



NEXT-GENERATION IMMUNOMEDICINES

DECEMBER 2019

Forward-Looking Statements

To the extent that statements contained in this presentation are not descriptions of historical facts, they may be deemed to be forward-looking statements under the Private Securities Litigation Reform Act of 1995. Words such as “may,” “will,” “expect,” “anticipate,” “estimate,” “intend,” “next,” “near-term,” “future” and similar expressions, as well as other words and expressions referencing future events, conditions, or circumstances, are intended to identify forward-looking statements. Examples of forward-looking statements in this presentation may include, among others, statements regarding: (i) the timing, progress and results of our preclinical and clinical trials; (ii) the timing or likelihood of regulatory filings for our product candidates; (iii) our manufacturing capabilities and strategy; (iv) the potential benefits and activity of our product candidates; (v) our expectations regarding the nature of the biological pathways we are studying; (vi) our expectations regarding our FIND-IO platform; and (vii) the potential benefits of our relationships with Dr. Lieping Chen, Yale University and Eli Lilly and Company.

Various factors could cause actual results to differ materially from those projected in any forward-looking statement. Such risks and uncertainties include, among others: our limited operating history and no products approved for commercial sale; our history of significant losses; our need to obtain additional financing; risks related to clinical development, marketing approval and commercialization; and the unproven approach to the discovery and development of product candidates based on our FIND-IO platform. No forward-looking statement is a guarantee of future results or events, and one should avoid placing undue reliance on such statements. For further discussion of these and other factors that could affect the outcome of our forward-looking statements, see our filings with the Securities and Exchange Commission, including in "Risk Factors" and "Special Note Regarding Forward-Looking Statements" in our prospectus dated November 14, 2019. Except as otherwise indicated, this presentation speaks as of the date indicated herein. Except as required by law, we assume no obligation to update any forward-looking statements, or to update the reasons why actual results could differ materially from those anticipated in the forward-looking statements, even if new information becomes available in the future. The information in this presentation is not complete and may be changed.

NEXTCURE HIGHLIGHTS

Pipeline of Immuno-medicines

- NC318 (S15): Phase 1 data presented at SITC 2019
- NC410 (LAIR-1): IND expected Q1 2020
- Manufacturing: dedicated, state-of-the-art facility

Platform for Novel Target Discovery

- FIND-IO functional screening discovery engine
- Oncology partnership with Lilly
- Expanding into autoimmune diseases

Proven Abilities

- Experienced management team
- Scientific founder Dr. Lieping Chen: discovered PD-L1 & other key targets
- Strong balance sheet to deliver on objectives

THE UNMET NEEDS OF CANCER PATIENTS ARE SIGNIFICANT

NON-RESPONDERS

RAPID PROGRESSION

LIMITED TREATMENTS



We Need New Solutions

COMMITTED TO DISCOVERING & DEVELOPING NOVEL, FIRST-IN-CLASS IMMUNOMEDICINES TO IMPROVE LIVES

NEW

therapeutic options

POSITIVE

clinical responses

IMPROVED

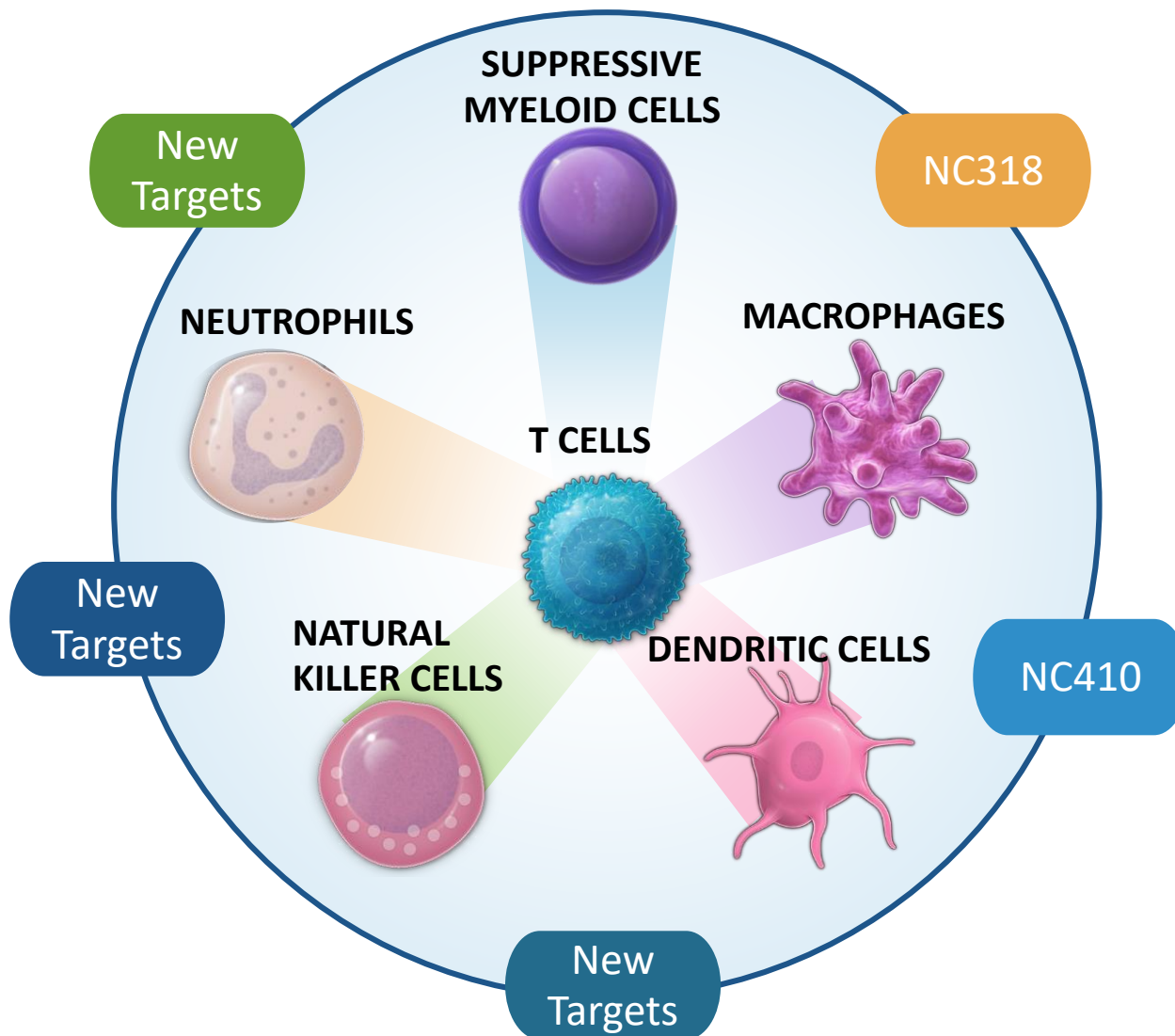
quality of life



Focused on Cancer Patients Not Adequately Addressed Today

NextCure

EXPANDING TARGETS BEYOND T CELLS



EXPERIENCED TEAM WITH STRONG TRACK RECORD

HISTORY AND SUCCESS OF WORKING TOGETHER

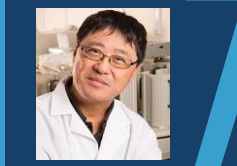
Michael Richman CEO				
Timothy Mayer, PhD COO				
Steve Cobourn, CPA CFO				
Kevin N Heller, MD CMO				
Sol Langermann, PhD CSO				
Jim Bingham, PhD CDO				
Linda Liu, PhD SVP, Research				
Sebastien Maloveste, PhD VP, Business Development				
Dallas Flies, PhD VP, Discovery Research				

WORLD-RENOWNED SCIENTIFIC FOUNDER AND KEY COLLABORATION WITH YALE

YALE COLLABORATION

LIEPING CHEN, MD, PhD

Discovered multiple
key immune pathways,
including PD-L1



WORLD-RENOWNED INSTITUTION

Sponsored research, clinical
samples, cell lines & models

TEAM OF COLLABORATORS

Roy Herbst, MD, PhD
David Rimm, MD, PhD
Mario Sznol, MD

NATURE MEDICINE PUBLICATION

nature
medicine

ARTICLES

<https://doi.org/10.1038/s41591-019-0374-x>

Siglec-15 as an immune suppressor and potential target for normalization cancer immunotherapy

Jun Wang^{1,5}, Jingwei Sun^{1,5}, Linda N. Liu², Dallas B. Flies², Xinxin Nie¹, Maria Toki³, Jianping Zhang¹,
Chang Song², Melissa Zarr², Xu Zhou¹, Xue Han¹, Kristina A. Archer², Thomas O'Neill², Roy S. Herbst⁴,
Agedi N. Boto^{1,3}, Miguel F. Sanmamed¹, Solomon Langermann², David L. Rimm^{3,4} and Lieping Chen^{1,4*}

NEXTCURE HAS DELIVERED A ROBUST PRODUCT PIPELINE

PROGRAMS	CELLS	DISCOVERY	PRECLINICAL	PHASE 1	PHASE 2	PHASE 3	NEXT MILESTONE	WORLDWIDE RIGHTS
PRODUCT CANDIDATES								
NC318 (S15)	Tumors and macrophages	ONCOLOGY					Phase 2 data by end of 2020	NextCure
NC410 (LAIR-1)	Dendritic & T cells	ONCOLOGY					IND filing in Q1 2020	NextCure
DISCOVERY AND RESEARCH PROGRAMS								
Multiple Programs	Immune cells						First IND filing in early 2021	NextCure
FIND-IO Platform	Multiple cell types						First IND filing in late 2022	<div>Lilly</div> <div>NextCure</div>

NC318

HUMANIZED
MONOCLONAL
ANTIBODY



Phase 1/2 Clinical Trial

TARGET

Siglec-15 (“S15”)

CELL TYPES

Tumors &
macrophages

MOA

Designed to block
S15-induced
immunosuppression

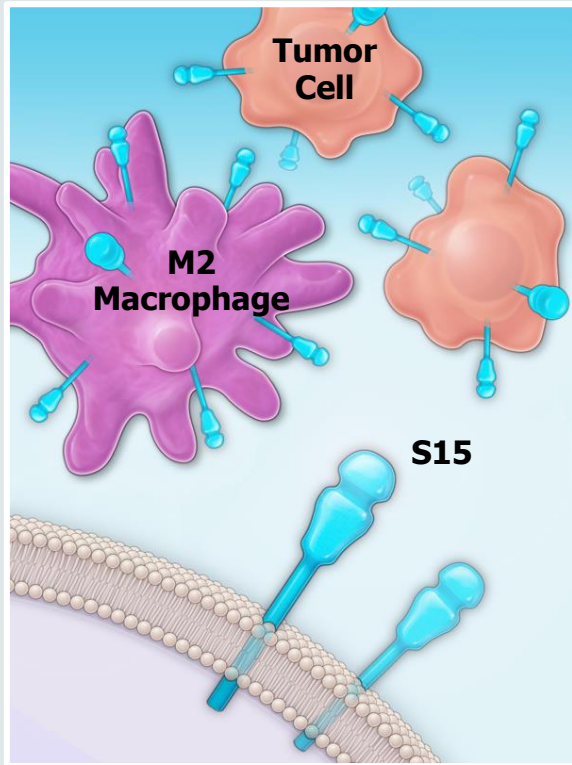
INDICATIONS

Advanced or metastatic
solid tumors, including
NSCLC, ovarian, head &
neck and triple negative
breast cancers

S15 AS A TARGET

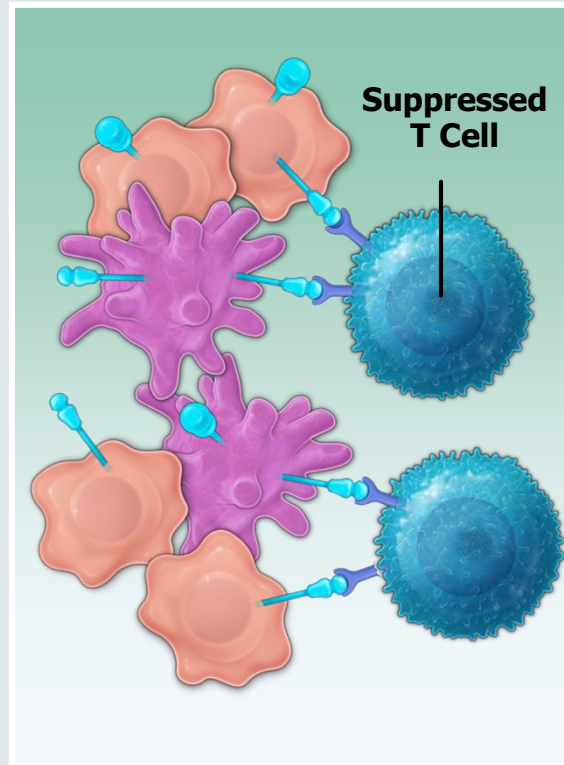
EXPRESSION

Tumors &
Macrophages



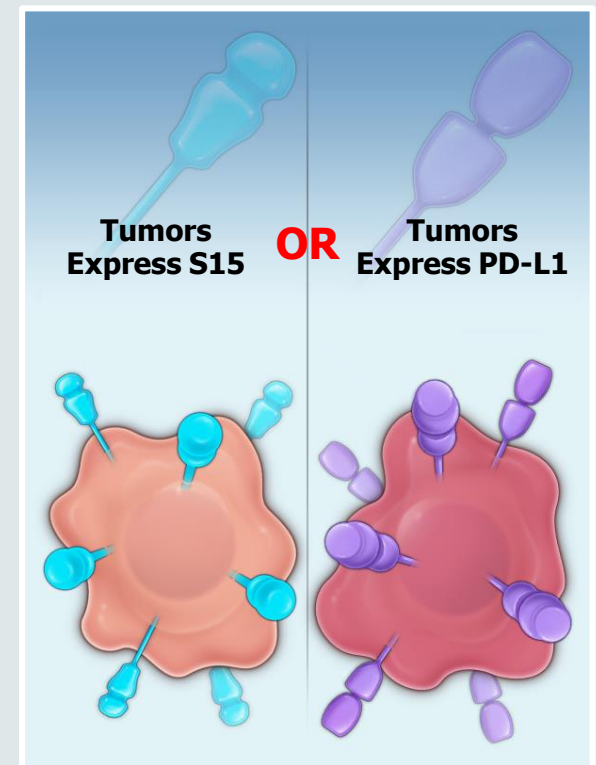
FUNCTION

Potently Suppresses
T Cell Function

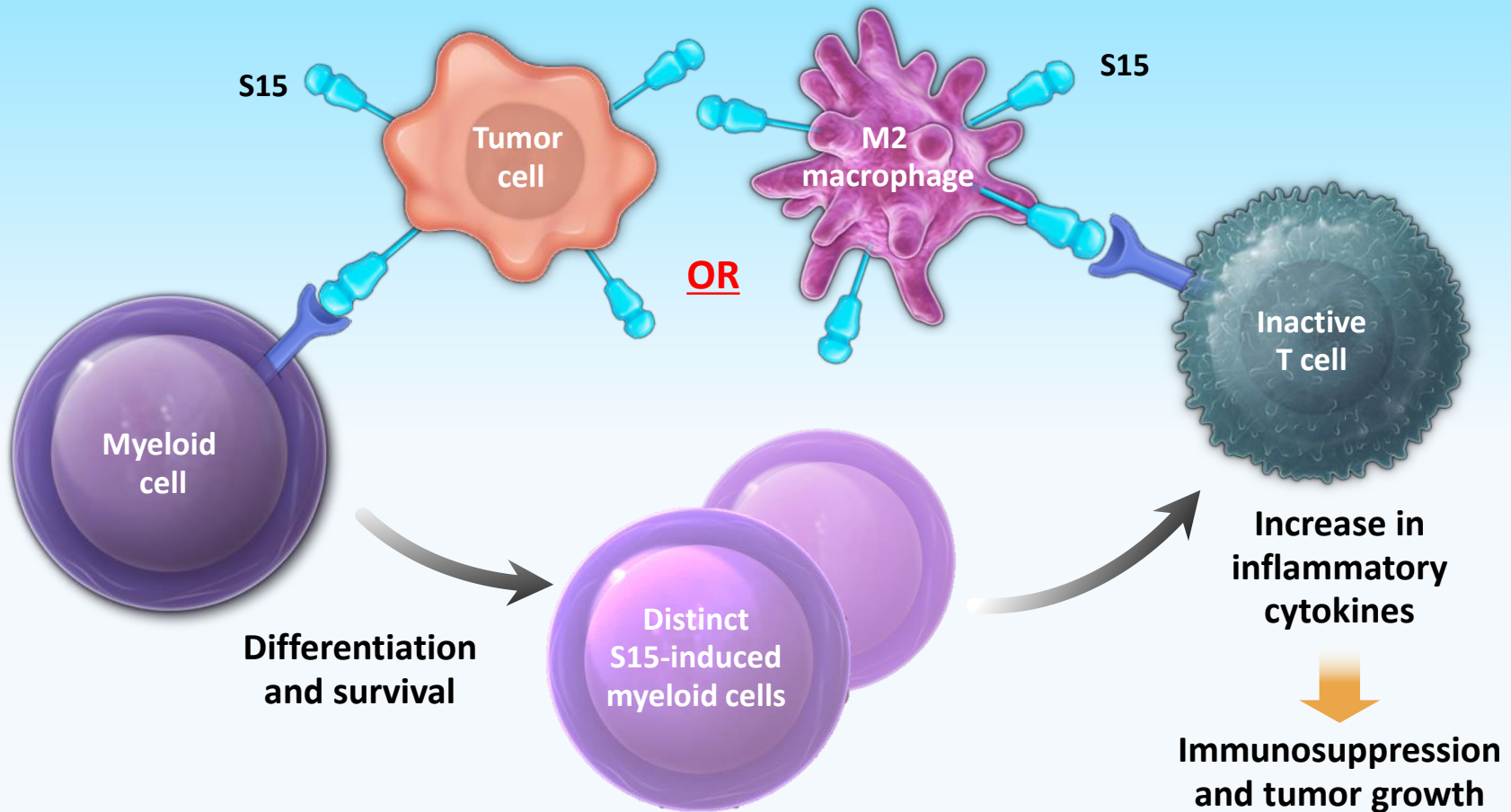


NON-RESPONDERS

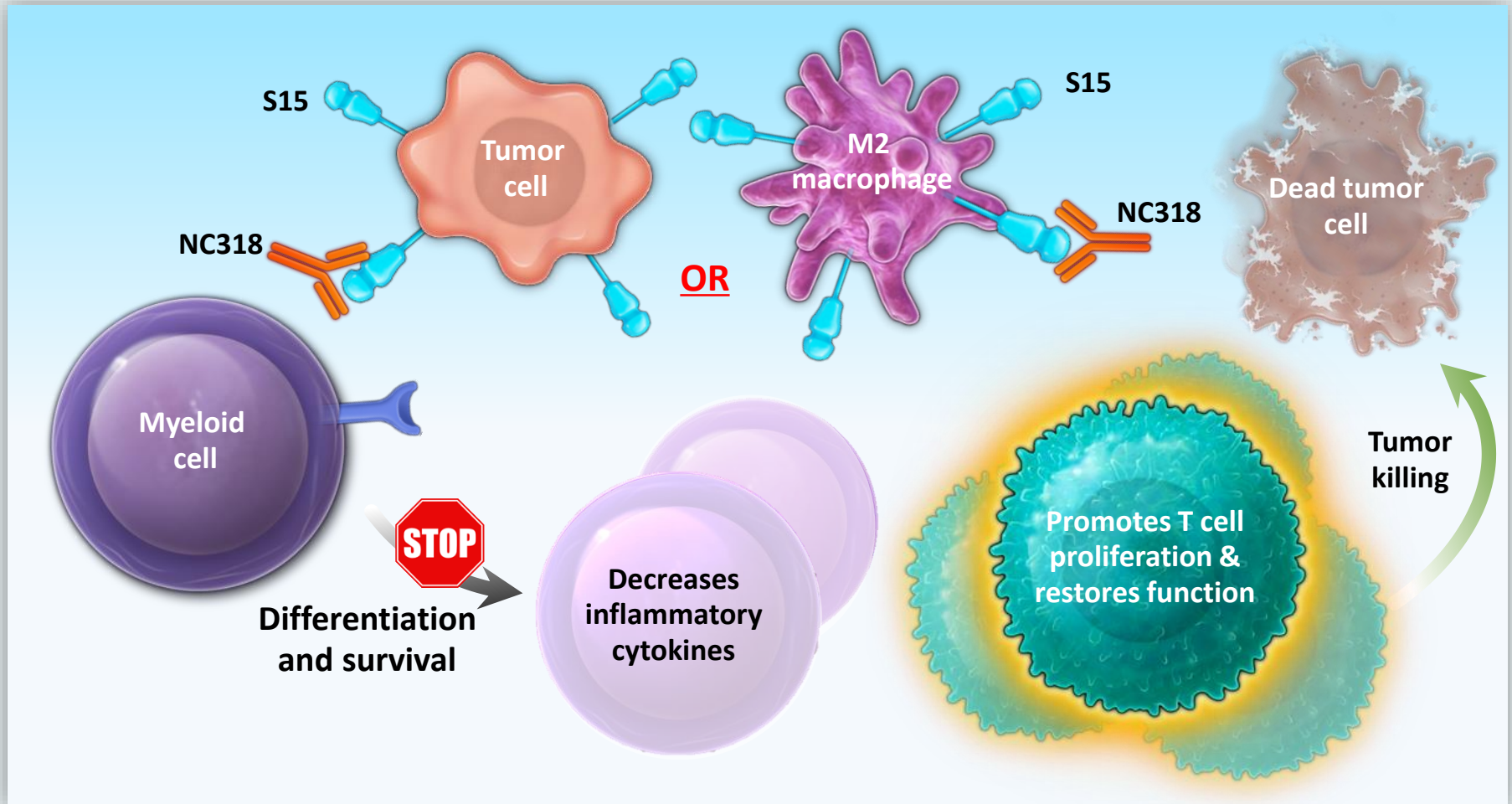
Generally Non-Overlapping
with PD-L1 Expression



S15 IS HIGHLY IMMUNOSUPPRESSIVE IN THE TME OF MULTIPLE TUMOR TYPES



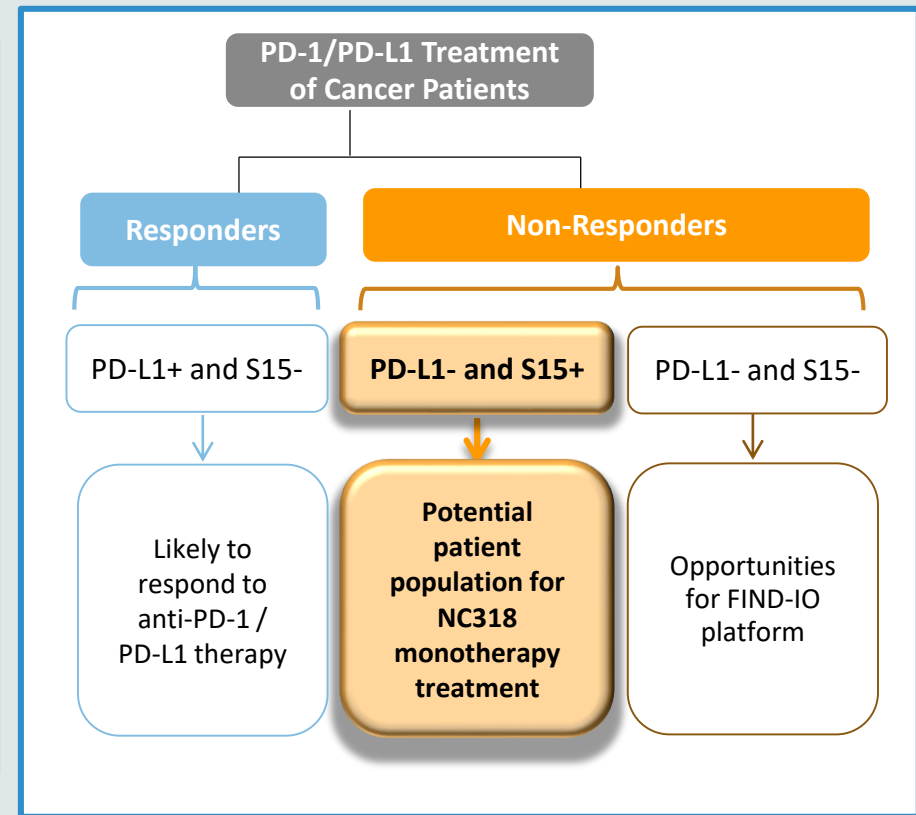
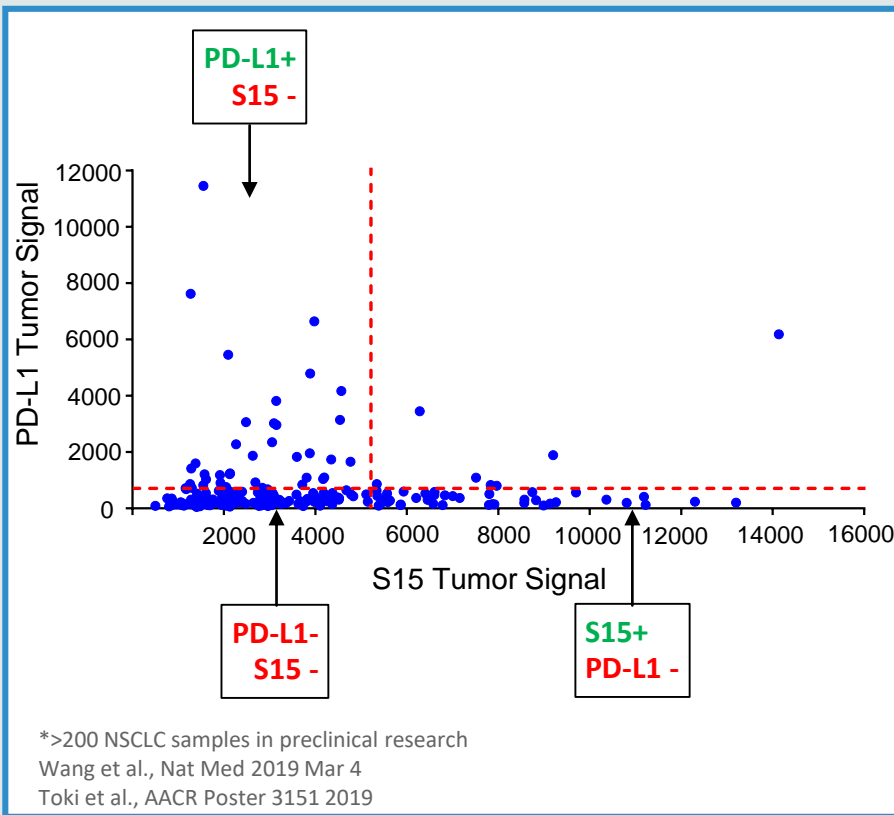
NC318 IS DESIGNED TO BLOCK IMMUNOSUPPRESSIVE ACTIVITY INDUCED BY S15



NC318: A POTENTIAL TREATMENT OPTION FOR PD-1/PD-L1 NON-RESPONDERS

S15 AND PD-L1 EXPRESSION GENERALLY DO NOT OVERLAP IN NSCLC TUMORS*

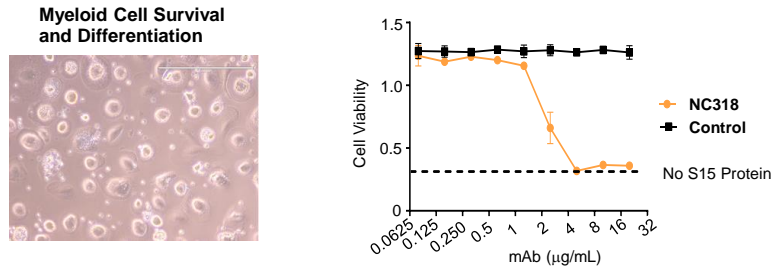
POTENTIAL NEW TREATMENT OPTIONS FOR PD-1/PD-L1 NON-RESPONDERS



NC318 RESTORED IMMUNE FUNCTION *IN VITRO*

INHIBITED

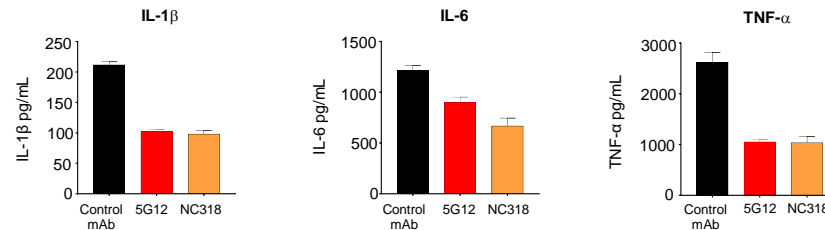
Myeloid Cell
Differentiation
and Survival



Blocked survival of
myeloid cells

DECREASED

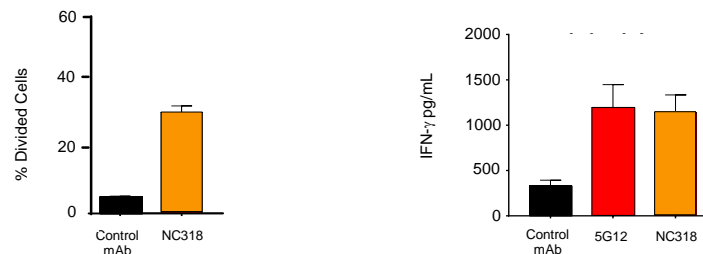
Pro-Inflammatory and
Pro-Tumorigenic
Cytokines



Decreased IL-1 β ,
IL-6 & TNF- α

PROMOTED

Tumor-Specific
T Cell Function



Increased T cell
proliferation &
IFN- γ production

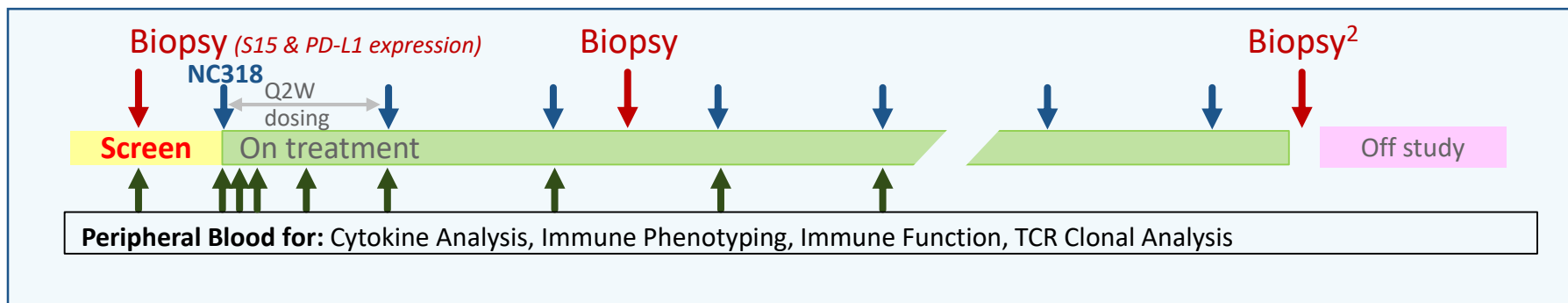
NC318 MONOTHERAPY TRIAL UNDERWAY DESIGNED FOR RAPID PROOF-OF-CONCEPT

PHASE 1

- Opened in 4Q 2018; preliminary data announced November 2019
- Dose-escalation¹
- Safety, tolerability & biomarker readouts
- Advanced or metastatic solid tumors
- All comers regardless of PD-L1 or S15 expression status

PHASE 2

- Opened in 4Q 2019; initial data by end of 2020
- Efficacy assessment
- Tumor types shown to have elevated S15 expression, including NSCLC, ovarian, head & neck and TNBC
- PD-L1 TPS <50%



(1) Dose escalation evaluated 7 dose cohorts (8 mg – 1,600 mg or approximately 0.1 - 20 mg/kg) administered every 2 weeks

(2) In Phase 2 portion of trial

NC318 PHASE 1 TRIAL STATUS AS OF NOVEMBER 9, 2019

Dose Escalation – Enrollment Complete

ENROLLMENT

- 49 patients dosed
- 15 different tumor types
- Median of 3 prior therapies
- All comers regardless of PD-L1 or S15 expression status

SAFETY

- No DLTs through 800 mg
- 1 DLT at 1600 mg: Grade 3 pneumonitis
- Common irAEs observed, including diarrhea, rashes, vitiligo, arthralgias

RESPONSES

- Evaluations every 8 weeks
- 1 confirmed CR (55+ weeks)
- 1 confirmed PR (28+ weeks)
- 14 durable SD (≥ 16 weeks)

• Angeles Clinic

• MSKCC

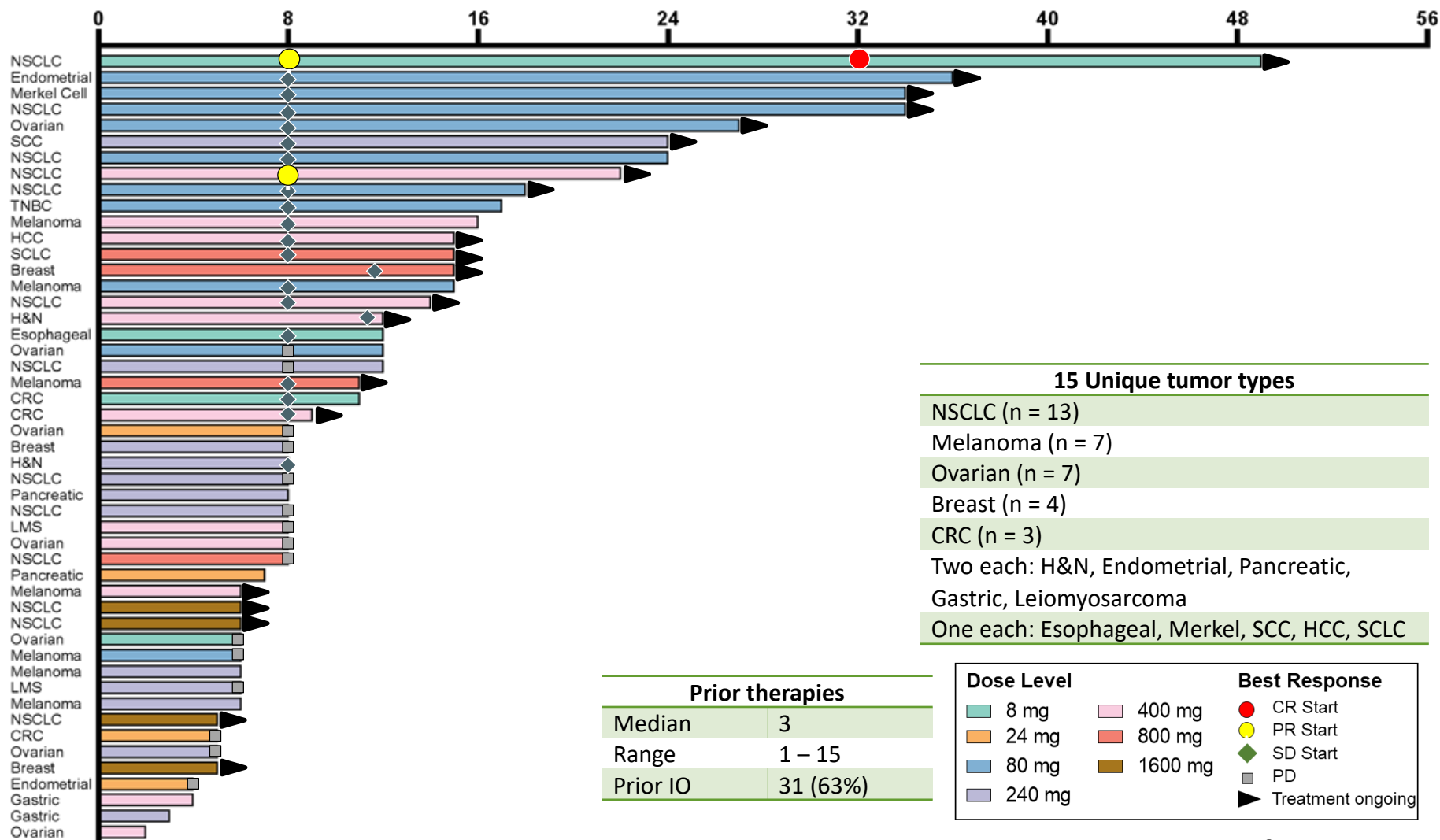
• Next Oncology

• NYU

• Yale University

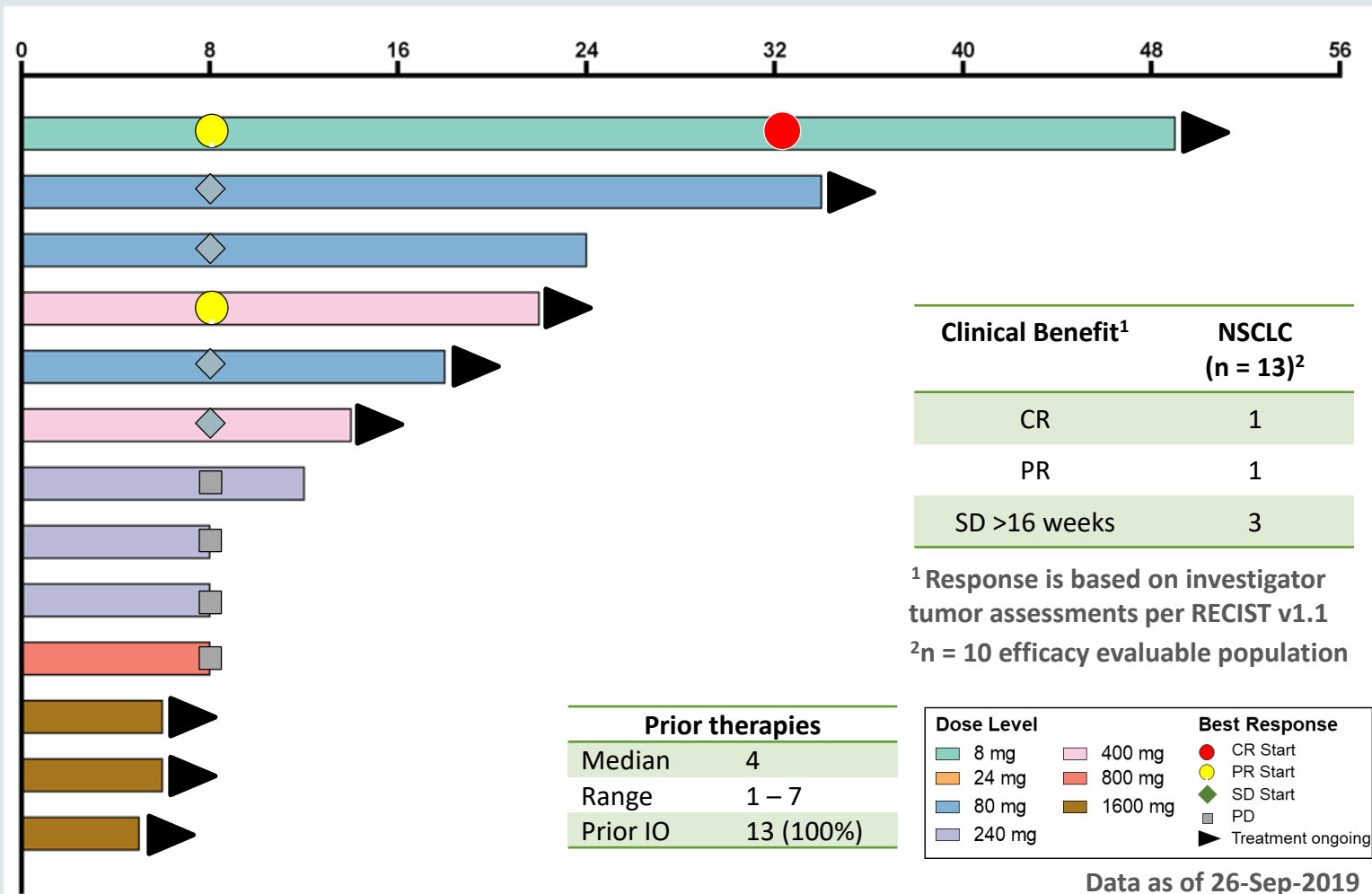
Most common AEs: diarrhea, infusion reactions, fatigue, headaches, pruritis, elevated amylase and elevated lipase

TREATMENT DURATION IN WEEKS FOR ALL PHASE 1 PATIENTS



Data as of 26-Sep-2019

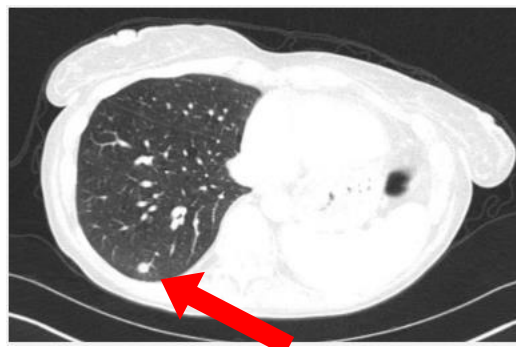
DURABLE CLINICAL BENEFIT FOR PD-1 REFRACTORY NSCLC PATIENTS



NC318: SINGLE AGENT ACTIVITY IN PD-1 REFRACTORY NSCLC

CONFIRMED COMPLETE RESPONSE

BASELINE



Target lesion

Week 16



Target lesion gone

56 y/o NSCLC with
multiple lesions
(PD-L1 TPS <50%)
8 mg every 2 weeks

Prior therapies:
Nivolumab (best
response stable disease
then progression)

CONFIRMED PARTIAL RESPONSE

BASELINE



Week 16



Target lesions -71%

74 y/o NSCLC
(PD-L1 TPS <50%)
400 mg every 2 weeks

Prior therapies:
Immunotherapy:
LAG3/PD-1 (best
response stable disease
then progression)

CONCLUSIONS FROM PHASE 1 PORTION OF NC318 TRIAL

- NC318 has been well tolerated across multiple dose levels
- Adverse event profile consistent with other approved immunotherapies
- Predictable pharmacokinetic profile
- NC318 has shown encouraging single-agent anti-tumor activity
 - PD-1 refractory NSCLC: 1 CR, 1 PR, and stable disease in 3 patients
 - Durable stable disease (>24 weeks observed in multiple tumor types)
- Phase 2 enrollment underway

NC318 PHASE 2 TRIAL STATUS AS OF NOVEMBER 9, 2019

Dose Expansion – Enrolling

TUMOR TYPES

NSCLC

H&N

Ovarian

TNBC

DESIGN

- Biopsies required
- PD-L1 TPS <50%
- S15 evaluated retrospectively
- Monotherapy
- 400 mg every 2 weeks

DELIVERABLES

Initial Phase 2 data by the end of 2020

• Angeles Clinic

• MSKCC

• Next Oncology

• NYU

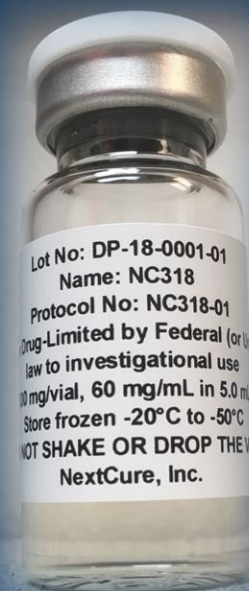
• Yale University

NC318

DESIGNED TO RESTORE
IMMUNE FUNCTION IN A
HIGHLY SUPPRESSIVE TUMOR
MICROENVIRONMENT



- ☒ MOA / Preclinical studies complete
 - Relieved S15-mediated inhibition of T cells
 - Increased IFN- γ production
 - Decreased inflammatory cytokines
- ☒ Completed enrollment of Phase 1
- ☒ Reported preliminary data at SITC 2019
- ☐ Initiate Phase 2 combination trial with SOC chemotherapies
- ☐ Report initial Phase 2 data by end of 2020



NC410

DECOY HUMAN FUSION PROTEIN
TARGETING THE TME



IND Filing Expected Q1 2020

TARGET

Leukocyte-
Associated
Immunoglobulin-
like Receptor-1
(LAIR-1)

CELL TYPES

Dendritic cells
and T cells

MOA

Promotes T cell
function and
dendritic cell activity

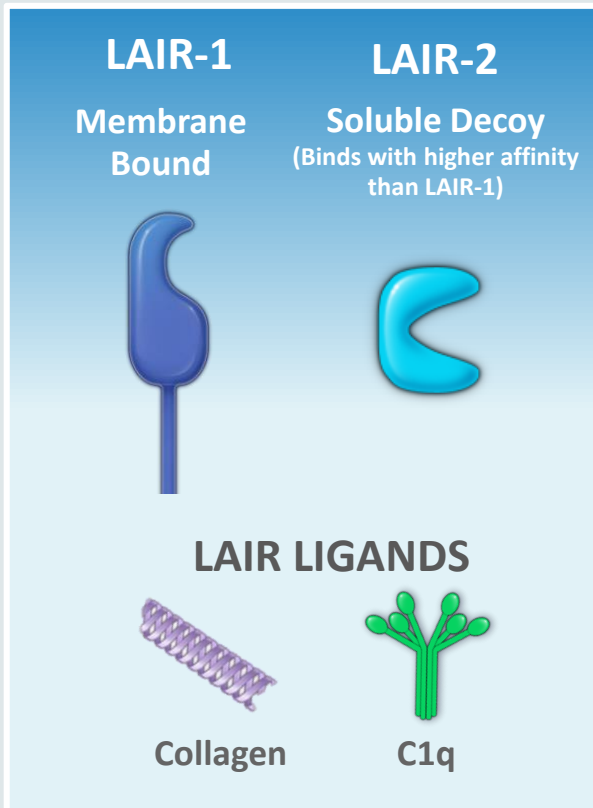
INDICATIONS

Advanced or
metastatic solid
tumors

LAIR-1 & LAIR-2 FUNCTIONAL RELATIONSHIP

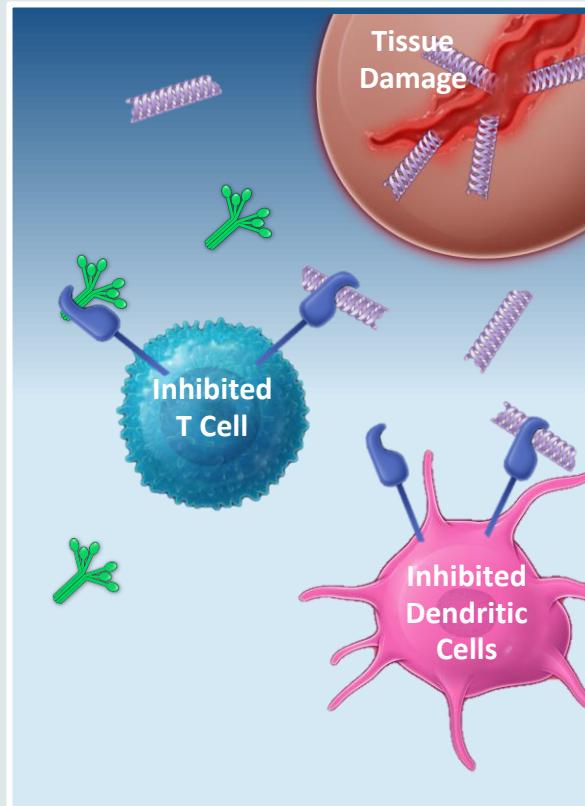
LAIR & LIGANDS

LAIR-1 and LAIR-2 bind collagen and C1q



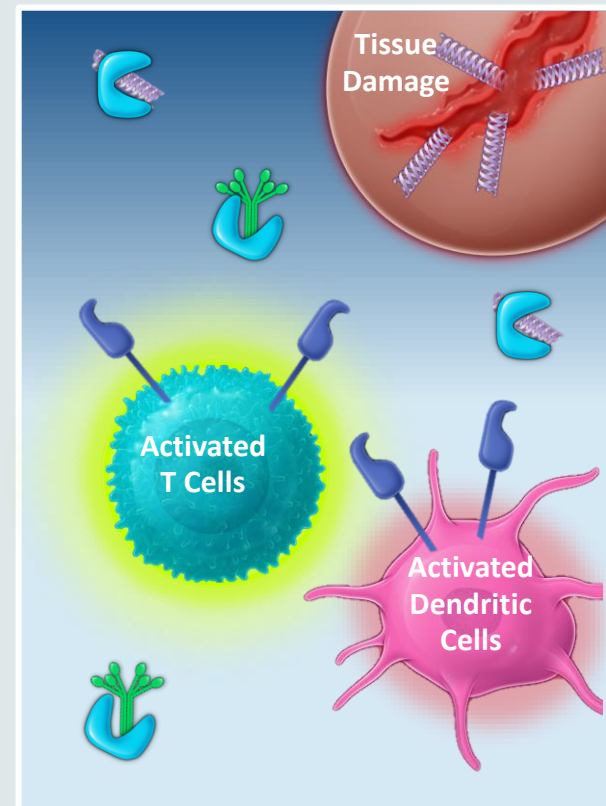
LAIR-1

Ligands are expressed in response to inflammation & inhibit immune function



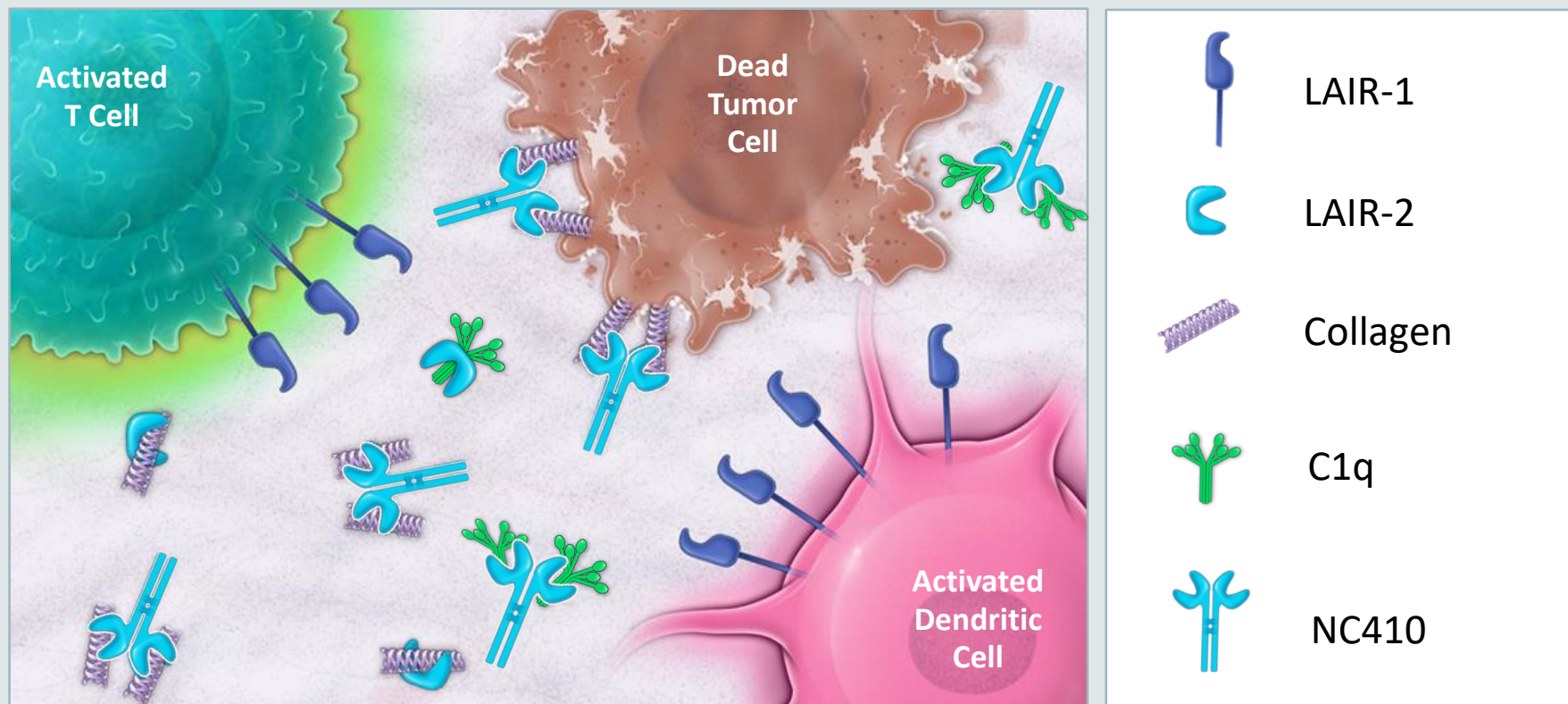
LAIR-2

LAIR-2 modulates LAIR-1 mediated inhibition



NC410 IS DESIGNED TO PREVENT IMMUNE SUPPRESSION CAUSED BY LAIR-1

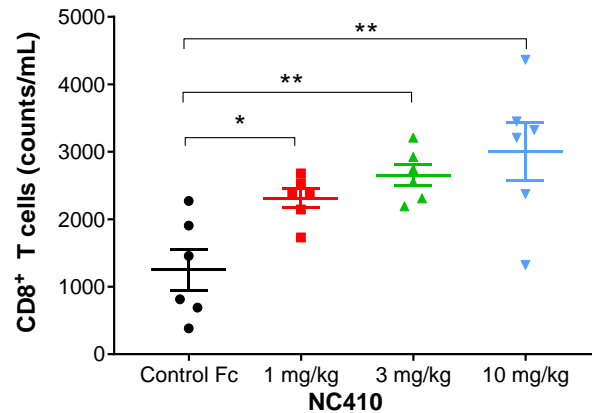
NC410 is a Fusion Protein of LAIR-2 and a Decoy for LAIR-1



NC410 Promotes T Cell Function and Dendritic Cell Activation

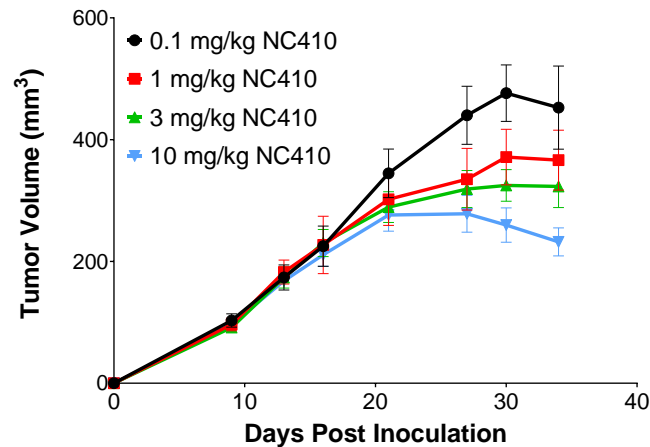
NC410 ENHANCED T CELL EXPANSION AND RELIEVED IMMUNOSUPPRESSION

BLOCKED
Suppression



Human CD8⁺ T cell
expansion *in vivo*

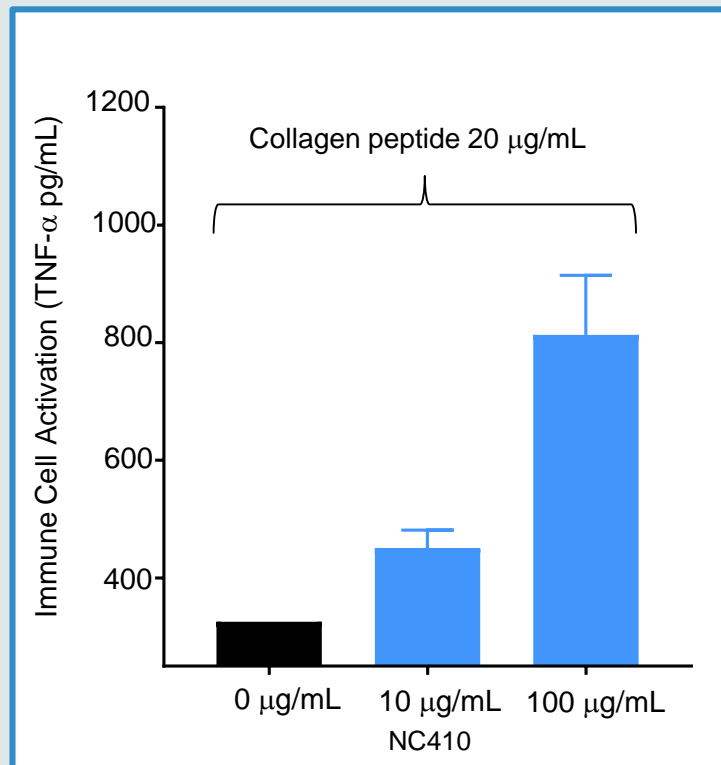
DECREASED
Tumor Volume



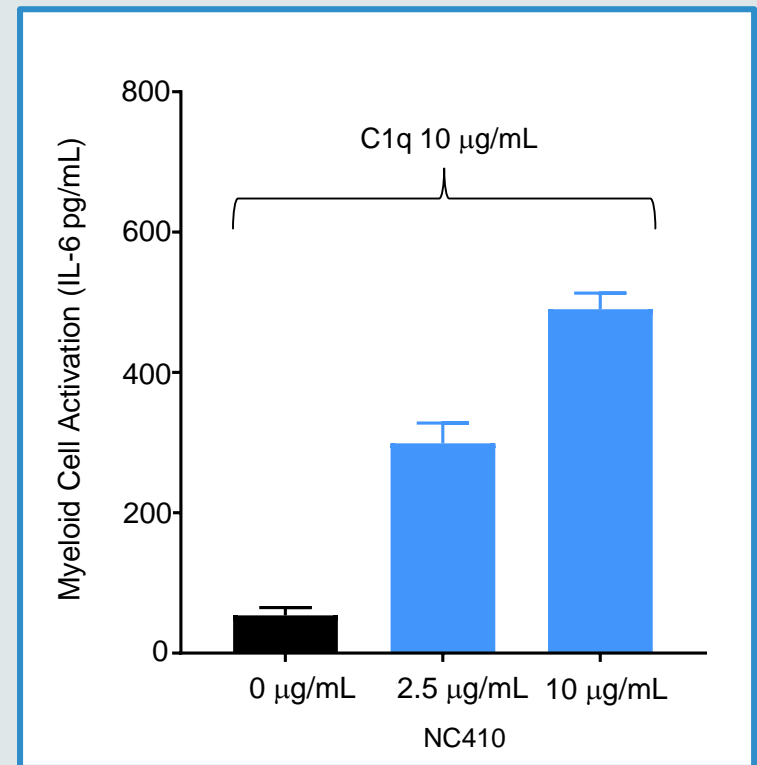
Human PBMCs in mice:
CD8⁺ T cell activity
decreased tumor volume
in HT29 model

NC410 PROMOTED IMMUNE CELL ACTIVATION IN THE PRESENCE OF COLLAGEN AND C1Q

REVERSED COLLAGEN SUPPRESSION OF HUMAN PBMCS



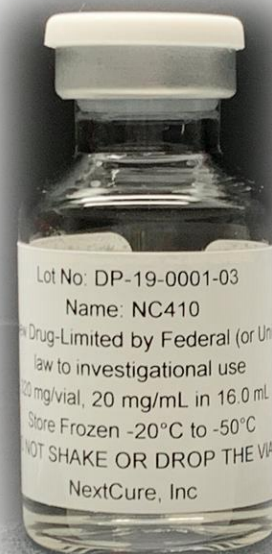
REVERSED C1Q SUPPRESSION OF HUMAN MYELOID CELLS



NC410 SUMMARY

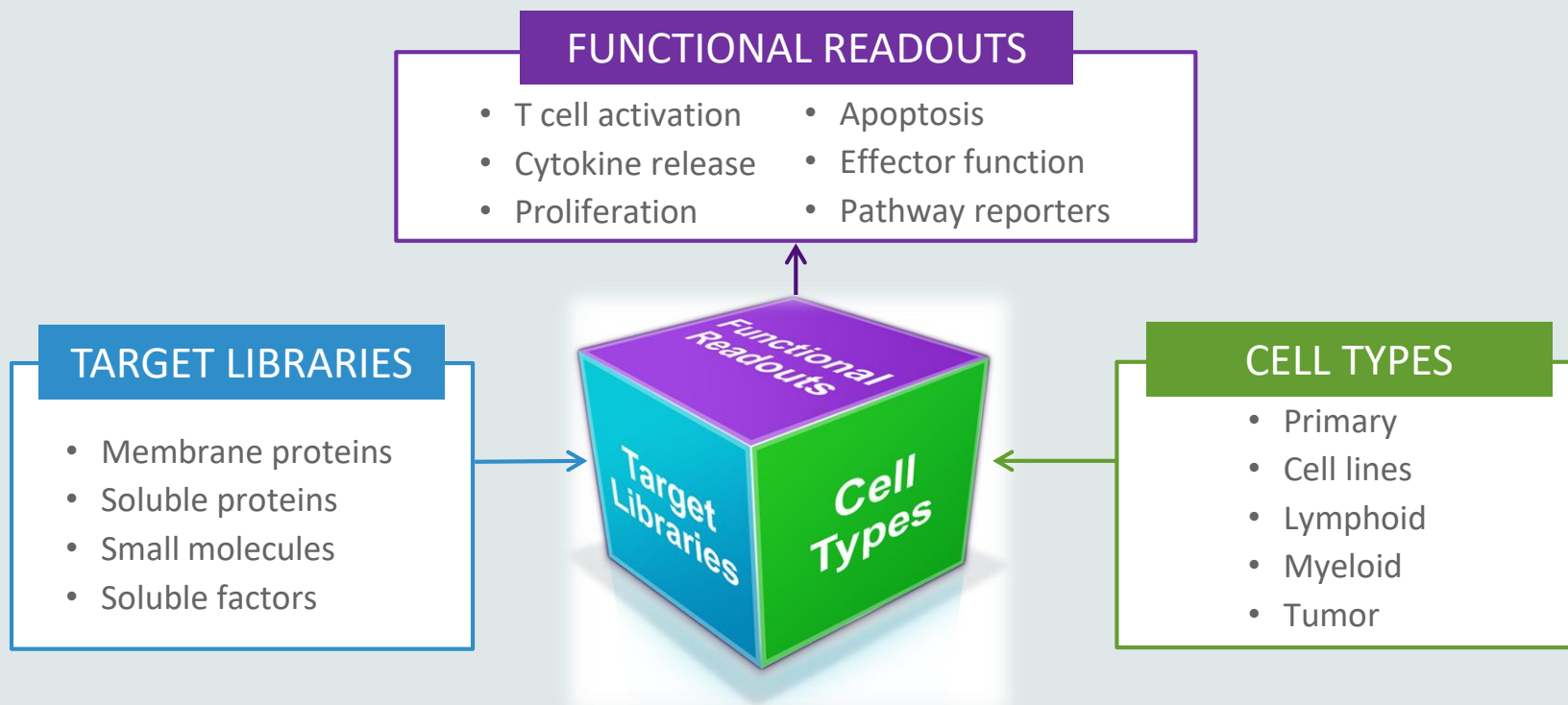


- ☒ Based on normal immune regulatory mechanism
- ☒ Promoted T cell function and dendritic cell activity in preclinical studies
- ☒ Designed to alleviate tumor-mediated immunosuppression
- ☒ IND-enabling tox studies complete
- ☒ cGMP manufacturing
- ☐ IND filing expected Q1 2020

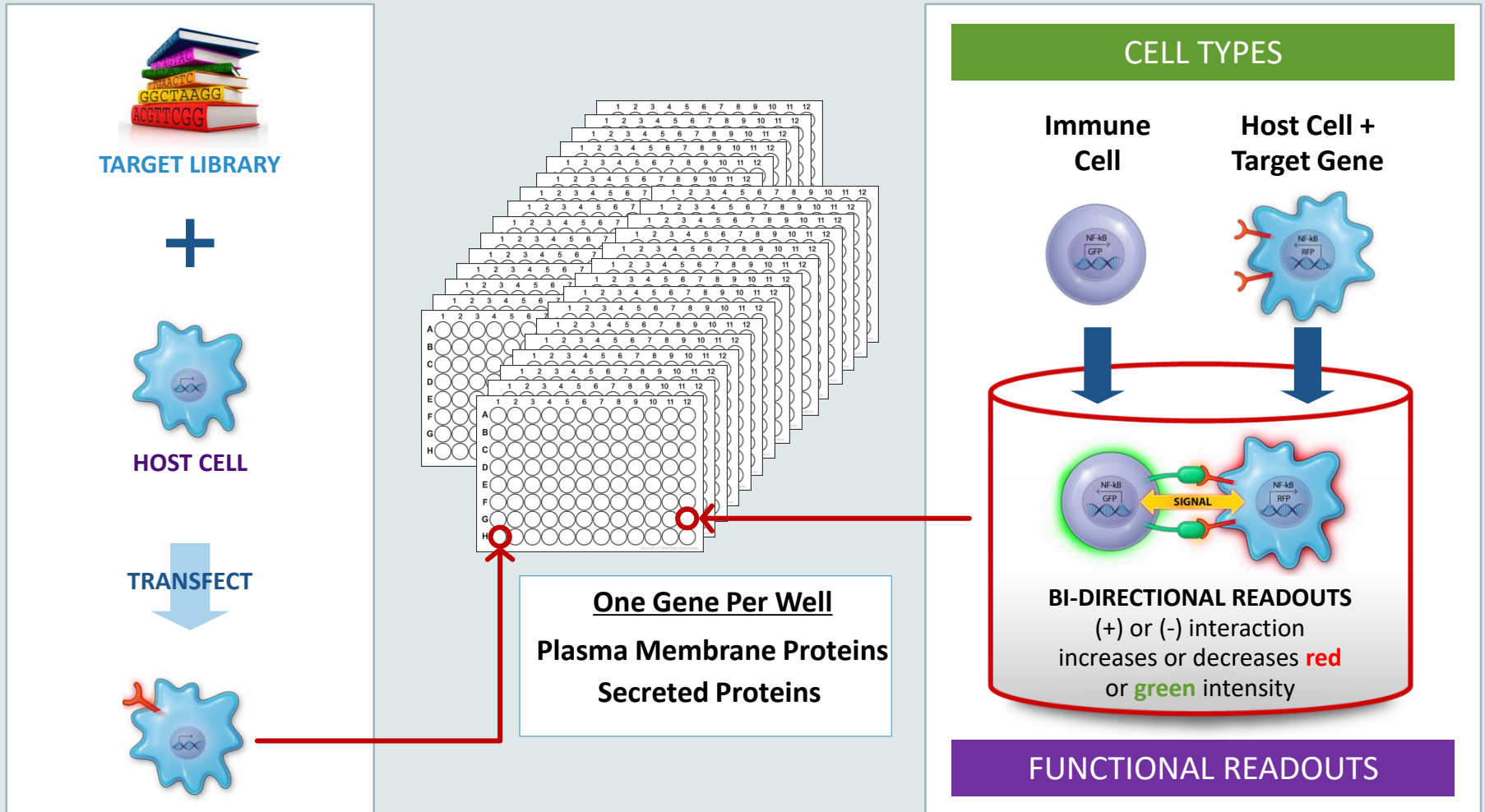


FINDING SOLUTIONS WITH A POWERFUL DISCOVERY ENGINE

Functional, Integrated, NextCure Discovery in Immuno-Oncology

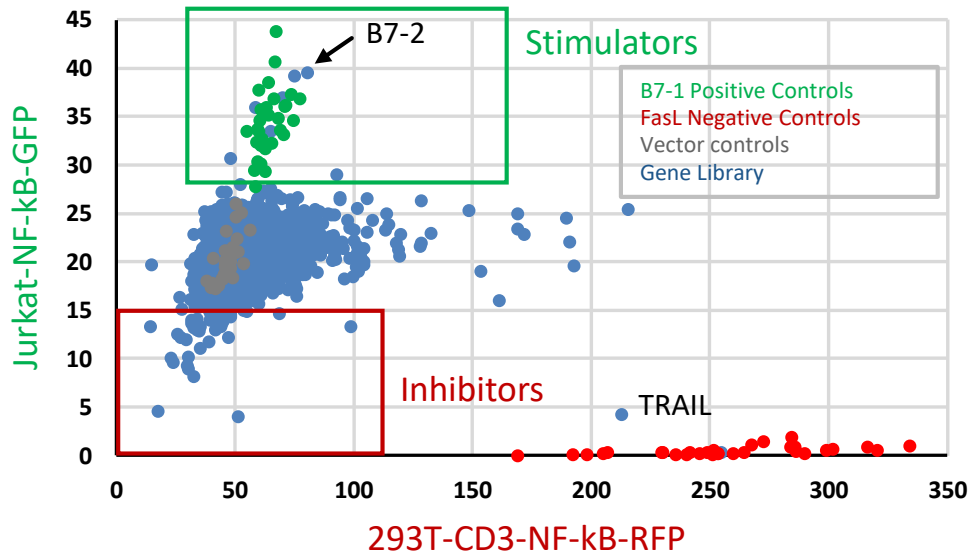


FIND-IO SCREENING METHODOLOGY

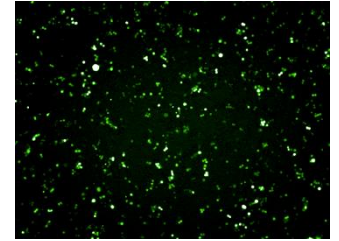


JURKAT “T CELL LINE” SCREENING AND VALIDATING FIND-IO HITS

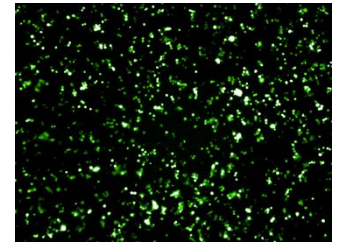
Identified Novel T Cell Stimulators and Inhibitors



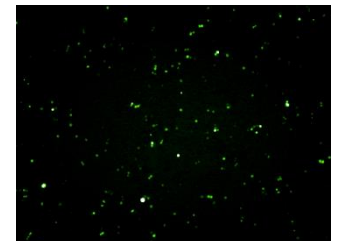
Vector Control



Stimulatory B7-2



Inhibitory TRAIL



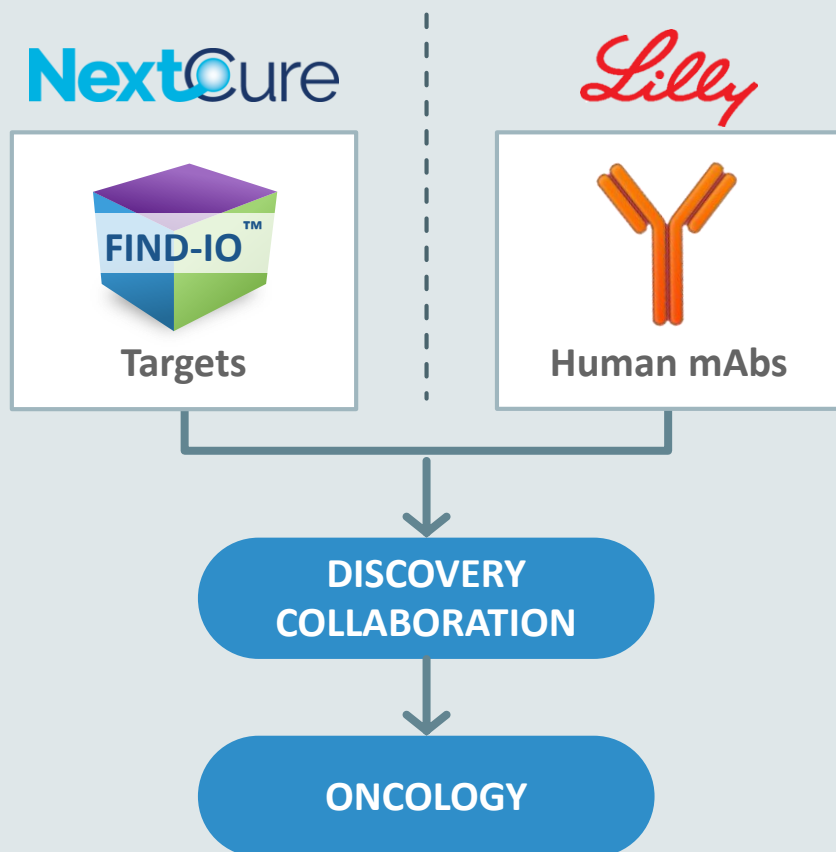
REPRODUCIBILITY

ROBUSTNESS

RELEVANCY

LILLY – NEXTCURE PARTNERSHIP TO VALIDATE PLATFORM AND APPROACH

Synergies



Overview

Structure

- Each party has options to exclusively license certain antibodies

Terms

- Upfront: \$25M
- Equity investment: \$15M
- R&D support
- Option payments
- Development & sales milestones
- Royalties

DIVERSIFICATION BEYOND ONCOLOGY

TODAY



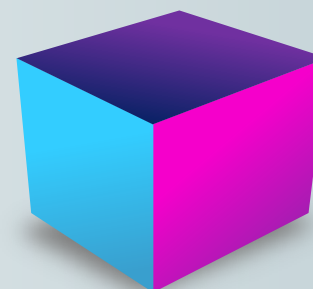
Oncology

TOMORROW



Autoimmunity

FUTURE



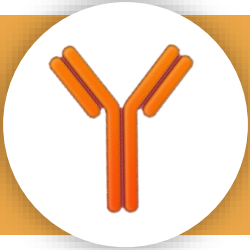
Neuro-Inflammation

Gene Libraries

Immune Cells

Functional Screening

ANTICIPATED NEAR-TERM MILESTONES



NC318

- Initiate Phase 2 combination trial with standard of care chemotherapies in 1H 2020
- Report initial Phase 2 data by end of 2020



NC410

File IND and initiate clinical development in Q1 2020



DISCOVERY

Identify novel targets and initiate validation



Committed to Addressing the Unmet Needs of Cancer Patients
With New Solutions

FOCUSED
Approach

PROVEN
Momentum

INNOVATIVE
Platform

EXPERIENCED
Team

FUTURE
Deliverables