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): January 13, 2025

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

Delaware (State or other jurisdiction of incorporation)

001-37465 (Commission File Number)

27-4326290 (IRS Employer Identification No.)

101 Cambridgepark Drive Cambridge, MA (Address of principal executive offices)

02140 (Zip Code)

	Registrant's telepho	ne number, including area code: (617) 945-9626	
	(Former Name	Not Applicable or Former Address, if Changed Since Last	Report)	
	cck the appropriate box below if the Form 8-K filing is into owing provisions:	ended to simultaneously satisfy the f	iling obligation of the registrant under any of the	
	Written communications pursuant to Rule 425 under the	e 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 1	3e-4(c) under the Exchange Act (17	CFR 240.13e-4(c))	
Sec	urities 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)	
	cate by check mark whether the registrant is an emerging pter) or Rule 12b-2 of the Securities Exchange Act of 1934		405 of the Securities Act of 1933 (§ 230.405 of this	
Em	erging growth company			
	n emerging growth company, indicate by check mark if the or revised financial accounting standards provided pursua			

tem 7.01. Regulation FD Disclosure.

On January 13, 2025, Seres Therapeutics, Inc. (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 99.1 to this Current Report on Form 8-K (this "Current Report") and incorporated herein by reference.

The information in Item 7.01 of this Current Report, including Exhibit 99.1 attached hereto, is intended to be furnished and shall not be deemed "filed" for purposes 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.

Item 9.01. Financial Statements and Exhibits.

(d) Exhibits.

The following Exhibit 99.1 relates to Item 7.01 and shall be deemed to be furnished, and not filed:

Exhibit No.

99.1 <u>Seres Therapeutics, Inc. Corporate Presentation as of January 2025</u>

104 Cover Page Interactive Data File - the cover page XBRL tags are embedded within the Inline XBRL document.

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.

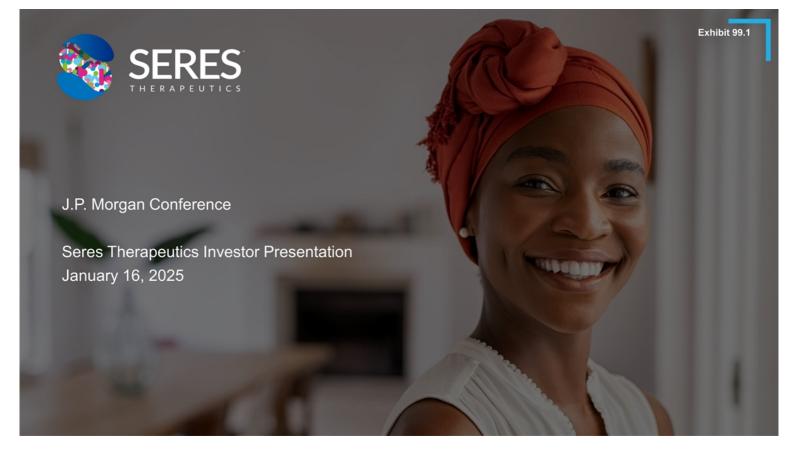
Date: January 13, 2025

SERES THERAPEUTICS, INC.

By: // Thomas J. DesRosier

Name: Thomas J. DesRosier

Title: Chief Legal Officer and Executive Vice President



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: our anticipated financial performance, including cash and cash equivalents, for any period of time, including for the year ended December 31, 2024; the timing and results of our clinical studies and data readouts; our clinical development plans; the anticipated timing of communications with or feedback from the FDA; the impact, value or potential benefits of Breakthrough Therapy designation, Fast Track designation or any other regulatory designations; our ability to secure a partnership and/or generate additional capital; the potential market and commercial opportunity for SER-155 and other product candidates, if approved; projected cash runway; 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) our novel approach to therapeutic intervention; (5) our reliance on third parties to conduct our clinical trials and manufacture our product candidates; (6) the competition we will face; (7) our ability to protect our intellectual property; (8) our ability to retain key personnel and to manage our growth; (9) the effect of the VOWST sale on our ability to retain and hire key personnel and maintain relationships with our customers, suppliers, advertisers, partners and others with whom we do business, or on our operating results and businesses generally; (10) the risks associated with the disruption of management's attention from ongoing business operations due to the obligation to provide transition services; (11) our failure to receive the installment payments or the milestone payments in the future; (12) the uncertainty of impact of the 50/50 profit and loss sharing arrangement on our reported results and liquidity; and (13) we may not be able to realize the anticipated benefits of the VOWST sale. These and other important factors discussed under the caption "Risk Factors" in our Quarterly Report on Form 10-Q filed with the Securities and Exchange Commission (SEC), on November 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 st



Transforming patient outcomes using proprietary consortia of live biotherapeutics

Strong Foundation

- Validated platform highlighted by VOWST® FDA approval as first ever oral microbiome therapy with outstanding clinical results
- VOWST asset sale strengthened balance sheet and streamlined organization
- Cash into Q1 2026

Positive SER-155 Phase 1b Data in Allo-HSCT

- 77% relative risk reduction for bloodstream infections
- Well tolerated safety profile; no treatment-related SAEs and engraftment
- Biomarker data supports impact on epithelial barrier integrity
- FDA Breakthrough Designation

Blockbuster SER-155 Opportunity

- Accelerate SER-155 development in allo-HSCT
- Potential to initiate multiple clinical trials in next 12-18 months for additional indications
- SER-155 represents multibillion net sales opportunity across indications (e.g., autologous-HSCT, blood cancers, CAR-T recipients)

Expansive Platform Potential

- Systemic inflammation and immune homeostasis biomarker data support the potential in inflammatory and immune diseases, such as IBD (e.g., ulcerative colitis and Crohn's disease)
- SER-147 designed to prevent infections in chronic liver disease

Pursuing SER-155 strategic partnership to accelerate next study in allo-HSCT and expand to multiple target populations



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



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 (completed September 2024) provided capital to support pipeline advancement and resulted in a more streamlined, focused organization



VOWST asset sale completed September 30, 2024: transformational for Seres – provides resources to support SER-155 advancement







- VOWST asset purchase agreement provided infusion of capital and supports SER-155 development
- Asset sale extended operational runway into Q4 '25; now into Q1 '26
- · Retires 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)

Transaction results in a more streamlined, focused Seres organization and lower cash burn rate

*SPN: Société des Produits Nestlé S.A.



The gut microbiome has substantial untapped therapeutic potential to both prevent and treat diseases



The genetic content of the microbes living within and on your body is at least

100-fold greater than that contained in the human genome



Disrupted gastrointestinal microbiome is mechanistically linked to infections and disease

Disrupted & Disease Susceptible **Not Disrupted** Intestinal lumen Outer mucus layer Inner mucus layer **Epithelial** barrier Lamina propria Pathogenic Commensal Bacteria Presenting Cells Bacteria

Impacted Biology & Disease Relevance



Pathogen or "undesirable" microbe abundance



Lost epithelial barrier & mucosal integrity



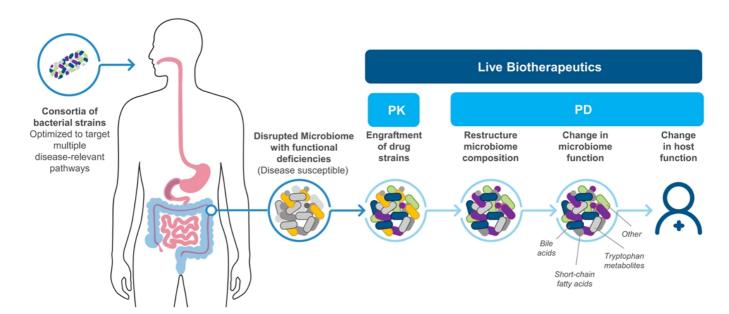
Unfavorable immune responses

Increased pathogen
abundance,
in combination with
impaired barrier integrity
and decreased immune
function...

...enables bacterial and antigen translocation leading to infections & inflammation



Consortia of live commensal bacteria can be used as therapeutics





Seres' biotherapeutics and pipeline candidates are expected to 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

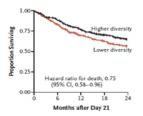
Sources: Nanayakkara et al, *CA Cancer J Clin* 2021; Penack et al, *Blood Adv* 2020; Zheng et al, *Infect Dis Ther* 2021



Seres' initial focus is preventing infections in allo-HSCT; potential is broad

Allo-HSCT

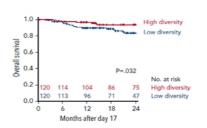
Survival by microbiome diversity



Lower microbiome diversity post allo-HSCT is associated with increased blood stream infections and decreased overall survival

Autologous HSCT

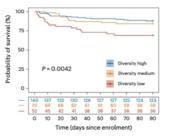
Survival by microbiome diversity



Lower microbiome diversity post auto-HSCT is associated with worse overall and progression-free survival

Chronic Liver Disease

Survival by microbiome diversity



Microbiome diversity decreases with disease severity and is associated with increased risk of infection & death

Common biology across vulnerable populations: pathogen domination in the GI microbiome, loss of epithelial barrier integrity, impacted immune function

Sources: Peled, et al, NEJM 2020; Khan et al, Blood 2021; Bajaj et al, Clin Gastroenterol Hepatol 2019; Odenwald, et al. Nature Micro. 2023



Near-term focus on SER-155 as anchor biotherapeutic program; achieved Breakthrough and Fast Track designations



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

provides clinical proof of concept

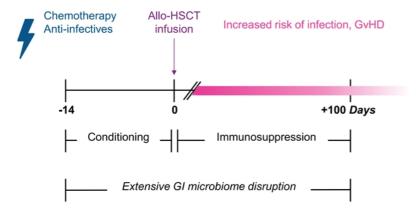
	Program	Lead Indication & Development Stage	Therapeutic Objectives	Potential Additional Indications
	SER-155	Allogeneic HSCT: Phase 1b Cohort 2 (placebo controlled) data announced Sept. '24 and exploratory biomarker data in Jan '25	Reduce incidence of serious bacterial infections (e.g., BSIs), febrile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

Briefing Book Submitted – engaging with FDA in Q1 '25 to explore **potential for SER-155 to have single** registrational study for efficacy, following successful precedent from VOWST



Allo-HSCT regimen can result in life-threatening complications

Allo-HSCT treatment regimen



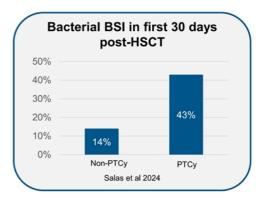
- Only ~60% survival 3 years posttransplant
- Significant immune compromise
- ~10% transplant mortality for adults in first 100 days posttransplant
- Infections are leading cause of death in first 100 days posttransplant for adults
- Other leading causes of death are disease relapse and organ failure



Bloodstream infections (BSIs) are a leading cause of death post-transplant and are increasing in incidence



- BSI risk increasing due to recent adoption of post-transplant cyclophosphamide (PTCy) for GvHD prophylaxis
- · ~50% of infections are gut-seeded
- 50-80% febrile neutropenia incidence



Impact

- Infection is leading cause of death in first 100 days post-HSCT for adults
- ~7.5% mortality rate from bloodstream infections
- Complications including infection associated with longer hospital stay and ICU utilization, driving substantial cost increase



Sources: Gill et al, Microorganisms 2023; Sava et al, Bone Marrow Transplantation 2022; Carreira et al, Transplant Cell Ther 2022; Youssel et al, Pediatric Transplantation 2020; Song et al, Infect Drug Resist 2023; Bolanos-Meade J et al, NEJM 2023; Salas et al, Transplant Cell The 2024; Rearigh et al, Annals of Hematology 2020; CIBMTR 2023 US summary slidies; Pertales et al Biol Blood Marrow Transplant 2017; 2024 Seres physician market research

Standards of care to prevent bloodstream infections in allo-HSCT patients are poor and decreasing in efficacy



- Fluoroquinolones (FQ) recommended for antibacterial prophylaxis
- Prophylaxis recommended during window of expected neutropenia



EBMT guidelines and Infectious Disease Working Party recommendations

- 2022 Workshop recommends move to "targeted" antibacterial prophylaxis
- Multiple analyses suggest reduced infections from antibacterial prophylaxis but no overall survival benefit

Trends

- High BSI rate happening despite broad FQ prophylaxis in the US
- Prophylaxis efficacy decreasing as bacteria become resistant to antibiotics
- PTCy adoption for GvHD prophylaxis is increasing BSI rates



* ASCO: American Society of Clinical Oncology; IDSA; Infectious Diseases Society of America; EBMT: formerly known as the European Society for Blood and Marrow Transplantation; PTCy: post-transplant cyclophosphamide Sources: Taplitz et al, *J Clin Oncol* 2018; EBMT 2022 report; Neuerberg et al *Transplant Cell Ther* 2024; Bolaños-Meade et al, *NEJM* 2023

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



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SER-155 Safety: 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)
 - o 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



SER-155 Efficacy: SER-155 associated with 77% relative risk reduction in bacterial BSIs and reduction in systemic antibiotic exposure

Bloodstream infections

Significant decrease in bacterial bloodstream infections in SER-155-treated subjects vs. placebo with **77% relative risk reduction**

Antibiotic exposures

Significantly lower mean cumulative exposure (days) and exposure rate to systemic antibacterials / antimycotics for SER-155-treated subjects vs. placebo

Febrile neutropenia

Numerically lower incidence rate of febrile neutropenia in 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

- 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



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)
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	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

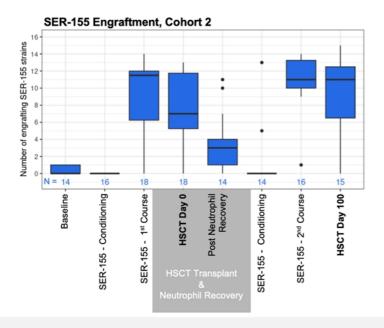
Mean Difference (95% CI)	-0.2 (-0.38, -0.05)	
p-value	0.0163	

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

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



- The majority of SER-155 strains engrafted and were durable through chemotherapy exposure
- · Engraftment decreased but was detectable postneutrophil recovery, suggesting sustained engraftment through period of greatest BSI susceptibility
- · The second course of SER-155 was effective at increasing strain engraftment following neutrophil recovery
- · Engraftment durable out to day 100 following transplant



SER-155 Pharmacodynamics: the Ph1b study included multiple measures of drug pharmacodynamics



Preclinical Optimization

Measurements on Ph1b clinical study



Inhibit Pathogens or "Undesirable" Microbes



Genomic measurements of pathogen abundance in gastrointestinal tract, including domination by pathogenic bacteria



Repair & Protect Mucosal Epithelial Barrier Integrity



Concentration of fecal albumin in subject stool, reflecting barrier damage and leakage of serum albumin into the GI tract



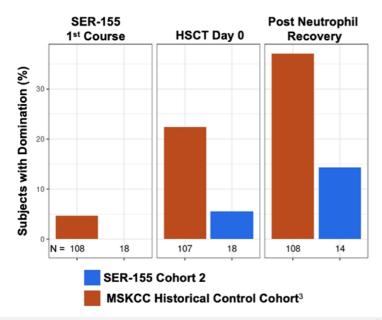
Modulate Immune function



Concentrations of plasma cytokines, chemokines and signaling molecules reflecting GI inflammation, systemic inflammation, and immune homeostasis



SER-155 Pharmacodynamics (pathogen domination): prevalence in SER-155 Cohort 2 was substantially lower relative to Historical Control Cohort



- Pathogen domination (i.e., relative abundance in the GI ≥30%) has been associated with risk of BSIs and other negative clinical outcomes^{1, 2}
- Observed pathogen domination events were low in both the placebo and SER-155 arms with no significant differences observed
 - The ability to detect pathogen domination in the placebo arm was constrained due to the imbalance in the number of available stool samples between the arms.
- 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 ≥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



US HCPs seek significant reduction of infection risk without compromising patient safety and would incorporate such a therapeutic into standard of care

Profile of SER-155
Safety
Etticacy

- Reduce incidence of bloodstream infections: target 20% BSI rate vs. current 40% rate
- Reduce systemic antibiotic use that is contributing to rising resistance
- Reduce incidence of febrile neutropenia: target 60% FN rate vs. current 80% rate
- No major adverse event signals
- Expect translation to fewer deaths in the active study arm

Health economics

Shorten hospital stay, reduce readmissions & reduce ICU utilization

SER-155 Phase 1b data suggest potential to deliver this profile in a pivotal study

SERES THERAPEUTICS**

HCPs see SER-155 as a potentially transformative means to eliminate complications that get in the way of achieving transplant success

Primary Value Driver for SER-155

Reducing the risk of HSCT-related complications, thus ensuring successful engraftment and long-term health of the patient

A relative risk reduction of 50% in BSIs is seen as "transformative" and would support broad inclusion in standard protocols for allo-HSCT patients



Health Care Providers

Streamlines the transplant process so they can spend more time treating the patient's underlying conditions and less time dealing with potential morbidities



Patients

One less thing to worry about for patients already dealing with a lot; additional financial and QoL benefits due to shortened hospital stays



Healthcare System

Reduced healthcare costs due to shorter hospital stays, fewer ICU visits, fewer antibiotic days and lower incidence of severe negative outcomes



The benefit would be massive because people die from these infections and so preventing them, the biggest benefit is mortality. The rest of the stuff with ICU admits and sepsis protocols and all...I think some of that also gets averted. That would be huge."



"This would probably be standard of care. It would be all eligible patients minus those who cannot tolerate it or are allergic."



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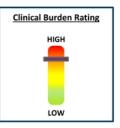
Source: Seres primary market research 2024

US payers confirm high unmet clinical need for infection prophylaxis in allo-HSCT recipients and robust profile of SER-155



High clinical burden and unmet need

- High perceived clinical burden of BSIs within the allo-HSCT patient population is driven by high frequency of occurrences and poor associated outcomes
- Lack of an efficacious prophylactic therapy that successfully limits incidence of BSIs is considered a key unmet need and current driver of clinical burden
- Top unaided clinical concerns raised by payers include risk of febrile neutropenia, sepsis, and antibiotic-resistant infection



Profile of SER-155 positively received

- Proposed risk reduction of BSIs and related endpoints were seen as clinically meaningful and supportive of value proposition
- "The reductions in primary and secondary endpoints are encouraging and brings hope that you are going to escape this risky scary complication of cancer therapy."

Site of dispensing drives reimbursement pathway

hospital setting

· Oral administration is viewed as convenient by payers, with most expecting coverage

under outpatient pharmacy benefit given oral administration likely outside of the



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 has blockbuster commercial potential, driven by a robust SER-155 profile and substantial unmet needs in allo-HSCT and additional groups

- ✓ High unmet need to prevent frequent and serious infections
- √ ~40K annual transplants worldwide; 3% annual growth from aging population and transplant success rates
- ✓ Costly procedure (~\$400K US year 1 allo-HSCT per patient cost) with high incremental costs of infections (incremental ~\$180K/patient)
- ✓ SER-155 has potentially "transformational" profile with robust efficacy and safety
- Highly concentrated universe of transplant centers conducting stem ell transplant procedures allows for an efficient commercial model with rapid education on 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 CMS.gov; Merck SEC filings

Indication expansion: potential 10x addressable population

WW annual diagnoses or transplants

US annual diagnoses or transplants

Unmet needs addressed by SER-155

Allo-HSCT

~40,000

~9,300

Prevent mortality and cost of post-transaction infections

Autologous HSCT

~60,000

~13,500

Prevent mortality and costs of post-transplant infections

Broader leukemia & lymphoma population*

~500,000

~87,000 initial focus

Reduce morbidity, mortality, and cost of infections and febrile neutropenia from chemotherapy

*Includes acute myeloid leukemia, multiple myeloma, and aggressive B cell non-Hodgkin lymphomas (diffuse large B-cell lymphoma, mantle cell, Burkitt's lymphoma Sources: CIBMTR, US NCI SEER, Thandra et al 2021 report of WHO data, Niederwieser et al Haematologica 2022; WHO Global Cancer Observatory; American Cancer Society



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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 with single registrational study for efficacy

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

- Received Breakthrough Therapy designation in December 2024
- · Applied for Orphan Drug designation

Intend to evaluate SER-155 in **additional patient populations** with high risk of serious bacterial infections

Seeking SER-155
strategic partnership to
accelerate next study in
allo-HSCT
and expand to multiple
target populations



30

Advancing SER-147 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%

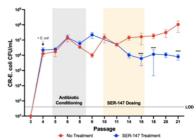
of infections are spontaneous bacterial peritonitis (SBP) and bloodstream infections likely to be gut-seeded

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

Declining E. coli titers

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



SERES

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 of live biotherapeutics going forward

Additional Opportunities

- Prevent life-threatening bacterial infections, including antimicrobial resistant infections in additional populations
- Treat immune-related diseases (e.g., IBD, GvHD, checkpoint colitis)

SER-147

- Infections in Chronic liver disease: Progressing towards IND readiness
- Indication expansion (e.g., radiation enteritis, ICU and long-term care patients, organ transplant)

SER-155

- Bloodstream infections in Allo-HSCT: Engaging with FDA to accelerate; received Breakthrough Therapy designation and filed for Orphan Drug designation
- Evaluate in additional cancer treatment populations with high risk of serious bacterial infections

VOWST

· rCDI: Proven clinical and regulatory success; asset sale to Nestlé; Seres to participate in future milestones



Summary and path forward

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

- · 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 while advancing SER-147
- VOWST approval validates using live biotherapeutics to prevent life-threatening infections

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

- Administration associated with 77% relative risk reduction for BSIs, significant reduction in systemic antibiotic exposure and lower incidence of febrile neutropenia vs placebo; Exploratory biomarker data support epithelial barrier integrity and the potential role for Seres' platform to provide clinical benefit to patients with inflammatory and immune diseases, such as IBD, including ulcerative colitis and Crohns disease
- · FDA Breakthrough Designation obtained
- Company pursuing SER-155 strategic partnership to accelerate next study in allo-HSCT and expand to multiple target populations

VOWST asset sale strengthens financial position

- ~\$31M in cash/cash equivalents at Dec 31, 2024; cash runway projected into Q1 2026
- · Fully retired debt
- VOWST asset sale closed in Sept 2024; \$50M expected to be received in January 2025 and \$25M (less ~\$1.5M in employment-related payments) due in July 2025 + \$275M potential future milestones



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