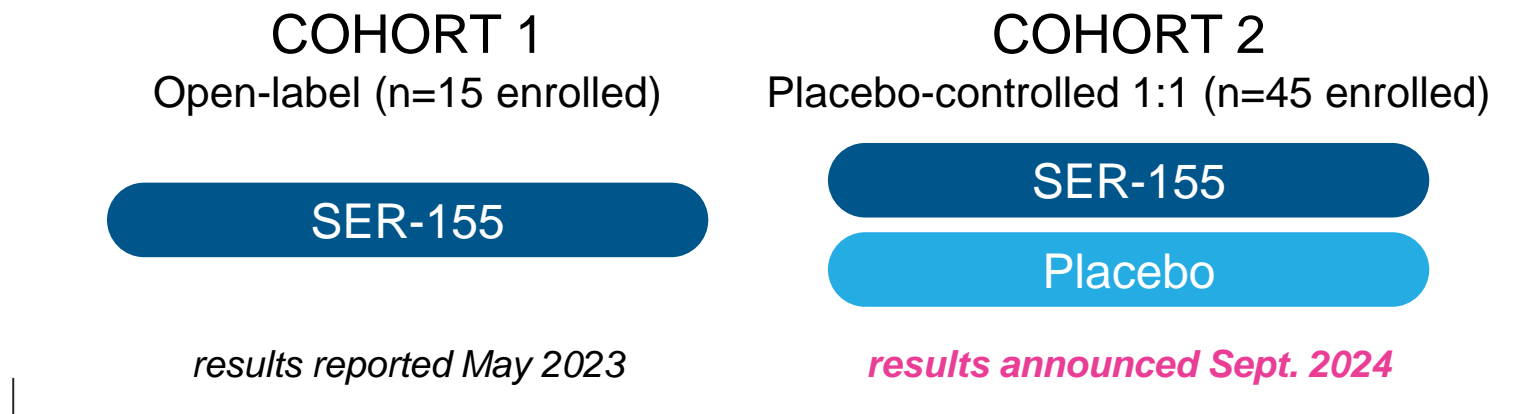
- Investigational live oral biotherapeutic cultivated from clonal master cell banks
- Designed to prevent GI-derived bacterial bloodstream infections (BSIs) and other pathogen-associated complications

Allo-HSCT treatment regimen and SER-155



- Only ~60% survival 3 years post-transplant
- ~10% transplant mortality in first 100 days post-transplant
- ~80% of adult deaths in first 100 days caused by complications of procedure; half of these due to infections and GvHD
- Complications have substantial impact: mortality, cost, hospital stay

SER-155 Phase 1b study evaluated safety, pharmacology, and efficacy in adult allo-HSCT recipients



Primary Endpoints:

- Safety and tolerability
- SER-155 bacterial strain engraftment

Key Secondary Endpoints through HSCT Day 100:

- Incidence of bloodstream infections (BSI), GI infections, and acute GvHD \geq Grade 2
- Incidence and duration of febrile neutropenia
- Bacterial pathogen abundance

**Received US FDA Fast Track Designation in December 2023;
Intend to pursue Breakthrough Therapy designation**

Patient Safety: Cohort 2

SER-155 was generally well tolerated with no treatment-related SAEs

Treatment-emergent adverse events (TEAEs)

- All but one subject in the placebo arm experienced at least 1 TEAE
- Most common for SER-155 treated subjects ($\geq 50\%$ and with $\Delta \geq 5\%$ greater than placebo): diarrhea (86% vs. 74% placebo), nausea (62% vs. 53% placebo)
- 1/40 (3%) subject experienced a TEAE leading to treatment discontinuation (active = 0; placebo = 1)
- 3/40 (8%) subjects experienced a TEAE leading to study discontinuation (active = 1; placebo = 2)

Serious adverse events (SAEs)

- 19/40 (48%) subjects experienced an SAE: 11/21 (52%) SER-155-treated subjects vs. 8/19 (42%) placebo-treated subjects; none considered related to SER-155 (no SUSARs)
 - Most common SAE SOC: infections & infestations (24% active vs. 37% placebo)
 - 3 deaths prior to Day 100 (active = 1; placebo = 2), 1 death after Day 100 (active), none considered related to SER-155

Adverse events of special interest (AESIs)

- AESIs (bloodstream infections, GI infection, invasive infection): 14/40 (35%) subjects
- Rates of AESIs were lower in SER-155 arm vs placebo arm (29% vs 42% respectively)
- No SER-155 species were identified in culture from any subject

Efficacy: SER-155 administration favorable with significant* reduction in both bacterial BSIs and systemic antibiotic exposure; lower febrile neutropenia

Secondary Efficacy	Bloodstream infections	Significant decrease in bacterial bloodstream infections in SER-155-treated subjects vs. placebo
	Febrile neutropenia	Numerically lower incidence rate of febrile neutropenia in SER-155-treated subjects vs. placebo
	GI infections	All GI infections were CDI** ; 4 subjects in SER-155-treated (20%) and 2 subjects in placebo (14.3%) developed GI infections from HSCT Day 0-100
	Acute GvHD	No subjects in either arm developed \geq Grade 3 acute GvHD; 2 subjects in each arm developed Grade 2 acute GvHD
Exploratory Efficacy	Antibacterial / antimycotic exposure	Significantly lower mean cumulative exposure (days) to systemic antibacterials / antimycotics for SER-155-treated subjects vs. placebo Significantly lower cumulative exposure rate to systemic antibacterials / antimycotics for SER-155-treated subjects vs. placebo

Bloodstream infections from HSCT Day 0 to Day 100: Lower incidence in SER-155 treated subjects vs. placebo

Bloodstream infections from Day 0 to Day 100 (# patients)	SER-155 n=20 n (%)	Placebo n=14 n (%)
Subjects with confirmed BSI	2 (10.0%)	6 (42.9%)
95% confidence interval	(1.2, 31.7)	(17.7, 71.1)

mITT-1 population

Odds ratio	0.15
95% confidence interval	(0.01, 1.13)
p-value	0.0423

Organisms in SER-155 patients: Finegoldia magna; E. coli/Strep mitis

Organisms in placebo patients: E.coli; Enterococcus faecium/staph haemolyticus/Candida krusei; Staph aureus; Staph haemolyticus; Pseudomonas aeruginosa; E coli

Cumulative exposure to systemic antibacterials / antimycotics through HSCT Day 100: Lower incidence in SER-155 treated subjects vs. placebo

Cumulative Antibacterial or Antimycotic Exposure (HSCT Days)	SER-155 n=20 n (%)	Placebo n=14 n (%)
Mean (SD)	9.2 (5.44)	21.1 (20.31)
Median	9.0	14.0
Min, Max	0, 19	0, 74

Mean Difference (95% CI)

-11.9 (-23.85, -0.04)

mITT-1 population

p-value

0.0494

- Cumulative exposure is the sum of all days a subject received systemic antibacterials and/or antimycotics between HSCT Day 0 through Day 100; counting once per day regardless of number of agents taken
- 95% confidence interval and p-value based on independent samples t-test of the difference in mean days between SER-155 and placebo

Cumulative exposure rate to systemic antibacterials / antimycotics through HSCT Day 100: Lower incidence in SER-155 treated subjects vs. placebo

Cumulative Antibacterial or Antimycotic Exposure Rate (% of days)	SER-155 n=20 n (%)	Placebo n=14 n (%)
Mean (SD)	0.090 (0.0530)	0.305 (0.2898)
Median	0.089	0.244
Min, Max	0.00, 0.18	0.00, 0.90

Mean Difference (95% CI)

-0.2 (-0.38, -0.05)

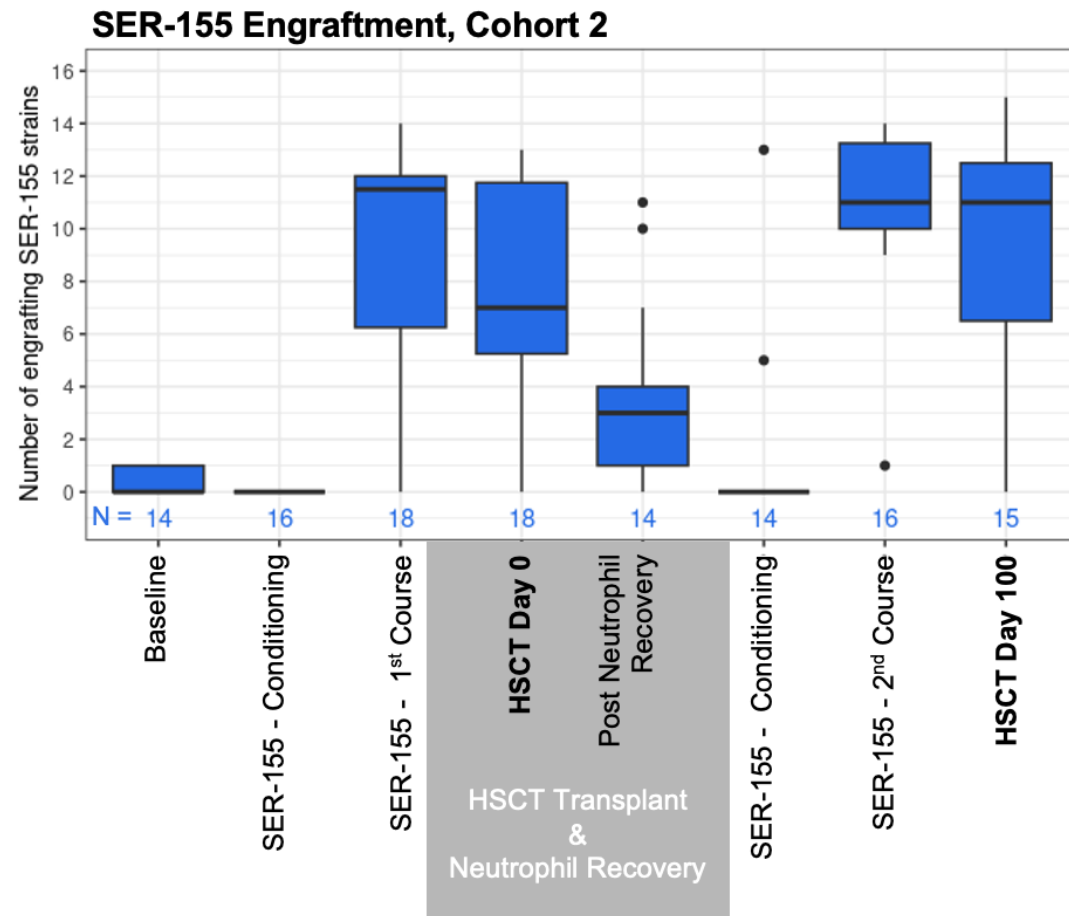
mITT-1 population

p-value

0.0163

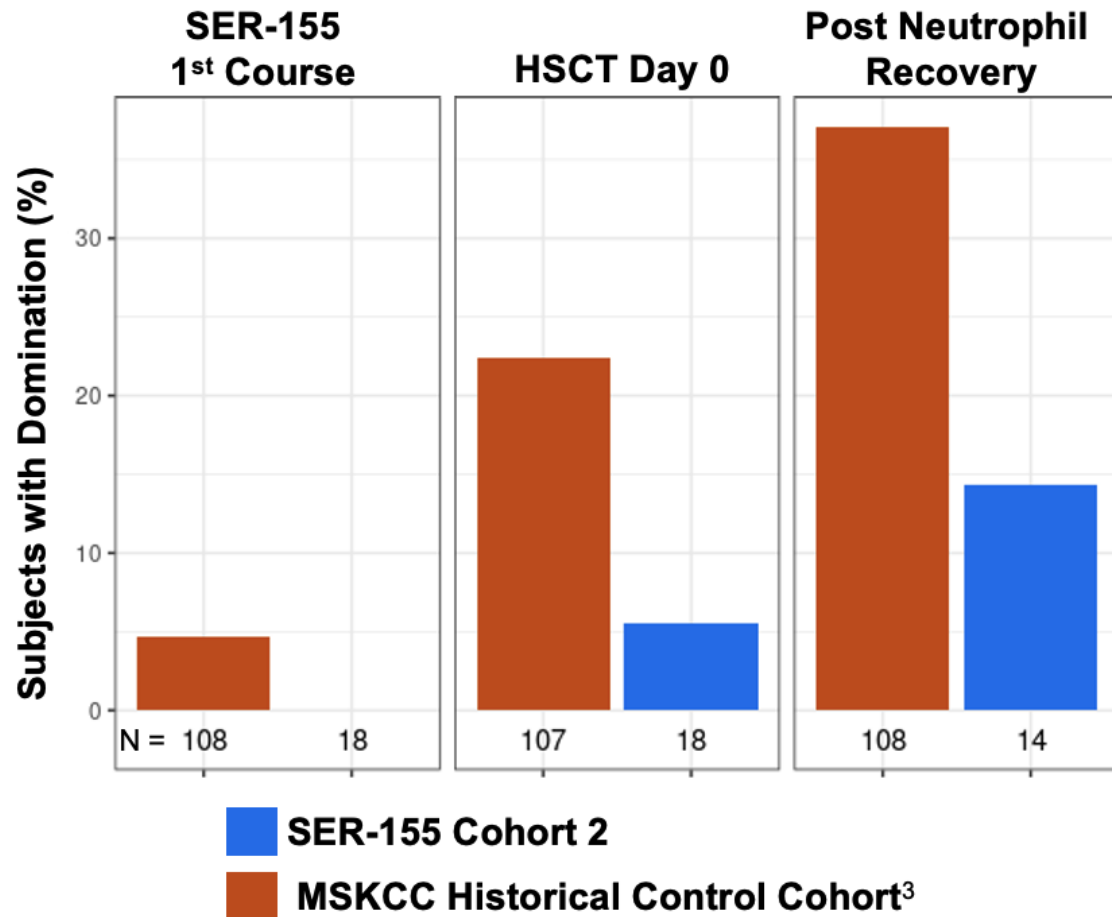
- Cumulative exposure rate is calculated as the sum of all days a subject received systemic antibacterials and/or antimycotics on or after HSCT Day 0 (counting once per day, regardless of number of antibacterial/antimycotic medications taken in a day) through HSCT Day 100 over the total number of days a subject was on the study from HSCT Day 0 to the earliest of EOS, or HSCT Day 100
- 95% confidence interval and p-value are based on independent samples t-test of the difference in mean days or mean rate of cumulative exposure between SER-155 and Placebo

SER-155 Strain Engraftment: Primary objective achieved - drug bacteria strain engraftment was robust and as expected



- The majority of SER-155 strains were present at start of HSCT conditioning and durable through chemotherapy exposure
- Engraftment decreased but was detectable post-neutrophil recovery, suggesting sustained engraftment, even under unfavorable GI conditions (e.g., antibiotic exposure), and through period of greatest BSI susceptibility
- The second course of SER-155 was effective at increasing strain engraftment following transplant & neutrophil recovery, with engraftment durable out to day 100 following transplant
- Cohort 1 and Cohort 2 engraftment magnitude and kinetics had high congruence

Pathogen Domination: Prevalence in SER-155 Cohort 2 was substantially lower relative to Historical Control Cohort



- SER-155 was designed to reduce pathogen domination that has been associated with risk of BSIs and other negative clinical outcomes¹
- In Cohort 2, the ability to detect pathogen domination² (i.e., relative abundance in the GI $\geq 30\%$) in the placebo arm, and differences between the study arms, was constrained due to the limited number of placebo stool samples and an imbalance in the number of available stool samples between the arms
- Observed pathogen domination events were low in the placebo and SER-155 arms with no significant differences observed
- Pathogen domination was substantially lower in SER-155 Cohort 2 compared to Historical Control Cohort³

1 - Peled et al, NEJM 2020; Stein-Thoeringer et al, Science 2019; Kusakabe et al, BBMT 2020

2 - Bacteria in the families: ESKAPE (Enterococcaceae, Enterobacteriaceae & Staphylococcaceae) & Streptococcaceae; domination defined as $\geq 30\%$ relative abundance in the GI

3 - Subjects that are sampled at similar time points as SER-155 Phase 1b subjects; microbiome data produced using same protocols as SER-155 Phase 1b subjects

Viral prophylaxis provides precedent in medically vulnerable patients

Prevymis - increasingly used for viral infection prophylaxis (e.g., allo-HSCT and solid organ transplant populations)



**\$605M '23
WW sales**

- Reduces CMV infection in allo-HSCT recipients
- Lowers mortality rate

- Overall cost of allo-HSCT is high (~\$400K US year 1 allo-HSCT costs)
- Transplant-related complications (e.g., infections) raise cost by ~\$180K
- Infections result in longer hospital stays, readmissions, increased ICU utilization

SER-155 allo-HSCT commercial opportunity is meaningful

- ✓ Serious bacterial infections are frequent, creating a strong medical rationale for prophylaxis to prevent infections and complications
- ✓ ~40K transplants per year worldwide
- ✓ Very high overall cost of allo-HSCT and related cost of complications supports financial rationale to treat
- ✓ Well-defined treatment centers could rapidly adopt as a new standard of care

Accelerating SER-155 clinical development with positive Ph1b outcomes

Aim to **accelerate SER-155 development** in allo-HSCT

- Potential to follow successful precedent from VOWST development

Engage with FDA on advancement of SER-155 allo-HSCT program

- Seek Breakthrough Therapy designation
- Pursue additional designations (Orphan Drug, Qualified Infectious Disease Product)

Evaluate SER-155 in **additional patient populations** with high risk of serious bacterial infections

Potential global SER-155 development and commercialization in broad range of indications in medically vulnerable populations

Anticipated SER-155 expansion in biologically adjacent populations

Population	Transplants / diagnoses per year (US + EU)
Autologous HSCT	~30K
Blood cancers with high neutropenia rates (acute myeloid leukemia, multiple myeloma, B cell non-Hodgkin's lymphomas)	~190K
Solid organ transplant (liver, kidney)	~65K

Potential to initiate multiple clinical studies within the next 12-18 months

SER-147 in development to prevent infections in chronic liver disease patients

Substantial unmet need

0.5M  2.1M 

~50% experience bacterial infections in a 6 month period

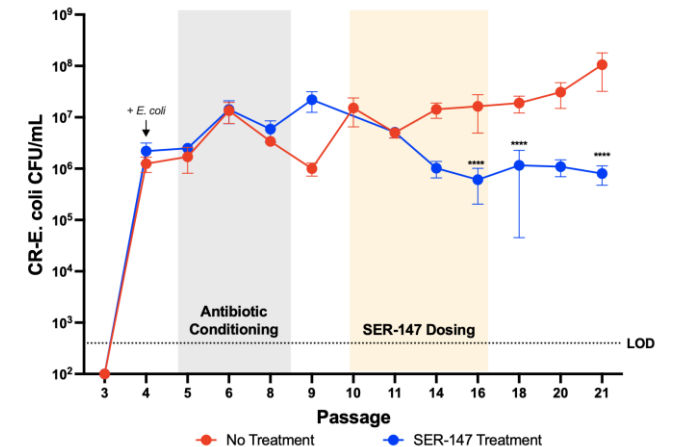
~20-25% of infections are spontaneous bacterial peritonitis and bloodstream infections likely to be gut-seeded

Promising preclinical data

SER-147 is an investigational live oral biotherapeutic designed to reduce pathogens causing gut-seeded SBP and BSIs in liver disease patients

Example: 1-3 log reduction of *E. coli* in *in vivo* models, plus reduction of other pathogens

Declining *E. coli* titers



End-to-end capabilities & expertise for discovery and development of bacterial live biotherapeutics

GI microbiome biomarker & drug target identification

Lead candidate design, screening, optimization, & drug pharmacology

Novel biotherapeutic GMP manufacturing & quality

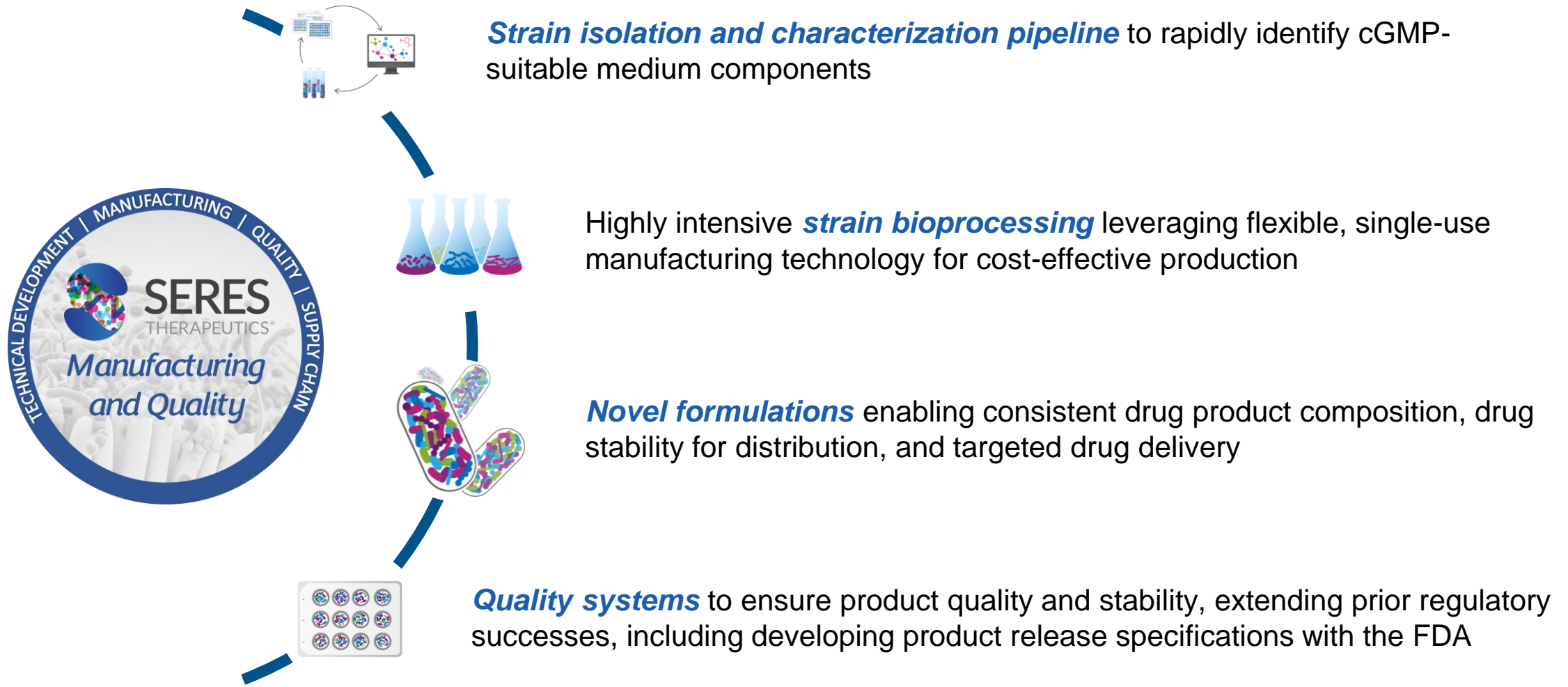


Clinical translation & patient subpopulation insights

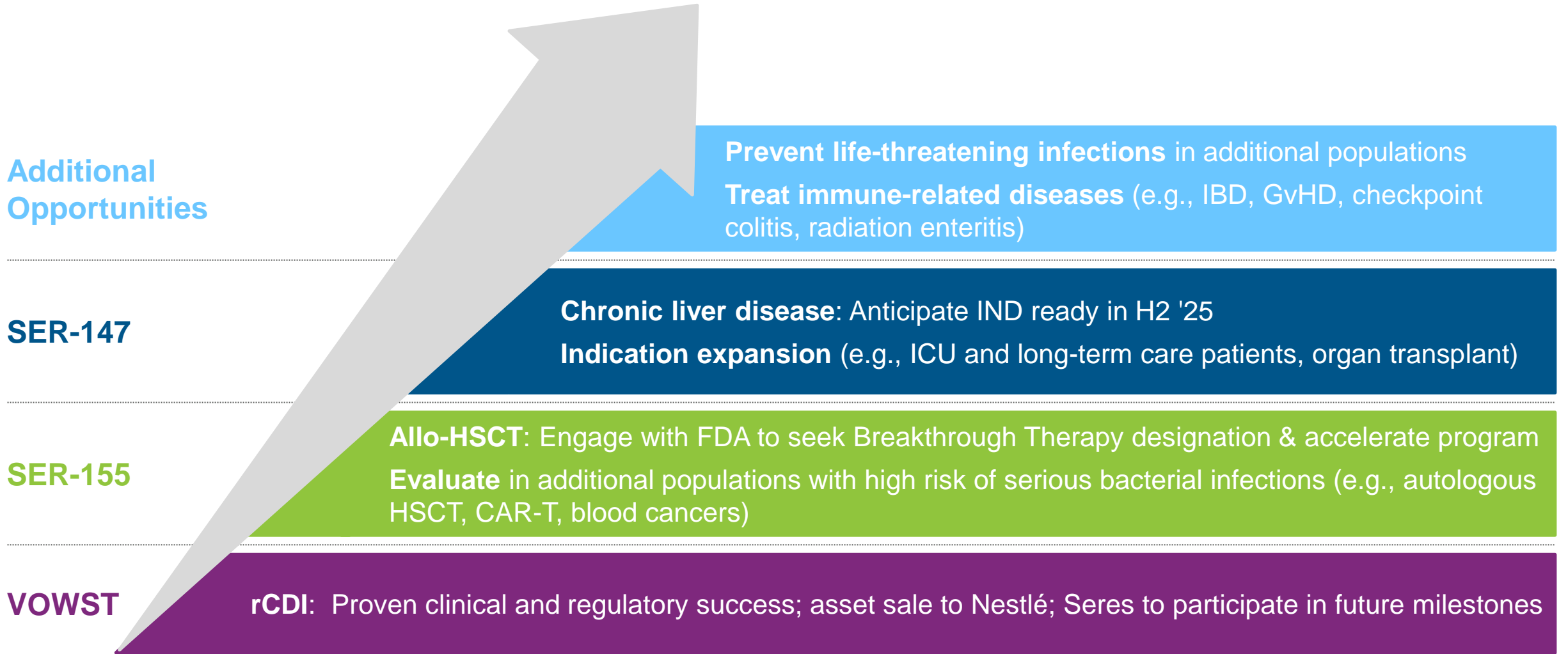
Proprietary know-how on clinical trial design and execution

Regulatory expertise pioneering a novel biotherapeutic class

Manufacturing platform delivers defined consortia in oral formulation using cost-effective production



Maximizing opportunity going forward



Summary and path forward

Developing a pipeline of novel live biotherapeutics in areas of high unmet need

- Successful VOWST development validates using live biotherapeutics to prevent life-threatening infections
- SER-155 Phase 1b placebo-controlled clinical efficacy data further support Seres' strategy
- Pipeline aims to bring transformative medicines to a wider set of patients, led by SER-155 with SER-147 anticipated to be IND ready in H2 '25

SER-155 Phase 1b placebo-controlled clinical results promising

- SER-155 demonstrated generally well tolerated safety profile and confirmed drug bacteria strain engraftment
- SER-155 administration associated with significant reduction in both bacterial bloodstream infections and systemic antibiotic exposure, as well as lower incidence of febrile neutropenia, as compared to placebo through day 100 post HSCT

VOWST asset sale strengthens financial position

- VOWST asset sale expected to close shortly after September 26 shareholder meeting, with \$175M due at closing less an ~\$20M settlement of net obligations, and \$75M (less ~\$1.5M in employment-related payments) in installment payments due in 2025 + \$275M potential future milestones
- \$71.2M in cash at end Q2 2024; asset sale expected to extend cash runway into Q4 2025
- 151.5M shares of MCRB outstanding as of May 6, 2024; additional ~14M issued at closing