

Q1-FY 2020 Update Call

Immune Responses, On Cue™

Nasdaq: CUE | Tuesday, May 19, 2020

Forward-Looking Statements

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Agenda

Introduction and Q1 Highlights

Dan Passeri, CEO

CUE-101 Clinical Update

Dr. Ken Pienta, Acting CMO

Pipeline and Platform Progress

Dr. Anish Suri, President and CSO

Financials

Kerri-Ann Millar, VP Finance

Concluding Remarks

Dan Passeri

Q&A

All



Overview of Progress in Q1

Validate

- Data from first four CUE-101 patient cohorts demonstrates favorable characteristics and drug properties, including safety, tolerability and dose proportional PK that aligns with projections
- Growing evidence supporting that the CUE-101 appears to be "clinically active"
- Recent announcement of collaboration with Merck to study CUE-101 with anti-PD-1 combination in H&N front line setting

Expand

- Presentation of data from CUE-102 demonstrating ex vivo expansion of WT-1-specific human T cells, polyfunctionality and their killing of target cells at NYAS Frontiers in Cancer Immunotherapy
- PoC data supporting feasibility of Immuno-STATs to target KRAS-mutant cancers
- Continue to expand Immuno-STAT applications in autoimmune diseases via Merck collaboration focused on two indications; and early assessments and interest in additional indications

Accelerate

- Generation of early proof of concept data supporting the biological activity of molecules generated via the Neo-STAT platform
- Opportunity to rapidly expand the IL-2-based CUE-100 series using the Neo-STAT platform

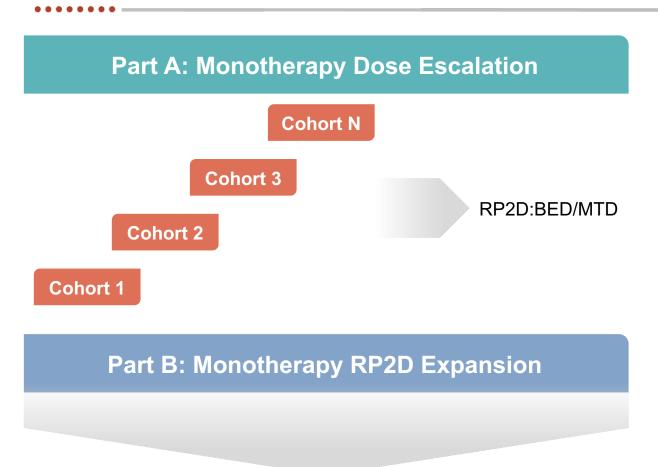
CUE-101: Phase 1 Clinical Development Network

- Emory Winship Cancer Institute: Nabil Saba
- Karmanos Cancer Institute: Elizabeth Heath and Ammar Sukari.
- MD Anderson Cancer Center: Bonnie Glisson
- MGH/Harvard and Dana Farber Cancer Institute: Sara Pai and Lori Wirth
- Moffitt Cancer Center: Christine Chung
- Memorial Sloan Kettering Cancer Center: Lara Dunn
- Stanford Cancer Center: A. Dimitrios Colevas
- University of Arizona Center: Julie Bauman
- University of Michigan Rogel Cancer Center: Frank Worden
- University of Washington Fred Hutch Cancer Center: Cristina Rodriguez
- Vanderbilt-Ingram Cancer Center: Jill Gilbert and Mike Gibson
- Washington University Siteman Cancer Center: Doug Adkins
- Yale Cancer Center: Barbara Burtness

Cue Biopharma has engaged a network of nationally recognized clinical investigators and 13 Phase 1 sites are now open



CUE-101: Ongoing First-In-Human Study



Late Line Accelerated Monotherapy
Approval Opportunity in H&N

Eligibility

- Part A & B: HPV+ H&N Cancer, R/M 2L+
- Design (CUE-101 Q3W)
 - Part A: Dose Escalation (3+3)
 - Part A: Safety Expansion (Up to 9 Patients)
 - Part B: Dose Expansion (10-20 Pts at RP2D)

Objectives

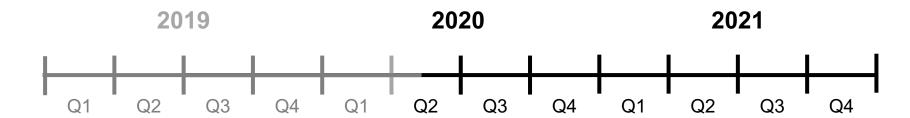
- Primary: Safety and Tolerability
- Secondary: PK/PD, Anti-Tumor Activity
- Biomarkers (Pre/Post CUE-101 Dose)
 - HPV E7-specific CD8+ T cell counts
 - HPV E7-specific CD8+ T cell functionality
 - Immunophenotyping, cytokine release, and TCR sequencing

CUE-101: Phase 1 Dose Cohorts

Cohort	CUE-101 Dose	CUE-101 Dose relative to approved Proleukin dose (0.037 mg/kg)	CUE-101 IL-2 content (nmol/kg)	CUE-101 IL-2 content relative to the approved Proleukin dose (2.4 nmol/kg)
Cohort 1 √	0.06 mg/kg (starting dose)	~ 1.6x	1.1	~ 0.46x
Cohort 2 √	0.18 mg/kg	~ 4.9x	3.4	~ 1.4x
Cohort 3 ✓	0.54 mg/kg	~ 14.6x	10.3	~ 4.3x
Cohort 4 √	1 mg/kg	~ 27.0x	19.1	~ 8.0x
Cohort 5	2 mg/kg	~ 54.0x	38.2	~ 16.0x
Cohort 6	4 mg/kg	~ 108.0x	76.4	~ 32.0x
Cohort 7	8 mg/kg	~ 216.0x	152.9	~ 64.0x



CUE-101: Clinical Development Plan



H&N R/M 2L+

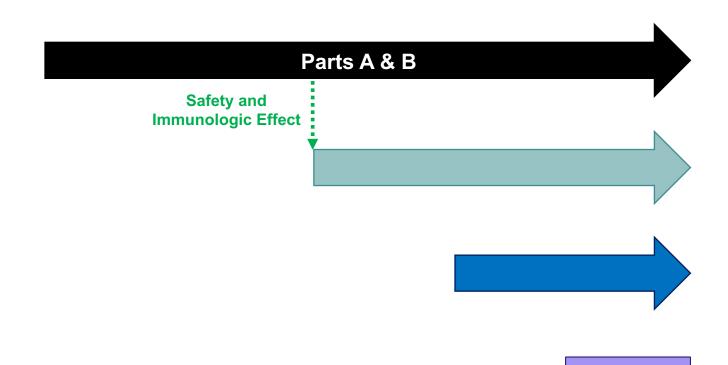
CUE-101 Monotherapy

H&N R/M 1L

CUE-101 + anti-PD-1 Combotherapy

H&N Neoadjuvant

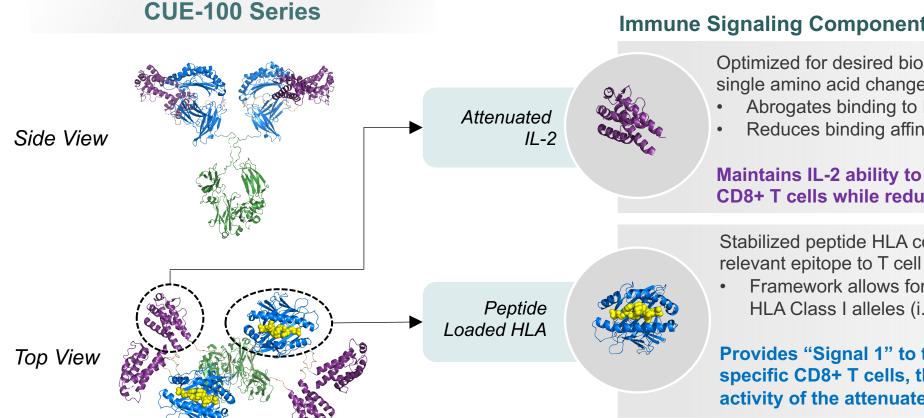
Other HPV-Driven Cancers





TBD

CUE-100 Series: Exploiting IL-2 via Rational Protein Design



Immune Signaling Components

Optimized for desired biological activity through two single amino acid changes

- Abrogates binding to IL-2R alpha
- Reduces binding affinity to IL-2R beta

Maintains IL-2 ability to stimulate antigen-specific CD8+ T cells while reducing Treg expansion

Stabilized peptide HLA complex to present diseaserelevant epitope to T cell receptor

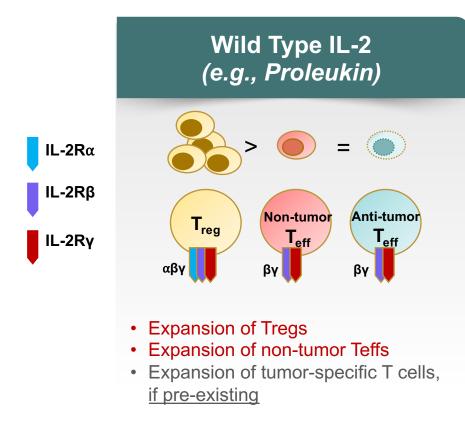
Framework allows for incorporation of an array of HLA Class I alleles (i.e., A02, A11, A24)

Provides "Signal 1" to the targeted antigenspecific CD8+ T cells, thereby enhancing the activity of the attenuated IL-2

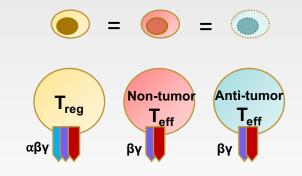
Therapeutic framework is not dependent on barriers of antigen processing & presentation, and is designed to avoid systemic immune activation



CUE-100 Series: Mechanistic Differentiation Over Emerging "Not Alpha" IL-2 Landscape

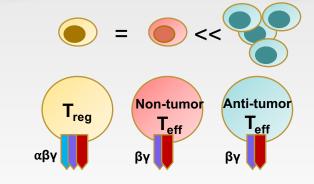


"Not Alpha" IL-2 (e.g., THOR-707)



- Reduced expansion of Tregs
- Expansion of non-tumor Teffs
- Expansion of tumor-specific T cells, <u>if pre-existing</u>

CUE-100 IL-2 Series (e.g., CUE-101)

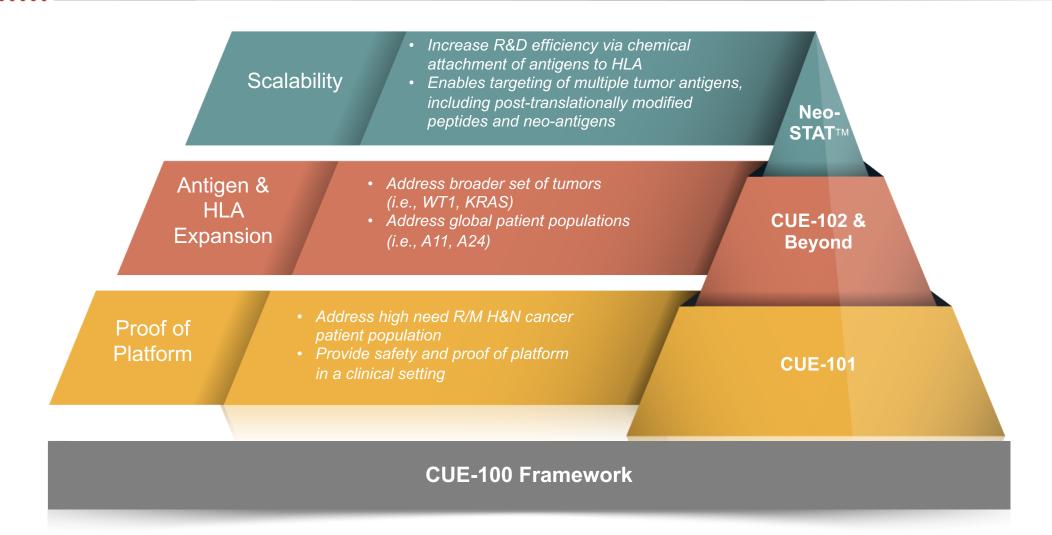


- Reduced expansion of Tregs
- Reduced expansion of non-tumor Teffs
- Expansion of pre-existing tumorspecific T cells and priming of naïve T cells

CUE-100 series is designed for selective induction and expansion of tumor-specific CD8+s without reliance on a pre-existing repertoire



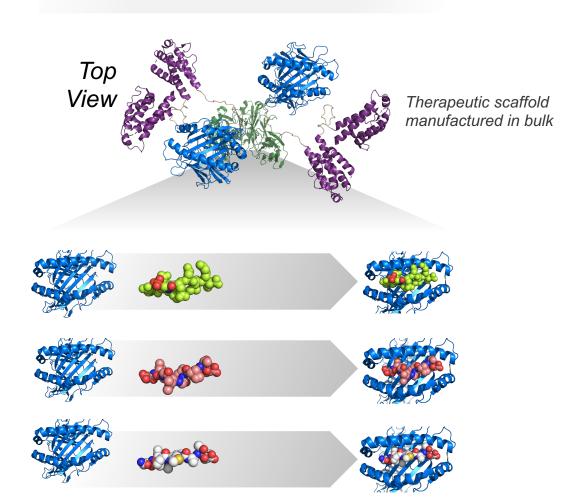
Building Blocks of IO Growth Strategy





Neo-STAT: Next-gen Evolution of the Immuno-STAT Framework

CUE-100 Neo-STAT

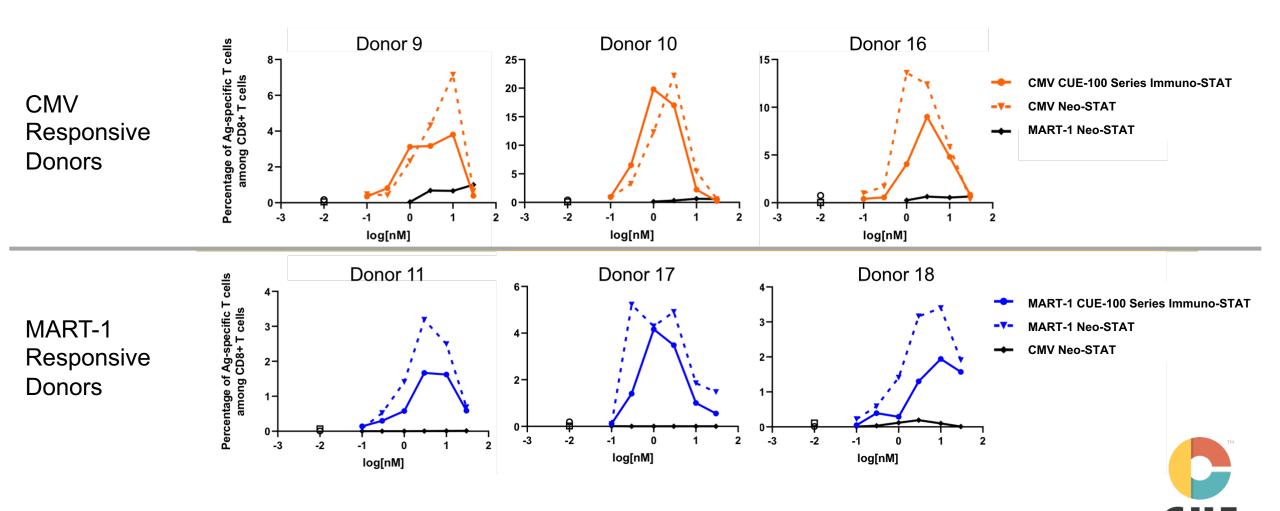


Therapeutic scaffold receptive for chemical conjugation of peptides, that potentially:

- Increases R&D efficiency and reduces cost of the generation of clinical grade material on the CUE-100 framework
- Enables targeting of multiple tumor antigens including post-translationally modified peptides and neo-antigens for personalized therapy



Primary Human T Cell Expansion: Neo-STAT = Immuno-STAT



Pipeline

PARTNER LATE CLINICAL **TARGET SELECTION** PRE-CLINICAL PHASE 1 CUE-101 (HPV E7 / A02) CUE-102 (WT1 / A02) LG Chem **CUE-100** IL-2 Asia Rights to **CUE-102 (WT1 / A24)** CUE-101, CUE-102, and CUE-103 **CUE-103 (Undisclosed)** KRAS / A11 **CUE-200** CD80 & 4-1BBL **CUE-201 (Undisclosed)** CUE-301 (Proins / DR4) MERCK **CUE-300** PD-L1 & Collaboration **CUE-302 (Undisclosed)** for Autoimmune Disease Undisclosed



Balance Sheet and Statement of Operations Summary

(in thousands)	As of March 31, 2020	As of December 31, 2019
Cash and Cash Equivalents	\$23,432	\$44,290
Marketable Securities	\$25,298	\$15,120
Total Current Assets	\$51,190	\$61,025
Working Capital	\$39,100	\$49,370

(in thousands)	Q1 2020	Q1 2019
Collaboration revenue	\$900	\$370
General & Administrative	\$3,989	\$3,444
Research & Development	\$9,906	\$8,353
Total Operating Expenses	\$13,895	\$11,797

Our current cash position is estimated to take us into the fourth quarter of 2021



Key 2020 Milestones

- PK/PD results from the Phase 1 CUE-101 clinical trial in 2Q20
- 2 Clinical responses in Phase 1 CUE-101 via RECIST criteria in 2H20
- 3 Initiate CUE-101 combination trial with Keytruda in 1L SCCHN in 2H20
- Initiate and extend IND-enabling activities for CUE-102 in 2H20
- 5 Select target for CUE-103 in 2Q20
- 6 Demonstrate Neo-STAT manufacturability and efficiencies in 2H20
- 7 Identify potential clinical candidates in autoimmune collaboration with Merck in 2H20

Key objectives met in 2019 and early 2020 have set the stage for data flow from multiple programs in the remainder of 2020

