



gritstone  
bio

CORPORATE PRESENTATION

# The Promise of Potent and Durable Immune Responses

October 2024

# Legal Disclaimer: Safe Harbor and Forward-Looking Statements

This presentation contains forward-looking statements including, but not limited to, statements related to Gritstone bio, Inc.'s ("Gritstone", "we" or "our") proprietary drug candidates, including GRANITE, SLATE and CORAL, the timing of the start, conclusion and status of ongoing or planned clinical trials, including the timing of, and our ability to achieve, anticipated milestones, the likelihood of preliminary data to accurately reflect complete trial data, the sufficiency of our cash, cash equivalents and short-term investments, availability of funding, business strategy, the timing and outcome of regulatory decisions, future availability of pre-clinical and clinical trial data, our collaborations for our product candidates and the maintenance of those collaborations; business and results from operations; and other matters. Forward-looking statements generally contain words such as "believes," "expects," "may," "will," "should," "seeks," "approximately," "intends," "plans," "estimates," "anticipates," and other expressions that are predictions of or indicate future events and trends and that do not relate to historical matters. Because forward-looking statements are inherently subject to 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. The forward-looking statements in this presentation are based on information available to Gritstone as of the date this presentation. Except as required by applicable law, we do not plan to publicly update or revise any forward-looking statements contained herein, whether as a result of any new information, future events, changed circumstances or otherwise. For a further description of the risks and uncertainties that could cause actual results to differ from those expressed in these forward-looking statements, as well as risks relating to our business in general, see Gritstone's most recent Annual Report on Form 10-K filed on March 5, 2024 and any subsequent current and periodic reports filed with the Securities and Exchange Commission.

This presentation also contains estimates and other statistical data made by independent parties and by us relating to market size and growth and other data about our industry. This data involves a number of assumptions and limitations, and you are cautioned not to give undue weight to such estimates. In addition, projections, assumptions, and estimates of our future performance and the future performance of the markets in which we operate are necessarily subject to a high degree of uncertainty and risk.

This presentation concerns drugs that are under clinical investigation and which have not yet been approved for marketing by the U.S. Food and Drug Administration. They are currently limited by Federal law to investigational use, and no representation is made as to their safety or effectiveness for the purposes for which they are being investigated. Certain data in this presentation are based on third-party study or cross-study comparisons and are not based on any head-to-head clinical trials. Cross-study or third-party comparisons are inherently limited and may suggest misleading similarities and differences. The values shown in the cross-study or third-party comparisons are directional and may not be directly comparable.

This presentation shall not constitute an offer to sell or the solicitation of an offer to buy, nor shall there be any sale of these securities in any state or other jurisdiction in which such offer, solicitation or sale would be unlawful prior to registration or qualification under the securities laws of any such state or other jurisdiction.

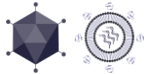
# ABOUT GRITSTONE

Developing Next-Generation Vaccines for Oncology and Infectious Disease

## Platforms Drive More Potent and Durable Immunity



Best-in-class antigen prediction



Proprietary, next-gen vectors drive response

## Fully-integrated Biomanufacturing



Pleasanton, CA

Vaccine  
Manufacturing

Boston, MA

GMP Sequencing  
and Prediction

Potential best-in-class neoantigen-based personalized cancer vaccine program (GRANITE) in randomized Phase 2 study for MSS-CRC

Self-amplifying mRNA (samRNA) platform has distinct potential benefits versus currently-approved infectious disease vaccines

Upcoming data readouts could further de-risk clinical platforms and potentially enable expansion into additional disease types

### Recent & Anticipated Upcoming Milestones

Add'l 12-month data from COVID-19 Phase 1 studies (Apr 2024)

Interim data from Phase 2 GRANITE study (3Q 2024)

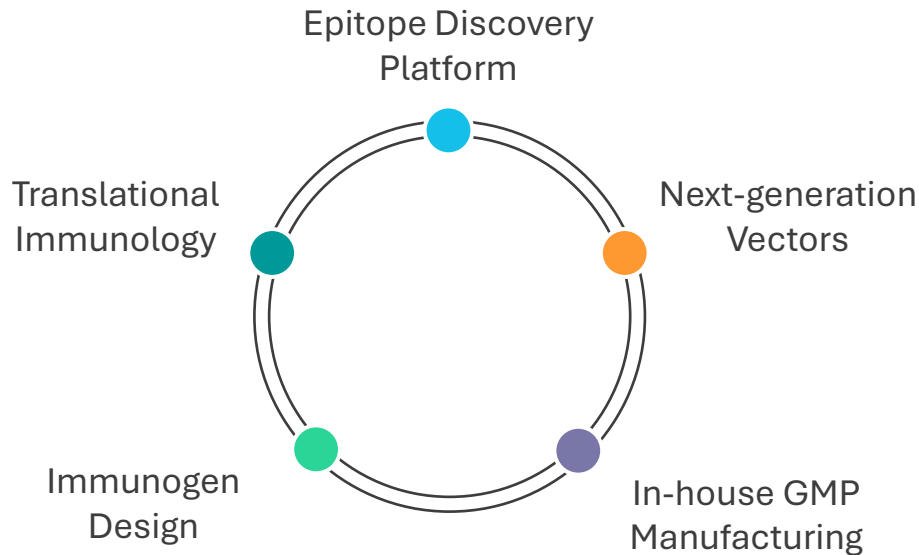
Overall Survival Data from Ph2 GRANITE study (2H 2025)



# OUR CAPABILITIES

## Pursuing More Potent and Durable Immune Responses

Capabilities uniquely designed to address current challenges in cancer and infectious disease



**Proprietary artificial intelligence platform (EDGE™)** to identify critical T cell targets



**Next-gen vectors, ChAd and self-amplifying mRNA\***, to drive potent and durable immune responses suited to the clinical context



**In-house GMP manufacturing** enables personalized and off-the-shelf products (clinical stage and scale-up)






















**Immunogen design** is key component of novel vaccine formats – must be studied in clinical trials





**Bench-to-bedside-to-bench:** innovative product development pushes scientific boundaries



# Solid Tumor Programs Ready for Expansion & Partnership

GRANITE and SLATE programs are currently wholly-owned by Gritstone bio, Inc

Sponsor	Target/Approach	Indication	Preclinical	Phase 1	Phase 2	Phase 3	Anticipated Milestones
 POC in MSS-CRC enables Phase 2 expansion in different tumor types	Individualized Neoantigens	Front-line microsatellite-stable colorectal cancer (MSS-CRC)	 		<b>GRANITE</b>		Mid-2025: OS data
	TBD		 		GRANITE		
	TBD		 		GRANITE		
	TBD		 		GRANITE		
 Off-the-shelf SLATE platform is ready for plug and play application in many solid tumors	Shared Antigens	KRAS <sup>mut</sup> -driven tumor types; front-line metastatic	 		<b>SLATE</b>		Initiate Randomized Ph 2
	TBD		 		SLATE		
	TBD		 		SLATE		
	Neoantigen Cell Therapy-Vaccine Combination	KRAS <sup>mut</sup> -driven tumor types	 		<b>SLATE</b>		IND Cleared in October 2023



  Ongoing and/or fully planned studies

  Potential expansion opportunities

# Oncology

*Redefining Survival in  
Solid Tumors*

**Neoantigen Targeting Immunotherapy**

***GRANITE (Individualize) &  
SLATE (Off-the-Shelf)***

# Individualized Cancer Vaccines are a New Era of Immunotherapy

Potential POC for neoantigen targeting immunotherapies is growing, with multiple randomized studies ongoing

- Genomic Sequencing
- Epitope Identification
- Immunogen Payloads
- Delivery Vectors
- “n of 1” Manufacturing

	gritstone <sup>1</sup> <small>bio</small>	BIONTECH	moderna <sup>®</sup>
Tumor Type	MSS-CRC (1L) <i>Cold</i>	Melanoma (1L) <i>Hot</i>	Melanoma (Adj) <i>Hot</i>
Neoantigen Training Data	Proprietary	Proprietary	Proprietary
Regimen/Vector	ChAd prime + samRNA boosts (heterologous, intramuscular)	mRNA <sup>2</sup> prime + boosts (homologous, intravenous)	mRNA <sup>2</sup> prime + boosts (homologous, intramuscular)
CD8+ T cell Priming <sup>3</sup>	++ <sup>4</sup>	+	+
Data	Interim Phase 2 data reported 3Q 2024	Randomized Phase 2 data expected in 2024	✓ Recurrence-free survival benefit (HR=0.56)

<sup>1</sup>GRTS vaccine candidates have not been studied head-to-head with those listed.

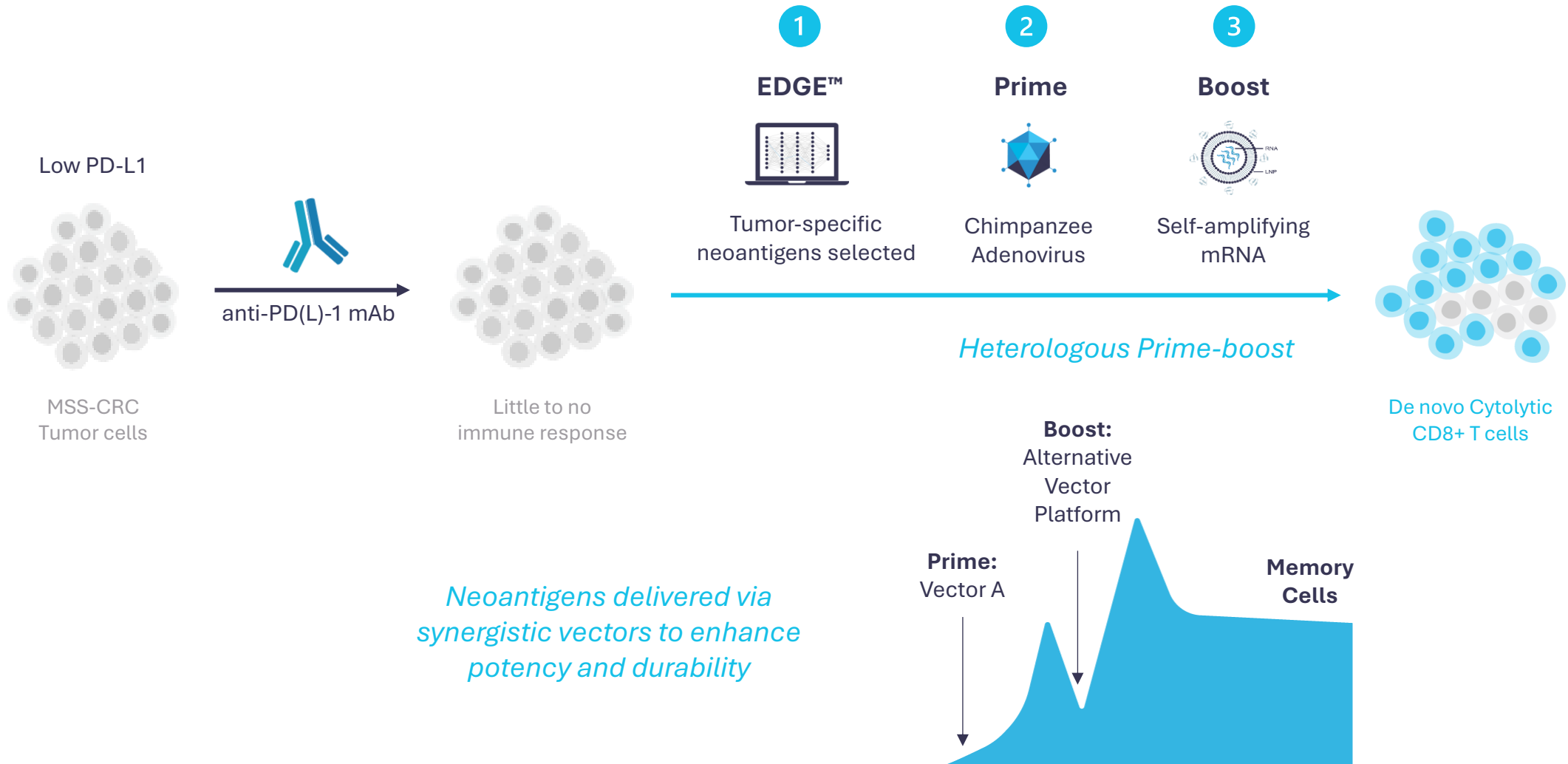
<sup>2</sup>BioNtech uses optimized Uridine mRNA. Moderna uses Modified Uridine mRNA.

<sup>3</sup>Semi-quantitative assessment of strength and breadth of human T cell immune response to neoantigen vaccine based on cross-study comparisons of published data

<sup>4</sup>CD8 T cell priming: Miao et al., Molecular Cancer 20, 41 (2021)

# Gritstone's Approach: Induce CD8+ T cells Against "Cold" Solid Tumors

Antigen selection + prime-boost regimen may be effective in tumors unresponsive to anti-PD(L)-1 therapy





# GRANITE Phase 1/2: GRANITE Induces Potent Tumor-Specific T Cell Responses That Appear To Drive Clinical Benefit

*Vaccine-induced neoantigen-specific cytotoxic T cells kill tumor cells and appeared to prolong OS*



**nature medicine** ARTICLES  
<https://doi.org/10.1038/s41591-022-08974-6>

**Individualized, heterologous chimpanzee adenovirus and self-amplifying mRNA neoantigen vaccine for advanced metastatic solid tumors: phase 1 trial interim results**

Christine D. Palmer<sup>1</sup>, Amy R. Rappaport<sup>1</sup>, Matthew J. Davis<sup>1</sup>, Meghan G. Hart<sup>1</sup>, Claran D. Scallan<sup>1</sup>, Sue-Jean Hong<sup>1</sup>, Leonid Giltin<sup>1</sup>, Lauren D. Kraemer<sup>1</sup>, Sonia Kounlavouth<sup>1</sup>, Aaron Yang<sup>1</sup>, Lindsey Smith<sup>1</sup>, Desiree Schenk<sup>1</sup>, Mojca Skoberne<sup>1</sup>, Kiara Taqeschel<sup>1</sup>, Martina Marrali<sup>1</sup>, Jason R. Jaroslavsky<sup>1</sup>, Charmaine N. Nganje<sup>1</sup>, Elizabeth Maloney<sup>1</sup>, Rita Zhou<sup>1</sup>, Daniel Navarro-Gomez<sup>1</sup>, Adrienne C. Greene<sup>1</sup>, Gijbert Grotenborg<sup>1</sup>, Renee Greer<sup>1</sup>, Wade Blair<sup>1</sup>, Minh Duc Cao<sup>1</sup>, Shawn Chan<sup>1</sup>, Kyounghwa Bae<sup>1</sup>, Alexander I. Spira<sup>1</sup>, Sameek Roychowdhury<sup>1</sup>, David P. Carbone<sup>1</sup>, Brian S. Henick<sup>1</sup>, Charles G. Drake<sup>1</sup>, Benjamin J. Solomon<sup>1</sup>, Daniel H. Ahn<sup>1</sup>, Amit Mahipal<sup>1</sup>, Steve B. Maron<sup>1</sup>, Benny Johnson<sup>1</sup>, Raphael Rousseau<sup>1</sup>, Roman Yelensky<sup>1</sup>, Chih-Yi Liao<sup>1</sup>, Daniel V. T. Catenacci<sup>1</sup>, Andrew Allen<sup>1</sup>, Andrew R. Ferguson<sup>1</sup> and Karin Jooss<sup>1,2</sup>

Volume 28 Issue 8, August 2022

**nature biotechnology** ARTICLES

**Deep learning using tumor HLA peptide mass spectrometry datasets improves neoantigen identification**

Brendan Balk-Sullivan<sup>1</sup>, Jennifer Busby<sup>1</sup>, Christine D Palmer<sup>1</sup>, Matthew J Davis<sup>1</sup>, Tyler Murphy<sup>1</sup>, Andrew Clark<sup>1</sup>, Michele Busby<sup>1</sup>, Fujiko Drake<sup>1</sup>, Aaron Yang<sup>1</sup>, Lauren Young<sup>1</sup>, Noelle C. Ojo<sup>1</sup>, Kamillah Caldwell<sup>1</sup>, Jesse Abhyankar<sup>1</sup>, Thomas Buecher<sup>1</sup>, Meghan G Hart<sup>1</sup>, Vladimir Makarov<sup>1</sup>, Vincent Thomas De Montpreville<sup>1</sup>, Olaf Mercier<sup>1</sup>, Timothy A Chan<sup>1</sup>, Giorgio Scagliotti<sup>1</sup>, Paolo Rironzo<sup>1</sup>, Silvia Novella<sup>1</sup>, Niki Karachaliou<sup>1</sup>, Rafael Rosell<sup>1</sup>, Ian Anderson<sup>1</sup>, Nohar Gabrail<sup>1</sup>, John Fromm<sup>1</sup>, Chaturang Limvarapasa<sup>1</sup>, Karin Choquette<sup>1</sup>, Alexander Spira<sup>1</sup>, Raphael Rousseau<sup>1</sup>, Cynthia Young<sup>1</sup>, Naiver A. Rivera<sup>1</sup>, Eric Fadel<sup>1</sup>, Mark Frattini<sup>1</sup>, Karin Jooss<sup>1</sup>, Mojca Skoberne<sup>1</sup>, Joshua Francis<sup>1</sup> & Roman Yelensky<sup>1,2</sup>

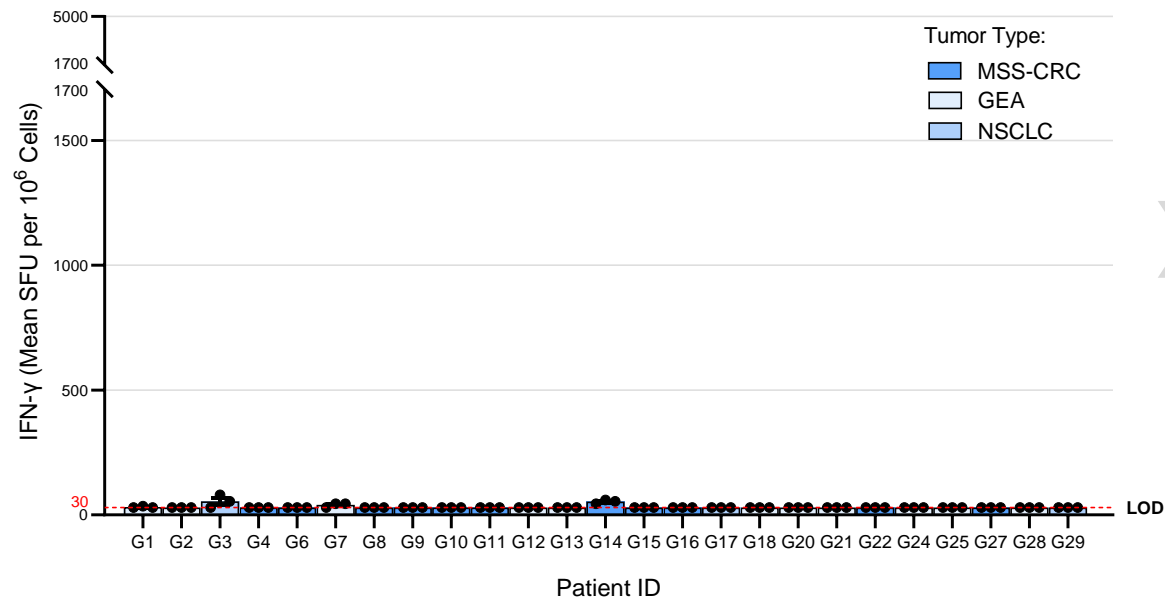
Neoantigens, which are expressed on tumor cells, are one of the main targets of an effective antitumor T-cell response. Cancer immunotherapies to target neoantigens are of growing interest and are in early human trials, but methods to identify neoantigens either require invasive or difficult-to-obtain clinical specimens, require the screening of hundreds to thousands of synthetic peptides or tandem mass spectrometry, or are only relevant to specific human leukocyte antigen (HLA) alleles. We apply deep learning to a large (N = 74 patients) HLA peptide and genomic dataset from various human tumors to create a computational model of

Volume 36 Issue 37, December 2018

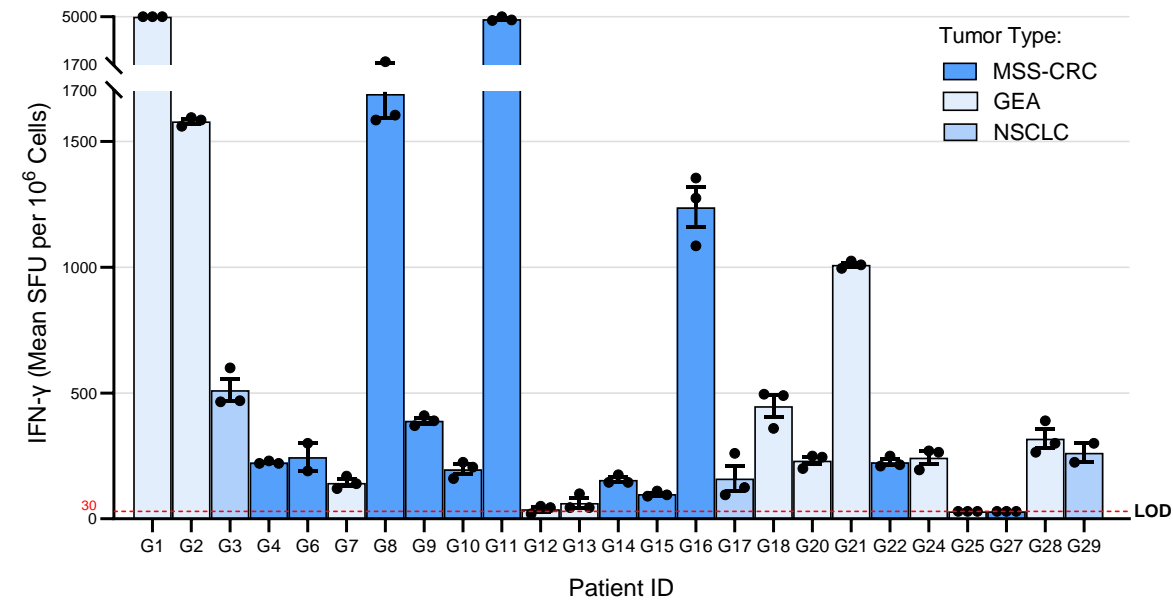
# GRANITE Generates Neoantigen-Specific T Cells in Majority of Patients

Results from Phase 1/2 study in late-line solid tumors

## Baseline



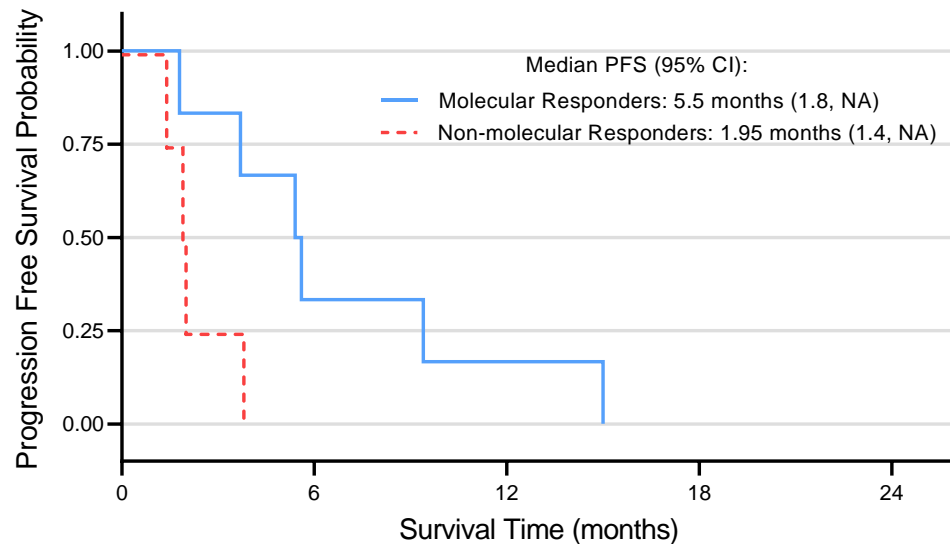
## Peak Response Post-GRANITE<sup>1</sup>



# In Previously Treated Metastatic MSS-CRC, the Benefit of GRANITE is More Pronounced for OS Compared to PFS (GRANITE First-in-Human Trial)

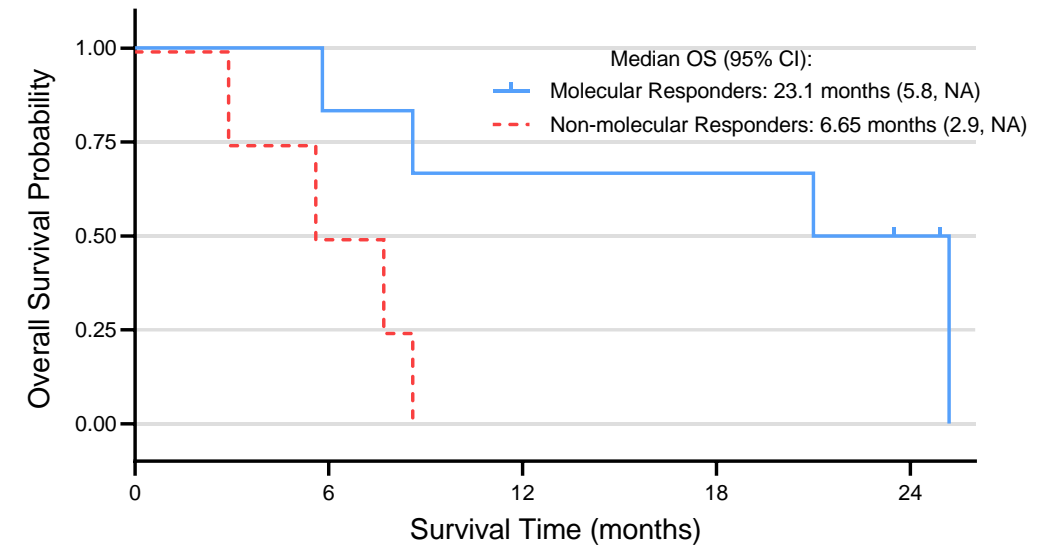
Results from Gritstone's Phase 1/2 study in with previously treated metastatic MSS-CRC

## Progression-free Survival Probability



	0	6	12	18	24
Molecular Responders	6	2	1	0	0
Non-molecular Responders	4	0	0	0	0

## Overall Survival Probability



	0	6	12	18	24
Molecular Responders	6	5	4	4	2
Non-molecular Responders	4	2	0	0	0

Data as of March 06, 2023; patients received two prior lines of therapy; baseline ctDNA taken at time of first vaccination

Median PFS in this population is third line treatment setting is ~2 months; Grothey et al. Lancet 2013

Molecular response defined as >30% reduction in ctDNA from baseline (initiation of study treatment)

# GRANITE Aims to Address Lack of Immune Reactivity in Solid Tumors with High Unmet Need

First Indication = MSS-CRC

**2<sup>nd</sup>** leading cause of cancer-related deaths <sup>1</sup>

~153,000+ diagnoses expected in US in 2023 <sup>1</sup>  
~53,000+ deaths expected in US in 2023 <sup>1</sup>

**~11** months

Median progression-free survival<sup>2</sup>

**~24** months

Median overall survival <sup>2</sup>





# Phase 2: Emerging Benefit for GRANITE vs. Standard of Care in 1L MSS-CRC

## Encouraging randomized Phase 2 data

~20% improvement in PFS with GRANITE (HR = 0.79 [0.422, 1.497]) in all treated population  
13/39 (33%) GRANITE and 7/30 (23%) of control patients remain on study and free of progression

- Last ctDNA assessment is stably below assay LoQ in 12/13 GRANITE and 4/7 controls
- A progression free survival “tail” may emerge in GRANITE recipients – not expected for chemotherapy

Favorable safety and tolerability profile – no Grade 3/4 AEs related to GRANITE and no treatment discontinuation for AEs

## Notable efficacy in patients with low disease burden

More robust improvement in PFS seen in GRANITE patients with low ctDNA at study entry vs. control arm  
Low baseline ctDNA is a likely predictive factor  
Consistent with the field showing vaccine benefit in adjuvant (ie, low-risk) settings)

## Immune data reinforce clinical activity

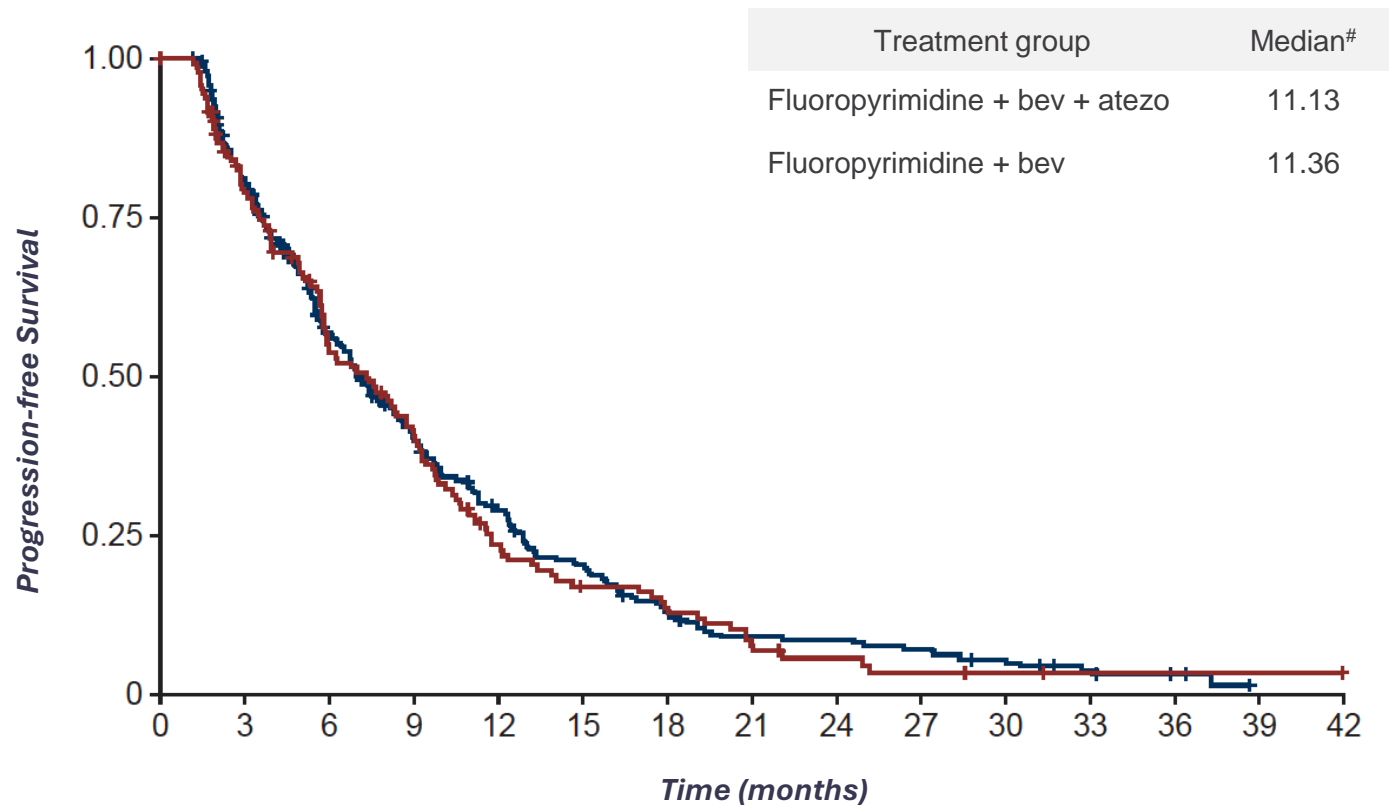
Functional neoantigen-specific T cells observed in all 16/16 GRANITE patients tested (by ex vivo and/or IVS ELISPOT)  
Expansion & increased breadth of neoantigen-specific T cell clones post GRANITE, with persistence after each boost (by TCR sequencing)  
Association of PFS and peak ex vivo ELISPOT responses emerging, ex vivo ELISPOT may be a correlate of PFS

## Future GRANITE studies

These data define population of metastatic CRC patients expected to derive clinical benefit from GRANITE  
A prospective Phase 3 trial is warranted using ctDNA to select suitable patients  
(Neo)adjuvant studies in other solid tumor types are also attractive options

# There is No Substantial Group of Long-Term Progression-Free Survivors in Metastatic MSS-CRC Patients Treated with Chemo +/- Atezolizumab

MODUL study: atezolizumab as 1st line maintenance with fluoropyrimidine + bevacizumab after FOLFOX+bev induction chemotherapy



*“If (the PFS hazard ratio) really solidifies in the mature data set, gets down into the sub-0.8 (call it) area, that’s a really remarkable result in an indication where immune approaches have been completely dead in the past”*

**Jon Miller, Evercore Equity Research**

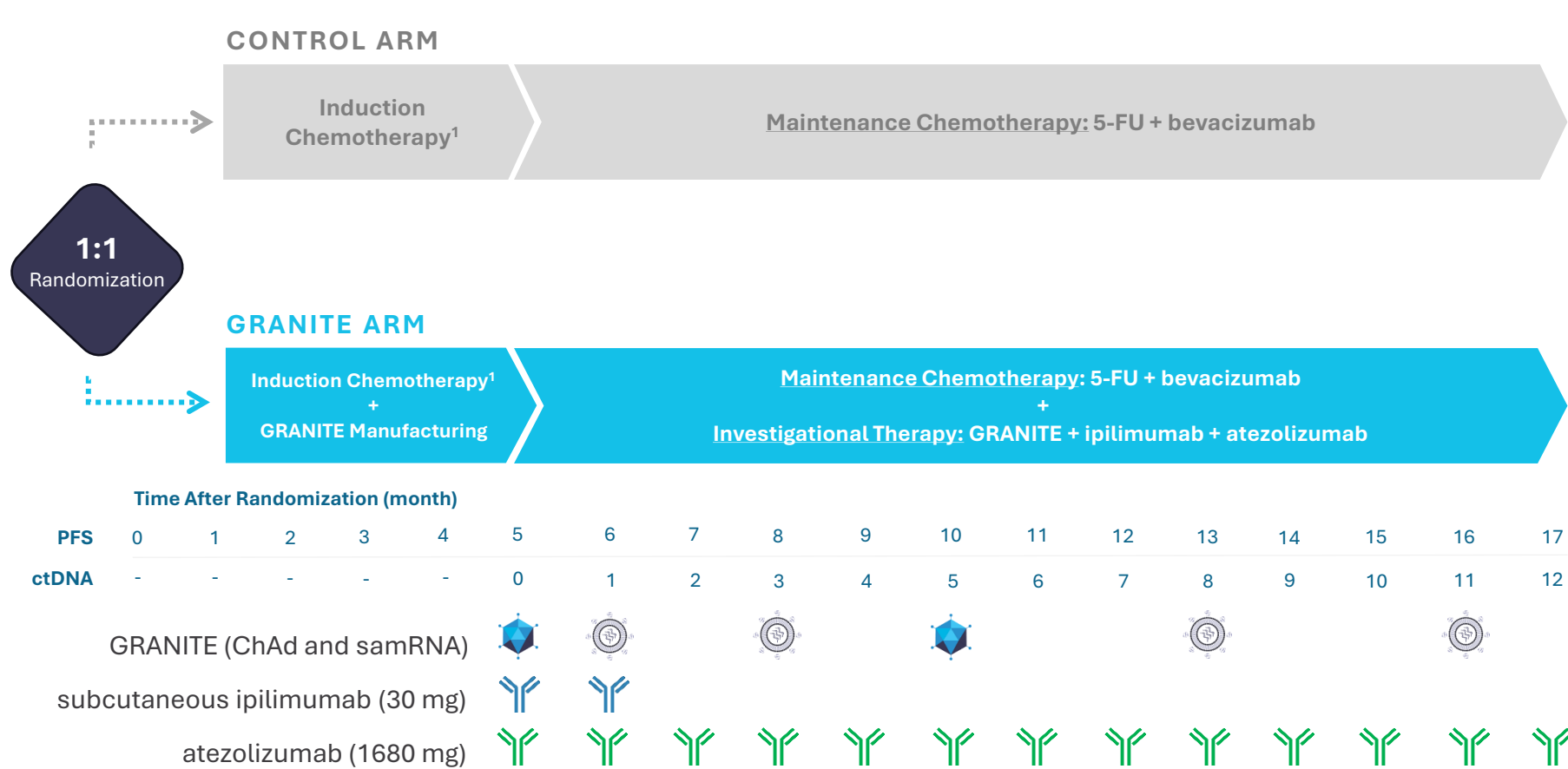
# values reflect additional 4 months as patients in the study were treated after 4 months of FOLFOX as 1L therapy; bev = bevacizumab; atezo = atezolizumab

Modified from Taberero et al ESMO Open 2022

# Ongoing GRANITE Randomized Phase 2: 1L Metastatic MSS-CRC (GO-010)

## Inclusion Criteria

- Untreated metastatic CRC
- Microsatellite-stable tumor status
- SOC with FOLFOX or FOLFOXIRI + bevacizumab (bev)



## Key Endpoints

- Molecular response (Primary)
- Progression-free survival (PFS)
- Overall survival (OS)
- Safety

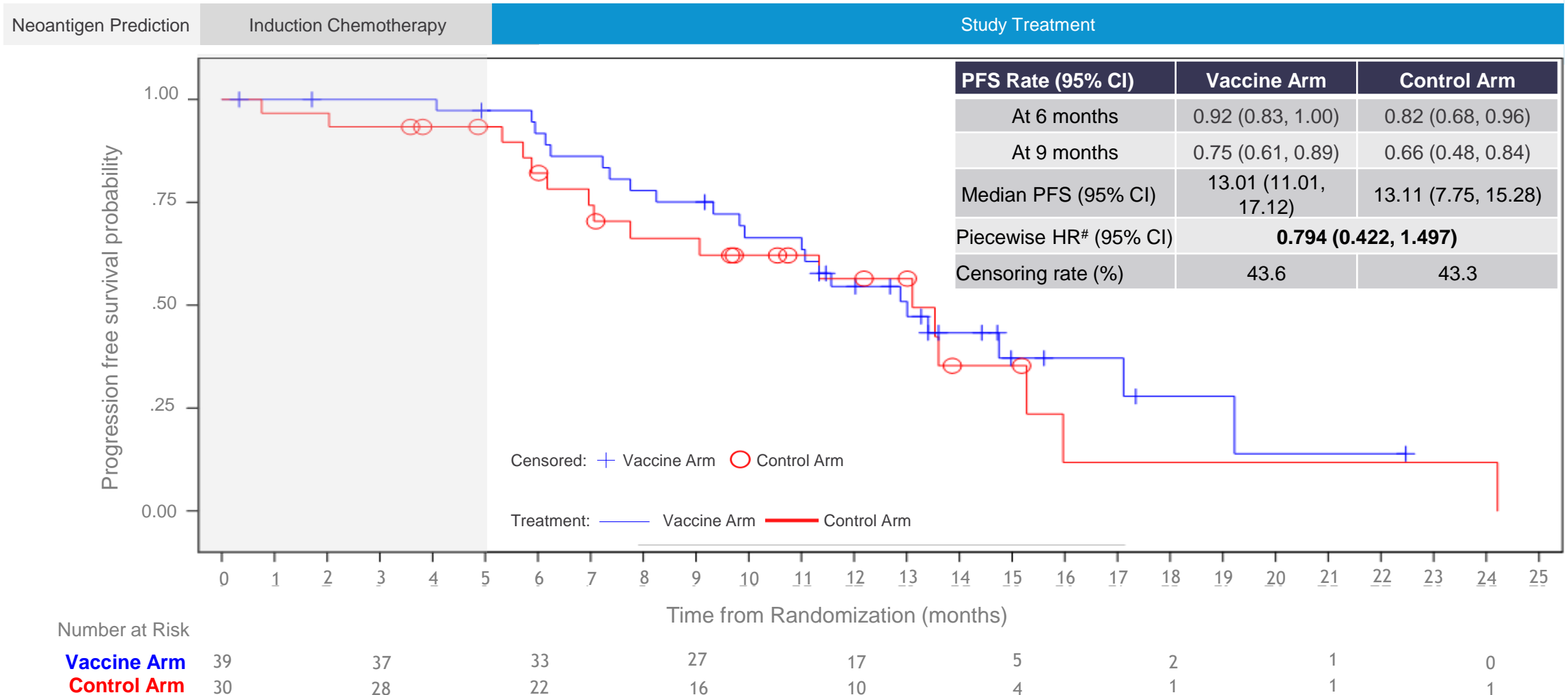
# Patients are Well Balanced for Multiple Prognostic Features

<i>Characteristics</i>	<i>Statistics</i>	GRANITE ARM (N=39) N (%)	CONTROL ARM (N=30) N (%)
<i>Sidedness</i>			
Right-side	n(%)	14 (35.9)	13 (43.3)
Left-side	n(%)	25 (64.1)	17 (56.7)
<i>KRAS Status</i>			
Wild Type	n(%)	18 (46.2)	15 (50.0)
Mutant	n(%)	19 (48.7)	14 (46.7)
Missing	n(%)	2 (5.1)	1 (3.3)
<i>Presence of Liver Disease</i>			
Yes	n(%)	33 (84.6)	22 (73.3)
No	n(%)	6 (17.9)	8 (26.6)
<i>Stage at Study Entry</i>			
Stage IVA (1 site of metastasis)	n(%)	19 (48.7)	12 (40.0)
Stage IVB (2 sites of metastasis)	n(%)	11 (28.2)	11 (36.7)
Stage IVC (peritoneal metastasis)	n(%)	9 (23.1)	7 (23.3)
<i>ctDNA at Start of 1L SOC</i>			
Mean VAF <sup>#</sup>		12.6%	8.7%
Median VAF <sup>#</sup>		7.6%	2.6%



# PFS Trend Apparent in GRANITE Patients with ~30% Continuing on Study

Progression Free Survival in All Patients Who Received Study Treatment

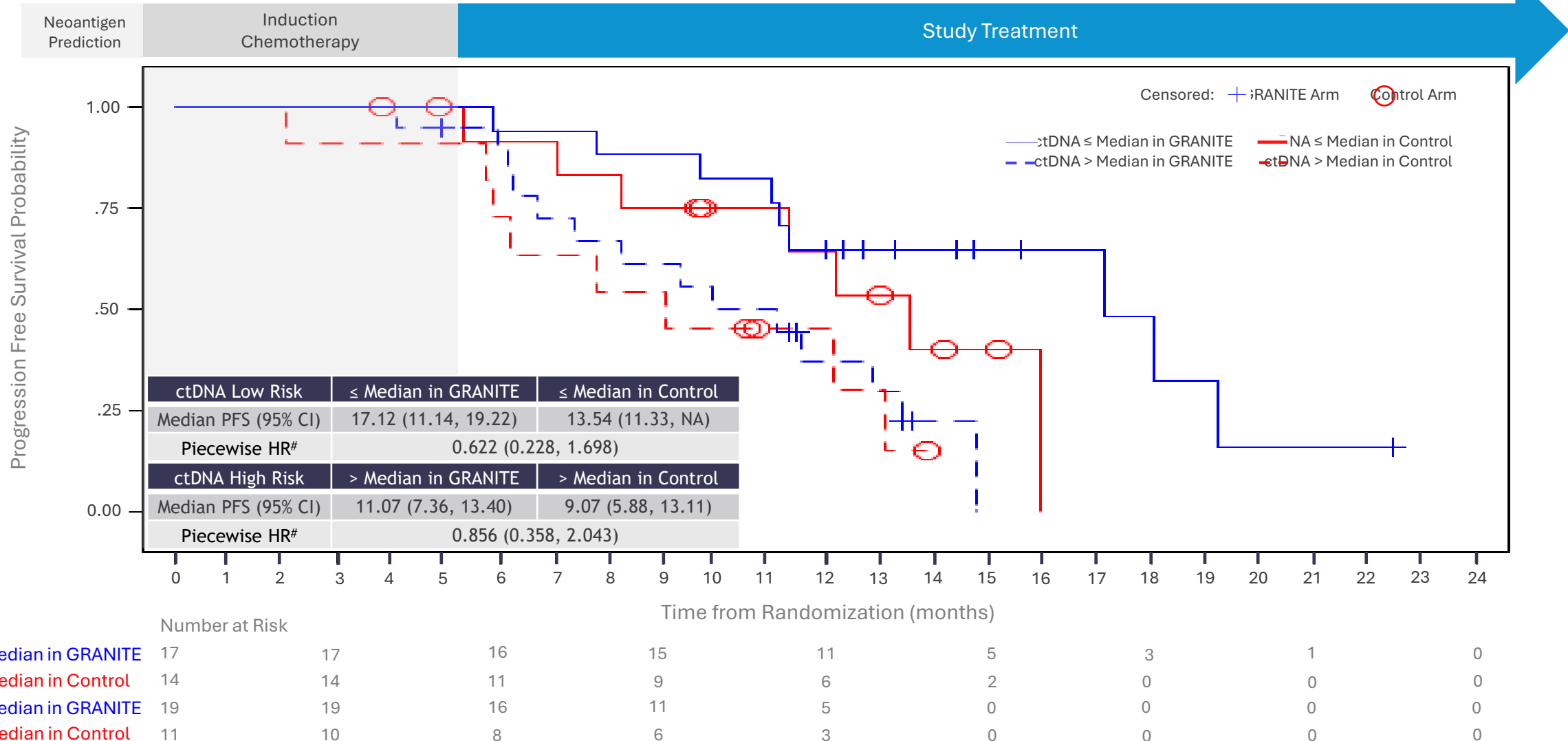


Data cut: 19 Aug 2024

# Patients with Low Disease Burden Demonstrate Greater Benefit

Further follow-up needed to quantify benefit more precisely

Progression Free Survival(PFS) in All Treated Patients



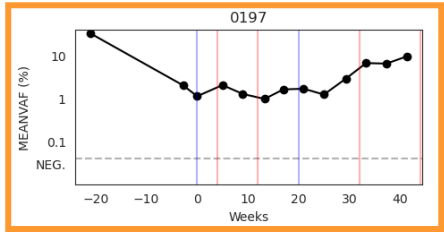
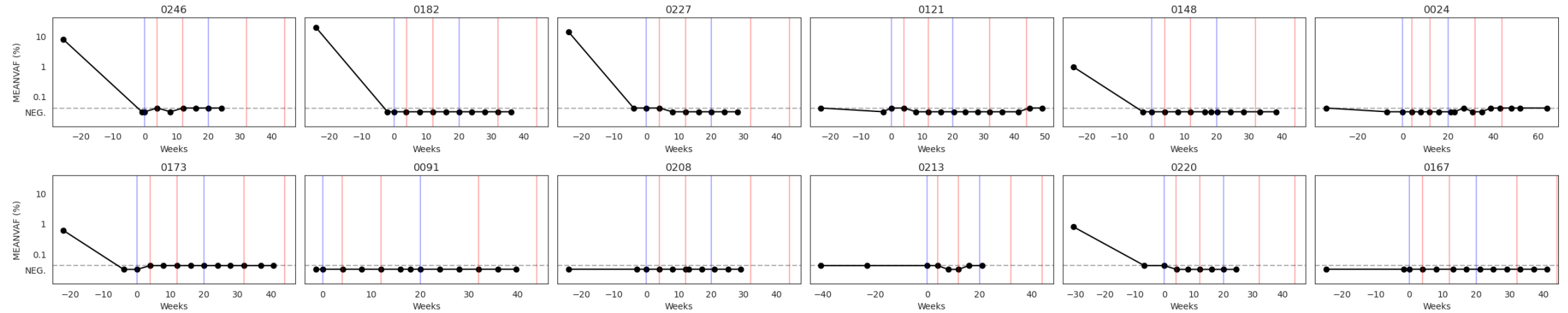
Note: 3.7% VAF is median ctDNA value at study enrollment in all patients

# HR calculated using Piecewise Cox- PH model for the following periods: 0 ≤ t ≤ 6 months and every 6 months up to 12 months and t > 12 months post-randomization

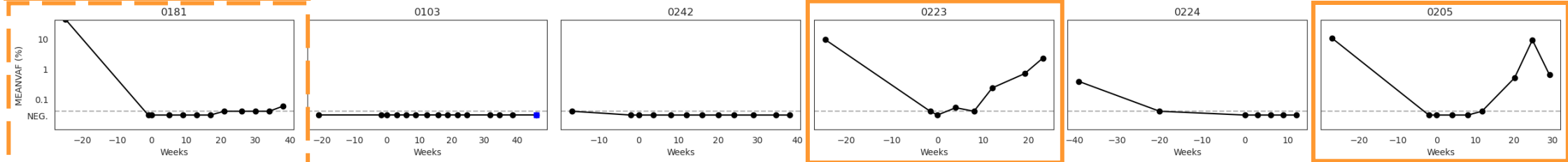
# Most Recent ctDNA Levels Are Very Low and Stable in 12/13 GRANITE Patients Remaining on Study (vs 4/7 Controls) – a PFS “Tail” May Emerge



GRANITE



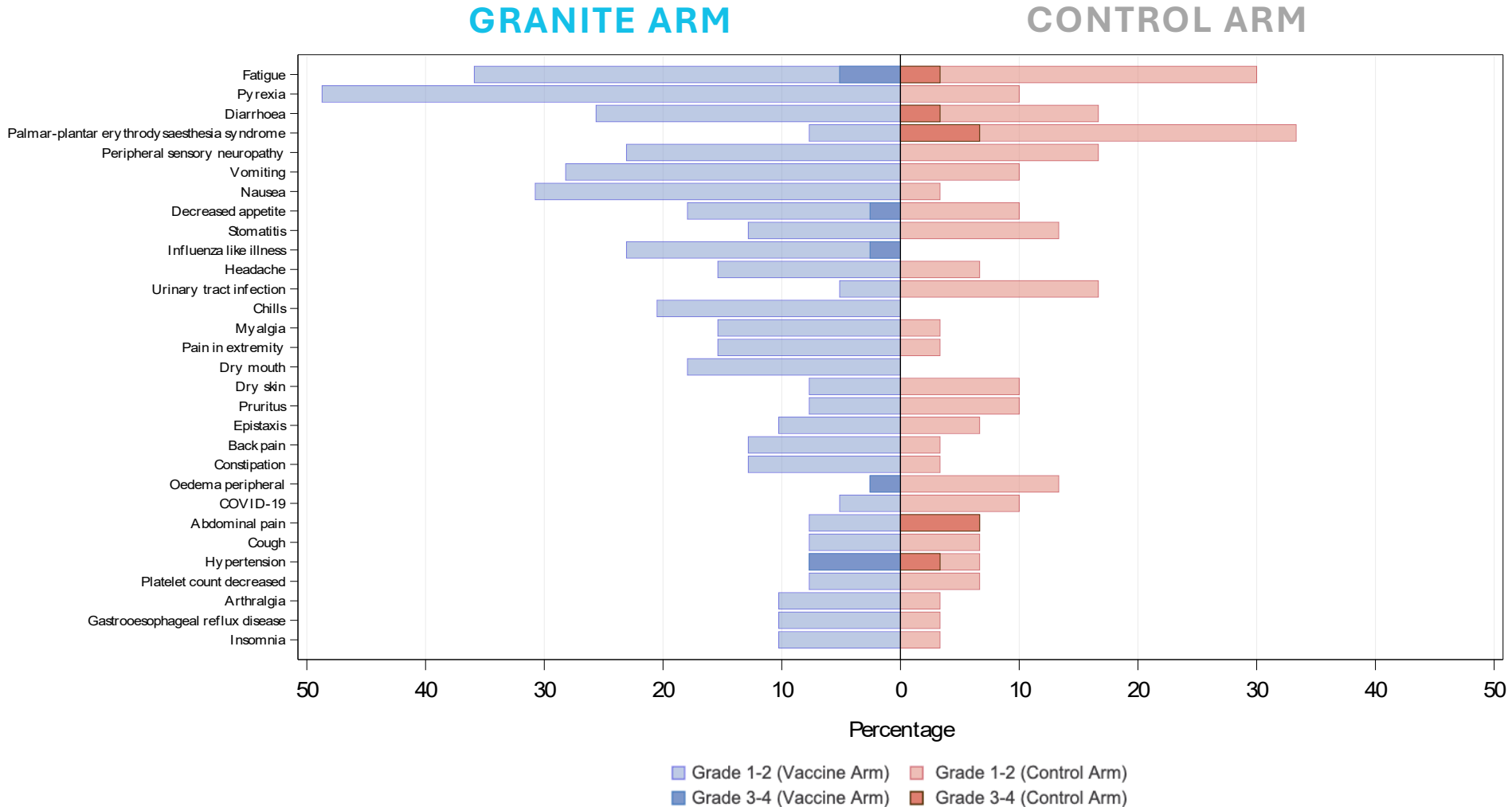
Control



— ChAd — samRNA ----- Limit of Quantification (meanVAF=0.04%)

# GRANITE Exhibits Favorable Tolerability Profile

## Treatment-Emergent Adverse Events



**No patients discontinued study treatment due to a treatment-related adverse event**

**Common adverse events are mild** systemic and local effects associated with any potent vaccine (i.e. transient flu-like illness)

**One treatment-related SAE-fatigue** (upon recovery, patient continued vaccinations without recurrence)

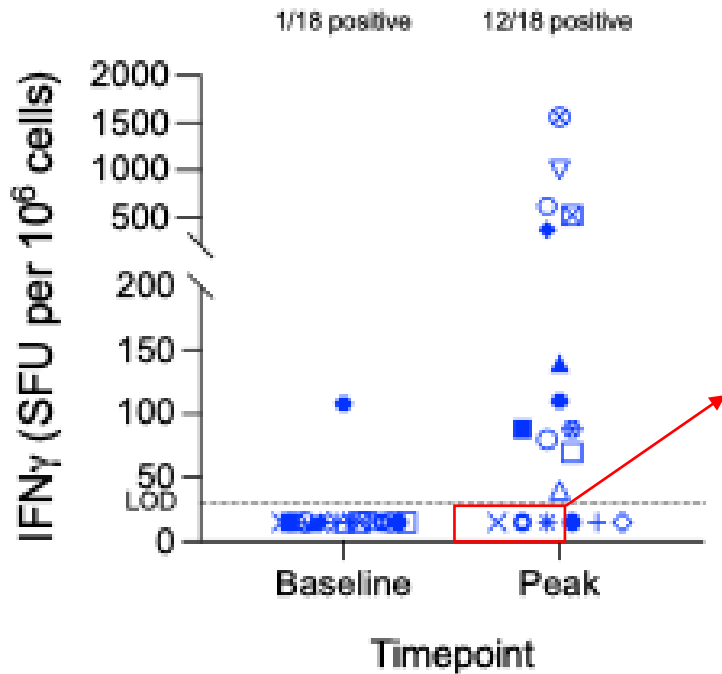
Data cut: 19 Aug 2024



# Neoantigen-specific T Cells Are Detectable By Ex Vivo ELISPOT In Majority Of GRANITE Patients Tested With Increased Breadth Compared To Pre-treatment

T cell responses were observed in 12/18 participants by ex vivo ELISpot. 4/4 ex vivo ELISPOT negative participants had a response by IVS ELISPOT. 7/14 (50%) tested patients had a response to >2 neoantigen post vaccination

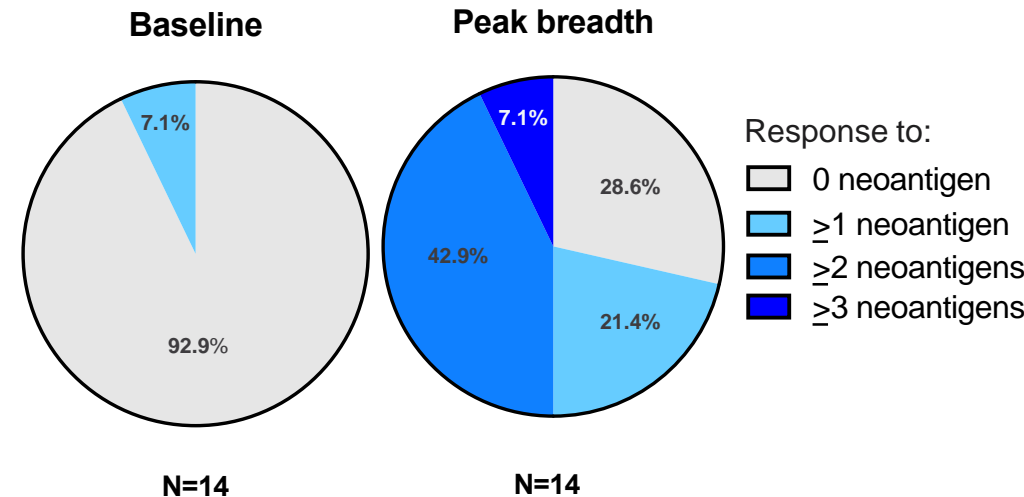
## Ex vivo ELISPOT Baseline vs. Peak



## IVS ELISPOT

Patient ID	Ex vivo ELISPOT	IVS ELISPOT
G10-025-0029	negative	positive
G10-023-0182	negative	positive
G10-030-0105	negative	positive
G10-066-0133	negative	positive

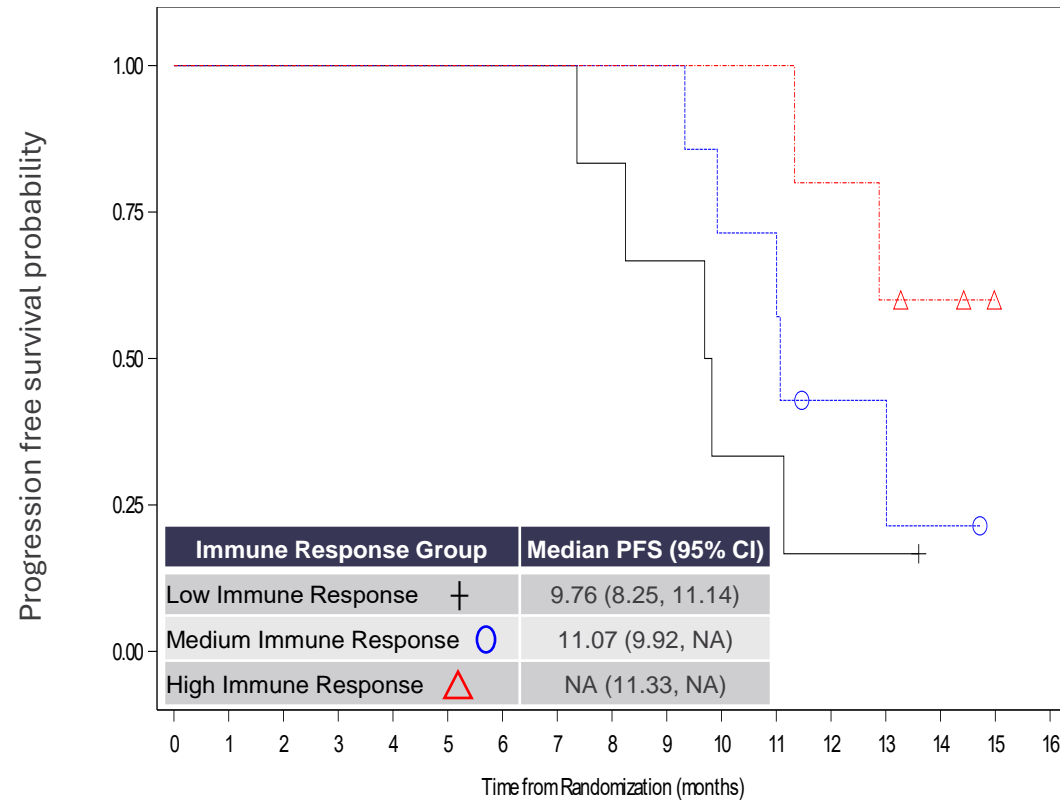
## Breadth of neoantigen-specific T cells by ex vivo ELISPOT



4 peptide pools were used per pt. Peptides representing a mutation are contained within the same pool, with 1 to 10 neoantigens per pool

# Association Between PFS And Peak Ex Vivo ELISPOT Responses Is Observed

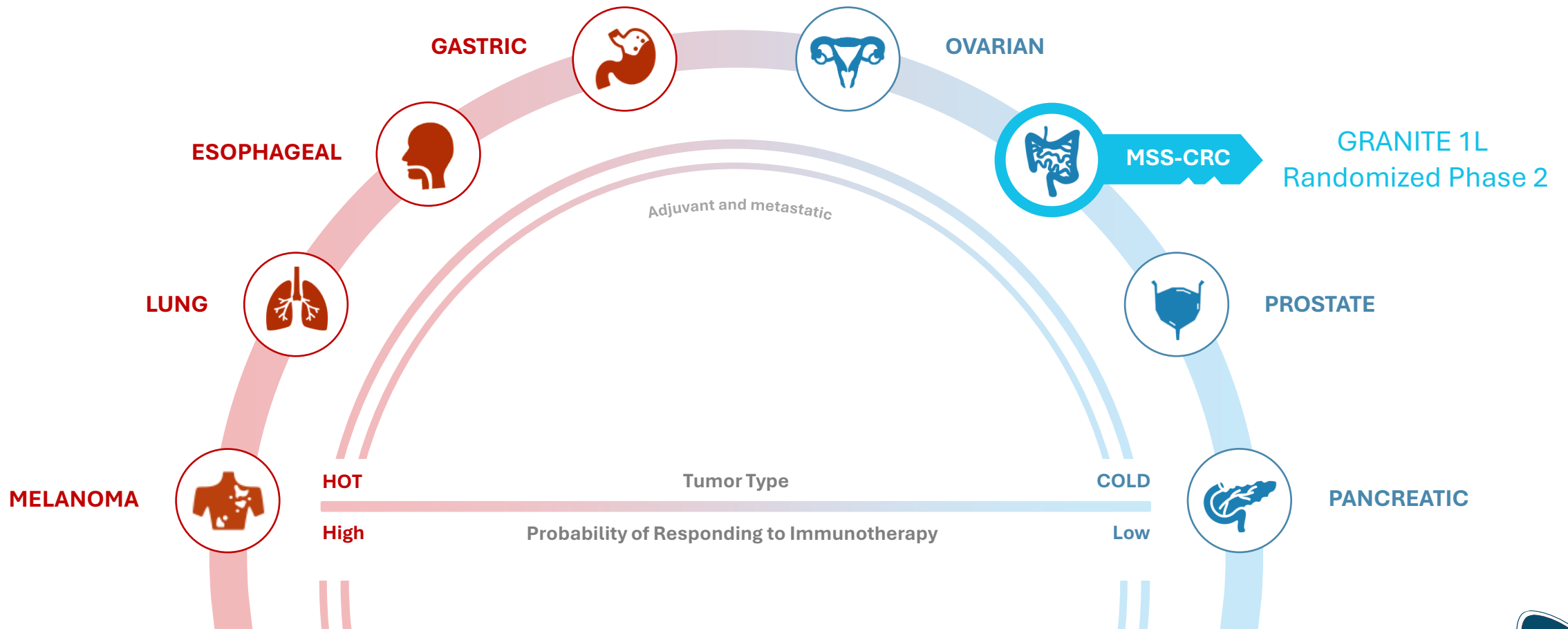
## Association: PFS and ex vivo ELISPOT positivity



	Number at Risk					
	0	1	2	3	4	5
Low Immune Response Group	6	6	6	4	1	0
Medium Immune Response Group	7	7	7	7	2	0
High Immune Response Group	5	5	5	5	4	0

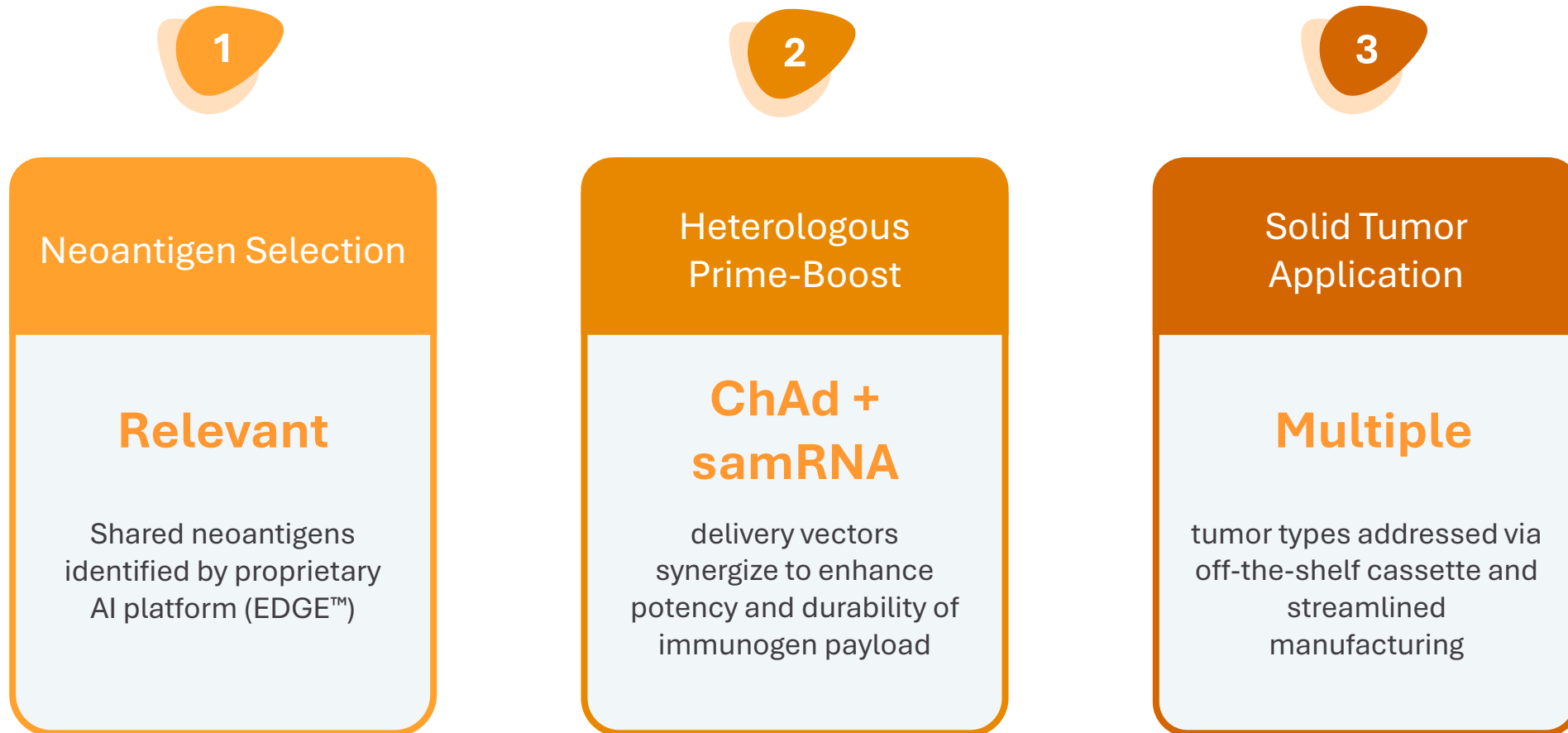
# Positive Phase 2 Results Could Unlock Additional Tumor Types

Success in MSS-CRC supports expansion to both “cold” and “hot” tumors



# SLATE: Off-the-Shelf Neoantigen Vaccines for Solid Tumors

Shared neoantigen program utilizing same antigen selection and vectors as GRANITE





# Molecular Response Associated with Prolonged OS in MSS-CRC and NSCLC

Phase 1/2 Proof-of-Concept: Median Overall Survival in Late-line MSS-CRC and NSCLC<sup>1</sup>



1

39% molecular response rate\* and favorable safety profile of SLATE reinforces therapeutic potential of neoantigen approach

2

Phase 2 data further supports the correlation seen between molecular response and overall survival in late-line solid tumors

3

Phase 2 data in late-line patients supports moving KRAS-directed candidate into earlier lines of treatment

*Data collected from Phase 1/2 SLATE study (NCT03953235); 7 of 18 subjects (MSS-CRC and NSCLC) demonstrated molecular response; molecular response defined as >30% reduction in ctDNA from baseline (initiation of study treatment).*

# SLATE: Serving Solid Tumor Patients via Shared Neoantigen Immunotherapy

## SLATE v1

Optimization

Refined payload of v1 cassette after initial studies indicated immunodominance of non-KRAS antigens

## SLATE-KRAS

Proof of Concept

KRAS-dedicated v2 cassette demonstrated Phase 2 proof of concept in MSS-CRC and NSCLC patients

## Plug and Play

Multiple shared tumor-specific antigen classes

- NeoAg ( $KRAS^{mut}$ )
- CTAs
- Gene fusions
- HERVs
- neoORFs
- Alt Splicing

Optimized and validated SLATE cassette now ready for “plug and play” application across solid tumor indications




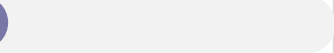



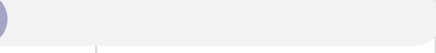



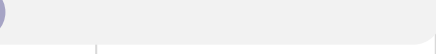




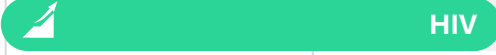
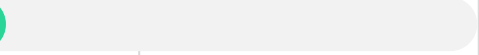


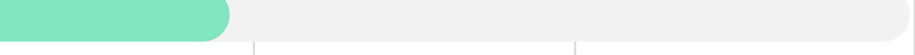

# Infectious Disease

*Leaving no strain behind*

Prophylactic & Therapeutic  
**Self-amplifying mRNA Vaccines**

# Infectious Disease Approach and Collaborations Provide Potential Upside

Collaborator	Indication	Preclinical	Phase 1	Phase 2	Collaboration Terms
	Prophylactic COVID-19 (Spike + TCE)				The BARDA Contract, as amended, consists of a base period (currently ending on or before March 31, 2025) and a total contract period-of-performance (base period + 2 stages gated at BARDA's discretion) of up to ~4 years.
	Prophylactic COVID-19 (Spike + TCE)				CEPI to provide up to \$25.6M in funding to conduct a Phase 1 study in South Africa (CORAL-CEPI).
	Prophylaxis COVID-19 (Spike + TCE)				Phase 1 trial conducted via NIAID-supported Infectious Disease Clinical Research Consortium (IDCRC). Gritstone has received multiple milestone payments.

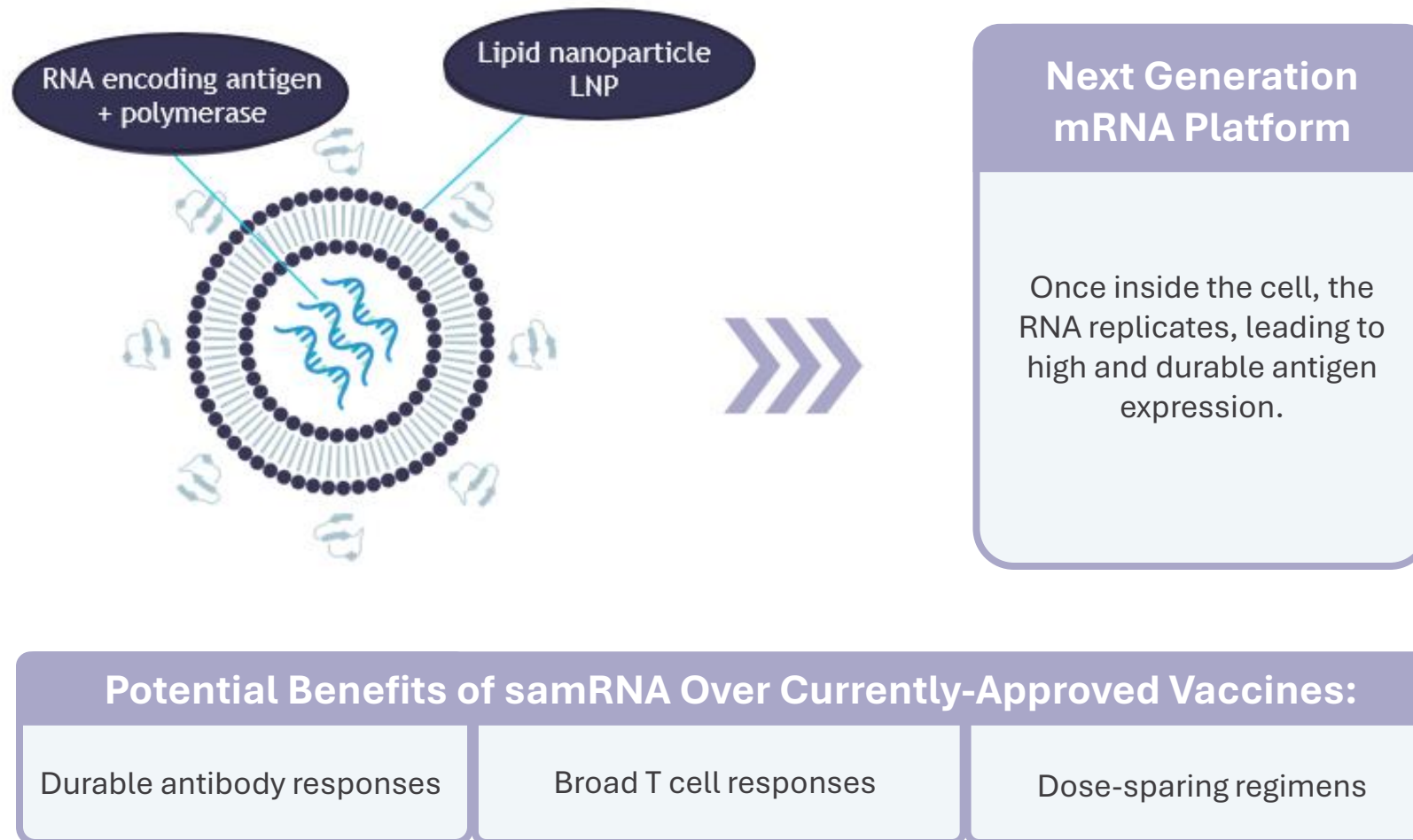
Collaborator	Indication	Preclinical	Phase 1	Phase 2	Collaboration Terms
	Therapeutic HIV Cure				Gilead to conduct Phase 1 and is responsible for all R&D. Gritstone to provide vaccine delivery platform and is eligible to receive up to \$785M in milestone payments, in addition to commercial royalties.
	Therapeutic HPV Cure				Undisclosed*



\*In late 2023, BARDA informed the Company that any potential funding beyond the base period is expected to be administered under a new award made by the Rapid Response Partnership Vehicle ("RRPV Consortium"). In early 2024, we applied to the RRPV Consortium for funding of our Phase 2b CORAL Study extending beyond the base period of the BARDA Contract. There is no certainty that the RRPV Consortium, which selects awardees at BARDA's discretion, will accept our application and on what terms. As of June 30, 2024, BARDA and Gritstone have amended the base period under the BARDA Contract to extent to March 31, 2025. Also, as of June 30, 2024, BARDA had not made the decision to proceed with either of the two stages, nor have we been awarded a new award by or entered into a new agreement with the RRPV Consortium, the terms and financials of any such new agreement may be different from the terms and financials of the BARDA Contract.

# Self-amplifying mRNA: Addressing Current Vaccine Limitations for ID

Well-tolerated, scalable platform technology offering potential advantages over first-generation mRNA



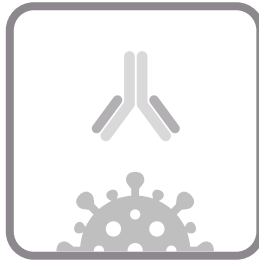


# Gritstone's Differentiated Approach to COVID-19

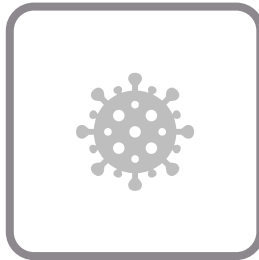
Novel "Spike-plus" approach designed to drive durable and broad immunity

## 1<sup>st</sup> Generation mRNA (Spikevax, Comirnaty)

nAbs Against Spike Only That Wane After 4-6 Months



Spike-specific Immunity Subject to Viral Mutation and Immune Evasion

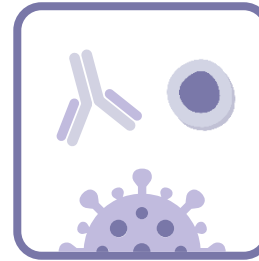


High-dose, Repeat Boosts

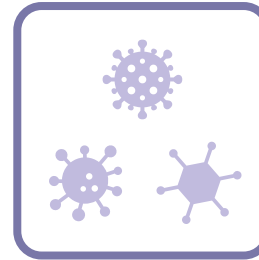


## Gritstone self-amplifying mRNA (CORAL)

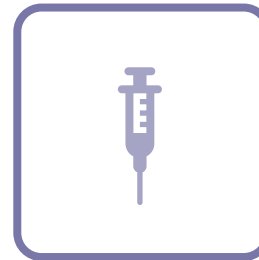
Durable nAbs Against Spike; T cells Against Conserved Viral Targets



Broad T cell Immunity May Enable Cross-Variant Protection



Low Dose, Durable Response

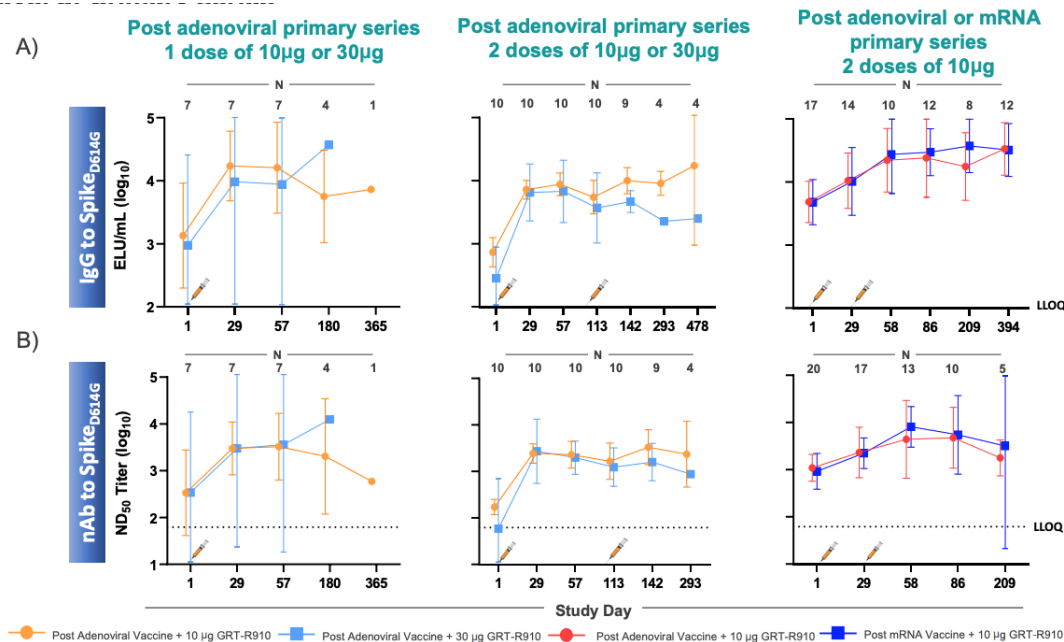


# Phase 1 Results Highlight Differentiated Long-lasting Antibody Response

Potent neutralizing antibody (nAb) levels sustained through 12 months across global SARS patient populations

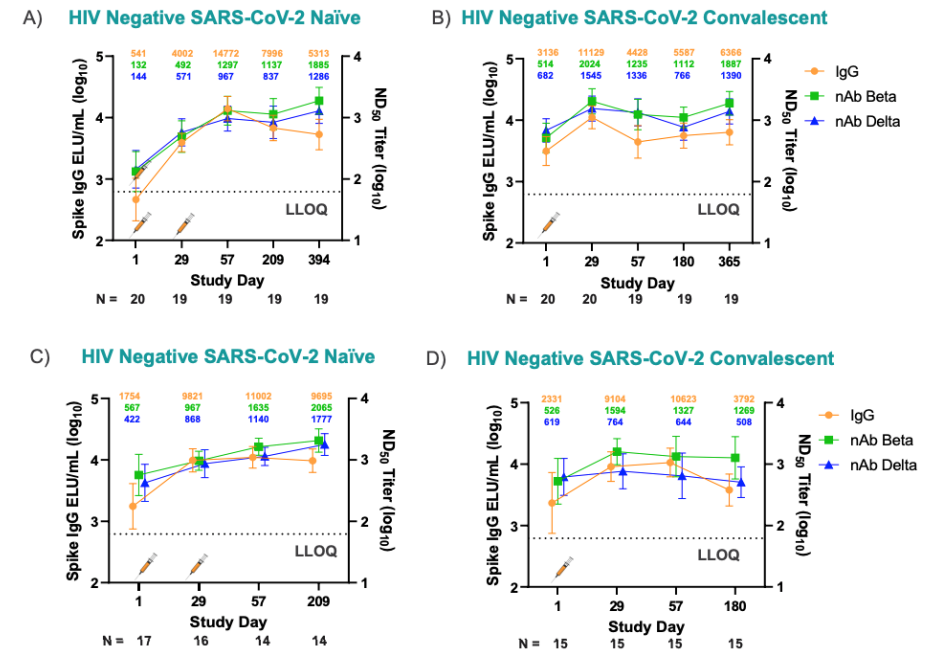
**CORAL-BOOST: Elderly UK Population, samRNA Following Adenoviral or mRNA Primary Series**

**CORAL-CEPI: Young, Unvaccinated S. African Population, samRNA as Single Dose or Homologous Prime Boost**



Binding antibodies (IgG) (A) and nAbs (B) against SpikeD614G measured in participants after receiving boost dose(s) of 10µg or 30µg of GRT-R910 post adenoviral and 30µg mRNA vaccine primary series. Geometric means with 95% confidence intervals are shown. Lower limit of quantitation (LLOQ): 50.3 ELU/mL; 62 ND50 Titer

CORAL-BOOST ID Week 2023 Poster



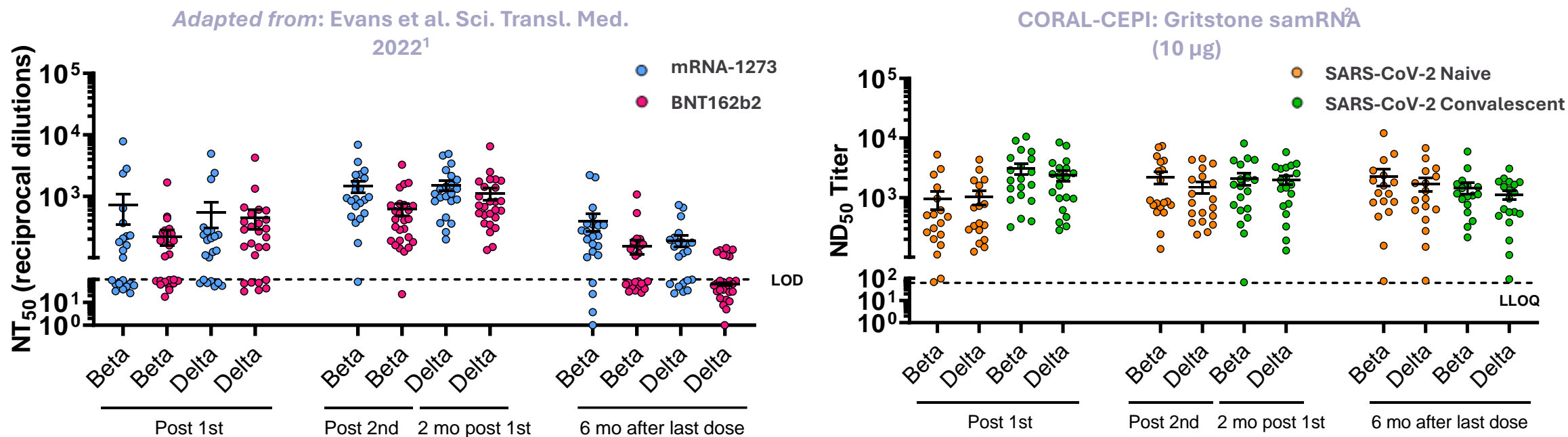
IgG bAb levels to Spike<sub>WT</sub> (ELU/mL) and nAb titers against Beta and Delta variants (ND50 titer) are shown for participants receiving 1 or 2 doses of 10µg GRT-R914 (A and B) or GRT-R912 (C and D). Geometric means with 95% confidence intervals are shown. Lower limit of quantitation (LLOQ): 62 ND50 Titer (nAb), 50.3 ELU/mL (IgG)

CORAL-CEPI ID Week 2023 Poster

# nAb Durability in Previously Unvaccinated Subjects (Beta and Delta VOCs)

Cross-study 6-month data vs. Moderna and Pfizer shown; vaccines not studied head-to-head directly

Gritstone's samRNA vaccine candidate elicits durable nAb responses against Beta and Delta variants, in contrast to FDA-approved Moderna and BioNtech/Pfizer mRNA vaccines



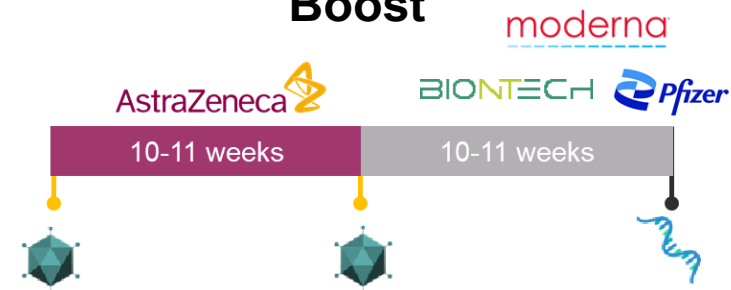
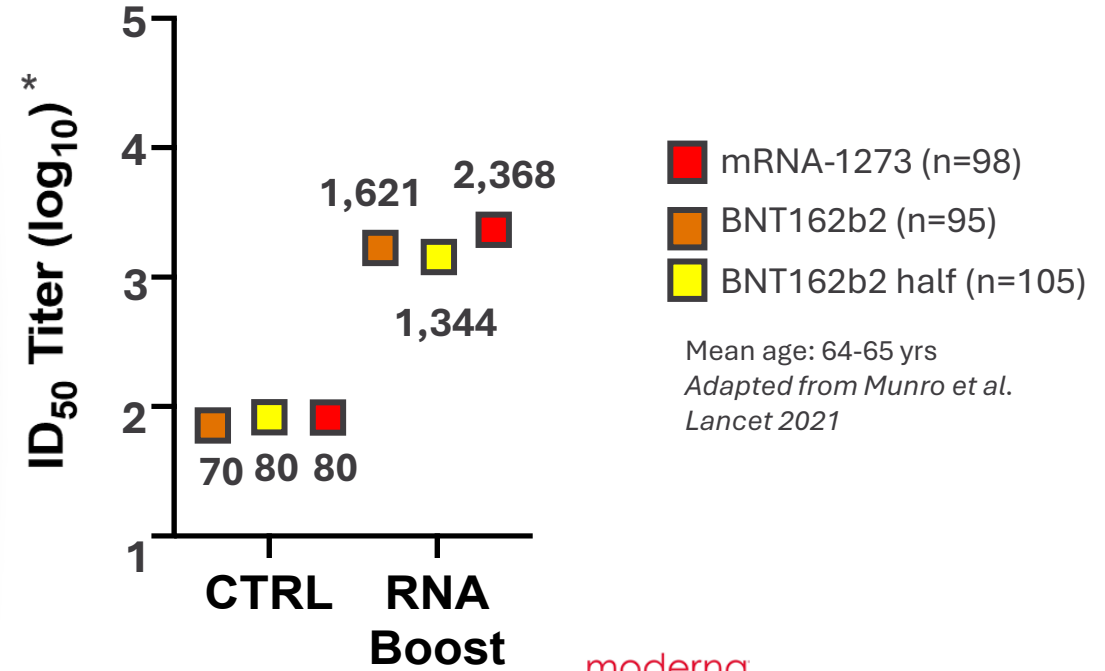
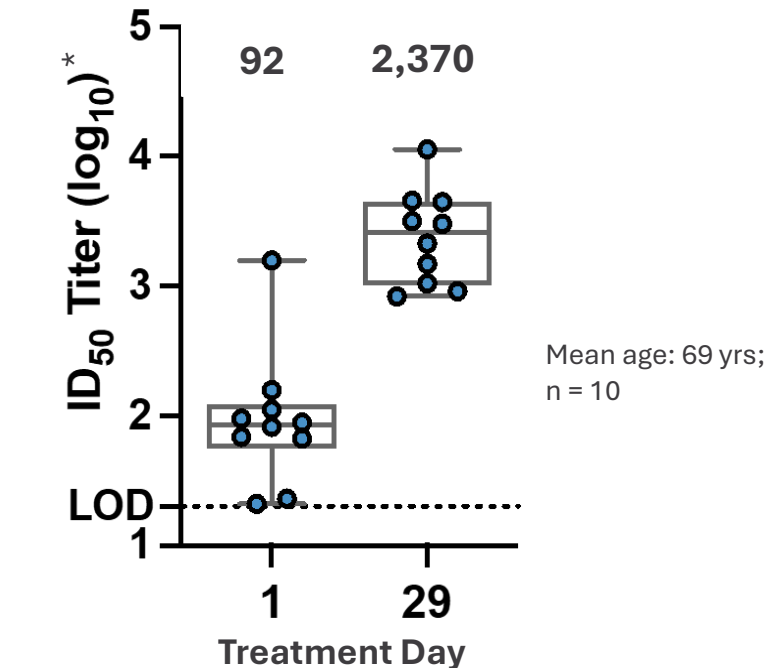
<sup>2</sup> SARS-CoV-2 naïve (orange): Participants received 2 doses with anti-N seronegative at baseline. nAb data were collected at Day 29 (Post 1<sup>st</sup> dose), Day 57 (Post 2<sup>nd</sup> dose), and Day 209 (6 month after last dose). SARS-CoV-2 convalescent (green): Participants received 1 dose with anti-N seropositive at baseline. nAb data were collected at Day 29 (Post 1<sup>st</sup> dose), Day 57 (2 months post 1<sup>st</sup> dose), and Day 180 (6 month after last dose). Error bars indicates means ± SEs; the dashed horizontal line indicates the limit of detection (NT<sub>50</sub> < 100) for Evans et al. Sci. Transl. Med. 2022. and it is lower limit of quantification (ND<sub>50</sub> < 62) for GO-012 data.

<sup>1</sup> Evans et al. Sci Transl Med. 2022 Mar

# CORAL-BOOST: samRNA Boost Elicited Similar nAbs at up to 1/10th the Dose

Cross-study comparison: 10µg of samRNA elicited similar nAbs as 100µg of Moderna (mRNA-1273) after AZ primary series\*

## Neutralizing antibodies (geomean) against Wild Type Variant



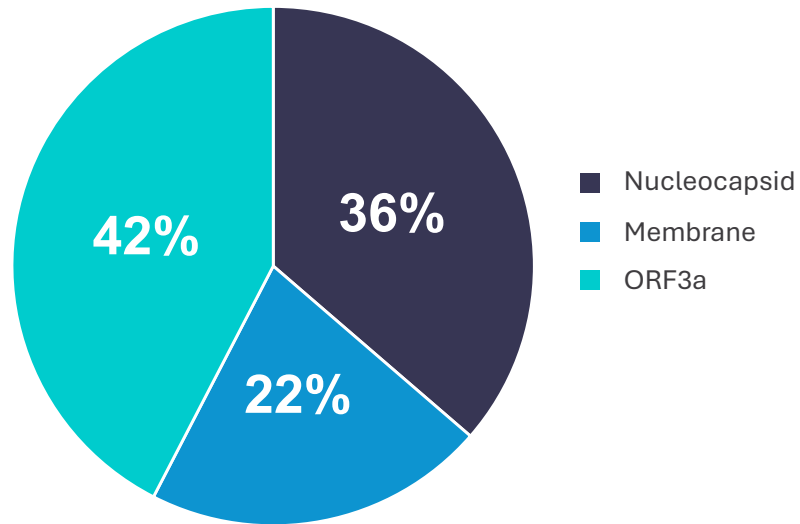
\*Not studied head-to-head directly. ID<sub>50</sub> = Median infective dose; Geomean ID<sub>50</sub> titer values notated. CTRL: Equivalent meningococcal conjugate vaccine; Treatment day = day 1 GRTS samRNA boost dose was administered. Boxes and horizontal bars denote interquartile range (IQR) and median neutralization, respectively. Whisker endpoints are equal to the maximum and minimum values below or above the median +/- 1.5 x IQR.

# Phase 1 Results: Robust and Broad CD8+ T cell Induction

samRNA has driven potent cytotoxic cellular responses against Spike and non-Spike SARS-CoV-2 viral epitopes

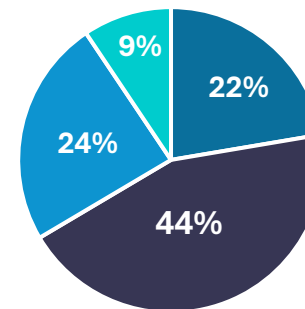
## CORAL-BOOST: Non-Spike CD8+ T cell responses after single 10ug samRNA boost following Vaxzevria (AstraZeneca) primary series

*Proportion of responses to TCE5 regions assessed by post-IVS ELISpot*



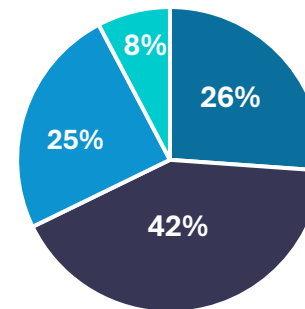
TCE5 overlapping peptide (OLP) pools to TCE5 Nucleocapsid, Membrane and ORF3a regions assessed by post-IVS ELISpot (post-treatment timepoint)

## CORAL-CEPI: Previous SARS-CoV-2 exposure does not affect T cell responses post-vaccination



### SARS-COV-2 naïve

- NSP
- Nucleocapsid
- Membrane
- ORF3a



### Convalescent

- NSP
- Nucleocapsid
- Membrane
- ORF3a

Frequency of peak T cell responses to TCE components at post-vaccination timepoint are shown for naïve (n=7) and convalescent (n=7) subjects following one or two doses of GRT-R914.

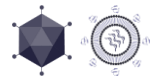
# ABOUT GRITSTONE

Developing Next-Generation Vaccines for Oncology and Infectious Disease

## Platforms Drive More Potent and Durable Immunity



Best-in-class antigen prediction



Proprietary, next-gen vectors drive response

## Fully-integrated Biomanufacturing



Pleasanton, CA

Vaccine  
Manufacturing

Boston, MA

GMP Sequencing  
and Prediction

Potential best-in-class neoantigen-based personalized cancer vaccine program (GRANITE) in randomized Phase 2 study for MSS-CRC

Self-amplifying mRNA (samRNA) platform has distinct potential benefits versus currently-approved infectious disease vaccines

Upcoming data readouts could further de-risk clinical platforms and potentially enable expansion into additional disease types

### Recent & Anticipated Upcoming Milestones

Add'l 12-month data from COVID-19 Phase 1 studies (Apr 2024)

Interim data from Phase 2 GRANITE study (3Q 2024)

Overall Survival Data from Ph2 GRANITE study (2H 2025)





[ir@gritstone.com](mailto:ir@gritstone.com)