

**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 10, 2022

Sensei Biotherapeutics, Inc.

(Exact Name of Registrant as Specified in its Charter)

Delaware
(State or Other Jurisdiction
of Incorporation)

001-39980
(Commission
File Number)

83-1863385
(IRS Employer
Identification No.)

1405 Research Blvd, Suite 125
Rockville, MD
(Address of Principal Executive Offices)

20850
(Zip Code)

Registrant's telephone number, including area code: (240) 243-8000

N/A
(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 Securities Exchange Act of 1934:

Title of each class	Trading symbol	Name of each exchange on which registered
Common Stock	SNSE	The Nasdaq Stock Market LLC

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.

During the week of January 10, 2022, members of management of Sensei Biotherapeutics, Inc. (the “**Company**”) will hold meetings to provide an overview of the Company. A copy of the presentation that will accompany the meetings is furnished as Exhibit 99.1 to this Current Report on Form 8-K.

The information in this Item 7.01 of this Current Report on Form 8-K (including Exhibit 99.1) is being 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 a filing.

Item 9.01 Financial Statements and Exhibits.

(d) Exhibits

<u>Exhibit Number</u>	<u>Exhibit Description</u>
99.1	Company Presentation.
104	The cover page from Sensei Biotherapeutics, Inc.’s Form 8-K filed on January 10, 2022, formatted in Inline XBRL.

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 10, 2022

Sensei Biotherapeutics, Inc.

/s/ John Celebi

John Celebi

President and Chief Executive Officer



Training the Immune System to Fight Cancer

John K. Celebi, MBA
President & Chief Executive Officer

JP Morgan Healthcare Conference
January 10, 2022

NASDAQ: SNSE

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2021



This presentation has been prepared by Sensei Biotherapeutics, Inc. (the "Company," "we," "us") and is made for informational purposes only. The information set forth herein does not purport to be complete or to contain all of the information you may desire. Statements contained herein are made as of the date of this presentation unless stated otherwise, and neither the delivery of this presentation at any time, nor any sale of securities, shall under any circumstances create an implication that the information contained herein is correct as of any time after such date or that information will be updated or revised to reflect information that subsequently becomes available or changes occurring after the date hereof.

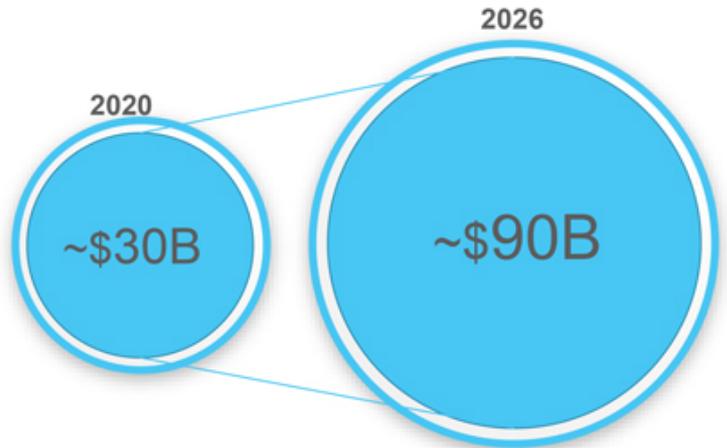
This presentation contains estimates and other statistical data made by independent parties and by us relating to market shares and other data about our industry. This presentation also contains "forward-looking" statements as that term is defined in the Private Securities Litigation Reform Act of 1995 that are based on our management's beliefs and assumptions and on information currently available to management. These forward-looking statements include, without limitation, statements regarding our industry, business strategy, plans, the preclinical and clinical development of our product candidates, and other financial and operating information. When used in this presentation, the words "may," "believes," "intends," "seeks," "anticipates," "plans," "estimates," "expects," "should," "assumes," "continues," "could," "will," "future" and the negative of these or similar terms and phrases are intended to identify forward-looking statements. Forward-looking statements involve known and unknown risks, uncertainties and other 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. Risks and uncertainties that may cause actual results to differ materially include uncertainties inherent in the development of therapeutic product candidates, such as preclinical discovery and development, conduct of clinical trials and related regulatory requirements, our reliance on third parties over which we may not always have full control, and other risk and uncertainties that are described in our Annual Report on Form 10-K filed with the SEC on March 30, 2021 and our other Periodic Reports filed with the SEC. Forward-looking statements represent our management's beliefs and assumptions only as of the date of this presentation and include all matters that are not historical facts. Our actual future results may be materially different from what we expect. Except as required by law, we assume no obligation to update these forward-looking statements publicly, or to update the reasons actual results could differ materially from those anticipated in the forward-looking statements, even if new information becomes available in the future.

Certain information contained in this presentation relates to, or is based on, studies, publications, surveys and other data obtained from third-party sources and the Company's own internal estimates and research. While the Company believes these third-party sources to be reliable as of the date of this presentation, it has not independently verified, and makes no representation as to the adequacy, fairness, accuracy or completeness of, any information obtained from third-party sources. In addition, all of the market data included in this presentation involves a number of assumptions and limitations, and there can be no guarantee as to the accuracy or reliability of such assumptions. Finally, while we believe our own internal research is reliable, such research has not been verified by any independent source.

Majority of patients don't respond to PD-1/PD-L1 monotherapy¹



Global PD-1/PD-L1 Market²



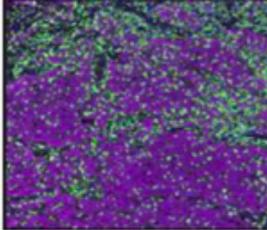
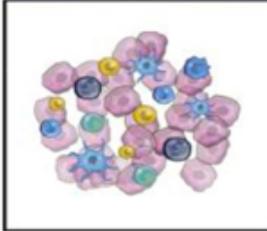
1. Gerber et al., Biochemical Pharmacology 2016
2. Market estimates from PD-1 and PDL-1 Inhibitors Market Size in 2021 – MarketWatch, 360 Research

Two Major Types of Non-Responders to PD-1 Blockade

Responders

T-cells Inside Tumor

Hot (inflamed) tumor

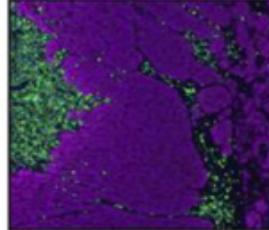
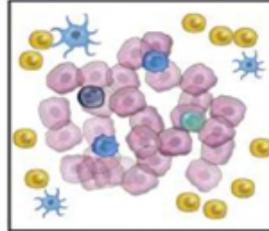


Green = T-cells
Purple = tumor

Non-Responders

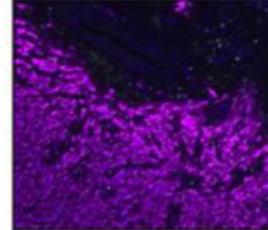
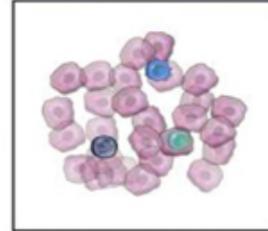
T-cells Inactive or
Outside Tumor

Cold (excluded) tumor



T-cells Absent

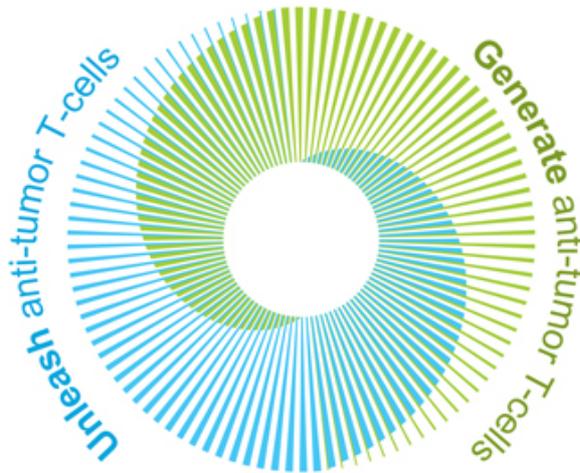
Cold (ignored) tumor





TMAb™ (Tumor Microenvironment Activated Biologics) Platform

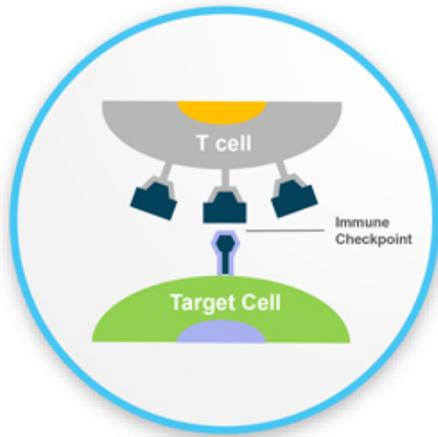
- Next-generation tumor activated mAbs
- Binding only in the low-pH tumor microenvironment
- Target checkpoints and/or other immune pathways
- Enable improved PK/PD and toxicity profiles



ImmunoPhage™ Platform

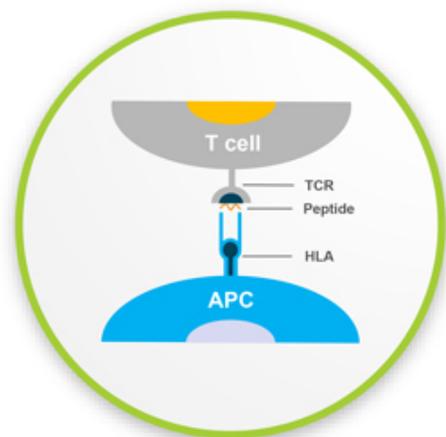
- Powerfully self-adjuvanted nanoparticle vaccine can drive B cell and T cell responses
- Multi-antigen vaccine enables personalized approach from "off-the-shelf" components
- Targets APCs
- Enhanced through addition of immunostimulatory nanobodies & cytokines

TMAb



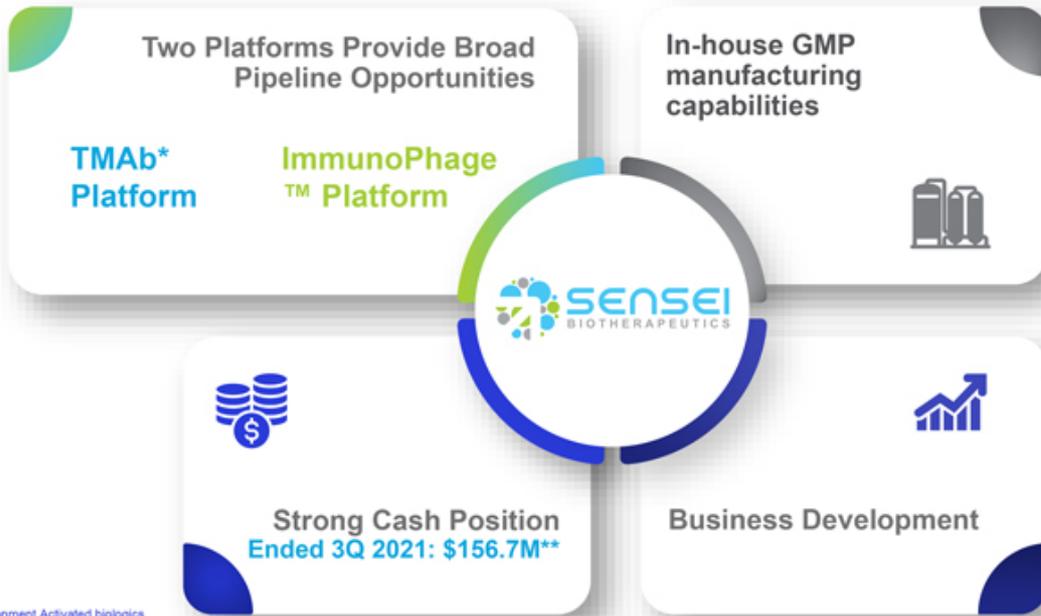
Focus on novel immune checkpoints to **UNLEASH** anti-tumor T-cells

ImmunoPhage™



Focus on multi-antigen approach to **GENERATE** anti-tumor T-cells

Positioned to Drive Value with Next Generation Product & Platform Development



*Tumor Microenvironment Activated biologics
**Consists of cash, cash equivalents and marketable securities

Pipeline Utilizing Pioneering ImmunoPhage Platform, TMAb Platform



	Program (Target)	Indication	Discovery	IND-enabling	Phase 1 / 2 Clinical
 TMAb	SNS-101 (VISTA)	Solid Tumors			
	SNS-102 (VSIG4)	Solid Tumors			
	SNS-103 (ENTPDase1/CD39)	Solid Tumors			
 ImmunoPhage	SNS-401-NG (Multiple Tumor Antigens)	Merkel Cell Carcinoma			
		Head and Neck Cancer			
		Lung Cancer			
		Melanoma			
		Breast Cancer			

TMAb (Tumor Microenvironment Activated biologics) Platform

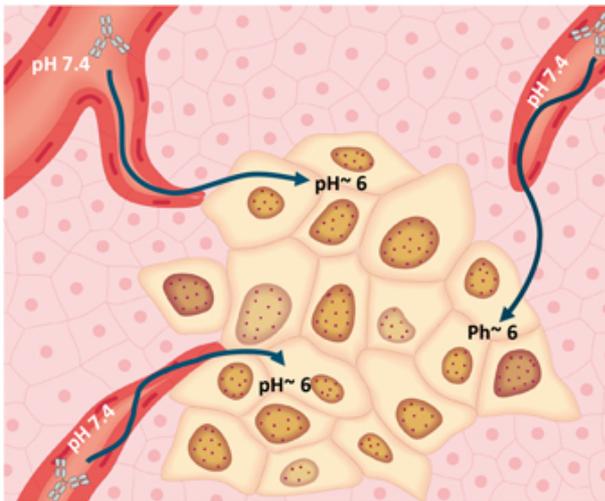


pH-sensitive Antibodies Selectively Bind Their Targets in the Low-pH Tumor Microenvironment

TMAb PLATFORM

The tumor microenvironment of pH ~6 is lower than physiological pH of 7.4

Sensei's technology identifies pH-sensitive antibodies that bind only at the tumor



- Antibodies that bind at physiological pH may encounter a “sink”
 - Prevents effective binding at the tumor and may lead to toxicity
- TMAb antibodies bypass tissue compartments other than the low-pH tumor microenvironment
- Potential for improved safety and clinical activity profile

Targeting Immunosuppressive myeloid cells is a promising strategy to overcome resistance to checkpoint Inhibitor therapy

THE PROMISE

- Using the body's own immune system to attack cancer
- Capitalizing on immunological specificity and long-term memory
- Achieving durable cures with minimal toxicity

THE CHALLENGE

- 70-80% of patients do not achieve increased survival with CPI monotherapy¹
- The immunosuppressive tumor microenvironment (TME) influences response to immune checkpoint blockade
- Innate immune cells such as myeloid cells are a key driver of immunosuppressive TME

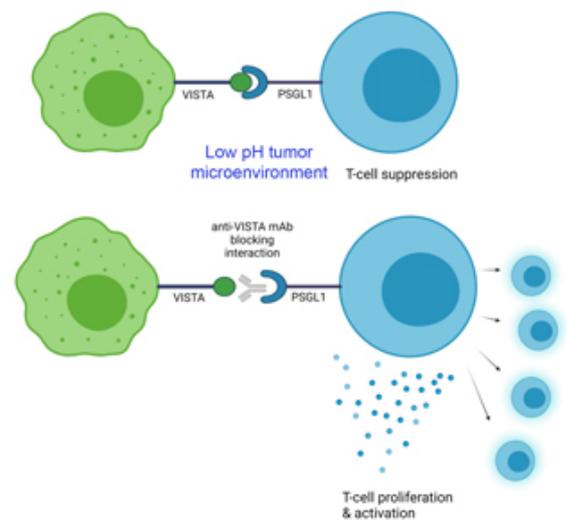
Target Overview:

- Large market opportunity
- B7 family ligand
- Extensive expression on myeloid cells¹
- Inhibition of VISTA may lead to activation of myeloid cells
- Excellent therapeutic combinability with CTLA-4 or PD-1/PD-L1 ICIs, especially in cold tumors²
- VISTA expression correlates with poor survival rates across multiple cancers
- Novel development program with no approved therapies

Sensei's Competitive Advantage:

- Extensive understanding of VISTA biology and differentiated candidate antibody

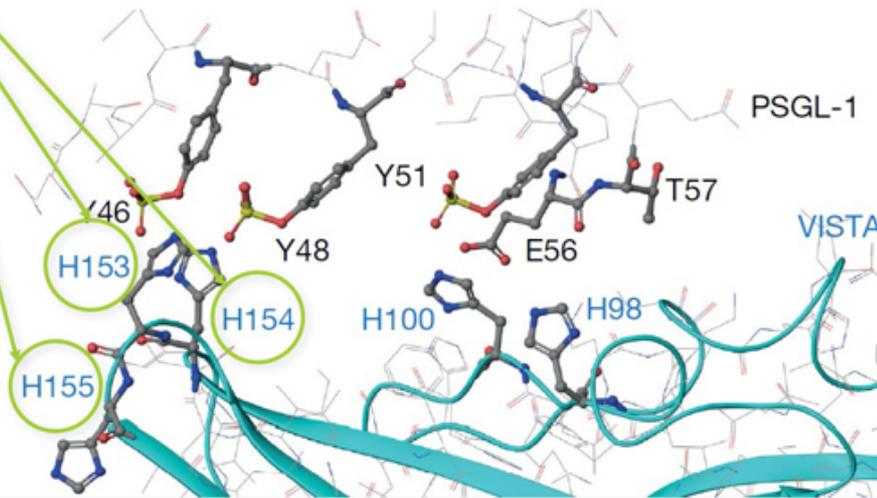
VISTA is a Negative Regulator of T cell Function



¹ Lines et al. Cancer research vol. 74,7 (2014)
² Gao et al. Nature medicine vol. 23,5 (2017)

VISTA Checkpoint is Activated at the Low pH of the Tumor Microenvironment

Antibodies that block protonated VISTA histidines interrupt PSGL-1 binding¹



VISTA's extracellular domain is uniquely rich in histidines¹

Histidines are protonated at low pH enabling VISTA to distinguish the active (acidic pH) and inactive (neutral pH) PSGL-1 binding interface

1. Block the pH-dependent binding of VISTA to PSGL-1 on T cells at low pH
2. Selectively bind VISTA at low pH to avoid:
 - target mediated drug disposition (TMDD)
 - on-target/off-tumor side effects
3. Utilize an Fc-competent IgG backbone to engage and activate Fc γ R on tumor-infiltrating myeloid cells

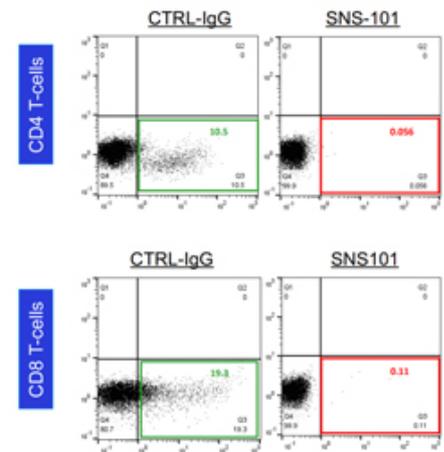
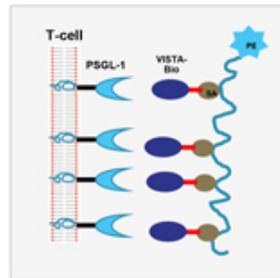


SNS-101 Inhibits Interaction of VISTA to its Receptor, PSGL-1, in CD4/CD8 T-Cells at Low pH 6.0

SNS-101:

- Fully human monoclonal antibody that selectively binds active (low pH) VISTA, but not inactive VISTA in the blood
- No significant binding to VISTA+ monocytes at pH 7.4
- Potent inhibitor of PSGL-1 binding to VISTA
- Fc-competent framework to deliver positive “kick” to help convert myeloid cells in the TME from an immunosuppressive to a proinflammatory state

PSGL-1: VISTA Interaction on Primary T-cells at pH 6.0

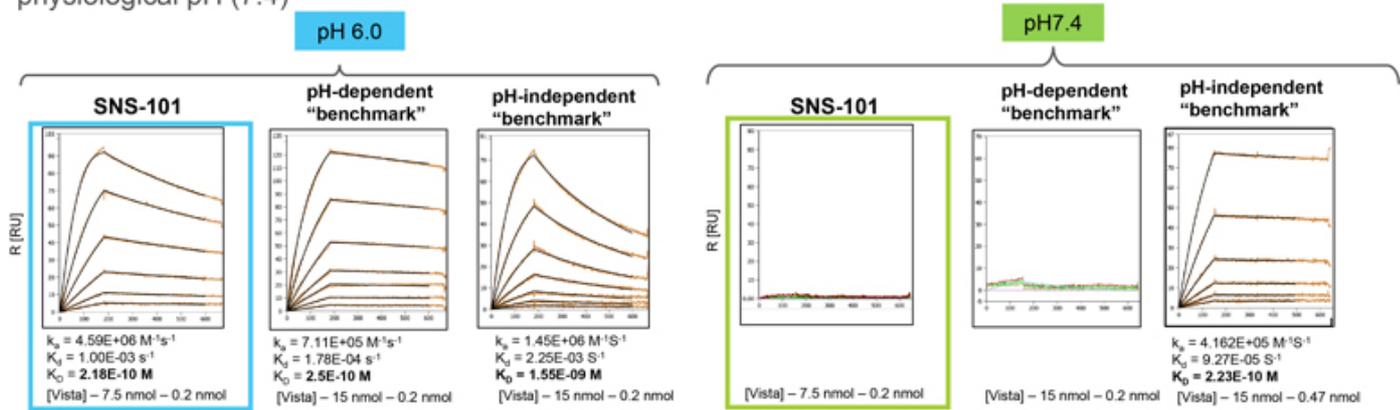


IND-Enabling Studies are Underway for SNS-101

SNS-101 Has >600-Fold Selectivity for VISTA^{pH6}

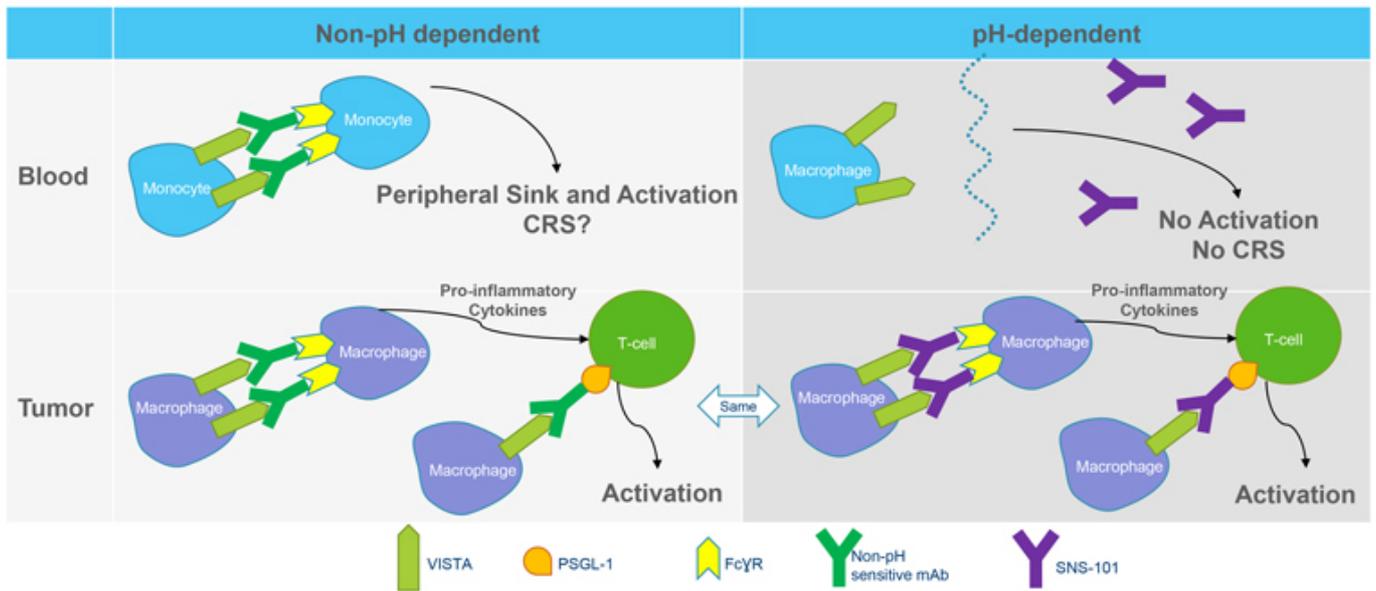
- Biophysical characterization demonstrates >600-fold selectivity for VISTA at pH 6.0
- Picomolar binding at low pH
- No significant binding observed at physiological pH (7.4)

	pH 6.0	pH 7.4
Monovalent Affinity (K_D) [nM]	0.218	132 (~No binding)



Proposed Mechanism of Action for SNS-101

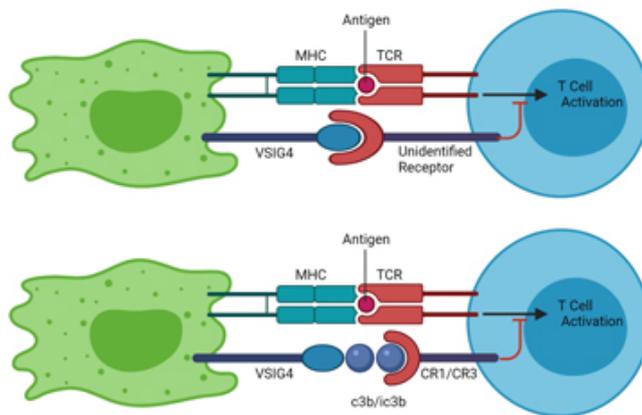
Fc-competent framework is required for optimal activity, but FcγR engagement in the blood may result in untoward “off tumor” activation (i.e. CRS)



SNS-101 Is a Differentiated Anti-VISTA Antibody

TMAb Platform

		VISTA.18 (BMS)	KVA12.1 (Kineta)	CI-8993; JNJ-61610588 (J&J/Curis)	K01401-020; W0180 (Pierre Fabre)	HMBD-002 (Hummingbird)
Inhibit PSGL-1 Binding	Yes	Yes	unknown	Yes	unknown	unknown
pH Sensitive Binding	Yes	Yes	No	No	No	No
Fc Active	Yes (IgG1)	No (IgG4)	Yes (IgG1)	Yes (IgG1)	N/A	No (IgG4)
Stage	Preclinical	Preclinical	Preclinical	Phase I	Phase I	IND submission
Clinical Data / Notes	<ul style="list-style-type: none"> Preclinical data presented at STIC IND-enabling studies underway 	<ul style="list-style-type: none"> N/A 	<ul style="list-style-type: none"> N/A 	<ul style="list-style-type: none"> JNJ initiated Phase I study in 2016 12 pts enrolled; initial dose 0.005 mg/kg Only patient treated at 0.3 mg/kg experienced grade 3 CRS-associated encephalopathy; trial was halted 	<ul style="list-style-type: none"> Ongoing 	



No approved therapies against VSIG4

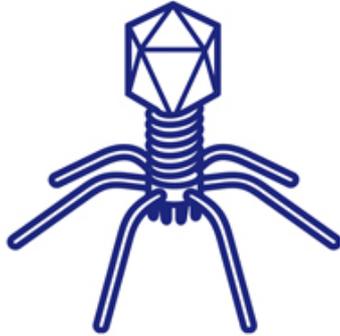
Adapted from Zang et al., J Clin Invest. 2006

- Second TMAb program
- B7 family related protein
- Expressed on macrophages
- Inhibits T-cell activation
- Novel therapeutic combinability with existing IO drugs

ImmunoPhage™ Platform

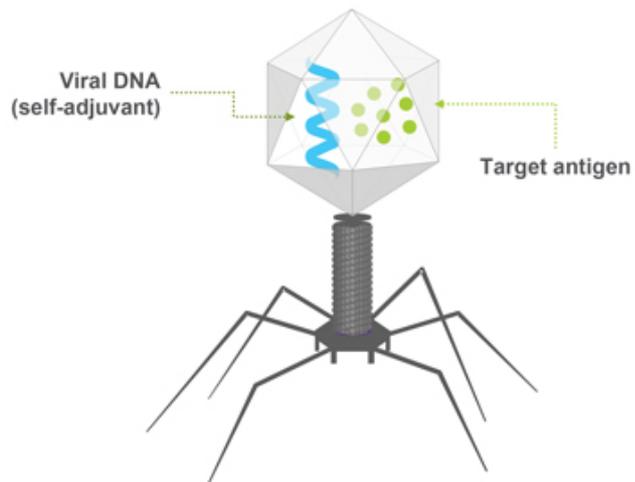


Bacteriophage



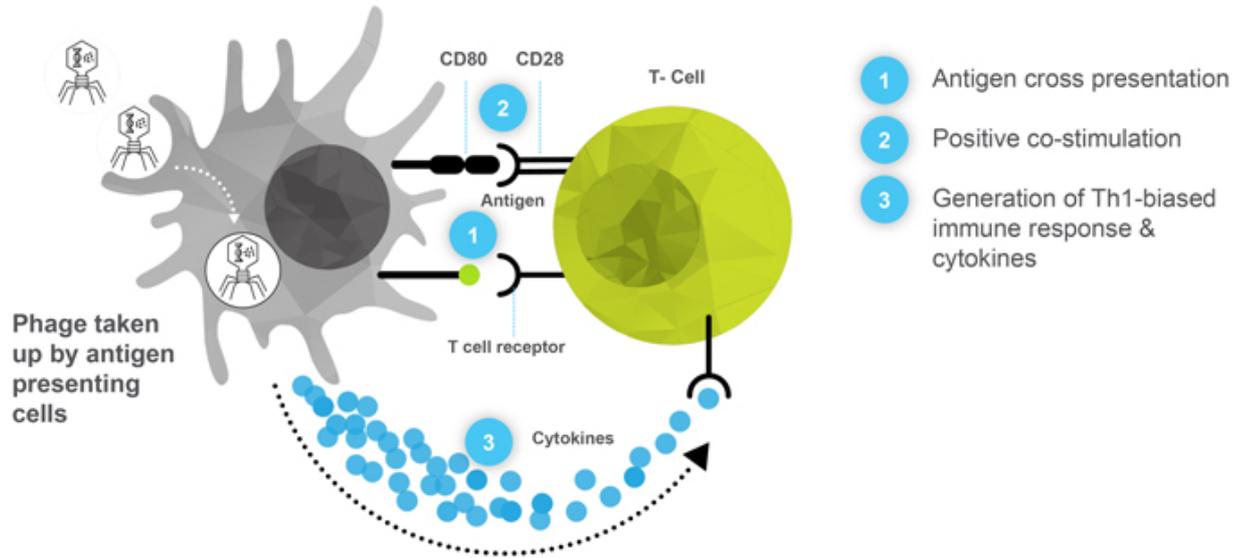
Ubiquitous viruses that infect bacteria but not mammalian cells. Adept at activating the human immune system in multiple unique ways

Bacteriophage virus is engineered and manufactured with both antigen and immune stimulatory viral DNA



The **ImmunoPhage™** bacteriophage is an icosahedron with a tail. This configuration can be viewed as an activating signal to the immune system

ImmunoPhages are taken-up by APCs and deliver three critical signals required to drive activation of T cells.



Our **ImmunoPhages** can mount a multi-modal attack on cancer, combining the benefits of a traditional vaccine with localized gene therapy

Targeted therapeutic vaccine

- MHC-mediated immunity
- Bacteriophage have natural tropism for APCs
- Can be further targeted to APCs with non-antigen capsid modifications



Phortress™ library

- Personalized - yet off the shelf - medicines
- Pre-manufactured cost effectively - then combined based on genetic profile

Gene therapy vehicle

- Phage containing self-replicating RNA
- Used to deliver payloads consisting of immunomodulatory proteins or nanobodies



Collaboration with University of Washington to build **first custom Merkel Cell Carcinoma (MCC) vaccine cocktail consisting of Merkel Cell Polyoma Virus epitopes** and other patient specific antigens

MCC is a rare, aggressive neuroendocrine skin cancer

- 33-46% disease-specific mortality
- 2,500 cases/yr with disease-specific mortality approaching 50%
- Vaccine combination therapy in adjuvant or neoadjuvant is attractive and feasible
 - PD-1/PD-L1 refractory MCC remains unmet medical need with aggressive clinical course
 - ~40% MCC patients recur <24 months following definitive local treatment

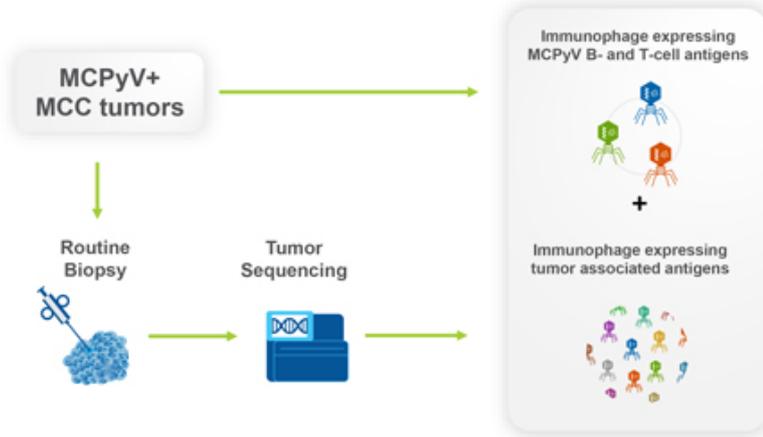
Integration of MCPyV is present in ~80% of U.S. cases

- In these cases, expression of a viral antigen (oncogenic T-antigen) **appears to be a strictly required tumor driver**
- Researchers at UW have mapped MCPyV epitopes and **determined CD8 T-cell, CD4 T-cell, and B-cell epitopes that are antigenic** in the context of MCPyV+ MCC tumors.

SNS-401-NG has Potential to be First Fully Customized, Yet Off-the-Shelf, Therapy

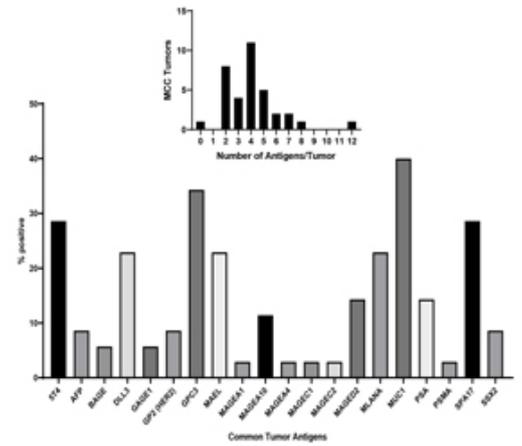
SNS-401-NG Development in Merkle Cell

Patients would receive a bespoke mixture of ImmunoPhage that included antigens from the MCPyV and a subset of TAA-expressing ImmunoPhage



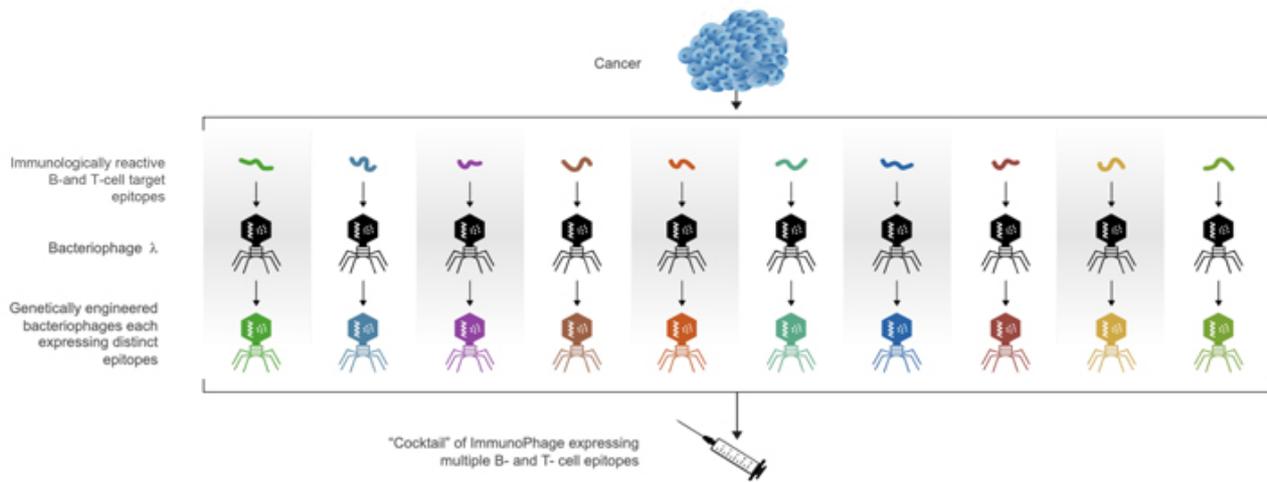
1. Based on internal data

Most MCC tumors contain multiple TAAs¹



Common Tumor Antigens

Phortress: Proprietary Library of Personalized Vaccine Cocktails with Off-the-Shelf ImmunoPhage “Ingredients”



- These “cocktails” are defined by the disease or patient genetics

- Combinations are customized to cover multiple epitopes, protein domains or targets

- Each *ImmunoPhage* is pre-manufactured to target a discrete antigen

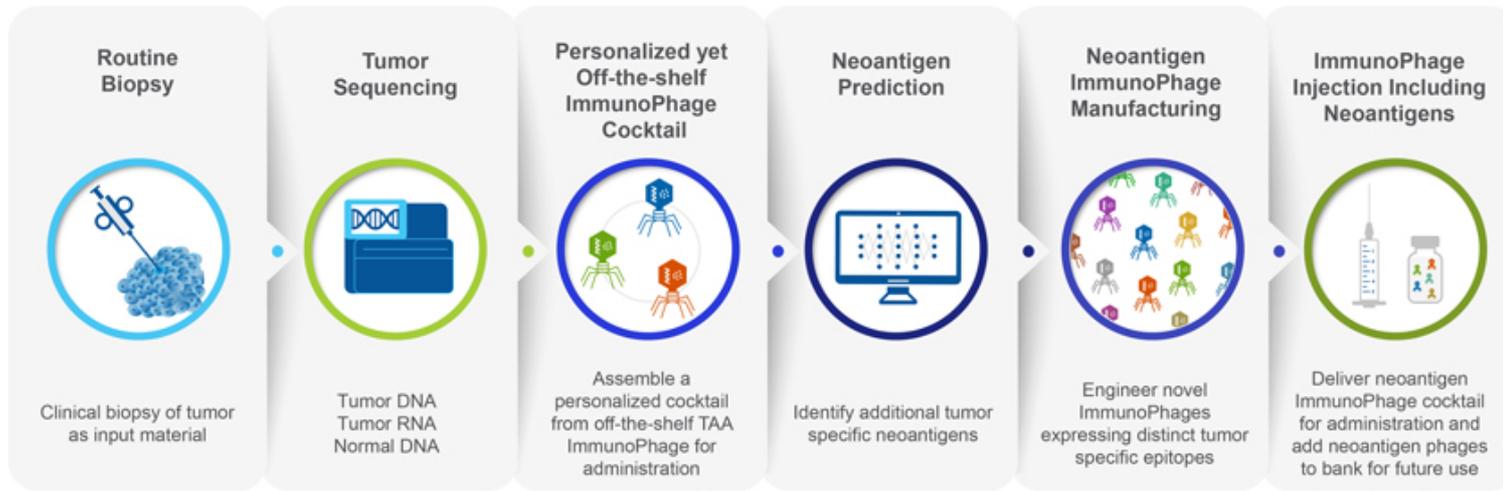
Personalized Immunotherapy Approach Could Accelerate Speed to Treatment



High speed and low cost-of-goods of ImmunoPhage allows a broader array of antigens

Personalized yet Off-the Shelf TAA Therapy

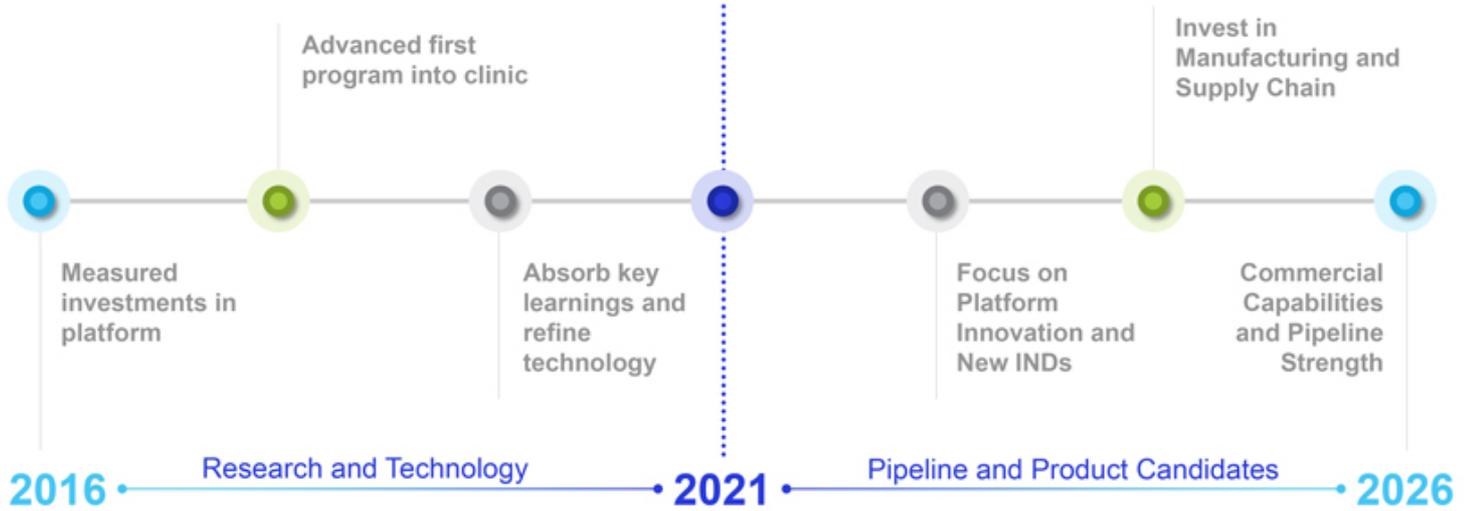
Off-the-Shelf + Patient-specific Neoantigen Therapy



Sensei's Vision to Capture Platform and Pipeline Value



Feb 2021: IPO



Proven Team With Deep Experience



John Celebi, MBA
President and CEO



Michael Boychyn, PhD
SVP, Technical Operations



Pauline Callinan, PhD
VP, Business Operations and Strategy



Jean Campbell, PhD
VP, Biologics Discovery



Robert Pierce, MD
Chief R&D Officer



Elisabeth Colunio
VP, Human Resources



Alice Drumheller
VP, Clinical Operations



Bao Le
VP, Regulatory



Erin Colgan
Chief Financial Officer



Edward van der Horst, PhD
SVP, TMAb Antibodies



SNS-101 (anti-VISTA)

- 1H 2023: IND filing
- 2022: Publish data demonstrating preclinical profile



SNS-401-NG

- 2H 2022: Initiate IND-enabling studies



SNS-102 (anti-VSIG4)

- 2023: Select product candidate



SNS-103 (anti-ENTPDase1/CD39)

- 2023: Select product candidate



Training the Immune System to Fight Cancer

John K. Celebi, MBA
President & Chief Executive Officer

JP Morgan Healthcare Conference
January 10, 2022

NASDAQ: SNSE

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