



NEXT-GENERATION IMMUNOMEDICINES

SEPTEMBER 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 May 8, 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: Phase 1/2 topline data in Q4 2019
- NC410 (LAIR-1): IND expected Q1 2020
- Additional research and development programs
- Manufacturing: dedicated, state-of-the-art facility

Platform for Novel Target Discovery

- FIND-IO functional screening discovery engine
- Oncology partnership with Lilly: \$40M upfront and equity
- 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

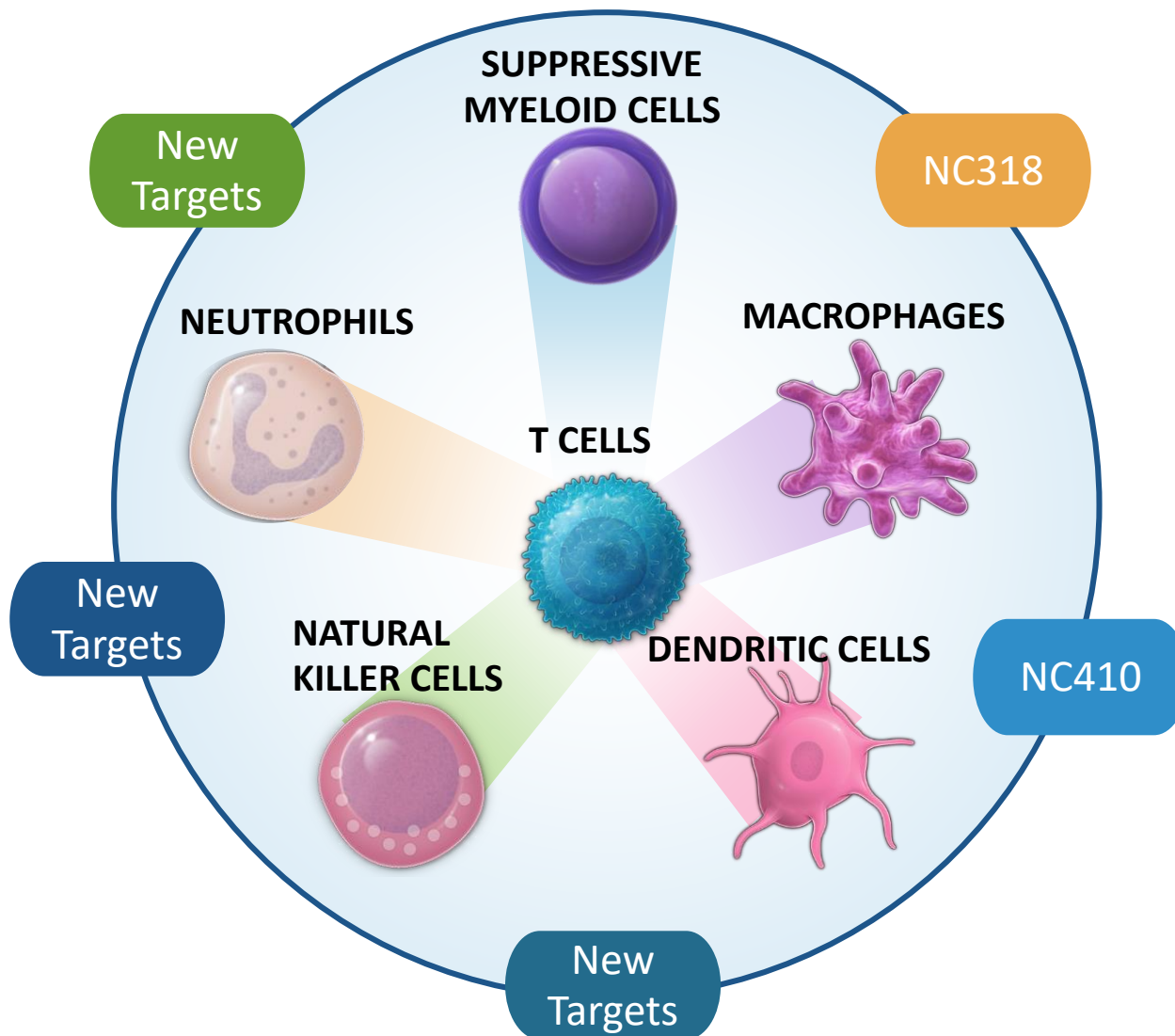
quality of life



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

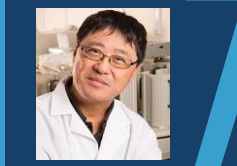
Michael Richman CEO				
Steve Cobourn, CPA CFO				
Kevin N Heller, MD CMO				
Sol Langermann, PhD CSO				
Jim Bingham, PhD CDO				
Timothy Mayer, PhD SVP, Corporate Development				
Linda Liu, PhD SVP, Research				
Sebastien Maloveste, PhD VP, Business Development				
Dallas Flies, PhD VP, Discovery Research				

WORLD-RENOWNED SCIENTIFIC FOUNDER AND KEY COLLABORATION

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 ROBUST PRODUCT PIPELINE IN *LESS THAN 3 YEARS*

PROGRAMS	CELLS	DISCOVERY	PRECLINICAL	PHASE 1	PHASE 2	PHASE 3	NEXT MILESTONE	WORLDWIDE RIGHTS
PRODUCT CANDIDATES								
NC318 (S15)	Tumors and macrophages	ONCOLOGY					Phase 1 complete in Q4 2019	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> NextCure

NC318

HUMANIZED
MONOCLONAL
ANTIBODY



Phase 1/2 Clinical Trial Initiated
October 2018

TARGET

Siglec-15 (“S15”)

CELL TYPES

Tumors &
macrophages

MOA

Designed to block
S15-induced
immunosuppression

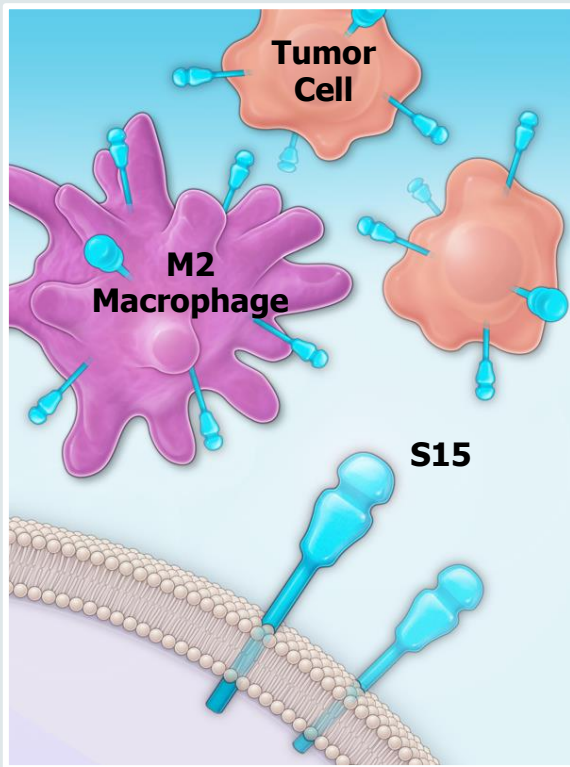
INDICATIONS

Advanced or
metastatic solid
tumors, which could
include ovarian,
NSCLC, and head &
neck cancers

S15 AS A TARGET

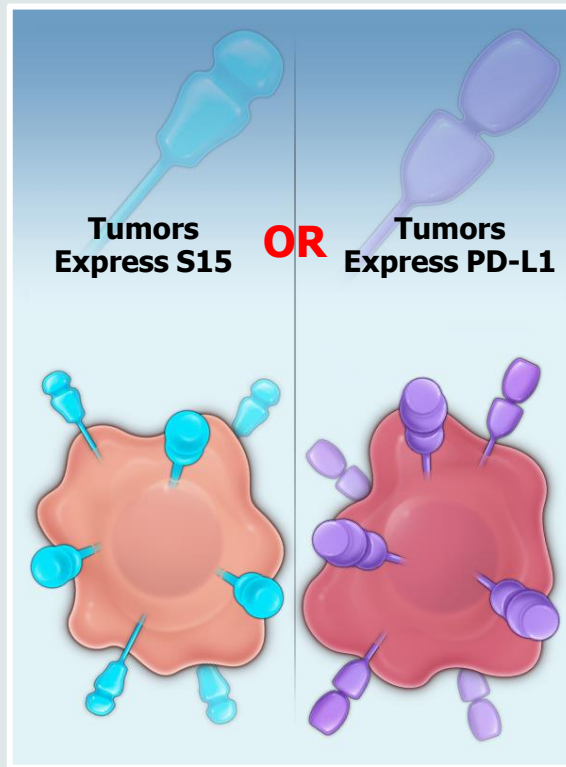
EXPRESSION

Tumors &
Macrophages



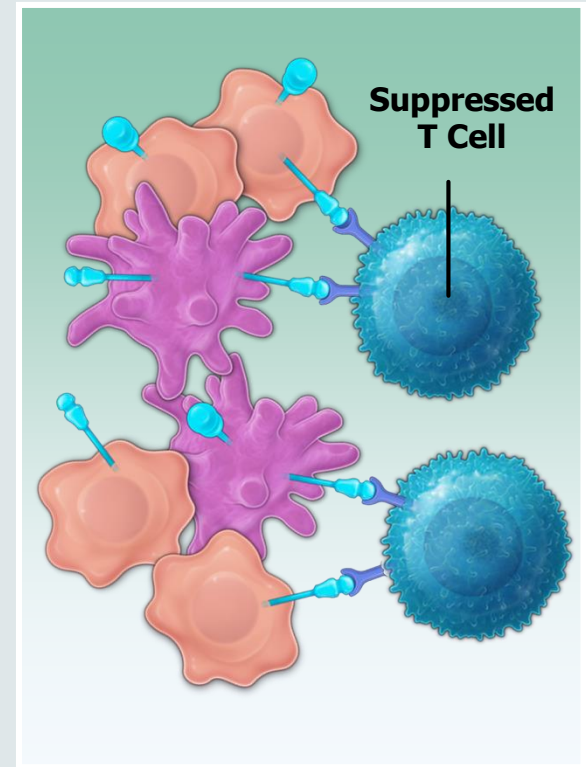
NON-RESPONDERS

Generally Non-Overlapping
with PD-L1 Expression

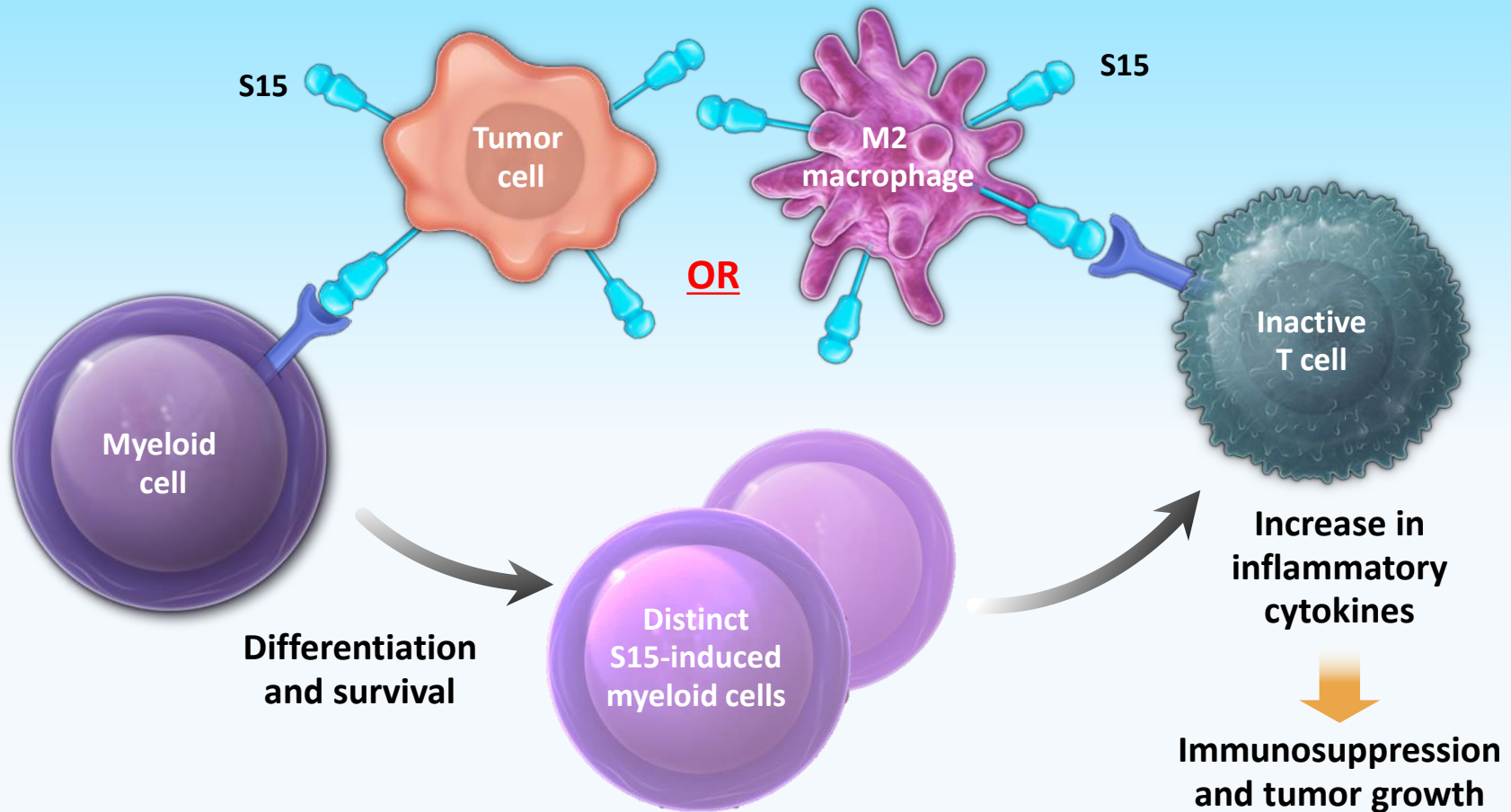


FUNCTION

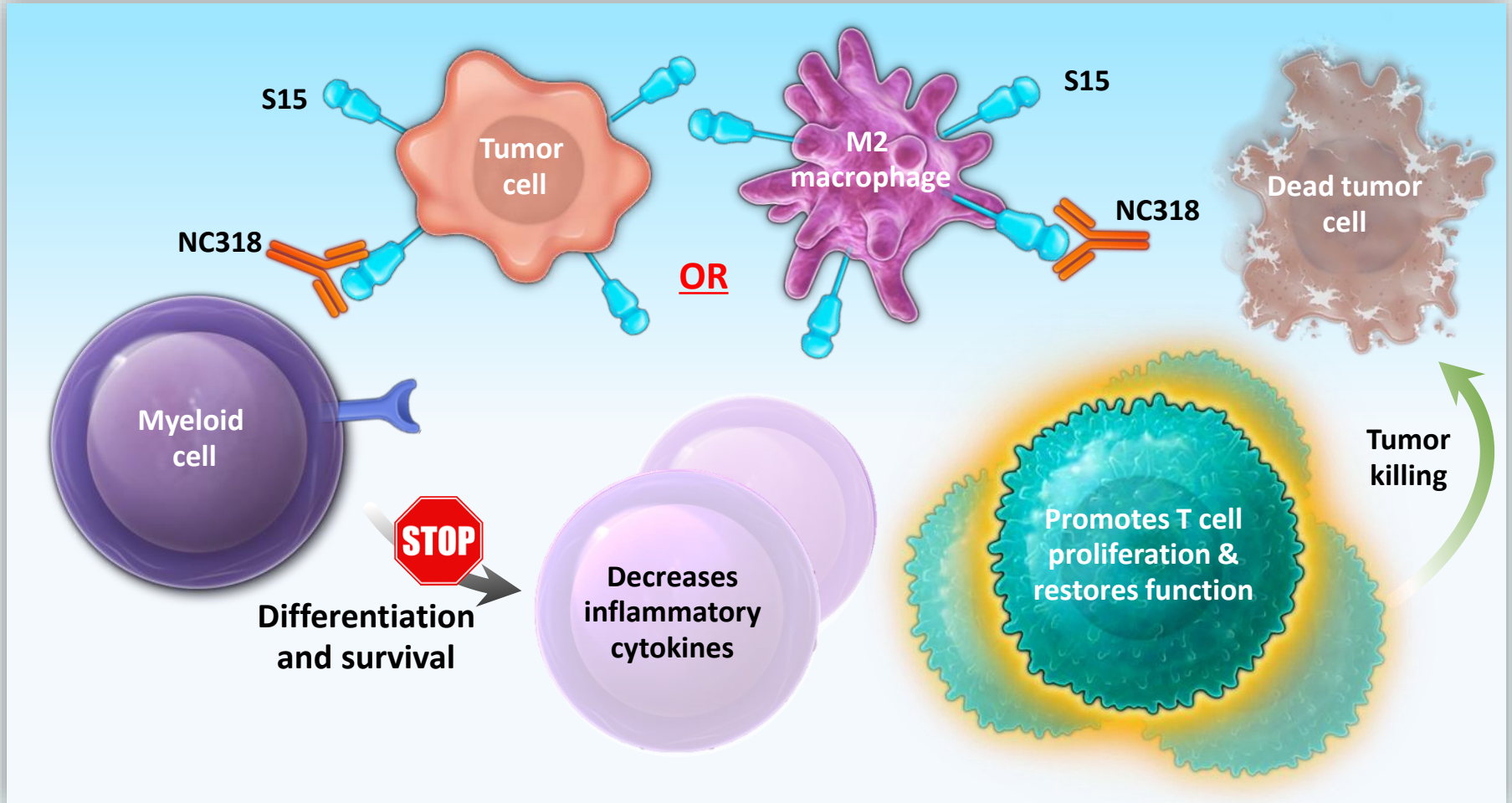
Potently Suppresses
T Cell Function



S15 IS HIGHLY IMMUNOSUPPRESSIVE IN THE TME OF MULTIPLE TUMOR TYPES



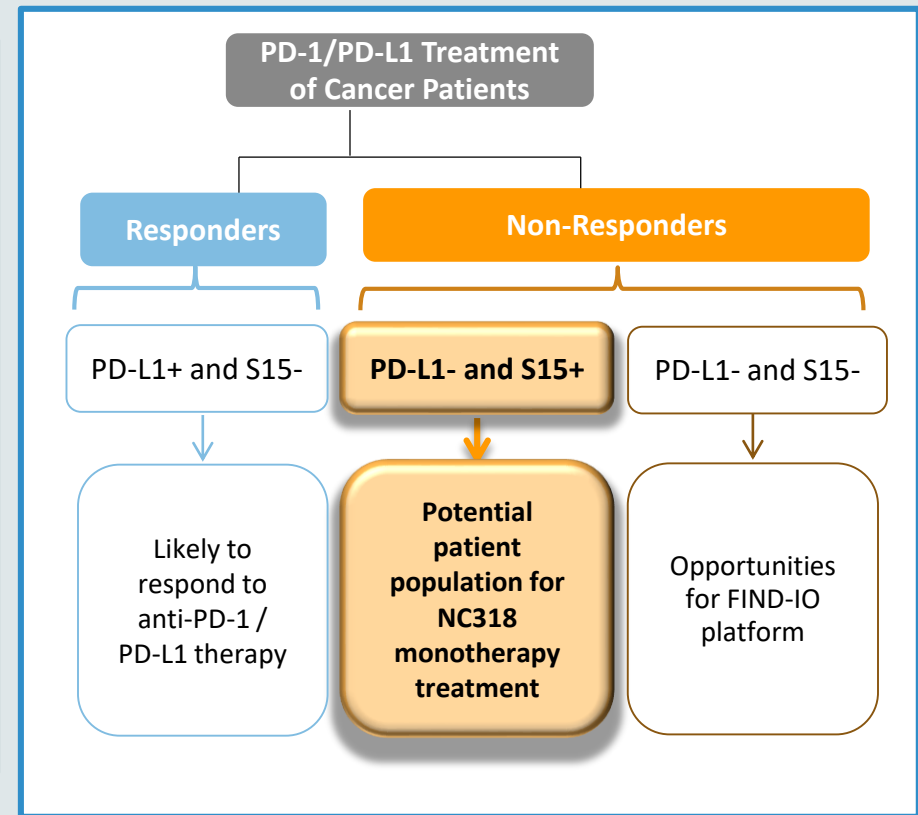
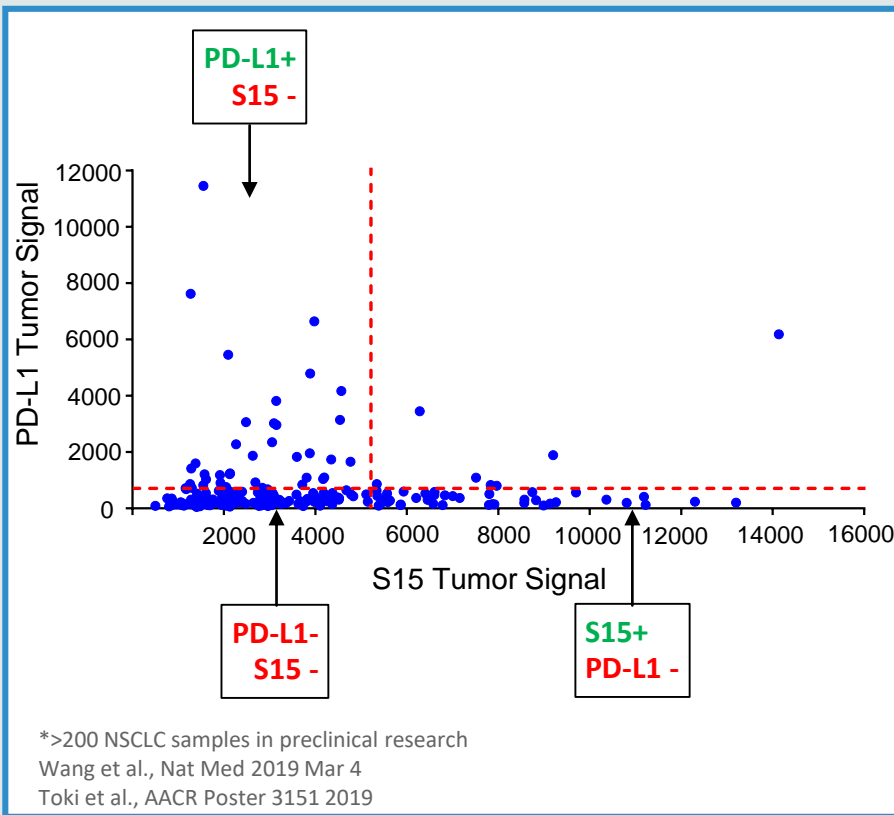
NC318 IS DESIGNED TO BLOCK IMMUNOSUPPRESSIVE ACTIVITY INDUCED BY S15



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

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

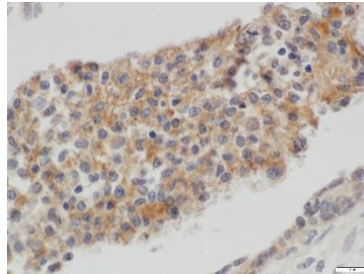
POTENTIAL NEW TREATMENT OPTIONS FOR PD-1/PD-L1 NON-RESPONDERS



S15 IS CLINICALLY AND FUNCTIONALLY RELEVANT

Clinical Relevance

S15 IN TUMORS



- Increased expression on tumor cells and immunosuppressive macrophages in multiple cancer types
- Minimal expression in normal tissues

Functional Relevance

S15 KNOCK-OUT MODEL

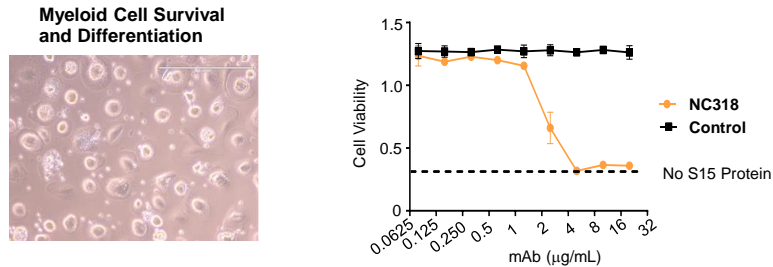


- S15-deficient mice showed
 - Enhanced antigen-specific T cell responses *in vivo*
 - Delayed tumor progression
 - Increase in survival

NC318 RESTORED IMMUNE FUNCTION *IN VITRO*

INHIBITED

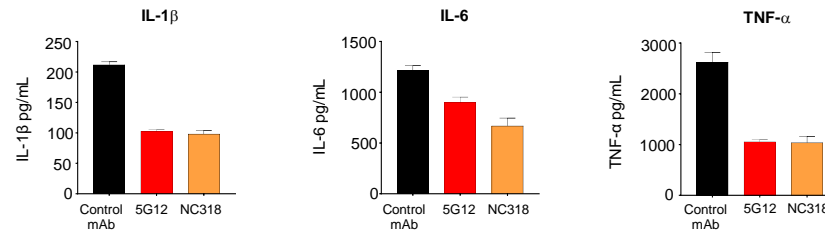
Myeloid Cell
Differentiation
and Survival



Blocked survival of
myeloid cells

DECREASED

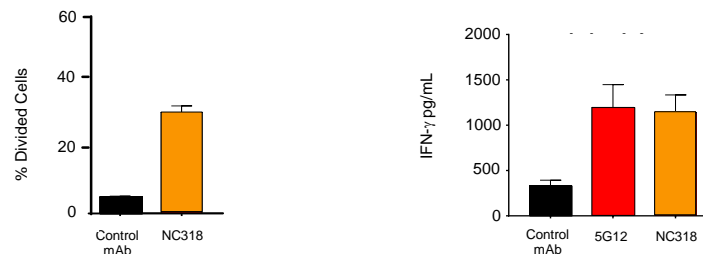
Pro-Inflammatory and
Pro-Tumorigenic
Cytokines



Decreased IL-1 β ,
IL-6 & TNF- α

PROMOTED

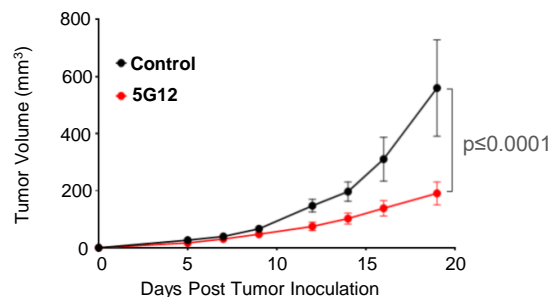
Tumor-Specific
T Cell Function



Increased T cell
proliferation &
IFN- γ production

NC318* HAS SHOWN MONOTHERAPY ACTIVITY IN A NUMBER OF ANIMAL MODELS

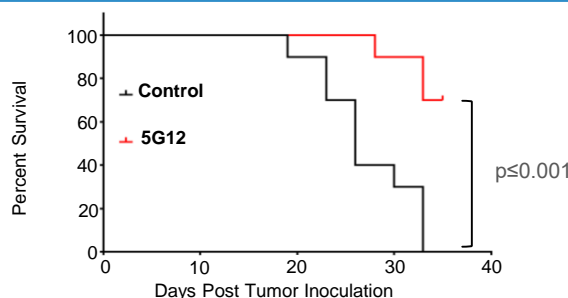
REDUCED
Tumor Growth



CT26 tumor model

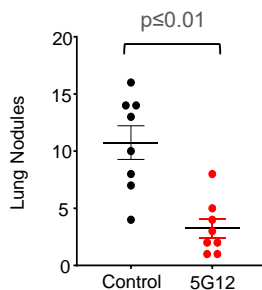
First 2 doses at 20 mg/kg followed by 10 mg/kg, Q4D, 7 doses

INCREASED
Survival



CT26 tumor model

DELAYED
Lung Metastasis



MC38 tumor model

30 mg/kg, Q7D, 4 doses

*Murine surrogate is 5G12

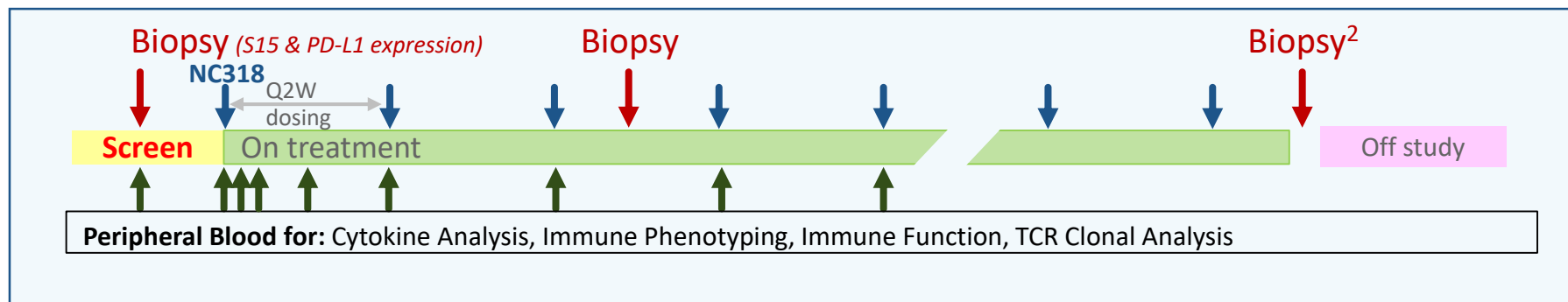
NC318 MONOTHERAPY TRIAL UNDERWAY DESIGNED FOR RAPID PROOF-OF-CONCEPT

PHASE 1

- Opened in 4Q 2018; Complete in 4Q 2019
- Dose-escalation¹
- Safety, tolerability, and biomarker readouts
- Advanced or metastatic solid tumors

PHASE 2

- Open in 1Q 2020; Complete in 4Q 2020
- Efficacy assessment
- Tumor types shown to have elevated S15 expression, including NSCLC, ovarian, and head & neck



(1) Dose escalation evaluates 6 dose cohorts (8 mg – 800 mg or approximately 0.1 - 10 mg/kg) administered every 2 weeks

(2) In Phase 2 portion of trial

NC318 PHASE 1 TRIAL STATUS AS OF MARCH 31, 2019

ENROLLMENT

- 21 subjects dosed
- 4th dose cohort open
- 10 different tumor types
- Recruitment on schedule

SAFETY

- No dose limiting toxicities or drug-related SAEs
- 1 transient elevation of amylase (grade 3) and lipase (grade 4) that was deemed probably related to NC318⁽¹⁾
- Possibly NC318-related AEs limited to transient asymptomatic lab findings or grade 1 events

RESPONSES

- Evaluations every 8 weeks
- 1 confirmed partial response
- 6 stable disease
- 6 progressive disease
- 8 subjects not yet evaluated

• Angeles Clinic

• MSKCC

• Next Oncology

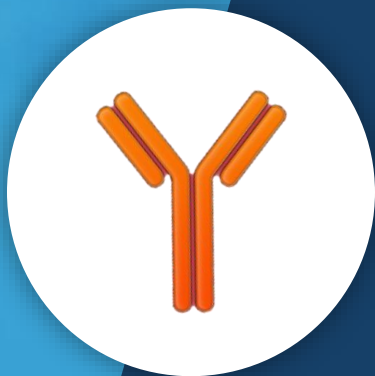
• NYU

• Yale University

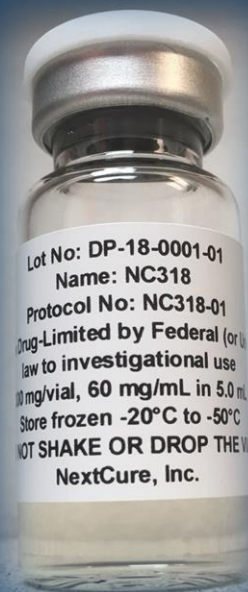
(1) The patient was asymptomatic and both elevations resolved without any interventions within 72 hours

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
- ☒ First-in-Human trial initiated in October 2018
- ☐ Complete Phase 1 and report topline data in Q4 2019
- ☐ Complete Phase 2 in Q4 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

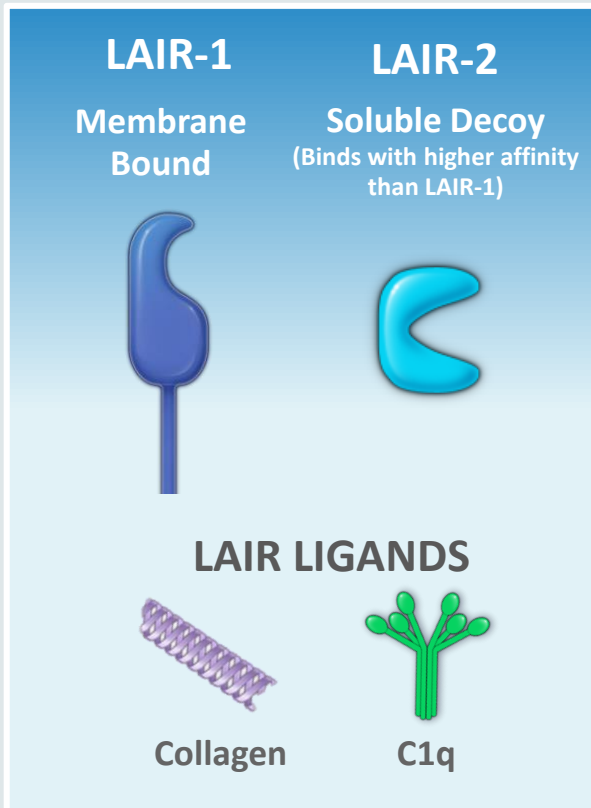
INDICATIONS

Advanced or
metastatic solid
tumors

LAIR-1 & LAIR-2 FUNCTIONAL RELATIONSHIP

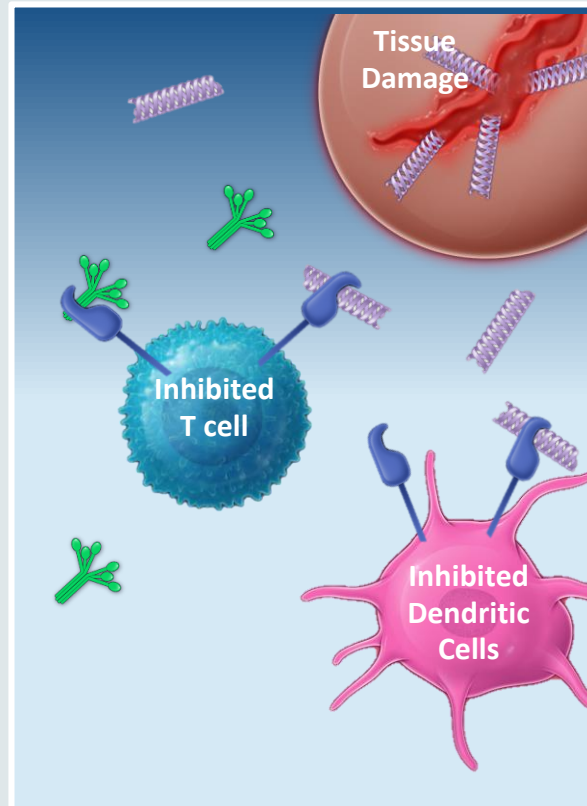
LAIR & LIGANDS

LAIR-1 and LAIR-2 bind collagen and C1q



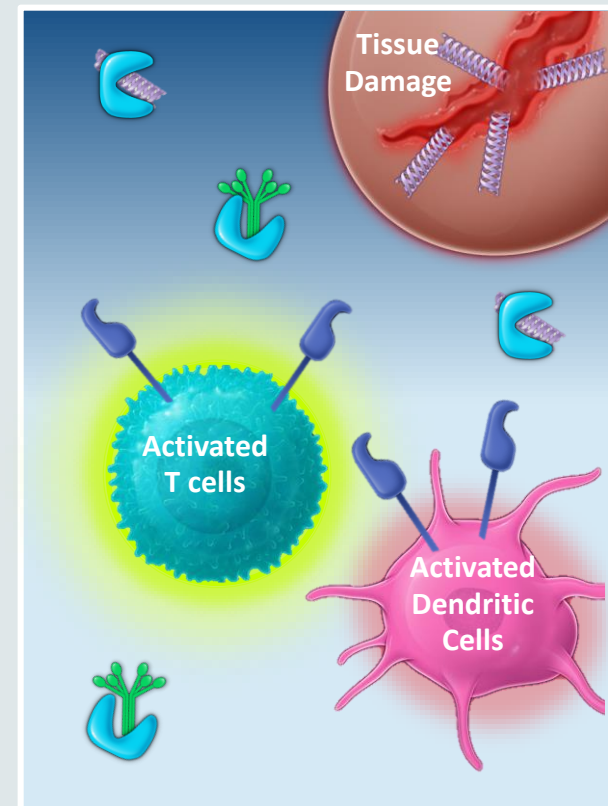
LAIR-1

Ligands are expressed in response to inflammation & inhibit immune function



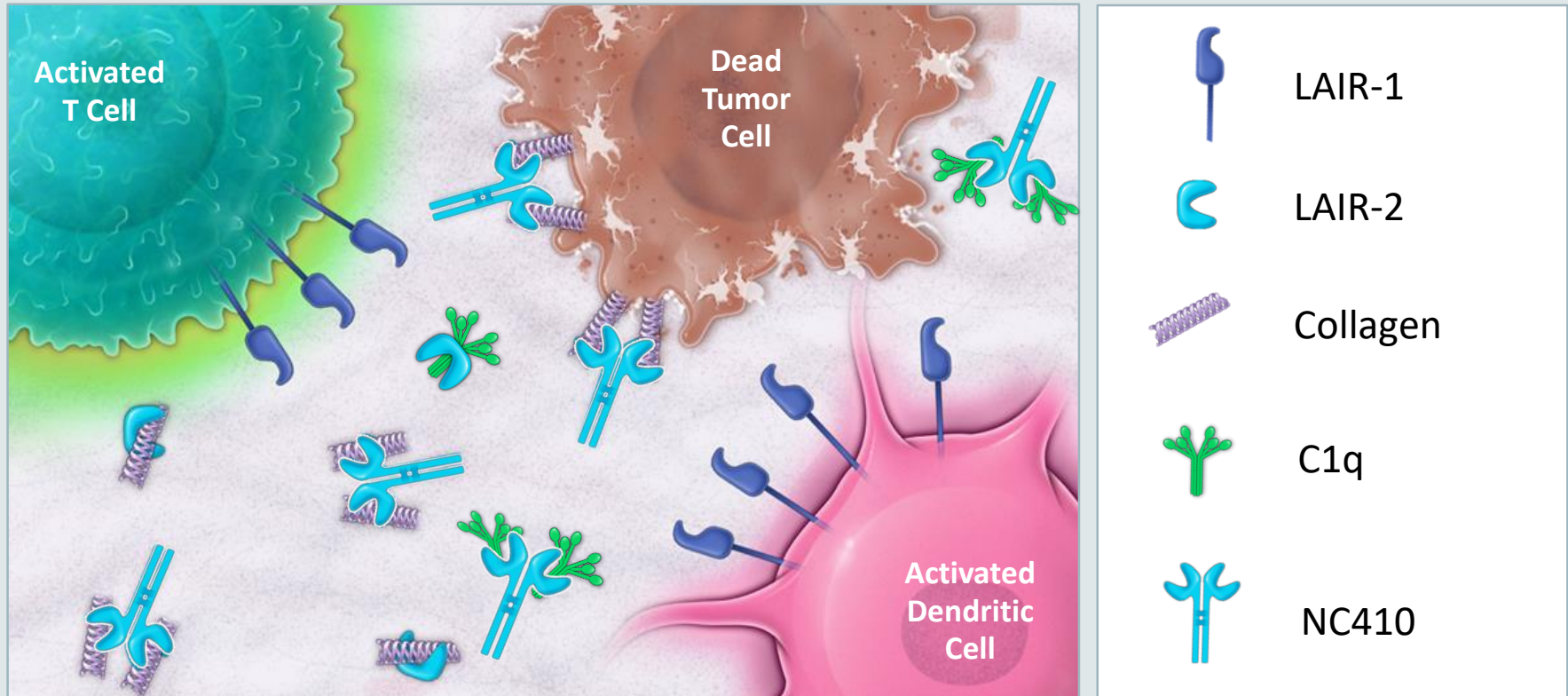
LAIR-2

LAIR-2 modulates LAIR-1 mediated inhibition



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

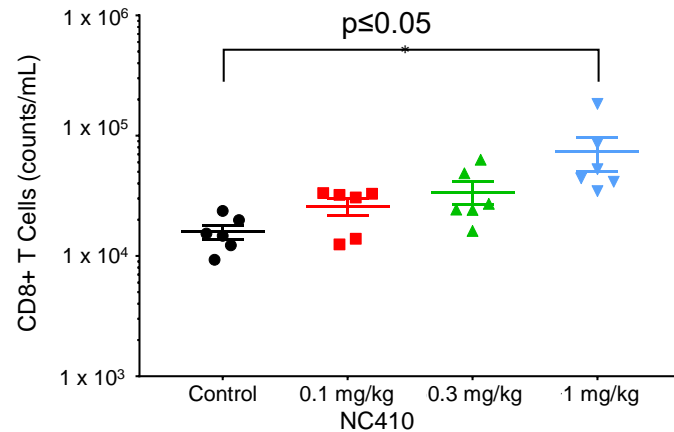
NC410 is a Fusion Protein of LAIR-2 and a Decoy for LAIR-1



NC410 Promotes T Cell Function and Dendritic Cell Activation

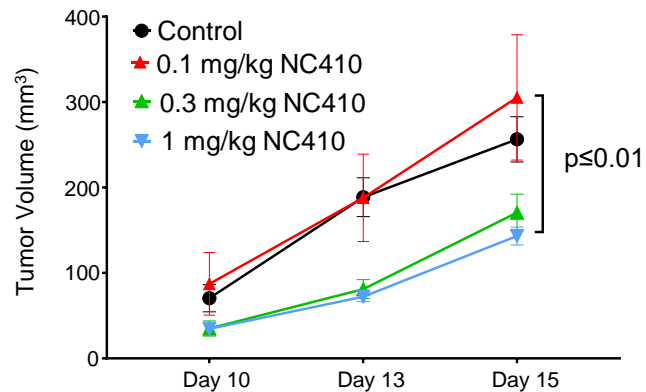
NC410 ENHANCED T CELL EXPANSION AND RELIEVED IMMUNOSUPPRESSION

BLOCKED
Suppression



Human CD8+ T cell
expansion *in vivo*

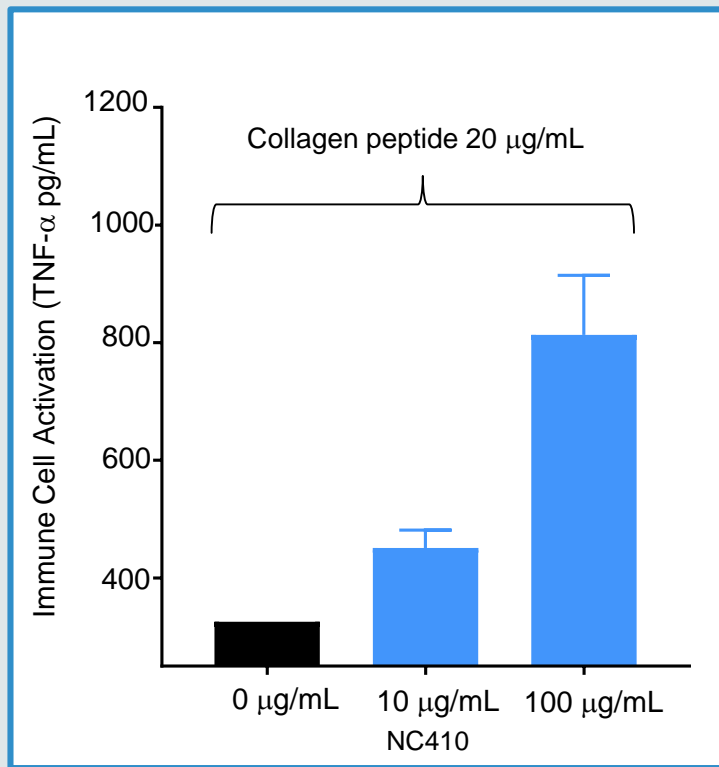
DECREASED
Tumor Volume



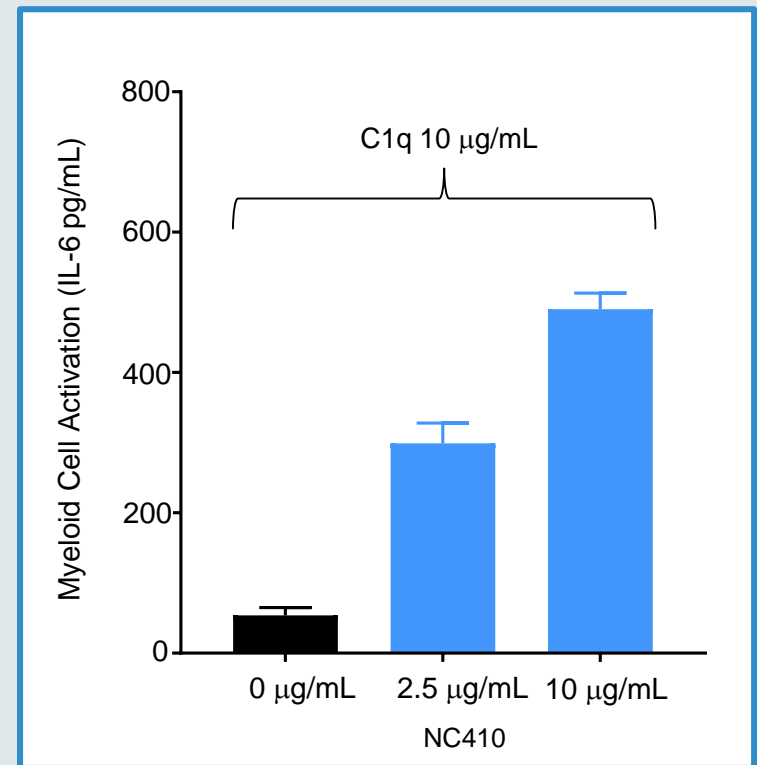
Human PBMCs in mice:
CD8+ T cell activity
decreased tumor volume

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



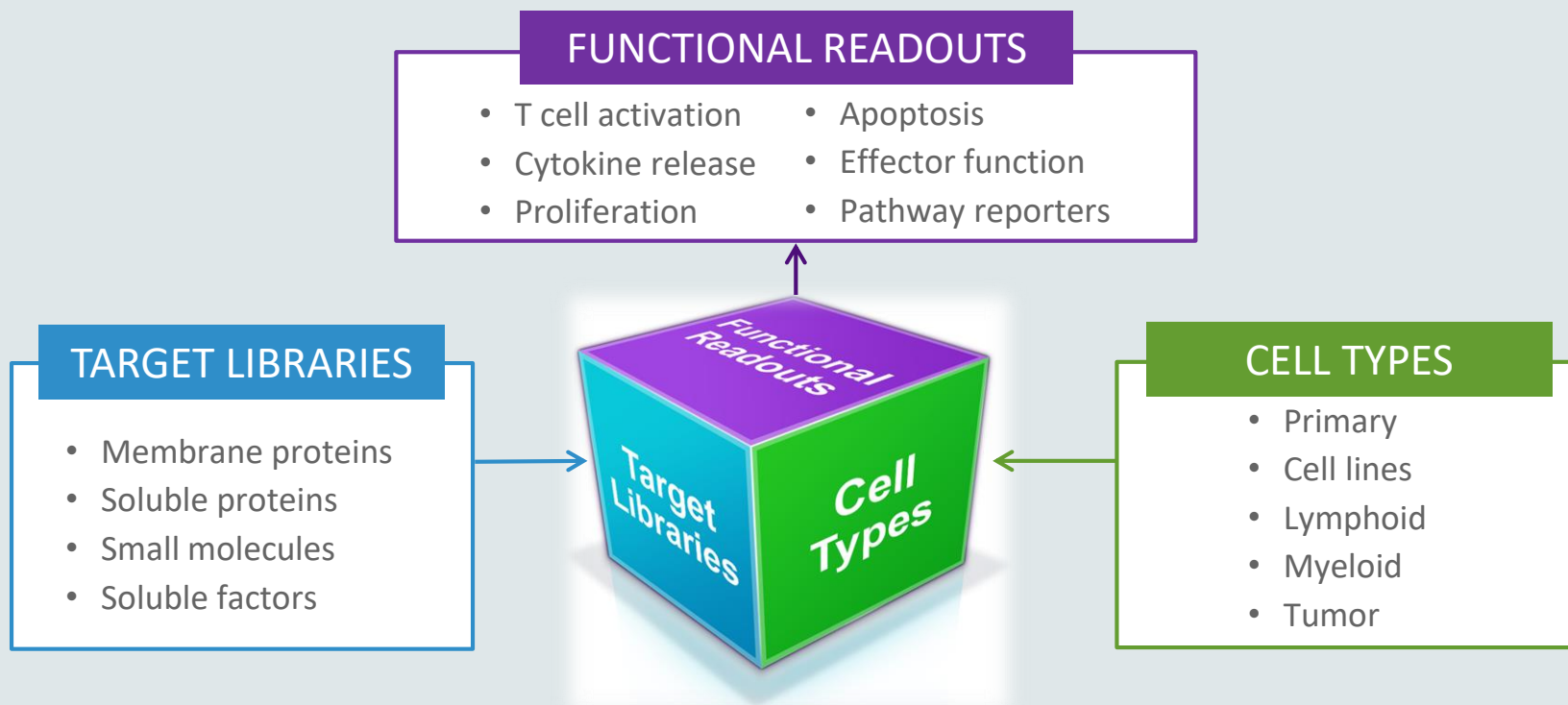
NC410 SUMMARY



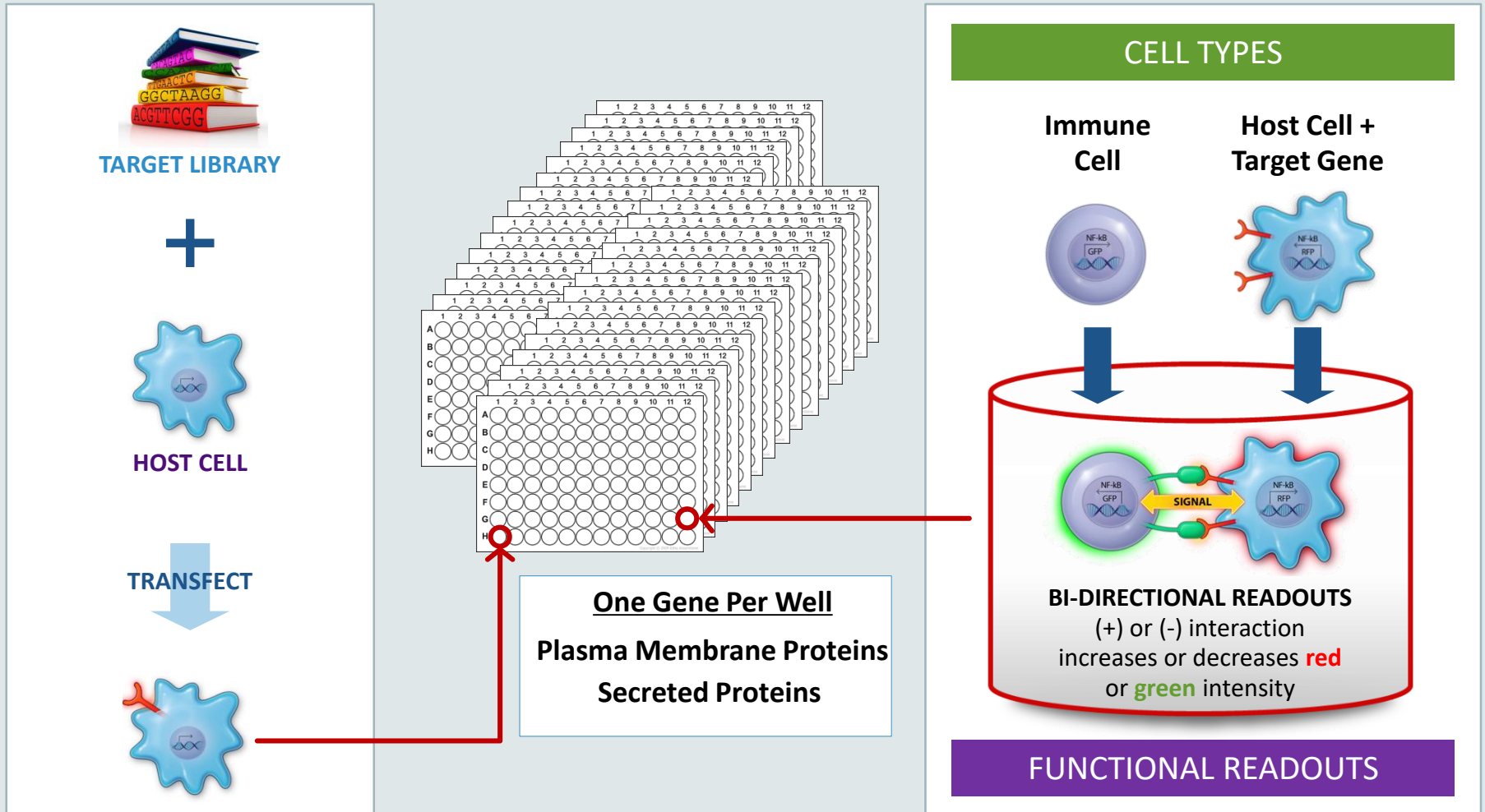
- ☒ Based on normal immune regulatory mechanism
- ☒ Promoted T cell function and dendritic cell activity in preclinical studies
- ☒ Designed to alleviate tumor-mediated immunosuppression
- ☒ IND-enabling tox studies in progress
- ☒ cGMP manufacturing
- ☐ IND filing expected Q1 2020

FINDING SOLUTIONS WITH A POWERFUL DISCOVERY ENGINE

Functional, Integrated, NextCure Discovery in Immuno-Oncology

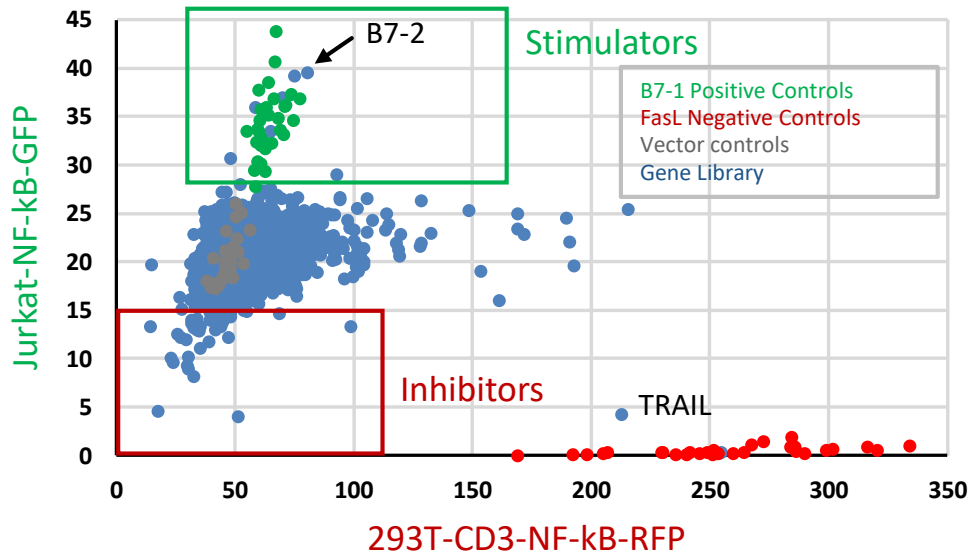


FIND-IO SCREENING METHODOLOGY

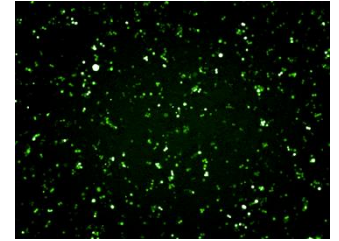


JURKAT “T CELL LINE” SCREENING AND VALIDATING FIND-IO HITS

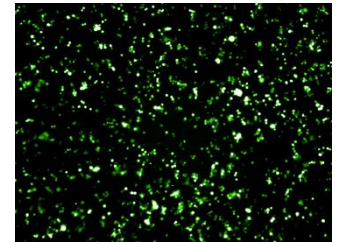
Identified Novel T Cell Stimulators and Inhibitors



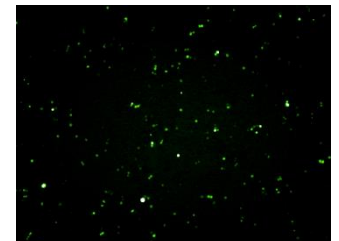
Vector Control



Stimulatory B7-2



Inhibitory TRAIL



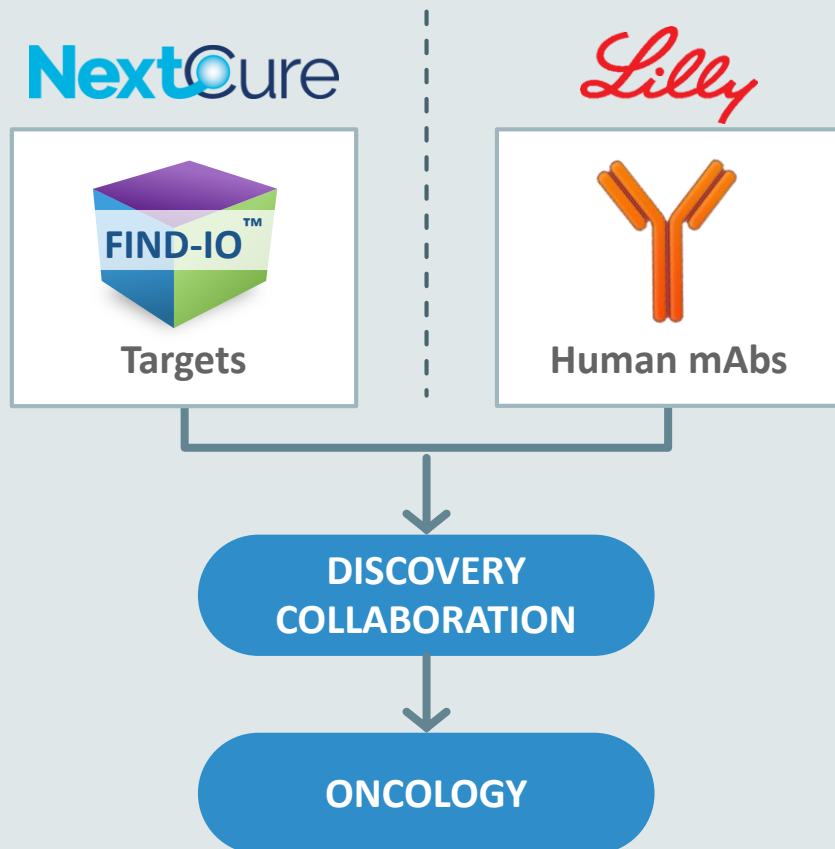
REPRODUCIBILITY

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



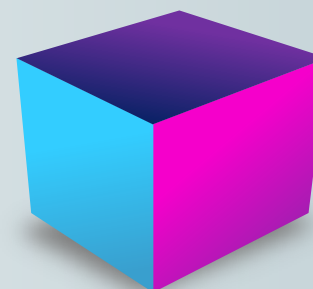
Oncology

TOMORROW



Autoimmunity

FUTURE



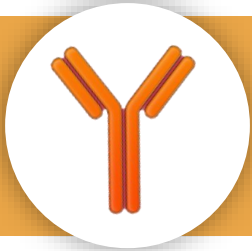
Neuro-Inflammation

Gene Libraries

Immune Cells

Functional Screening

ANTICIPATED NEAR-TERM MILESTONES



NC318

- Complete Phase 1 and report topline data in Q4 2019
- Complete Phase 2 in Q4 2020



NC410

- Complete IND-enabling tox studies
- File IND 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