



SERES
THERAPEUTICS

SER-155 Phase 1b Readout
September 12, 2024



Disclaimers

Forward Looking Statements

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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

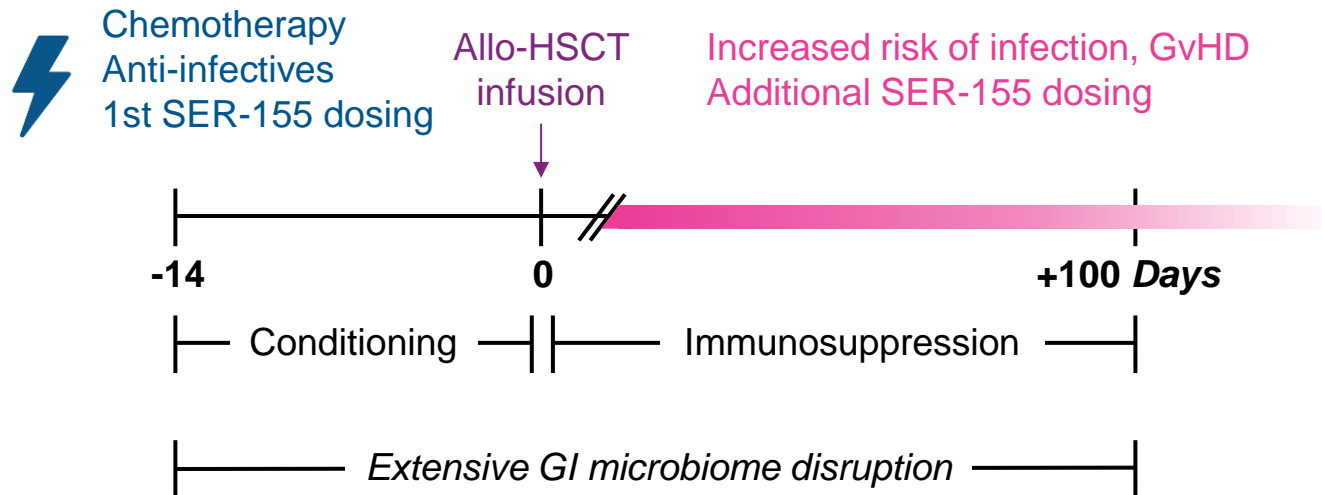
Seres and certain of its directors, executive officers and other members of management and employees may be deemed to be participants in soliciting proxies from its stockholders in connection with the proposed transaction. Information regarding the persons who may, under the rules of the SEC, be considered 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.

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

SER-155

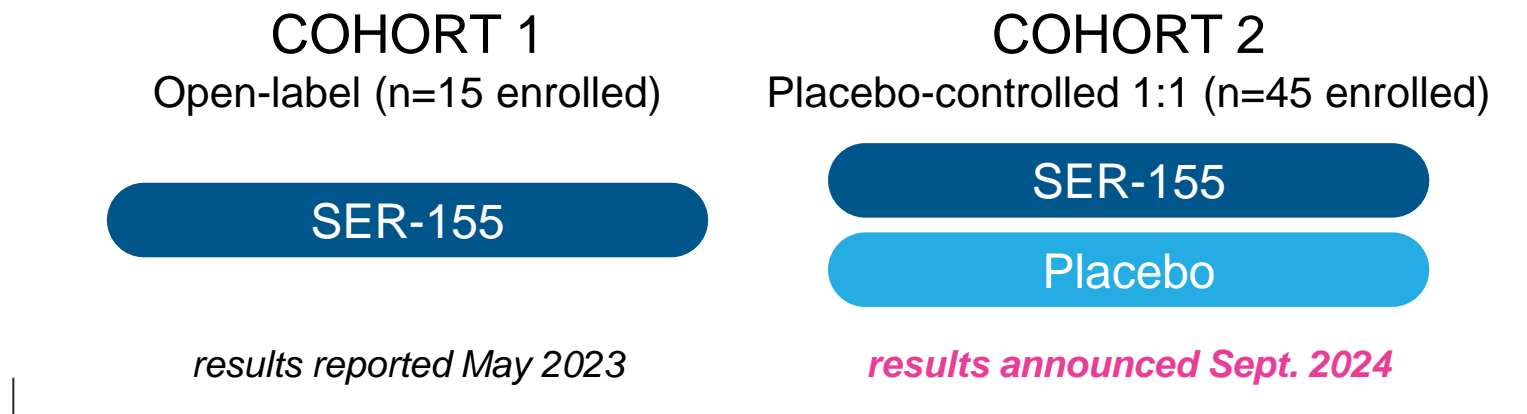
- Investigational live oral biotherapeutic cultivated from clonal master cell banks
- Designed to prevent GI-derived bacterial bloodstream infections (BSIs) and other pathogen-associated complications

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:

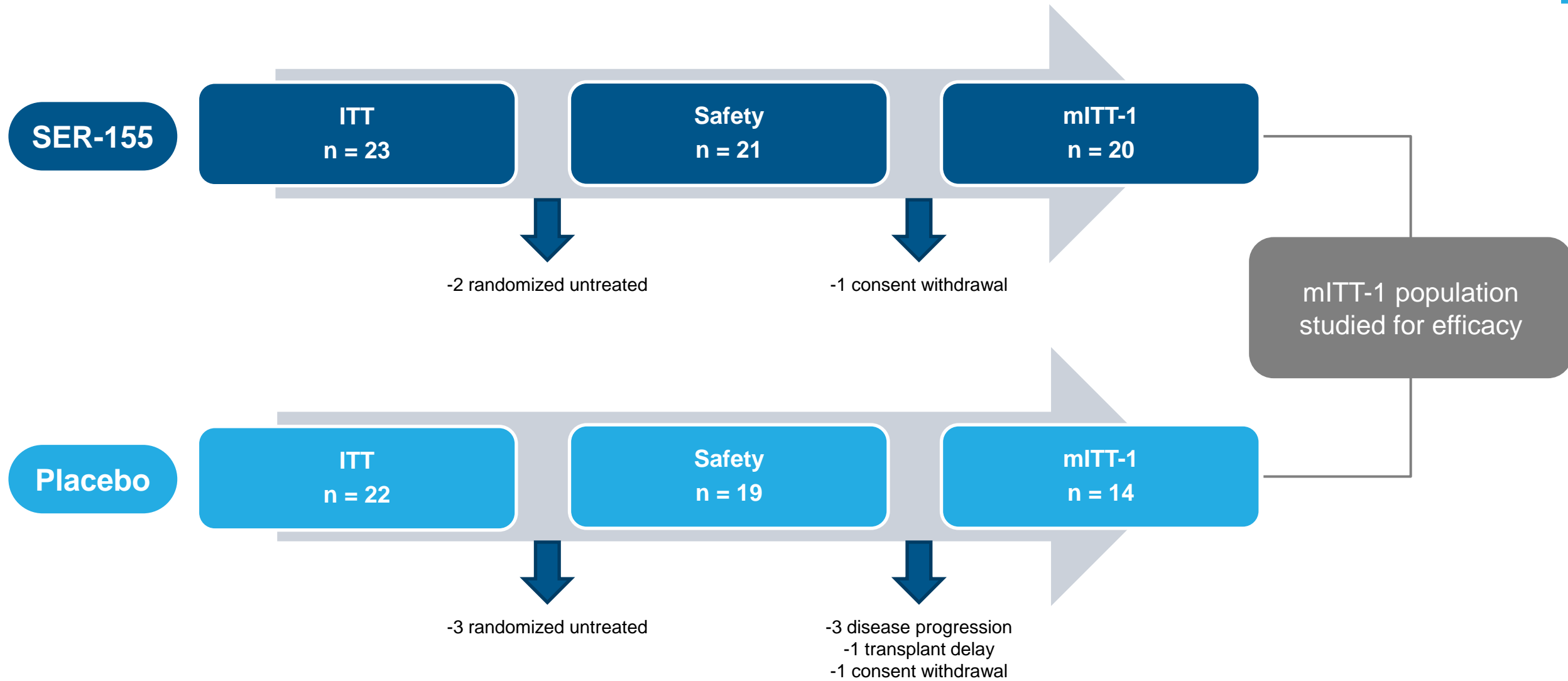
- Safety and tolerability
- SER-155 bacterial strain engraftment

Key Secondary Endpoints through HSCT Day 100:

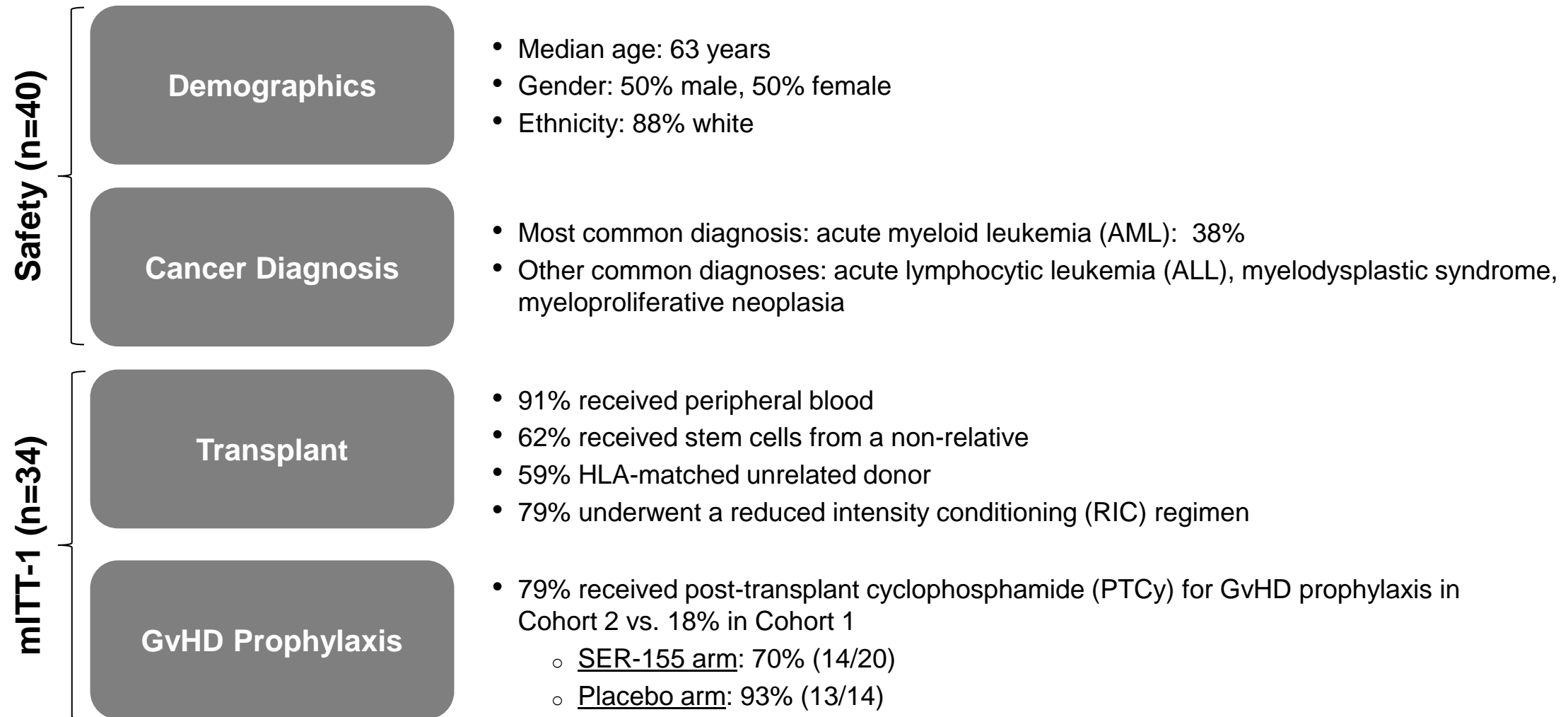
- Incidence of bloodstream infections (BSI), GI infections, and acute GvHD \geq Grade 2
- Incidence and duration of febrile neutropenia
- Bacterial pathogen abundance

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

Cohort 2 patient disposition



Cohort 2 demographics: Representative of the allo-HSCT population



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: *Finigoldia 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*

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

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 (SD)	Placebo n=14 n (SD)
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	SER-155 n=20 Rate (SD)	Placebo n=14 Rate (SD)
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)

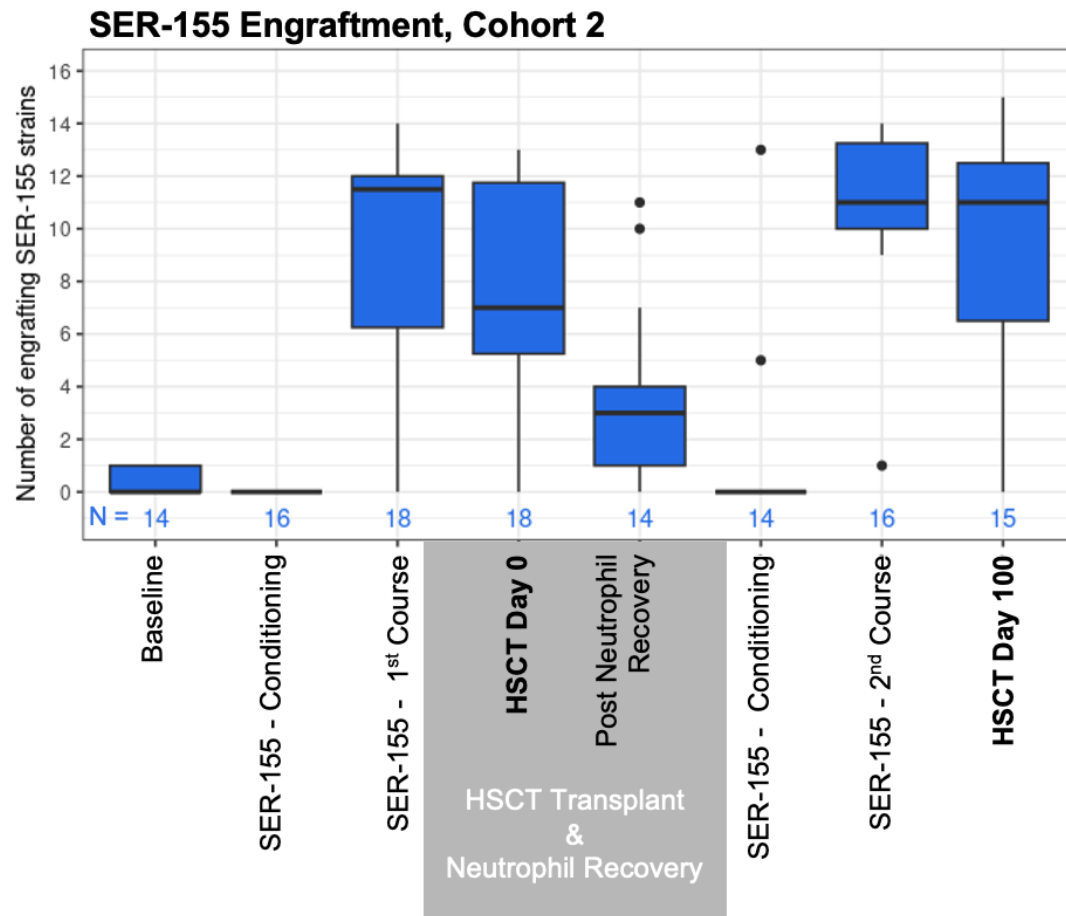
mITT-1 population

p-value

0.0163

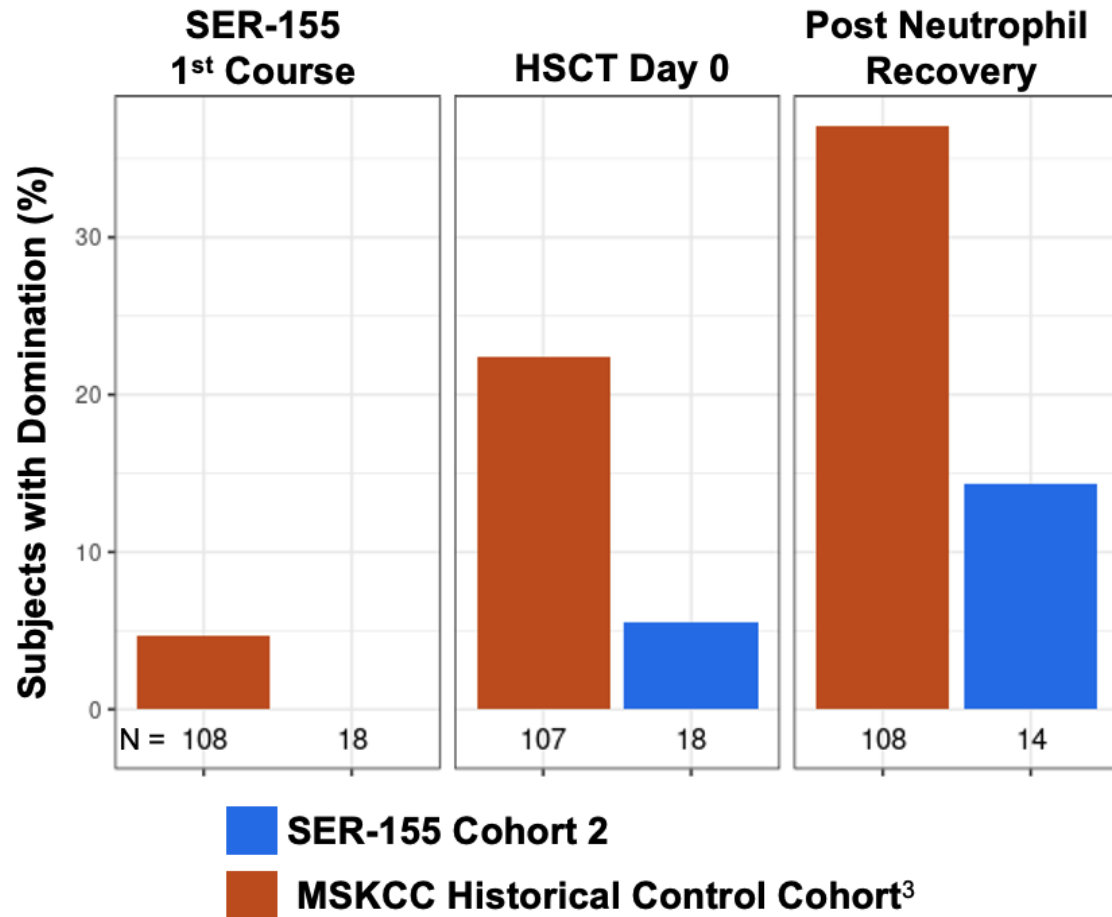
- Cumulative exposure rate is calculated as the sum of all days a subject received systemic antibacterials and/or antimycotics on or after HSCT Day 0 (counting once per day, regardless of number of antibacterial/antimycotic medications taken in a day) through HSCT Day 100 over the total number of days 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

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