

Corporate Presentation January 2025

Nasdaq: TSBX

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OUR MISSION

Profoundly transform the treatment paradigm for patients with a broad range of solid tumors with next-generation TIL therapies that overcome the limitations of current treatment options



Solid Tumors Represent a Serious Unmet Medical Need

Approximately 90% of all new cancers per year are solid tumors

In the U.S. Each Year

1.6M new cancer patients¹

500K deaths with low long-term survival¹

90%+

mortality in metastatic disease²

New Therapeutic Options Urgently Needed

Checkpoint inhibitors only benefit a fraction of cancer patients³ Targeted and other cell therapies have shown only limited success One FDA approved TIL therapy and only in advanced melanoma⁴



¹National Cancer Institute's Surveillance, Epidemiology, and End Results (SEER), accessed March 2024; ²Cancer Metastasis: Guan X. Cancer Metastases: Challenges and Opportunities. Acta Pharm Sin B. 2015 Sep;5(5):402-18. ³Haslam A, Prasad V. Estimation of the Percentage of US Patients With Cancer Who Are Eligible for and Respond to Checkpoint Inhibitor Immunotherapy Drugs. JAMA Netw Open. 2019 May 3;2(5):e192535.; ⁴United Stated Food and Drug Association (US FDA) approval granted on 02/16/2024; <u>News release</u>



Turnstone's Next-Generation TILs Are Designed to Address Solid Tumors of High Unmet Medical Need

Amtagvi 1st generation bulk TIL therapy is the first cell therapy approved for solid tumors

FDA NEWS RELEASE

FDA Approves First Cellular Therapy to Treat Patients with Unresectable or Metastatic Melanoma

For Immediate Release: February 16, 2024

Amtagvi is solely indicated for **metastatic melanoma** and bulk TILs have generally **failed to show success in other solid tumors** Next-generation selected TILs are designed to confer more targeted and potent tumor killing



Journal of

Oncoloav^{®3}

Clinical

Cancer Immunotherapy Based on Mutation-Specific CD4+ T Cells in a Patient with Epithelial Cancer

Eric Tran,¹ Simon Turcotte,¹* Alena Gros,¹ Paul F. Robbins,¹ Yong-Chen Lu,¹ Mark E. Dudley,¹† John R. Wunderich,¹ Robert P. Somerville,² Katherine Hogan,¹ Christian S. Hinrichs,¹ Maria R. Parkhurst,¹ James C. Yang,¹ Steven A. Rosenberg¹ ‡

Breast Cancers Are Immunogenic: Immunologic Analyses and a Phase II Pilot Clinical Trial Using Mutation-Reactive Autologous Lymphocytes

Niedoso Zacharakis, PhD⁺; Luffi M. Hug, BA⁺; Samantha J. Setter, DO⁺; Sanghyun P. Kim, PhD⁺; Jared J. Gartner, MSC⁺; Srasahi Sindri, MSC⁺; Vietoris K. Hill, PhD⁺; Yong F. Li, BS⁺; Biman C. Paria, PhD⁺; Salyaf Ray, PhO⁺; Billed Gaumi, MD⁺; Chychia Luce, MD, PhO⁺; Todo D. Pröckett, PhO⁺; Man R. Arnhouts, PhD⁺; Pani, F. Robies, PhD⁻; Michelle M. Langhan, BS⁺; Thomas E. Shetton, BS⁺; Atony Y. Pariki, MD⁺; Shoothan T. Levi, MD⁺; Jonathan M. Hernandez, MD⁺; Chuong D. Hoang, MD⁺; Richard M. Sherry, MD⁺; James C. Tang, MD⁺; Steven A Fedman, PhD⁺; Stephanel E. Gott, MD⁺; at Beren A. Rosehorser, MD, Pi

Turnstone's Selected TILs aim to harness a more reactive T cell population to drive efficacy in broader solid tumor types



Selected TILs Have Potential for More Targeted Tumor Killing





TIDAL-01 Process

Designed to select more potent population of T cells

The TIDAL-01 process is similar to standard bulk TILs but includes selection step designed to create product with significantly higher proportion of tumor-reactive T cells for more effective tumor killing



• Incubate TILs with presented tumor antigens and select tumor-reactive T cells based on activation markers

Turnstone Leadership Team

Proven experience across all areas and stages of drug development



Sammy Farah, MBA, PhD Chief Executive Officer

Sarvah Azmat **Chief Operating Officer**

- 20+ years of scientific, business and executive management experience in the biotech industry.
- Held senior positions at Merck, Immune Design, and Synthetic Genomics.
- Previously at Versant Ventures specializing in biotechnology investing and new company formation.

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VERSANT

- 10+ years of experience in biopharma business development, corporate strategy and capital formation.
- Former Global Lead for Oncology Search & Evaluation at Bristol-Myers Squibb, executing over 15 major transactions from preclinical to clinical development.

(^{III}) Bristol Myers Squibb^{**} putnam



Key expertise in cell therapy technical operations, CMC, and manufacturing.

Michael Fitch, PhD

SVP, Manufacturing

• Served as CMC Operations Lead at Kite Pharma/ **Gilead Sciences** supporting in the approval Yescarta and successful regulatory submissions for other cell therapies.





David Stojdl, PhD SVP, Research and Discovery

- 20+ years of research and drug development expertise spanning cell therapy, cancer biology, drug discovery, virology, and translational science.
- Co-founded Turnstone Biologics in 2014.
- Co-founded Jennerex **Biotherapeutics** which was later acquired by SillaJen **Biotherapeutics**.

ENNER X



Ines Verdon, MD SVP, Clinical Development

- 20+ years of academic and pharmaceutical industry experience.
- Joined from Rubius **Therapeutics** where she was VP of Clinical Development, developing cellular therapies for advanced solid tumors.
- Previously at Sierra **Oncology** as medical lead for several oncology programs across all stages of clinical development.





Wendy Worcester, CPA VP, Principal Finance and Accounting Officer

- 20+ years of financial and accounting industry experience.
- Joined from Gateway **Genomics** and played a key role in completing Turnstone's IPO in 2023 and building its Finance organization.
- Certified Public Accountant (CPA)(inactive) and has held the Chartered Financial Analyst (CFA) designation.

Gateway () Genomics **mesabiotech**



SYNTHETIC GENOMICS



Turnstone External Network

Supported by prominent scientific and corporate advisors and collaborators



Malcolm Brenner, MD, PhD Professor, Center for Cell and Gene Therapy Baylor College of Medicine



Alan Melcher, PhD Team Leader, Translational Immunology The Institute of Cancer Research



Vijay Chiruvolu, MBA, PhD Former CTO Turnstone Biologics / Instil Bio / Kite Pharma



Bernard Fox, PhD Chief, Laboratory of Molecular and Tumor Immunology Providence Cancer Institute



Tassos Gianakakos, MBA Former CEO MyoKardia



Adrian Hill, PhD Director, The Jenner Institute University of Oxford



James Mulé, PhD Associate Center Director of Translational Science Moffitt Cancer Center



Robert Seder, MD Chief, Cellular Immunology Section Vaccine Research Center National Institutes of Health



Eric Tran, PhD ACT Laboratory Lead Providence Cancer Institute



Simon Turcotte, MD, MSc Associate Professor of Surgery; Lead of Adoptive T Cell Cancer Immunotherapy Program University of Montreal Hospital Research Centre (CRCHUM)



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Turnstone Pipeline

Opportunity to address broad set of solid tumor patient populations

	Program	Product Overview	Key Indications	Preclinical	Phase 1	Phase 2	Phase 3	Next Anticipated Milestone
Selected TILs	TIDAL-01	Tumor-reactive Selected TILs	CRC; HNSCC; Uveal melanoma					Clinical program update in 1H 2025
			CRC; HNSCC; Uveal melanoma	Moffitt Collaborat	ion*			

*Two concurrent investigator sponsored trials at Moffitt Cancer Center

CRC = Colorectal cancer; HNSCC = Head and neck squamous cell carcinoma





TIDAL-01 Initial Clinical Data



Turnstone is Pioneering Advancements in Selected TIL Therapy

Turnstone is building on success of TIL therapy in melanoma by developing **next-generation Selected TILs** to address additional high-medical need solid tumors



TIDAL-01 is enrolling in multiple Phase 1 trials in solid tumors including CRC, head and neck cancer, and uveal melanoma

CRC = Colorectal cancer



Initial interim data in CRC demonstrated promising clinical evidence and supports our biological hypothesis



Results to date help to **focus our strategy** for manufacturing and future clinical development to advance new treatment for metastatic CRC



TIDAL-01 Phase 1 Clinical Trials in Advanced Solid Tumors



Phase 1 Study

Demonstrate the safety, biology, initial efficacy and manufacturing feasibility of TIDAL-01 in a Phase 1, first-in-human, non-randomized, open-label, single-dose study in patients with advanced solid tumors



Phase 1 Clinical Trials in Multiple Tumor Types Are Ongoing

TURNSTONE

Turnstone-sponsored trial (STARLING) enrolling across 10+ clinical sites

- Colorectal cancer (CRC)
- Head and neck cancer (HNSCC)
- Uveal melanoma



Two investigator-sponsored trials in collaboration with Moffitt Cancer Center

- Colorectal cancer (CRC)
- Head and neck cancer (HNSCC)
- Uveal melanoma



The initial clinical data comprises the **first 4 evaluable CRC patients** from the **STARLING** trial

Lead Indication: Colorectal Cancer



Difficult-To-Treat Tumor Unresponsive To Most Immune-Based Therapies

Immunologically "cold" tumor characterized by low tumor mutational burden (TMB) TURNST NE BIOLOGICS

We believe the key to overcoming challenges of CRC and other "cold" tumors is increasing the number of ontarget tumor-reactive T cells which is the foundation for Turnstone's Selected TIL therapy



Limited New Therapies in Development for Metastatic CRC

Dearth of Options in Development for Metastatic MSS CRC

- Little to no novel treatment options or new modalities in development for large population of metastatic MSS¹CRC patients
- Promising data for immunotherapies, particularly for anti-CTLA4 (including modified / enhanced versions) ± anti-PD-1 in advanced development for CRC have been limited to:
 - MSI² CRC (immune sensitive) where anti-PD-1 already approved as first line treatment
 - Local (non-metastatic) CRC in neoadjuvant setting
 - MSS mCRC without active liver metastases (which are highly prevalent in CRC patients)

Bulk TILs Have Not Demonstrated Clinical Benefit in CRC

()ITC

Efficacy and safety of autologous tumorinfiltrating lymphocytes in recurrent or refractory ovarian cancer, colorectal cancer, and pancreatic ductal adenocarcinoma

MDACC - Rodabe et al.; JITC 2024

- 2024 MDACC publication showed 0 responses in 8 patients with recurrent / metastatic CRC treated with bulk TILs
- Historical NCI data³ in 50+ patients treated with bulk TILs across various solid tumors (including CRC, Bile Duct, Pancreas, Breast, Gastric) demonstrated no clinical success

MSS indicates Microsatellite Stable form of CRC which is typically non-responsive to checkpoint inhibitors and represents up to 95% of metastatic colorectal cancer patients
MSI indicates Microsatellite Instable form of CRC which can be responsive checkpoint inhibitors and represents approximately 5% of metastatic colorectal cancer patients
Rosenberg AACR 2020 / NCT01585428



Academic Clinical Studies with Selected TILs in Metastatic CRC Provide Evidence of Activity



In the single patient NCI study published in NEJM in 2016 by Tran et al., selected TILs demonstrated **a deep and durable PR in metastatic CRC** NCI – Parkhurst et al; NatMed 2024 In NCI study published in NatMed in July 2024 by Parkhurst et al., selected TILs used to generate product for 7

metastatic MSS CRC patients resulted in 3 deep PRs

Adoptive transfer of personalized neoantigenreactive TCR-transduced T cells in metastatic

colorectal cancer: phase 2 trial interim results

Maria Parkhurst @ 🖂, Stephanie L. Goff O, Frank J. Lowery, Rachel K. Beye

Sivasish Sindiri. Sri Krishna 🛛 , Nikolaos Zacharakis, Lien Ngo, Satyajit Ray

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Article

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Accented: 4 June 2024

Turnstone's preliminary data combined with academic clinical studies with selected TILs create compelling emerging picture of the very promising potential of selected TILs to treat metastatic CRC



Colorectal Cancer Represents Large Medical Need and Significant Market Opportunity

Advanced metastatic MSS CRC (MSS mCRC) is typically treated with chemotherapy (FOLFIRI/FOLFOX) +/- combination with bevacizumab or anti-EGFR, but resistance to chemotherapy is inevitable^{1,2}

Targeted therapies are available, however most MSS mCRC patients do not have actionable mutations²

Biomarker	Frequency	
BRAF-V600E	5-10%	
HER2 Overexpression	2-3%	
KRAS-G12C	3%	

For patients lacking actionable mutations, OR that fail targeted therapies, treatment efficacy outcomes after exhausting chemotherapy are poor³⁻⁵



NCCN Guidelines Version 1.2024.
Am Soc Clin Oncol Educ Book 43, e389574(2023).
Grothey A, et al. Lancet. 2013; 381: 303-312.
Prager GW, et al. <u>N Engl J Med</u> 2023;388:1657-6
Dasari, A et al. Lancet 2023; 402: 41–53.



Promising Initial Results in MSS mCRC Patients Observed in TIDAL-01

4 MSS mCRC patients evaluable for response¹

- One patient with deep and durable confirmed Partial Response (PR) that converted into Complete Response (CR) at last scan at week 48
- One patient with confirmed Stable Disease (SD) at last scan at week 24

Patient	Best Response (based on independent central reads)	
CRC 1	Complete response (Week 48)	
CRC 2	Stable disease (Week 24)	
CRC 3	Progressive disease (Weeks 6 and 12)	
CRC 4	Stable disease (Week 4)	



Ongoing response and SD are notable in relapsed MSS mCRC population, where patients often have almost no viable treatment options⁴⁻⁶

Six MSS mCRC patients were dosed on STARLING but one was lost to follow-up (not evaluable for response), one patient scan data pending as of data cut-off on August 14 2024.
Patient is off study and started new treatment in consultation with treating physician.
Only one scan available for patient with a central scan reading of SD at week 4, patient has been off study since.
Grothey A, et al. Lancet. 2013; 381: 303-312.
Prager GW, et al. <u>N Engl J Med</u> 2023;388:1657-6
Dasari, A et al. Lancet 2023; 402: 41–53.



CRC Patient 1: Treating Advanced Disease in 3rd Line MSS mCRC

Progressed after 2 lines of prior standard chemotherapy regimens and in need of new treatment option



- 55-year-old female
- Diagnosed August 2021 with Stage IIIb node-positive CRC
- MSS CRC with no actionable mutations (KRAS/NRAS/BRAF wild type)
- 30 pack year smoker, shortness of breath, neuropathy from chemotherapy, PE, GERD

Date	Patient Treatment History
8/21	Neoadjuvant chemoradiotherapy
11/21	1L FOLFOX for 3 months until PD (new retroperitoneal nodes)
5/22	2L FOLFIRI + bevacizumab for 2 months with SD
9/22	Surgery (low anterior resection)
3/23	Mediastinal, liver and lung metastases
Note: patie	ent also received bridging therapy of 2 cycles FOLFIRI+

Note: patient also received bridging therapy of 2 cycles FOLFIRI+ panitumumab prior to washout period and receiving TIDAL-01

1L = First line; 2L = Second line; AE = Adverse event; CRC = Colorectal cancer; GERD = Gastroesophageal reflux disease; LAR = Low anterior resection; MSS = Microsatellite stable; PD = Progressive disease; PE = Pulmonary embolism; PY = Pack year; SD = Stable disease; TCR = T cell receptor; Wk = Week.



CRC Patient 1: Complete Response in MSS mCRC with Continuing Durability Through 48 Weeks





- Central scan reported PR starting at 12 weeks that deepened over time and converted into CR at 48 weeks
- Circulating tumor DNA declined to zero and has remained negative as of last ctDNA reading at week 36
- Response has been durable and patient has been progression free for one year
- Patient reported feeling clinically well and had her stoma reversed





CRC Patient 1: Product Characterization and Translational Data Support Biological Hypothesis for Selected TILs

Selected TIL product for CRC Patient 1 had **our targeted product characteristics and biological activity** and these **successfully translated** into strong clinical response



DP = Drug product; LD = Lymphodepletion; REP: Rapid expansion protocol.



TIDAL-01 Manufacturing Success Rate and Safety Are Consistent with Other Early Clinical Stage Cell Therapies



TIDAL-01 manufacturing success rate in CRC for patients with sufficient starting material¹



Target dose of 1E9 cells was exceeded in all CRC products manufactured

Patient	Dose		
CRC 1	3 x 10 ¹⁰ cells		
CRC 2	2 x 10 ¹⁰ cells		
CRC 3	6 x 10 ⁹ cells		
CRC 4	1 x 10 ¹⁰ cells		



No new or unexpected safety observations specific to Selected TILs



Safety profile was consistent with known AE profiles associated with the lymphodepletion regimen, IL-2 and pembrolizumab

Patient	Treatment regimen related SAEs
CRC 1	SAE ² G3 ³ pneumonitis related to pembro, resulted in pembro discontinuation
CRC 2	SAE G3 diabetic ketoacidosis related to pembro, resulted in pembro discontinuation
CRC 3	No treatment related SAEs observed
CRC 4	No treatment related SAEs observed

1. Insufficient starting material denotes patients that had very little or no TILs in their resected tissue at the time of starting the manufacturing process resulting in inability to make a TIL product - of note, this issue is not unique to the selected TIL process and the nature of the starting material would not yield a successful bulk TIL product either. 2. SAE = Serious Adverse Affect 3. G3 = Grade 3



Opportunity for Compelling Clinical Development Strategies



1 Build off preliminary success and continue to advance our Phase 1 study by **targeting enrollment** of CRC patients we believe are most likely to benefit from TIDAL-01 therapy

We also intend to develop TIDAL-01 in **earlier lines of treatment** in combination with chemotherapy to potentially treat an even more responsive patient population

We believe completion of Phase 1 study including expansion cohort has **potential to lead to registrational intent study in next 18-24 months**

Additional Indications

We are simultaneously advancing our Phase 1 studies in **head and neck cancer** and **uveal melanoma**



Selected TILs Have the Potential to be Transformative in the Treatment of CRC

TIDAL-01 shows promising clinical benefit in metastatic MSS CRC:

- In the first 4 patients we have seen a 25% overall response rate (ORR) with durable clinical benefit and 50% disease control rate (DCR) in a setting where patients are unresponsive to checkpoint inhibitors and have almost no treatment options
- No new safety signals were observed with TIDAL-01 and we have demonstrated the ability to successfully manufacture the product

TIDAL-01 findings are corroborated by academic clinical data for selected TILs in metastatic CRC:

 Clinical data from multiple studies at the NCI with selected TILs have also demonstrated responses in patients with metastatic CRC, strengthening the body of data for the approach in this indication

Taken together, the TIDAL-01 and academic clinical data demonstrate the potential for selected TILs to be a breakthrough treatment to address the large unmet medical need for patients with metastatic CRC



Building on the Promise of Selected TILs in Solid Tumors



Turnstone is building on success of recently FDA-approved TIL therapy in melanoma by developing **next-generation Selected TILs** to address additional high-medical need solid tumors



TIDAL-01 is currently enrolling Phase 1 trials in multiple solid tumor types including CRC, head and neck, and uveal melanoma



Initial TIDAL-01 interim data demonstrated promising clinical evidence for CRC supported by our biological hypothesis



Results to date help to **focus our TIDAL-01 strategy** for manufacturing and future clinical development to advance this potential new treatment for metastatic CRC





Thank You