

# **Bolt Biotherapeutics**

Nasdaq: BOLT

Boltbody™ ISAC Platform

October 2023

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# Bolt Bio: Dedicated to Generating Breakthroughs for Patients

# BDC-1001: Monotherapy Activity, Efficient Plan

#### Promising Phase 1 Results

- Monotherapy ORR<sup>1</sup> of 29% at RP2D of 20 mg/kg q2w
- Well tolerated

#### Phase 2 Program

- Option-based development
- 4 tumor types
- Upcoming data readouts

# Focused Pipeline, Proven Platform Technology

#### Pipeline

- BDC-1001 in Phase 2
- BDC-3042 in Phase 1

Collaborations validate Boltbody™ ISAC platform







## Well-Capitalized, Significant Upside Potential

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- 6 covering research analysts
- Consensus price target: \$5.00<sup>2</sup>

\$157M cash & equivalents<sup>3</sup>

Simple Corporate Structure

- 37.95 million shares of common stock oustanding<sup>4</sup>
- No debt
- No warrants



<sup>&</sup>lt;sup>1</sup>Objective Response Rate in evaluable patients with HER2+ tumors

<sup>&</sup>lt;sup>2</sup> \$5.00 is consensus price target of 6 covering analysts as of 8/31/23

<sup>&</sup>lt;sup>3</sup>\$157.1 million cash & cash equivalents

<sup>437,950,986</sup> shares outstanding as of 6/30/23

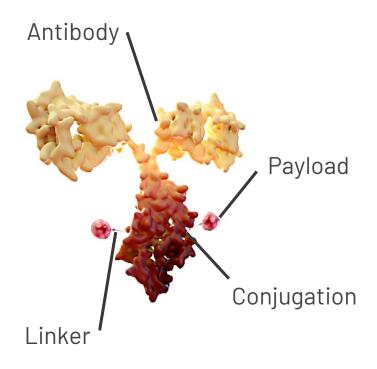
RP2D = Recommended Phase 2 Dose q2w = every other week dosing schedule





# Boltbody™ ISAC Platform

Best-in-class ISAC Platform with Emerging Clinical Validation



## Proven Ability to Achieve:

- ✓ Target-dependent activity
- ✓ Low systemic cytokine production
- ✓ Low immunogenicity

## **Antibody**

- Targets tumor antigen
- Active Fc essential for phagocytosis
- Impacts ISAC PK, activity & safety

## **Payload**

- 100s of payloads evaluated
- TLR7, TLR8, dual TLR7/8 & STING payloads available

#### Linker

- Non-cleavable for better safety
- Optimize physicochemical properties

## Conjugation

- Suite of modern & classical methods available
- Conjugation site impacts activity & safety

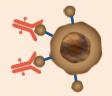


## Innate Immune Response

Myeloid Cells Kill Tumor Cells via ADCP

#### TUMOR MICROENVIRONMENT

Tumor
1 Antigen
Recognition



#### **Antigen Expression**

• High, medium, & low

FcRDependent
Phagocytosis
(ADCP)



#### **Myeloid Antigen-presenting Cells**

- Monocytes
- Macrophages
- pDCs and cDCs

TLRMediated
Activation



#### **Activated Myeloid Cells**

- Chemokine & cytokine secretion
- Enhanced antigen presentation

## Adaptive Immune Response

Engages T Cell-driven Tumor Killing

TUMOR-DRAINING LYMPH NODES



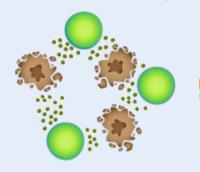
T cell Priming & Expansion



#### TUMOR MICROENVIRONMENT



T Cell Killing of Tumor Cells





#### **Result: An Immune "Hot" Tumor**

- Chemokines attract immune effector cells
- Cytokines lower immune activation threshold
- Increases myeloid APC phagocytosis
- Activated T cells migrate to tumor







Tumor Cell













MHC-Tumor Peptide Complex

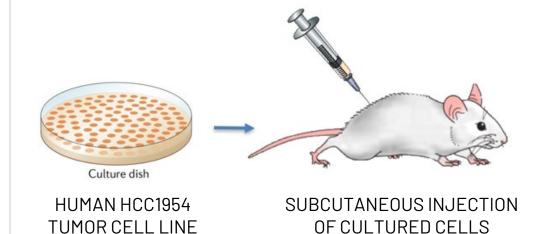




## ISACs Deliver Powerful Synergies

Covalent Attachment of TLR7/8 Agonist Dramatically Improves Anti-tumor Efficacy

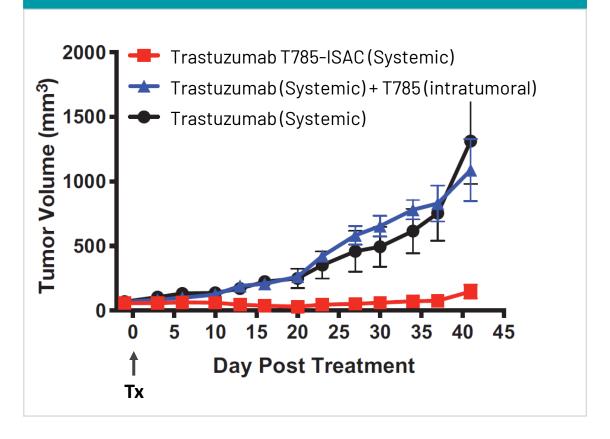
### HCC1954 Tumor Xenograft Model



- SCID/Beige lack T, B and functional NK cells, but retain a myeloid compartment
- Enables assessment of myeloid-mediated anti-tumor activity

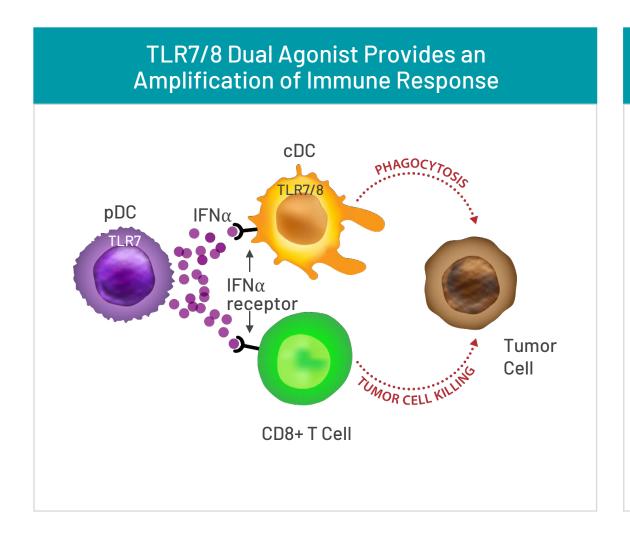
Adapted from Creative Biolabs

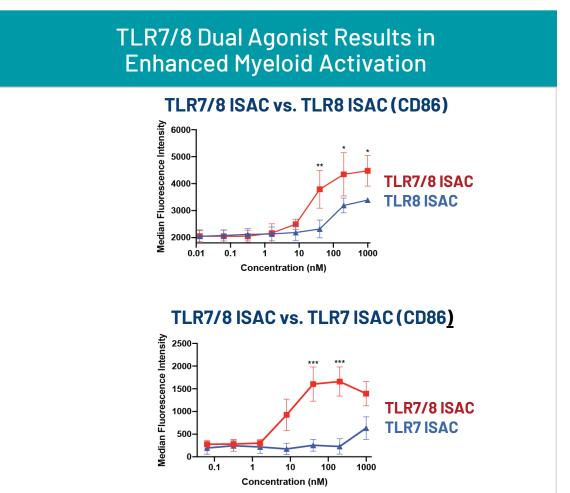
### Single ISAC Dose Mediates Tumor Regression





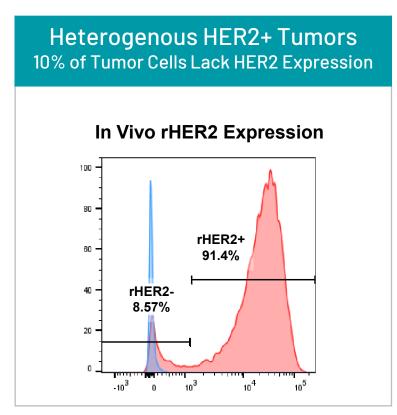
# Preclinical Boltbody™ ISAC Data Shows Advantage to Combining TLR7 <u>and</u> TLR8

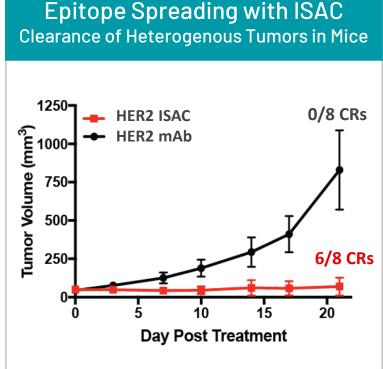


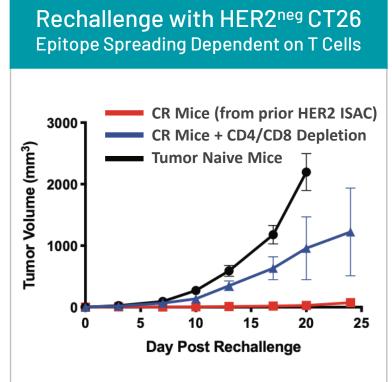




# Boltbody™ ISAC Induces Immune Memory that Extends to Tumors Lacking HER2 Immunity Extends to Tumor Neoantigens through Epitope Spreading







CT26-rHER2+ or CT26-rHER2<sup>neg</sup> Syngeneic Colorectal Cancer Models



# Strategic Collaborations Provide External Validation for Boltbody™ Platform

# Innovent

Fully integrated biopharma with large antibody library and strong presence in Greater China

- Innovent funds up to 3 Boltbody™ ISACs through early clinical development
- Bolt has option to co-develop & commercialize 2 candidates in certain regions
  - Bolt received \$5M upfront; eligible for future milestones & royalties



Innovative leader in antibody & bispecific development for oncology

- Genmab funds 3 bispecific Boltbody ISACs through early clinical development
- Bolt has option to co-develop & commercialize 1 candidate in certain regions
  - Bolt received \$25M upfront; eligible for up to \$285M in milestones plus tiered royalties per program exclusively developed & commercialized by Genmab

# **TORAY**

Global leader in innovative technologies, conducting research in cancer immunotherapeutics

- Toray funds Boltbody ISAC for specific & novel target through end of Phase 1
- Global co-development/co-commercialization





