UNITED STATES
SECURITIES AND EXCHANGE COMMISSION
WASHINGTON, D.C. 20549

FORM 8-K

CURRENT REPORT
Pursuant to Section 13 or 15(d)
of the Securities Exchange Act of 1934

Date of Report (Date of earliest event reported): September 12, 2024

SERES THERAPEUTICS, INC.
(Exact name of Registrant as Specified in Its Charter)

Delaware	**001-37465**	**27-4326290**
(State or other jurisdiction of incorporation)	**(Commission File Number)**	**(IRS Employer Identification No.)**

101 Cambridgepark Drive **Cambridge, MA**	**02140**
(Address of principal executive offices)	**(Zip Code)**

Registrant's telephone number, including area code: (617) 945-9626

Not Applicable
(Former Name or Former Address, if Changed Since Last Report)

Check the appropriate box below if the Form 8-K filing is intended to simultaneously satisfy the filing obligation of the registrant under any of the following provisions:

☐ Written communications pursuant to Rule 425 under the Securities Act (17 CFR 230.425)

☐ Soliciting material pursuant to Rule 14a-12 under the Exchange Act (17 CFR 240.14a-12)

☐ Pre-commencement communications pursuant to Rule 14d-2(b) under the Exchange Act (17 CFR 240.14d-2(b))

☐ Pre-commencement communications pursuant to Rule 13e-4(c) under the Exchange Act (17 CFR 240.13e-4(c))

Securities registered pursuant to Section 12(b) of the Act:

Title of each class	Trading Symbol(s)	Name of each exchange on which registered
Common stock, par value $0.001 per share	MCRB	The Nasdaq Stock Market LLC (Nasdaq Global Select Market)

Indicate by check mark whether the registrant is an emerging growth company as defined in Rule 405 of the Securities Act of 1933 (§ 230.405 of this chapter) or Rule 12b-2 of the Securities Exchange Act of 1934 (§ 240.12b-2 of this chapter).

Emerging growth company ☐

If an emerging growth company, indicate by check mark if the registrant has elected not to use the extended transition period for complying with any new or revised financial accounting standards provided pursuant to Section 13(a) of the Exchange Act. ☐

Item 7.01. Regulation FD Disclosure.

On September 12, 2024, Seres Therapeutics, Inc. (the "Company") posted a slide presentation on the SER-155 Phase 1b Cohort 2 Study results in the "Investors and News" portion of its website at www.serestherapeutics.com. The Company also issued a press release in connection with the foregoing. A copy of the slide presentation and press release are attached as Exhibit 99.1 and Exhibit 99.2, respectively, to this Current Report on Form 8-K (this "Current Report") and incorporated herein by reference.

On September 12, 2024, the Company posted an updated corporate presentation in the "Investors and News" portion of its website at www.serestherapeutics.com. A copy of the slide presentation is attached as Exhibit 99.3 to this Current Report and incorporated herein by reference.

The information in Item 7.01 of this Current Report, including Exhibit 99.1, Exhibit 99.2 and Exhibit 99.3 attached hereto, is intended to be furnished and shall not be deemed "filed" for purposes of Section 18 of the Securities Exchange Act of 1934, as amended (the "Exchange Act"), or otherwise subject to the liabilities of that section, nor shall it be deemed incorporated by reference in any filing under the Securities Act of 1933, as amended, or the Exchange Act, except as expressly set forth by specific reference in such filing. The Company undertakes no obligation to update, supplement or amend the materials attached hereto as Exhibit 99.1, Exhibit 99.2 and Exhibit 99.3.

Item 8.01. Other Events.

On September 12, 2024, the Company announced topline clinical data from Cohort 2 of its SER-155 Phase 1b placebo-controlled study in patients undergoing allogeneic hematopoietic stem cell transplantation (allo-HSCT). Study results demonstrate that SER-155 was associated with a significant reduction in both bloodstream infections (BSIs) and systemic antibiotic exposure, as well as a lower incidence of febrile neutropenia, as compared to placebo through day 100 post HSCT.

The Company believes that the SER-155 Phase 1 study results support the Company's corporate strategy to develop its platform, comprised of a pipeline of designed live biotherapeutics, in multiple medically vulnerable patient populations at high risk of life-threatening bacterial infections and associated negative clinical outcomes. The Company intends to seek Breakthrough Therapy designation, given the high unmet medical need associated with BSIs, and discuss advancing development of SER-155 for allo-HSCT with the U.S. Food and Drug Administration (FDA). The Company also intends to evaluate SER-155 in additional patient populations that have a high risk of serious bacterial infections.

SER-155 Phase 1b Study Design

The SER-155 Phase 1b study (NCT04995653) included two cohorts. Cohort 1 was designed to assess safety and drug pharmacology, specifically the drug strain engraftment in the gastrointestinal tract. Cohort 1 included 13 subjects who received any dosing of the SER-155 regimen, with 11 subjects subsequently receiving an allo-HSCT. Results from this cohort, announced in May 2023, showed SER-155 was generally well tolerated and resulted in successful drug strain engraftment and a reduction in pathogen domination in the GI microbiome relative to a historical control cohort.

Study Cohort 2 utilized a randomized, double-blinded 1:1 placebo-controlled design to further evaluate safety and drug strain engraftment, as well as key secondary and exploratory endpoints such as the incidence of bacterial bloodstream infections and related medical consequences such as febrile neutropenia and antibiotic use. Cohort 2 included 45 patients in the intention-to-treat (ITT) population. Of the ITT population, 20 received SER-155 and 14 received placebo, each of whom subsequently received an allo-HSCT, with data available for clinical evaluation through day 100, the study's prespecified primary observation point. Exploratory hypothesis testing was conducted at the two-sided α=0.05 level. Ninety-five percent (95%) 2-sided confidence intervals (CIs) were determined, where specified. No adjustment for multiplicity was done. A subset of patient samples was available for drug pharmacology analysis.

The median age in Cohort 2 was 63, and most subjects had acute myeloid leukemia, acute lymphocytic leukemia, myelodysplastic syndrome or myeloproliferative neoplasia as their primary disease and received reduced-intensity conditioning pre-transplant. Most patients received peripheral blood stem cells from a matched unrelated donor. A majority received post-transplant cyclophosphamide as part of their graft-versus-host disease (GvHD) prophylaxis.

Summary of Cohort 2 Study Results

Consistent with the observations from the Phase 1b study Cohort 1, SER-155 was generally well-tolerated, and no treatment-emergent serious adverse events related to drug were observed. SER-155 bacterial strains engrafted into the gastrointestinal tract of patients following the administration of SER-155.

The incidence of BSIs was significantly lower in the SER-155 arm compared with the placebo arm (2/20 (10%) vs. 6/14 (42.9%), respectively; [Odds Ratio: 0.15; 95% CI: 0.01, 1.13, p=0.0423]). In addition, while antibiotic starts were similar in each arm, patients administered SER-155 were treated with antibiotics for a significantly shorter duration compared to patients in the placebo arm (9.2 days vs. 21.1 days, respectively, with a mean difference of -11.9 days [95% CI: -23.85, -0.04; p=0.0494]). The incidence of febrile neutropenia was lower in patients administered SER-155 compared to placebo (65% vs. 78.6%, respectively; [Odds Ratio: 0.51; 95% CI: 0.07, 2.99; p=0.4674]). Six cases of gastrointestinal infections (*C. difficile* infections) were observed in the study, with four cases (20%) in the SER-155 arm and two cases (14.3%) in the placebo arm.

Recent changes in the allo-HSCT standard of care and the increasing use of post-transplant cyclophosphamide as part of prophylactic therapy for GvHD have reduced rates of GvHD overall in this patient population. The rates of GvHD in the study were low, with two cases of grade 2 GvHD observed in each arm, and no cases of grade 3 or 4 GvHD were observed.

In Cohort 2, the ability to detect pathogen domination (i.e., relative abundance in the GI ≥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 identified. In a comparison of the prevalence of pathogen domination versus a larger allo-HSCT historical control cohort, pathogen domination in SER-155 subjects was substantially lower, providing further evidence of SER-155 activity.

The overall safety results in the first 100 days showed that SER-155 was generally well-tolerated, with no observed treatment-related serious adverse events. In Cohort 2, all but one subject in the placebo arm experienced at least one treatment-emergent adverse event ("TEAE"). The most common TEAEs were diarrhea and nausea. Four subjects experienced TEAEs that led to discontinuation. Nineteen subjects, or 48% of subjects, experienced serious adverse events, none of which were considered related to SER-155. Three deaths were reported in the study, none of which were considered related to SER-155. Fourteen subjects, or 35% of the subjects, experienced adverse events of special interest, including bloodstream infections, GI infection and invasive infection.

Seres fully owns worldwide rights for the commercialization of SER-155.

Item 9.01. Financial Statements and Exhibits.

(d) Exhibits.

The following Exhibits 99.1 through 99.3 relate to Item 7.01 and shall be deemed to be furnished, and not filed:

Exhibit No.	Description
99.1	Seres Therapeutics, Inc. SER-155 Phase 1b Cohort 2 Study Results Slide Presentation as of September 2024
99.2	Seres Therapeutics, Inc. SER-155 Phase 1b Cohort 2 Study Results Press Release issued September 12, 2024
99.3	Seres Therapeutics, Inc. Corporate Presentation as of September 2024
104	Cover Page Interactive Data File - the cover page XBRL tags are embedded within the Inline XBRL document.

Important Additional 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.

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.

SIGNATURES

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned hereunto duly authorized.

Date: September 12, 2024

SERES THERAPEUTICS, INC.

By: /s/ Thomas J. DesRosier
Name: Thomas J. DesRosier
Title: Chief Legal Officer and Executive Vice President

Exhibit 99.1

SER-155 Phase 1b Readout

September 12, 2024

Disclaimers

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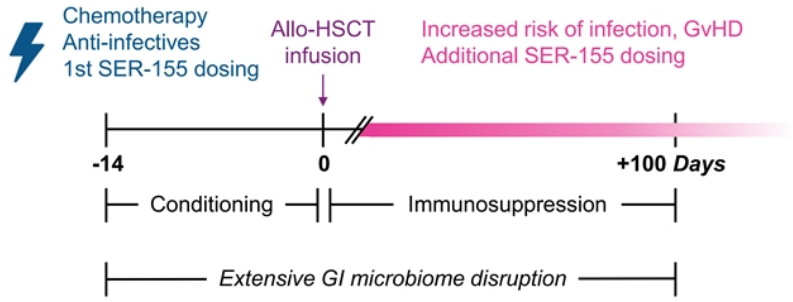
SER-155 is designed to reduce life-threatening complications of allo-HSCT

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

Sources: CIBMTR 2023 US summary slides; Penack et al, *Blood Adv* 2020; Khan et al, *Blood* 2021; Peled et al, *NEJM* 2020; Stein-Thoeringer et al, *Science* 2019; Bleakley & Riddell, *Nat Rev Cancer* 2004

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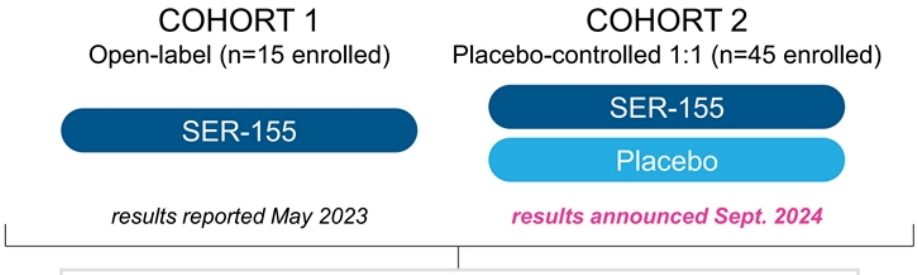
SER-155 Phase 1b study evaluated safety, pharmacology, and efficacy in adult allo-HSCT recipients



Primary Endpoints:

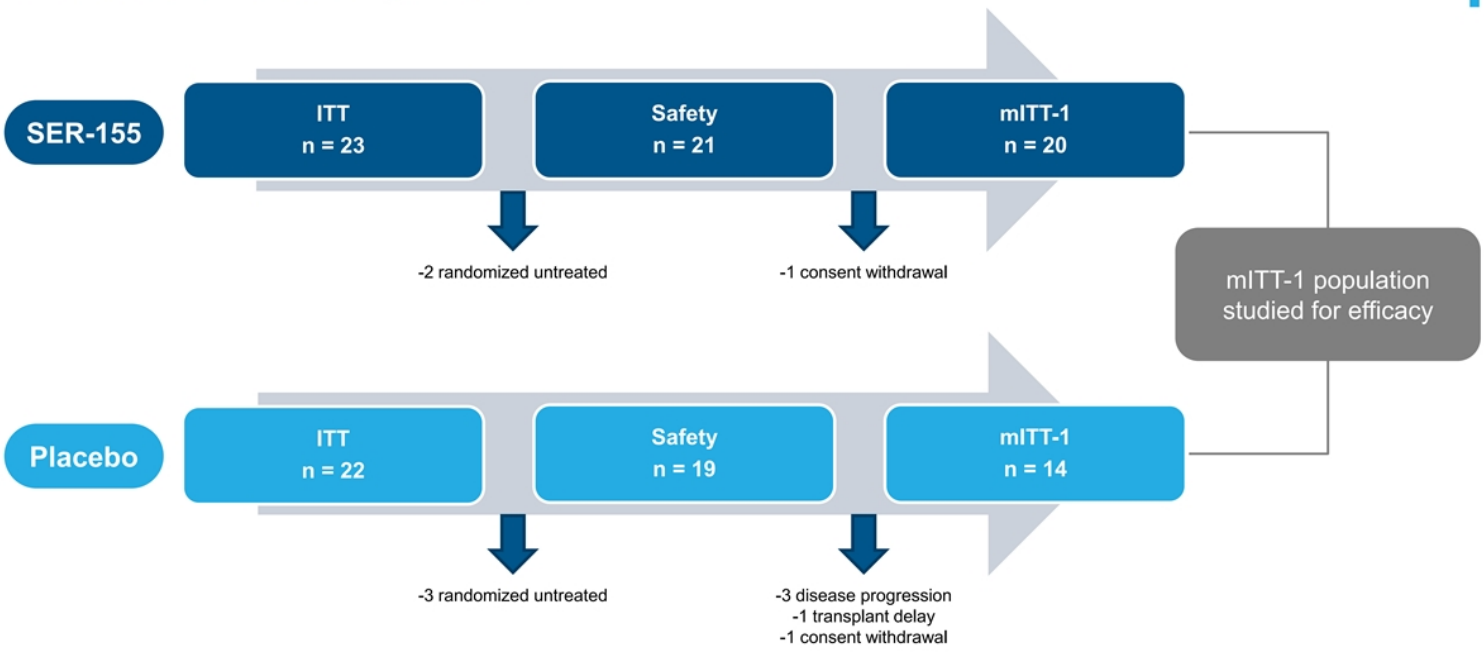
- Safety and tolerability
- SER-155 bacterial strain engraftment

Key Secondary Endpoints through HSCT Day 100:

- Incidence of bloodstream infections (BSI), GI infections, and acute GvHD ≥ 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

Cohort 2 patient disposition



ITT (intent-to-treat): all enrolled subjects
Safety: all subjects receiving any amount of drug
m-ITT1 (modified intent-to-treat 1): all subjects who received study drug in 1st course and underwent HSCT

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Cohort 2 demographics: Representative of the allo-HSCT population

Safety (n=40)

Demographics

- Median age: 63 years
- Gender: 50% male, 50% female
- Ethnicity: 88% white

Cancer Diagnosis

- Most common diagnosis: acute myeloid leukemia (AML): 38%
- Other common diagnoses: acute lymphocytic leukemia (ALL), myelodysplastic syndrome, myeloproliferative neoplasia

mITT-1 (n=34)

Transplant

- 91% received peripheral blood
- 62% received stem cells from a non-relative
- 59% HLA-matched unrelated donor
- 79% underwent a reduced intensity conditioning (RIC) regimen

GvHD Prophylaxis

- 79% received post-transplant cyclophosphamide (PTCy) for GvHD prophylaxis in Cohort 2 vs. 18% in Cohort 1
  - SER-155 arm: 70% (14/20)
  - Placebo arm: 93% (13/14)

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 (≥50% and with Δ ≥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)

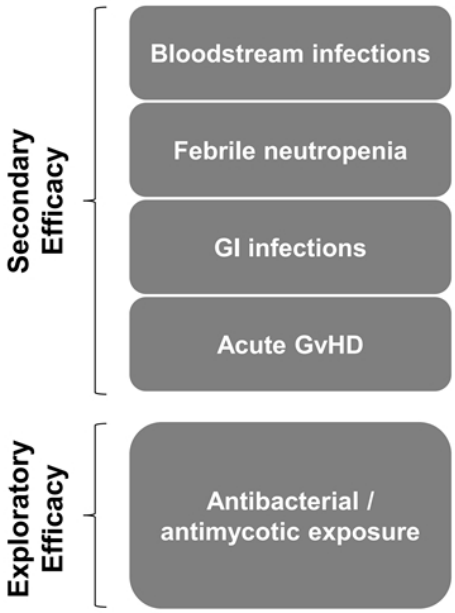
Serious adverse events (SAEs)

- 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



Significant decrease in bacterial bloodstream infections in SER-155-treated subjects vs. placebo

Numerically lower incidence rate of febrile neutropenia in SER-155-treated subjects vs. placebo

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

No subjects in either arm developed ≥ Grade 3 acute GvHD; 2 subjects in each arm developed Grade 2 acute GvHD

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

* no multiplicity adjustments were applied
** CDI: *C. difficile* infection

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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

- CI: 95% 2-sided Clopper-Pearson confidence interval of incidence is applied
- Odds ratio: for incidence between treatment groups (SER-155 and placebo) with 95% 2-sided confidence interval and the corresponding p-value calculated based on the Fisher's Exact test

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

Febrile neutropenia from Day 0 to Day 100 (# patients)	SER-155 n=20 n (%)	Placebo n=14 n (%)
Subjects with FN	13 (65.0)	11 (78.6)
95% confidence interval	(40.8, 84.6)	(49.2, 95.3)

mITT-1 population

Odds ratio	0.51
95% confidence interval	(0.07, 2.99)
p-value	0.4674

- CI: 95% 2-sided Clopper-Pearson confidence interval of incidence is applied
- Odds ratio: for the incidence between treatment groups (SER-155 vs. placebo) with 95% 2-sided confidence interval and the corresponding p-value are calculated based on the Fisher's Exact test

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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)**
p-value	**0.0494**

mITT-1 population

- 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)**
p-value	**0.0163**

mITT-1 population

- 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

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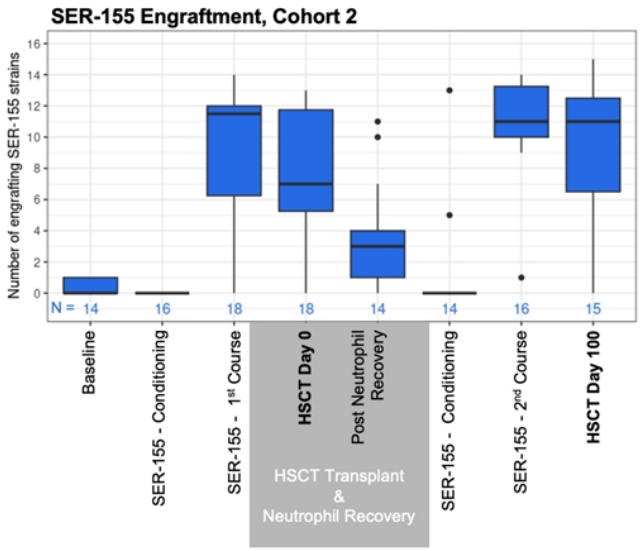
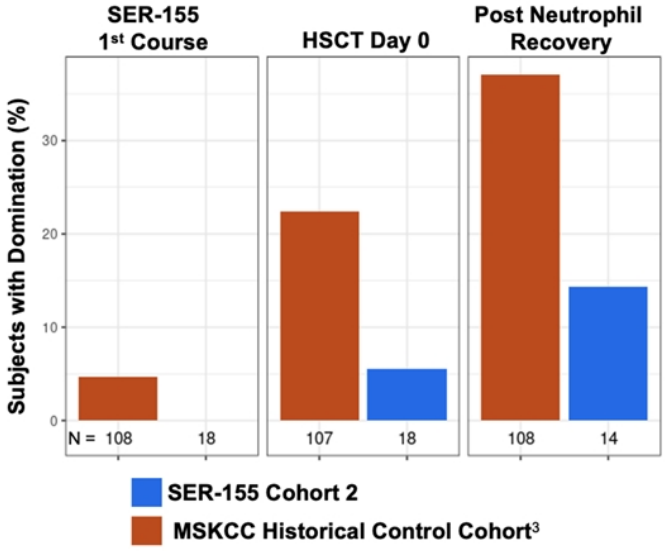
SER-155 Strain Engraftment:
Primary objective achieved - drug bacteria strain engraftment was robust and as expected



- 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 outcomes[1]
- In Cohort 2, the ability to detect pathogen domination[2] (i.e., relative abundance in the GI ≥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 Cohort[3]

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 ≥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

SERES THERAPEUTICS™

Summary and Next Steps

Seres to engage with FDA on advancement of SER-155 allo-HSCT program

- Seek Breakthrough Therapy designation given high unmet medical need associated with bloodstream infections
- Pursue additional designations *(Orphan Drug designation, Qualified Infectious Disease Product)*

Phase 1 results support Seres' strategy to pursue SER-155 and other live biotherapeutics for prevention of serious bacterial infections

- Intend to evaluate SER-155 in additional patient populations with high risk of serious bacterial infections
- Opportunities to address multiple medically vulnerable patient groups with SER-155 and additional pipeline programs

Seres Therapeutics Reports SER-155 Phase 1b Placebo-Controlled Cohort 2 Study Safety and Clinical Results in Patients Undergoing Allogeneic Hematopoietic Stem Cell Transplant (allo-HSCT)

SER-155 administration was associated with a significant reduction in both bacterial bloodstream infections (BSIs) and systemic antibiotic exposure, as well as lower incidence of febrile neutropenia, as compared to placebo through day 100 post HSCT

Demonstrated generally well tolerated safety profile and confirmed drug bacteria strain engraftment; no treatment-related serious adverse events

Company to seek Breakthrough Therapy designation from the FDA, given the high unmet medical need associated with BSIs, and discuss plans to advance development of SER-155 in allo-HSCT

Results support Seres' strategy to pursue SER-155 and other live biotherapeutics for prevention of a broader range of serious bacterial infections in multiple medically vulnerable patient populations

Conference call at 8:30 a.m. ET today

CAMBRIDGE, Mass.—September 12, 2024— Seres Therapeutics, Inc. (Nasdaq: MCRB), a leading live biotherapeutics company, today reports topline clinical data from Cohort 2 of its SER-155 Phase 1b placebo-controlled study in patients undergoing allo-HSCT. In this patient population, infections are frequent, severe and often life-threatening. BSIs are one of the three leading causes of death in allo-HSCT patients. Additionally, transplant related complications, such as infections, increase the recovery burden for patients as well as increase treatment costs due to readmissions, prolonged hospital stays, and increased time in intensive care units. Seres believes that current options to address infections are not sufficient and that SER-155 has the potential to bring significant value to patients, healthcare providers, and the healthcare system.

SER-155 is an investigational live oral biotherapeutic cultivated from clonal master cell banks designed to prevent GI-derived bacterial bloodstream infections and other pathogen-associated complications. Study results demonstrate that SER-155 was associated with a significant reduction in both bloodstream infections and systemic antibiotic exposure as well as a lower incidence of febrile neutropenia, as compared to placebo through day 100 post HSCT. SER-155 was generally well tolerated, with no observed treatment-related serious adverse events.

The Company believes that the SER-155 Phase 1 study results support Seres' corporate strategy to develop its platform, comprised of a pipeline of designed live biotherapeutics, in multiple medically vulnerable patient populations at high risk of life-threatening bacterial infections and associated negative

clinical outcomes. Seres intends to seek Breakthrough Therapy designation, given the high unmet medical need associated with BSIs, and discuss advancing development of SER-155 for allo-HSCT with the U.S. Food and Drug Administration (FDA). The Company also intends to evaluate SER-155 in additional patient populations that have a high risk of serious bacterial infections.

“The placebo-controlled Phase 1b study Cohort 2 results provide further evidence supporting the potential of SER-155 to reduce the risk of bacterial bloodstream infections, a leading cause of mortality and morbidity in patients undergoing allo-HSCT,” said Lisa von Moltke, M.D., Chief Medical Officer of Seres Therapeutics. “Given our encouraging clinical results and the severe consequences of bacterial infections, we will pursue Breakthrough Therapy designation with the FDA. We also look forward to discussing our plans to further develop SER-155 with the Agency.”

“Bacterial infections such as bacteremia (bacteria in blood) are a frequent and often life-threatening complication faced by patients undergoing HSCT as well as other patients with cancer.” said infectious diseases physician David Fredricks, M.D., Professor, Vaccine and Infectious Disease Division, and Professor, Clinical Research Division at the Fred Hutchinson Cancer Center in Seattle. “Many of these infections arise from bacteria in the gastrointestinal tract. Investigational live oral biotherapeutics such as SER-155 hold promise as a novel approach that could protect patients against these serious bacterial infections, resulting in improved patient outcomes, together with reduced use of antibiotics. The data from the SER-155 Phase 1b study, including results showing lower infection rates, less systemic antibiotic exposure, and reduced incidence of febrile neutropenia events, support continued development in allo-HSCT.”

SER-155 Phase 1b Study Design

The SER-155 Phase 1b study (NCT04995653) included two cohorts. Cohort 1 was designed to assess safety and drug pharmacology, specifically the drug strain engraftment in the gastrointestinal tract. Cohort 1 included 13 subjects who received any dosing of the SER-155 regimen, with 11 subjects subsequently receiving an allo-HSCT. Results from this cohort, announced in May 2023, showed SER-155 was generally well tolerated and resulted in successful drug strain engraftment and a reduction in pathogen domination in the GI microbiome relative to a historical control cohort.

Study Cohort 2 utilized a randomized, double-blinded 1:1 placebo-controlled design to further evaluate safety and drug strain engraftment, as well as key secondary and exploratory endpoints such as the incidence of bacterial bloodstream infections and related medical consequences such as febrile neutropenia and antibiotic use. Cohort 2 included 45 patients in the intention-to-treat (ITT) population. Of the ITT population, 20 received SER-155 and 14 received placebo, each of whom subsequently received an allo-HSCT, with data available for clinical evaluation through day 100, the study’s prespecified primary observation point. Exploratory hypothesis testing was conducted at the two-sided α=0.05 level. Ninety-five percent (95%) 2-sided confidence intervals (CIs) were determined, where specified. No adjustment for multiplicity was done. A subset of patient samples was available for drug pharmacology analysis.

The median age in Cohort 2 was 63, and most subjects had acute myeloid leukemia, acute lymphocytic leukemia, myelodysplastic syndrome or myeloproliferative neoplasia as their primary disease and received reduced-intensity conditioning pre-transplant. Most patients received peripheral blood stem cells from a matched unrelated donor. A majority received post-transplant cyclophosphamide as part of their graft-versus-host disease (GvHD) prophylaxis.

Summary of Cohort 2 Study Results

Consistent with the observations from the Phase 1b study Cohort 1, SER-155 was generally well tolerated, and no treatment-emergent serious adverse events related to drug were observed. SER-155 bacterial strains engrafted into the gastrointestinal tract of patients following the administration of SER-155.

The incidence of BSIs was significantly lower in the SER-155 arm compared with the placebo arm (2/20 (10%) vs. 6/14 (42.9%), respectively; [Odds Ratio: 0.15; 95% CI: 0.01, 1.13, p=0.0423]). In addition, while antibiotic starts were similar in each arm, patients administered SER-155 were treated with antibiotics for a significantly shorter duration compared to patients in the placebo arm (9.2 days vs. 21.1 days, respectively, with a mean difference of -11.9 days [95% CI: -23.85, -0.04; p=0.0494]). The incidence of febrile neutropenia was lower in patients administered SER-155 compared to placebo (65% vs. 78.6%, respectively; [Odds Ratio: 0.51; 95% CI: 0.07, 2.99; p=0.4674]). Six cases of gastrointestinal infections (*C. difficile* infections) were observed in the study, with four cases (20%) in the SER-155 arm and two cases (14.3%) in the placebo arm.

Recent changes in the allo-HSCT standard of care and the increasing use of post-transplant cyclophosphamide as part of prophylactic therapy for GvHD have reduced rates of GvHD overall in this patient population. The rates of GvHD in the study were low, with two cases of grade 2 GvHD observed in each arm, and no cases of grade 3 or 4 GvHD were observed.

In Cohort 2, the ability to detect pathogen domination (i.e., relative abundance in the GI ≥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 identified. In a comparison of the prevalence of pathogen domination versus a larger allo-HSCT historical control cohort, pathogen domination in SER-155 subjects was substantially lower, providing further evidence of SER-155 activity.

“The SER-155 Phase 1b results generate further evidence to support Seres’ strategy as SER-155 is our second live biotherapeutic, after VOWST, designed to prevent serious bacterial infections and associated negative clinical outcomes in medically vulnerable populations. An estimated 40,000 patients worldwide undergo allogeneic stem cell transplantation each year. Adding autologous stem cell transplants (auto-HSCT), a natural adjacent patient population, approximately doubles this figure. The potential to contract a bloodstream infection during the allogeneic transplant process is significant, with incidence reports in the literature reaching up to 45%. Allogeneic transplant-related complications, including infections, increase already significant treatment costs by approximately $180,000 per patient. Given this high unmet need, we believe SER-155 could provide meaningful value for patients and the healthcare system,” said Eric Shaff, Chief Executive Officer of Seres Therapeutics.

Mr. Shaff continued, “We are particularly encouraged by the consistency of related efficacy outcomes in this study, especially the significantly lower rates of bloodstream infections and systemic antibiotic exposure as well as fewer instances of febrile neutropenia, as compared to placebo. Supported by these data and our well-established clinical, pharmacological, CMC, and regulatory capabilities, we plan to engage with the FDA to seek Breakthrough Therapy designation and discuss advancing development of SER-155 in allo-HSCT. With SER-155 and additional pipeline programs, we believe we may have the opportunity to address multiple patient groups, including allo-HSCT, auto-HSCT, CAR-T, chronic liver disease, cancer neutropenia, and solid organ transplants, thereby potentially creating significant commercial opportunities.”

Seres fully owns worldwide rights for the commercialization of SER-155.

Conference Call Information

Seres’ management will host a conference call today, September 12, 2024, at 8:30 a.m. ET. The conference call may be accessed by calling 1-800-715-9871 (international callers dial 1-646-307-1963) and referencing the conference ID number 622932. To join the live webcast, please visit the “Investors and News” section of the Seres website at www.serestherapeutics.com. A webcast replay will be available on the Seres website beginning approximately two hours after the event and will be archived for at least 21 days.

About SER-155

SER-155 is an investigational live biotherapeutic designed to prevent GI-derived bloodstream infections, enhance epithelial barrier integrity to reduce the likelihood of bacterial translocation from the gut to the bloodstream, and induce immune tolerance responses to reduce the incidence of GvHD. SER-155 contains 16 bacterial strains selected using Seres’ reverse translation discovery and development platform technologies to optimize SER-155’s functional profile. The design incorporates biomarker data from human clinical data and screening data from nonclinical human cell-based assays and in vivo disease models. SER-155 has been evaluated in a Phase 1b placebo-controlled study in patients undergoing allo-HSCT. SER-155 has received FDA Fast Track designation for reducing the risk of infection and GvHD in patients undergoing HSCT. The early development of the program was supported by Combating Antibiotic-Resistant Bacteria Biopharmaceutical Accelerator (CARB-X), a global non-profit partnership accelerating antibacterial products to address drug-resistant bacteria.

About Seres Therapeutics

Seres Therapeutics, Inc. (Nasdaq: MCRB) is a commercial-stage company focused on improving patient outcomes in medically vulnerable populations through novel live biotherapeutics. Seres led the successful development and approval of VOWST™, the first FDA-approved orally administered microbiome therapeutic. The Company is developing SER-155, designed to prevent gastrointestinal-derived bloodstream infections, enhance epithelial barrier integrity, and induce immune tolerance responses to reduce the incidence of graft-versus-host-disease (GvHD). The Company is also advancing additional cultivated oral live biotherapeutics for medically vulnerable populations, including those with chronic liver disease, cancer neutropenia, and solid organ transplants. For more information, please visit www.serestherapeutics.com.

Important Additional 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.

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 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 Société des Produits Nestlé S.A. 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.

Investor and Media Contacts:

IR@serestherapeutics.com

Carlo Tanzi, Ph.D.
Kendall Investor Relations
ctanzi@kendallir.com

Exhibit 99.3

Seres Therapeutics Investor Presentation

September 12, 2024

Disclaimers

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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)

Sources: Feuerstadt P et al. *N Engl J Med.* 2022;386(3):220-229.

SERES THERAPEUTICS™

VOWST asset sale a transformational transaction for Seres

VOWST™ (fecal microbiota spores, live-brpk) → Nestlé HealthScience

- **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

*SPN: Société des Produits Nestlé S.A.

SERES THERAPEUTICS™

Potential to treat a range of vulnerable patient populations

Target population characteristics

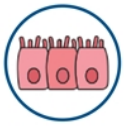

GI microbiome functional disruption


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

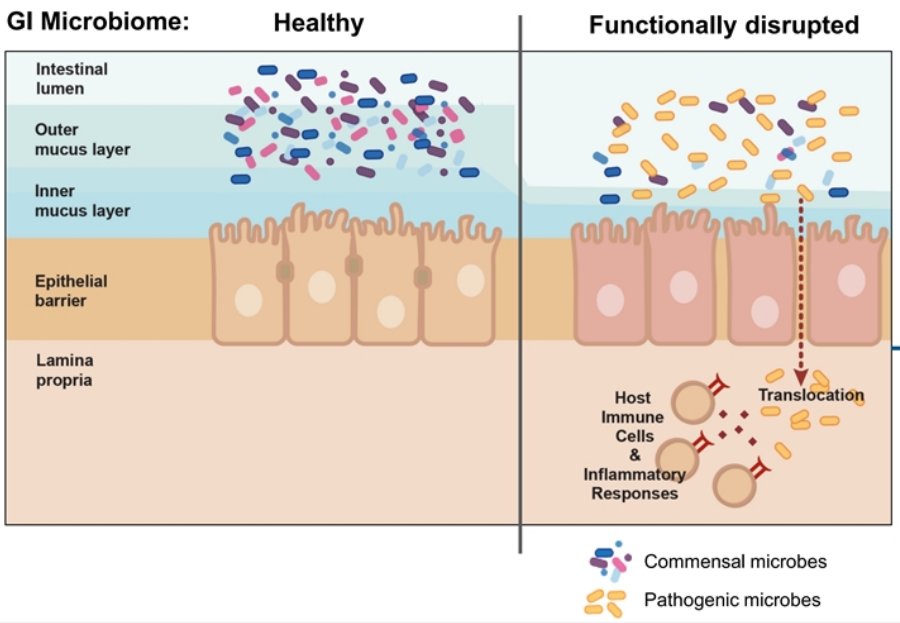
Treat immune-related diseases

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

HSCT: hematopoietic stem cell transplant; CAR-T: chimeric antigen receptor T-cell therapy

SERES THERAPEUTICS™

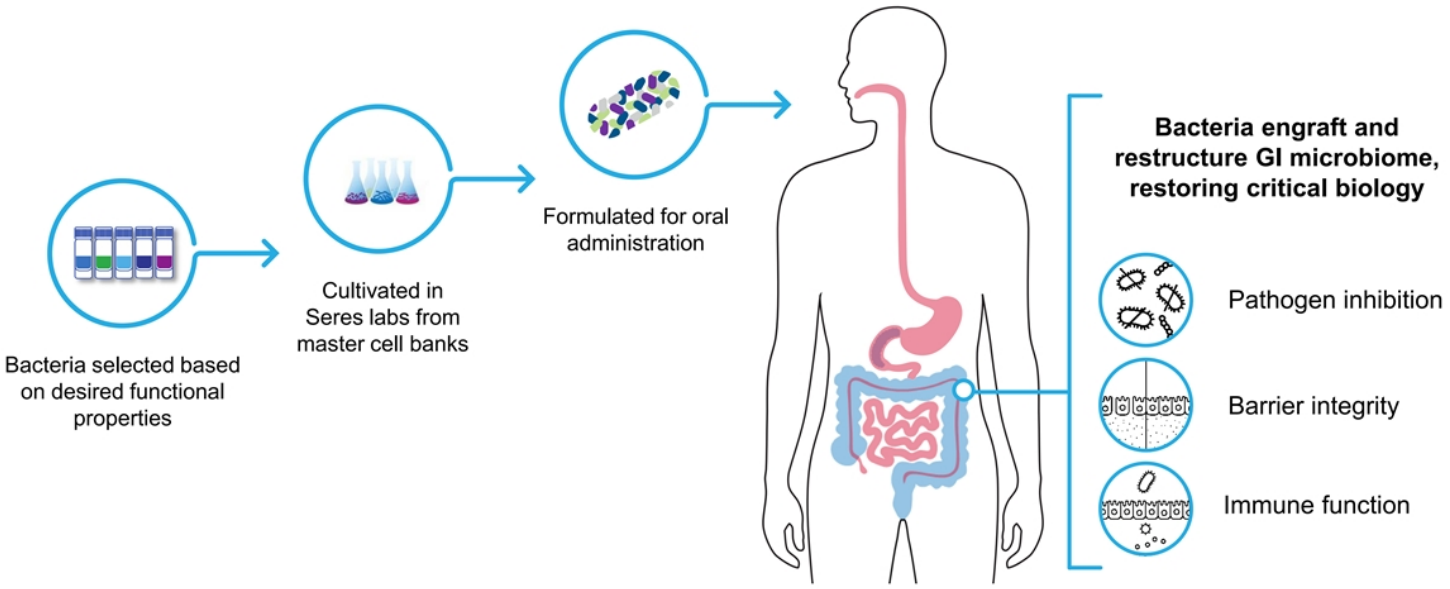
GI microbiome functional disruption leads to disease susceptibility



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

Sources: Nanayakkara et al, *CA Cancer J Clin* 2021; Penack et al, *Blood Adv* 2020; Zheng et al, *Infect Dis Ther* 2021

SERES THERAPEUTICS™

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

VOWST™ provides clinical validation

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

Program	Lead Indication & Development Stage	Therapeutic Objectives	Potential Additional Indications
SER-155	Allogeneic HSCT: Phase 1b Cohort 2 (placebo controlled) data announced Sept. '24	Reduce incidence of serious bacterial infections (e.g., BSIs), febrile neutropenia, and GvHD	• Autologous HSCT • CAR-T • Blood cancers • Solid organ transplant
SER-147	Chronic liver disease: Anticipate IND ready in H2 '25	Reduce incidence of serious bacterial infections (e.g., SBP, BSIs) and related complications	• ICU patients • Long-term care patients

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

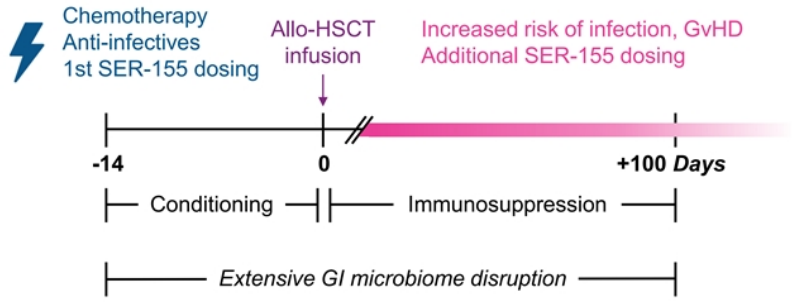
BSI: bloodstream infection; SBP: spontaneous bacterial peritonitis

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

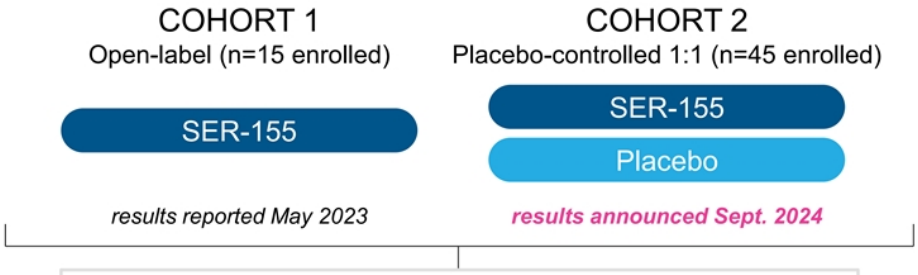


- **Only ~60% survival** 3 years post-transplant
- **~10% transplant mortality** in first 100 days post-transplant
- **~80% of adult deaths in first 100 days caused by complications** of procedure; half of these due to infections and GvHD
- Complications have substantial impact: mortality, cost, hospital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

Sources: CIBMTR 2023 US summary slides; Penack et al, *Blood Adv* 2020; Khan et al, *Blood* 2021; Peled et al, *NEJM* 2020; Stein-Thoeringer et al, *Science* 2019; Bleakley & Riddell, *Nat Rev Cancer* 2004

SERES THERAPEUTICS™

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 ≥ 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 (≥50% and with Δ ≥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)

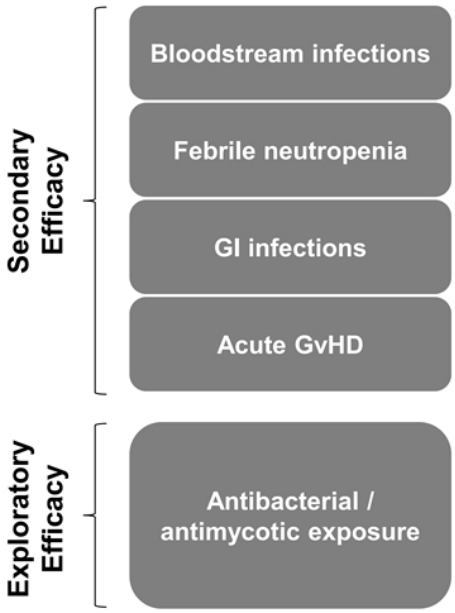
Serious adverse events (SAEs)

- 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



Significant decrease in bacterial bloodstream infections in SER-155-treated subjects vs. placebo

Numerically lower incidence rate of febrile neutropenia in SER-155-treated subjects vs. placebo

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

No subjects in either arm developed ≥ Grade 3 acute GvHD; 2 subjects in each arm developed Grade 2 acute GvHD

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

* no multiplicity adjustments were applied
** CDI: *C. difficile* infection

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

- CI: 95% 2-sided Clopper-Pearson confidence interval of incidence is applied
- Odds ratio: for incidence between treatment groups (SER-155 and placebo) with 95% 2-sided confidence interval and the corresponding p-value calculated based on the Fisher's Exact test

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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)**
p-value	**0.0494**

mITT-1 population

- 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)**
p-value	**0.0163**

mITT-1 population

- 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

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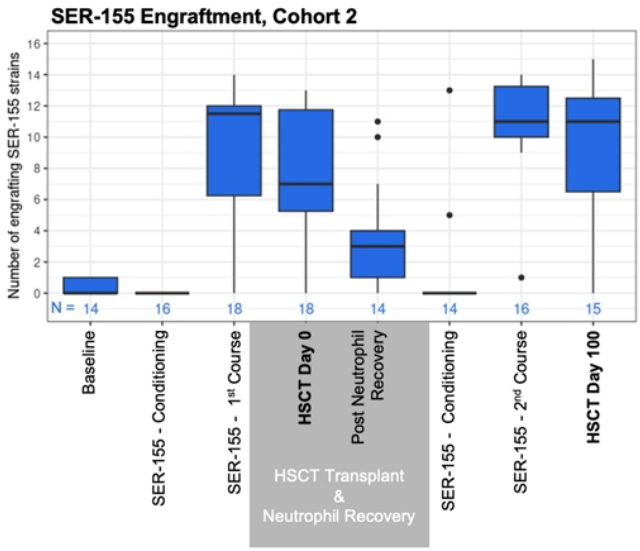
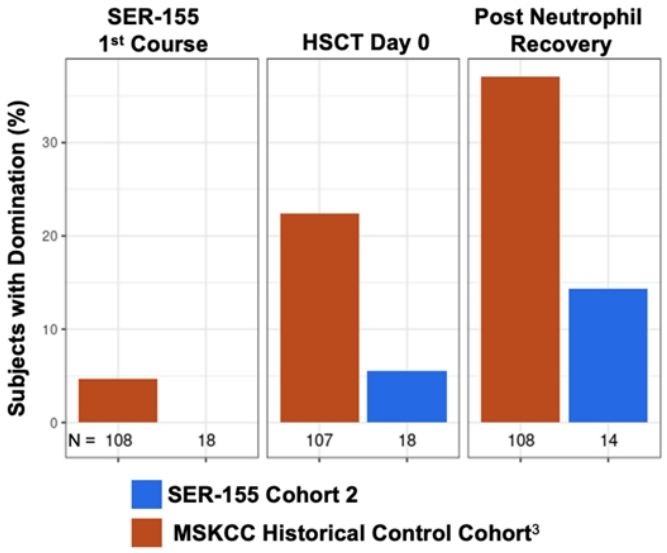
SER-155 Strain Engraftment:
Primary objective achieved - drug bacteria strain engraftment was robust and as expected



- 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 outcomes[1]
- In Cohort 2, the ability to detect pathogen domination[2] (i.e., relative abundance in the GI ≥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 Cohort[3]

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 ≥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

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Viral prophylaxis provides precedent in medically vulnerable patients

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

PREVYMIS® (letermovir)
240 mg, 480 mg tablets
Injection 20 mg/mL

$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

Sources: CMS.gov; Broder et al, *Am Drug Health Benefits* 2017; Perales et al, *Biol Blood Marrow Transplant* 2017; Merck SEC filings

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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

Sources: CIBMTR; HRSA; Passweg et al, *BMT* 2021; IQVIA; Broder et al, *Am Drug Health Benefits* 2017; Perales et al, *Biol Blood Marrow Transplant* 2017

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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

Sources: CIBMTR; Passweg et al, *BMT* 2021; US Organ Procurement & Transplantation Network; International Registry in Organ Donation and Transplantation; US National Cancer Institute SEER Program; American Cancer Society; European Cancer Information System

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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

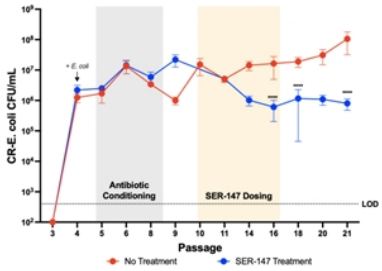
~20-25% of infections are spontaneous bacterial peritonitis and bloodstream infections likely to be gut-seeded

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



Sources: GBD 2017 Cirrhosis Collaborators, *Lancet Gastroenterology & Hepatology* 2020; United Nations world population data; Trebicka et al, *J Hepatol* 2020; Seres preclinical data from 2023 IDWeek

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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

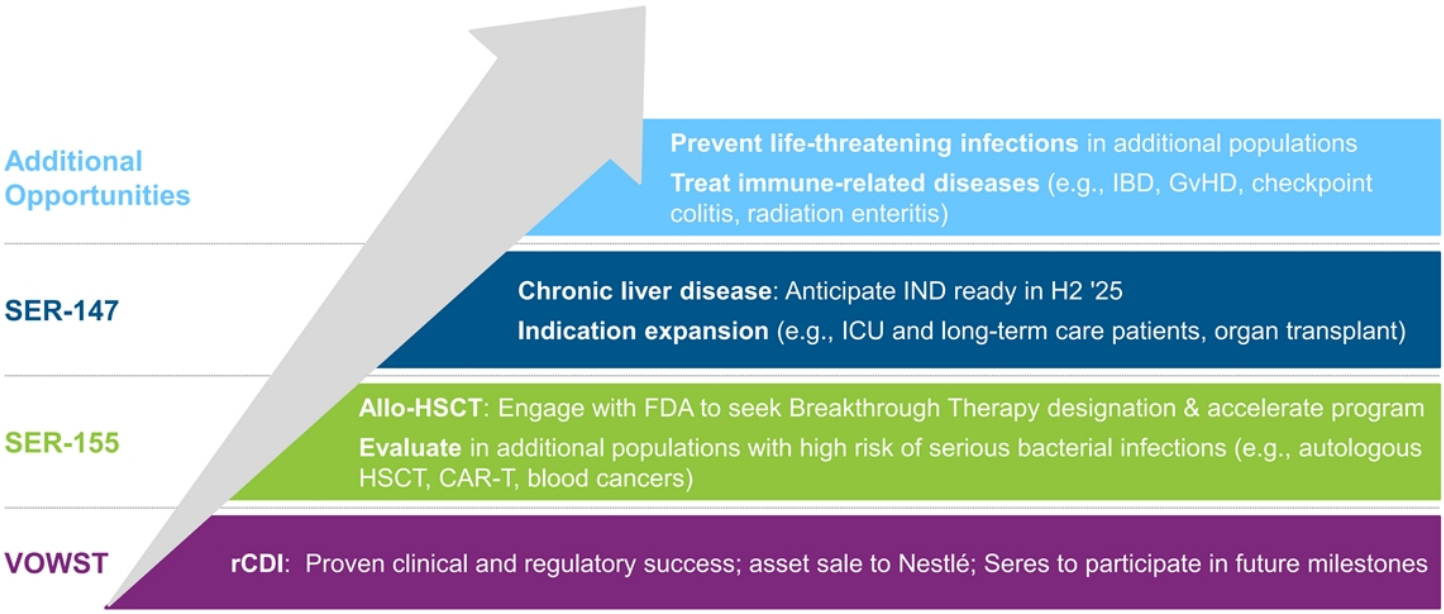
Strain isolation and characterization pipeline to rapidly identify cGMP-suitable medium components

Highly intensive ***strain bioprocessing*** leveraging flexible, single-use manufacturing technology for cost-effective production

Novel formulations enabling consistent drug product composition, drug stability for distribution, and targeted drug delivery

Quality systems to ensure product quality and stability, extending prior regulatory successes, including developing product release specifications with the FDA

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