

TELOMERE TARGETING IMMUNOTHERAPIES
FOR CANCER

**NYSE AMERICAN: MAIA** 

January 2024

#### **FORWARD-LOOKING STATEMENTS**



All statements in this presentation, other than those relating to historical facts, are "forward-looking statements." These forward-looking statements may include, but are not limited to, statements relating to our objectives, plans, and strategies; statements that contain projections of results of operations or of financial condition; statements relating to the industry and government policies and regulations relating to our industry; and all statements (other than statements of historical facts) that address activities, events, or developments that we intend, expect, project, believe, or anticipate will or may occur in the future. Forward-looking statements are not guarantees of future performance and are subject to risks and uncertainties. We have based these forward-looking statements on assumptions and assessments made by our management in light of their experience and their perception of historical trends, current conditions, expected future developments, and other factors they believe to be appropriate. Important factors that could cause actual results, developments, and business decisions to differ materially from those anticipated in these forward-looking statements include, among other things: the overall global economic environment; general market, political, and economic conditions in the countries in which we operate: projected capital expenditures and liquidity; changes in our strategy; government regulations and approvals; the application of certain service license; and litigation and regulatory proceedings. The Company has filed a registration statement on Form S-1, as may be amended (Registration No.: 333-269606). Before you invest, you should carefully read the registration statement, including the factors described in the "RISK FACTORS" section of the Registration Statement and other documents that we have filed, and will subsequently file, with the Securities and Exchange Commission to better understand the risks and uncertainties inherent in our business and industry and for more complete information about us and the offering. You may get these documents for free by visiting EDGAR on the Commission's website at www.sec.gov. We caution you that forward-looking statements are not guarantees of future performance and that our actual results of operations, financial condition and liquidity, and the development of the industry in which we operate may differ materially from the forward-looking statements contained in this presentation as a result of, among other factors, the factors referenced in the "Risk Factors" section of the Registration Statement. In addition, even if our results of operations, financial condition and liquidity, and the development of the industry in which we operate are consistent with the forward-looking statements contained in this presentation, they may not be predictive of results or developments in future periods. This presentation shall not constitute an offer to sell or the solicitation of an offer to buy any of our securities nor shall there be any sale of securities in any jurisdiction in which such offer, solicitation or sale would be unlawful prior to registration or qualification under the securities laws of any such jurisdiction. Any offering of securities can only be made in compliance with applicable securities laws. You should read carefully the factors described in the "Risk Factors" section of the Registration Statement to better understand the risks and uncertainties inherent in our business and underlying any forward-looking statements. These statements are only current predictions and are subject to known and unknown risks, uncertainties, and other factors that may cause our or our industry's actual results, levels of activity, performance, or achievements to be materially different from those anticipated by the forward-looking statements. You should not rely upon forward-looking statements as predictions of future events. Although we believe that the expectations reflected in the forward-looking statements are reasonable, we cannot guarantee future results, levels of activity, performance, or achievements. Except as required by law, we are under no duty to update or revise any of the forward-looking statements, whether as a result of new information, future events or otherwise, after the date of this prospectus. These forward-looking statements speak only as of the date of this presentation, and we assume no obligation to update or revise these forward-looking statements for any reason.

#### **INVESTMENT OVERVIEW**



- Telomere-Targeting Agents for Cancer Therapy:
  - THIO in clinic
  - Advancing pipeline
- Efficacy
- Safety
- U.S. FDA: 3 Orphan Drug Designations: Liver (HCC), Lung (SCLC), and Brain (GBM)
- REGN: Clinical Supply Agreement
- Phase 2 Go-to-Market Accelerated Approval THIO-101 in NSCLC underway
  - AUS and EU
  - US IND cleared
  - Disease Control Rates (DCR) ESMO 2023
  - Dose Selection
  - Upcoming Milestones: ORR, PFS, DoR, OS
- Phase 2 Go-to-Market Accelerated Approval THIO-102 planned to start
- Phase 2/3 Confirmatory/Expansion THIO-103 basket trial planned to start



#### **ROBUST PIPELINE**



THIO Telomere targeting agent	PHASE 1 PHASE 2 PH	RIGHTS
THIO-101 NSCLC-2 (THIO → Libtayo®)	Ph 2 Enrolling since July 2022	Worldwide rights owned by MAIA
THIO-102 CRC, HCC, SCLC, ST (THIO → CPI)	Ph 2 Planning	Worldwide rights owned by MAIA
<b>THIO-103</b> NSCLC-1, SCLC-1 (THIO $\rightarrow$ CPI)	Ph 2/3 Planning	Worldwide rights owned by MAIA
2 <sup>nd</sup> Generation Telomere targeting age	ents	
MAIA-2021-020 Multiple Ind. IND Enabling		Developed in-house
MAIA-2022-012 Multiple Ind. IND Enabling		fully-owned by MAIA
MAIA-2021-029 Multiple Indications		

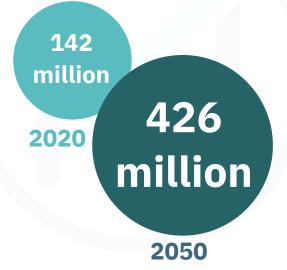


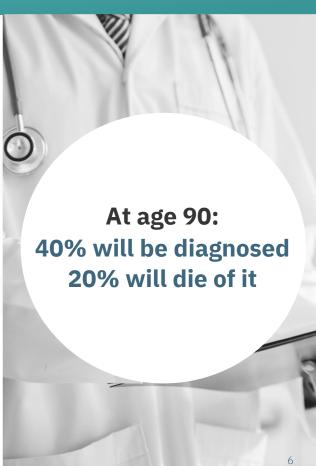
## MISSION AND APPROACH





Population aged >80 expected to triple by 2050







# THIO is the only direct telomere targeting agent currently in clinical development

#### **THIO - MECHANISMS OF ACTION**



#### **THIO**



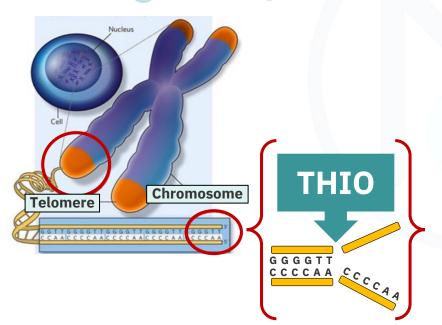


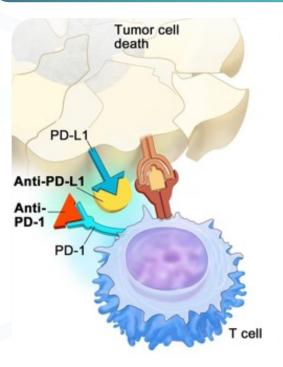
#### Followed by

#### Immune Checkpoint Inhibitor (CPI)



- Telomere targeting
- 2 Immunogenic effect

















#### REGENERON AGREEMENT



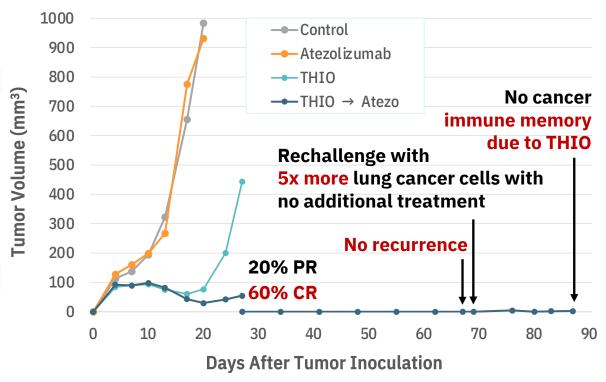


MAIA Biotechnology, Inc. Announces Clinical Supply Agreement with Regeneron for Phase 1/2 Clinical Trial Evaluating THIO in Sequential Administration with Libtayo<sup>®</sup> (cemiplimab) in Advanced Non-Small Cell Lung Cancer

#### THIO-101 - RATIONALE



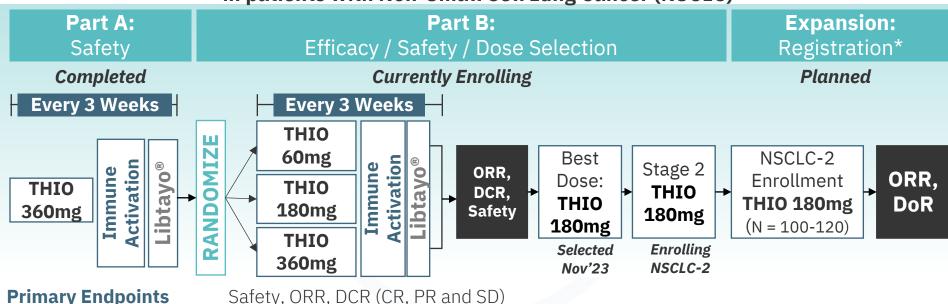
- THIO followed by CPI results in 60% complete response
- No recurrence after long-term follow-up
- Anticancer immune memory has been induced: no cancer after rechallenge with 5x more lung cancer (LLC) cells with no additional therapy



#### THIO-101 TRIAL: 60 PATIENTS DOSED TO-DATE



A Multicenter, Open-Label, Dose-Finding Phase 2 Trial Evaluating the Safety and Efficacy of THIO Administered in Sequence with LIBTAYO® (cemiplimab) in patients with Non-Small Cell Lung Cancer (NSCLC)



**Secondary Endpoints** 

DoR; PFS; OS

**Exploratory Endpoints** 

PK and PD (activity of THIO in circulating tumor cells measured by specific biomarkers)

#### **EXPECTED EFFICACY IN NSCLC-2**



Indication	CPI-Treated NSCLC Second-Line					
Treatment	Cyramza® (ran docet (REVEL ph I	axel	Ар	for Accelerated proval nts for NSCLC)	Estimated for TH (based on DCR fro 101)	
Efficacy Primary End Points	ORR: DOR:	23% N/A	ORR: DOR:	35% 6 months	ORR: 35- DOR: 6-8 mg	40% onths
Disease Control Rate	649	<b>%</b>			90%	

#### **FAVORABLE SAFETY PROFILE**



- Overall, THIO sequenced with cemiplimab has been generally well-tolerated to date in a heavily pretreated population, with the majority of Adverse Events (AEs) being Grade 1-2.
- Chemo (SOC): 70-80% incidence of Grade 3-5 side effect
- Best dose: to date, 60% of patients in the 180mg/dose reported no related AEs
- 3 out of 20 patients (15%) reported Grade 3 LFT increases, without clinical presentation; reverted to normal after treatment discontinuation
- No patients reported Grade 4 AEs in the 180mg/dose
- Of the 60 patients who received at least one dose of THIO, 5 subjects were recently enrolled with less than 1 full cycle of treatment and no AEs were entered in the database. They were removed from the safety analysis to not underestimate the AE incidence.

Treatment Related AEs	Cyramza® + docetaxel (n=627) REVEL Study*	THIO 180mg (n=20)
Grade 5	5.4%	
Grade 4	30%	
Grade 3	50%  Neutropenia: 49% Febrile Neutropenia: 16% Fatigue: 14% Leukopenia: 14%	15%  LFT Elevations (no clinical presentation):  15%
Any Grade	613 (97.8%)	8 (40%)

<sup>\*</sup> https://www.cyramza.com/hcp/nsclc-treatment/revel-trial-safety

#### DISEASE CONTROL RATE AS PRIMARY ENDPOINT IN **NSCLC-2+ TRIALS**



PMID: 34164275

- Disease control rate (DCR) has been demonstrated to be a better predictor of overall survival (OS) advantage compared to overall response rates (ORR) in NSCLC post first line of therapy
- Meta-analysis<sup>1</sup> aimed to evaluate the association of trial-level ORR and DCR with OS in randomized phase II NSCLC trials that evaluate second- or later-line chemotherapy (e.g. ramucirumab + docetaxel, bevacizumab + docetaxel, paclitaxel/carboplatin vs. vinorelbine/carboplatin)
- 74 trials sourced from PubMed, Cochrane database, Embase, and Web of Science as of May 15, 2020
- The correlation between trial-level odds ratio of objective response rate and hazard ratio of overall survival was weak (r=-0.29, 95% CI: -0.49 to -0.05, P=0.014), while odds ratio of disease control survival had moderate rank correlations with hazard ratio of overall survival (r=-0.53, 95% CI: -0.68 to -0.32, P<0.001)



Disease control and objective responsive rates in randomized phase II trials evaluating nonfirst-line chemotherapy for non-small cell lung cancer: a systematic review of 74 trials

Hiromi Matsumoto, <sup>1</sup> Nobuyuki Horita, <sup>1</sup> Kentaro Ito, <sup>2</sup> Risa Ebina-Shibuya, <sup>3</sup> Yu Hara, <sup>1</sup> Nobuaki Kobayashi, <sup>1</sup> and Takeshi Kaneko 1

"Since data on ORR and DCR are available simultaneously, we recommend using DCR instead of ORR as the primary endpoint in a randomized phase II trial evaluating secondor later-line chemotherapy for NSCLC."

#### DISEASE CONTROL RATES (DCR) BY LINE OF THERAPY



Treatment Line	Standard of Care DCR
NSCLC-1	71% KEYTRUDA (pembrolizumab; KEYNOTE 024)
NSCLC-2	<b>64%</b> CYRAMZA (ramucirumab) + docetaxel (REVEL) <b>53%</b> docetaxel monotherapy (REVEL)
NSCLC-3	25-35% chemotherapy (RWD)

THIO + Libtayo® (cemiplimab)*	
TBD	
90%	
83%	

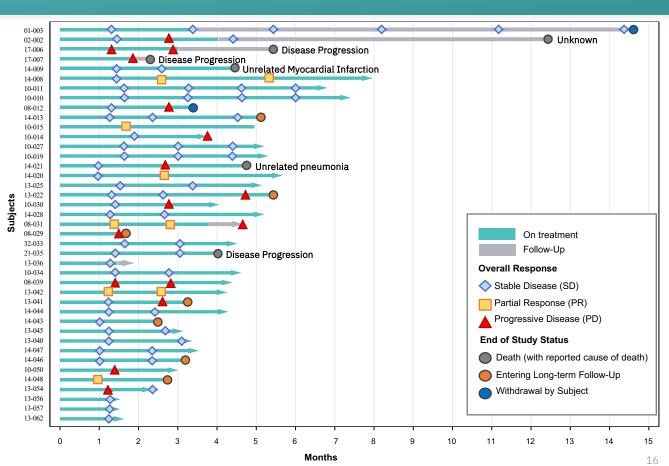
<sup>\*</sup>DCR as reported by PI through 13Nov2023 in THIO-101;

Patients who had at least one post-baseline scan.

#### SWIMMER PLOT: ALL SUBJECTS BY START DATE



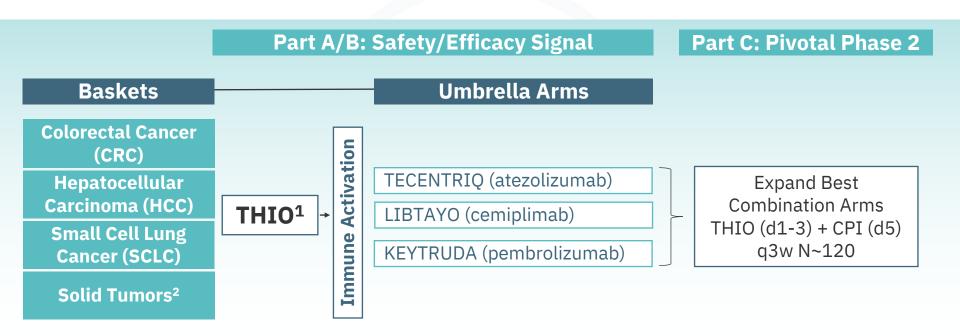
- 42 subjects completed at least 1 post baseline assessment completed at time of cut-off (13Nov2023)
- Partial Responses (PRs) per RECIST 1.1 were reported for 6 subjects, with 3 PRs confirmed by a 2nd scan (per Investigators' assessment). (\*Post cut-off update: 1 additional PR was recorded (7 total) and 14-020's PR was confirmed at next scan)
- The first 2 subjects dosed on trial (both receiving 3rd line of treatment) reported long term survival of 14.6 and 12.5 months respectively, with no new anti-cancer treatment initiated.



#### THIO-102 TRIAL (PLANNED)



A Multicenter, Open-label, Phase 2 Trial Evaluating the Safety and Efficacy of THIO Administered in Sequence with Anti-PD-1 or Anti-PD-L1



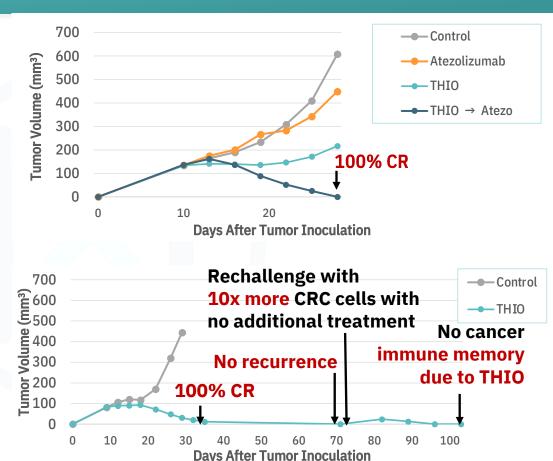
<sup>&</sup>lt;sup>1</sup> Dose to be selected from THIO-101 study results

<sup>&</sup>lt;sup>2</sup> E.g. Breast, Prostate, Gastric, Pancreatic, Ovarian, etc

#### THIO-102 - COLORECTAL RATIONALE



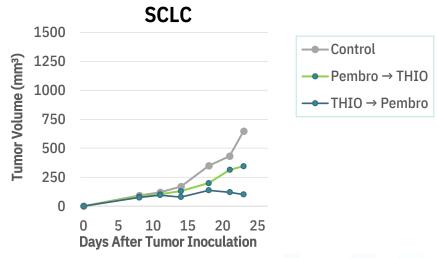
- THIO followed by CPI results in 100% complete response
- No recurrence after long-term follow-up
- Anticancer immune memory has been induced: no cancer after rechallenge with 10x more CRC cells with no additional therapy

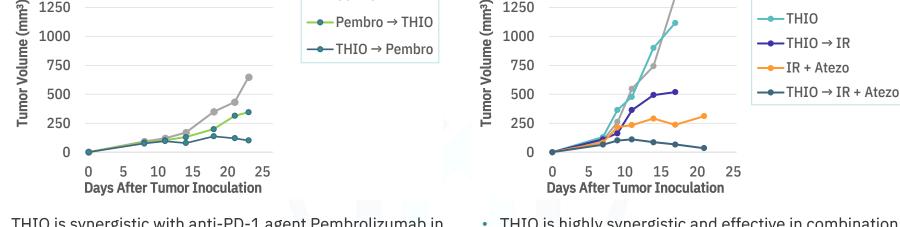


#### **SCLC & HCC – ORPHAN DRUG DESIGNATION**



----Control





1500

1250

- THIO is synergistic with anti-PD-1 agent Pembrolizumab in Small Cell Lung Carcinoma (H2081) in vivo in humanized murine cancer model.
- Treatment with THIO followed by Pembrolizumab results in highly potent anticancer effect, as compared to Pembrolizumab alone.
- THIO converts immunologically "cold non-responsive" SCLC tumor into "hot and responsive" to Pembrolizumab.

 THIO is highly synergistic and effective in combination with anti-PD-L1 agent Atezolizumab and Ionizing Radiation (IR 10Gy) in HCC53N Hepatocellular Carcinoma.

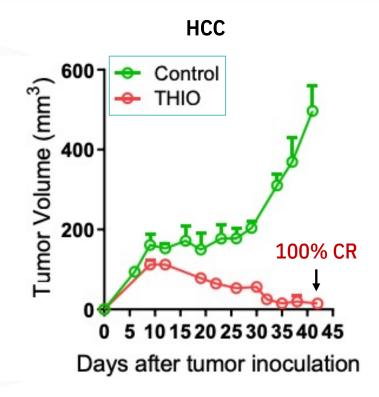
HCC

Treatment with THIO in combination with IR and Atezolizumab results in a complete regression of aggressive HCC tumors. The combination of IR and Atezolizumab is just partially efficacious.

#### **EXCELLENT EFFICACY IN HCC MODELS**



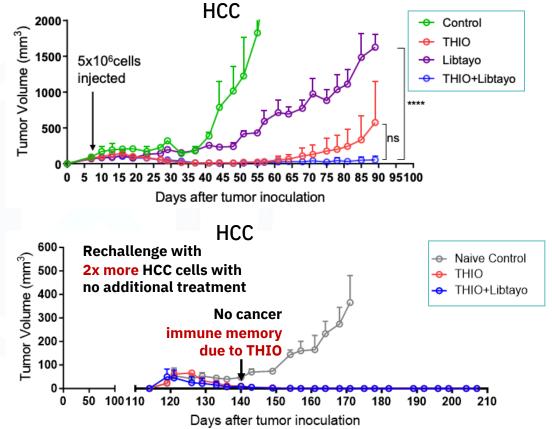
 THIO achieved complete and durable responses in Hepatocellular Carcinoma (HCC), the dominant histology in primary liver cancer (90%), in *in vivo* models



#### **HCC ANTI-CANCER IMMUNE MEMORY**



- When combined with immunotherapy checkpoint inhibitor Libtayo®, duration of response was further potentiated
- Upon rechallenge with two times more cancer cells and no additional treatment, tumor growth was completely prevented
- Administration of THIO alone and in combination with Libtayo® generated anti-cancer immune memory



#### THIO-103 TRIAL (PLANNED)



A Multicenter, Open-label, Phase 2 Trial Evaluating the Safety and Efficacy of THIO Administered in Sequence with Anti-PD-1 or Anti-PD-L1





## INVESTMENT OPPORTUNITY

#### EXCLUSIVITY AND INTELLECTUAL PROPERTY M MAIA



#### Goal: New Chemical Entity (NCE) Marketing Exclusivity

- THIO has never been previously approved by the FDA for commercialization
- Robust exclusivity
- US: 7 years; EU, Japan, other markets: 10 years

#### **Robust and Growing Patent Portfolio for THIO**

- 1 issued US patent
- 4 issued foreign patents
- 5 pending US patent applications
- 7 pending foreign patent applications

#### Current patents/provisional applications broadly cover the following key areas:

- Telomere targeting compounds (2034+)
- THIO's immunogenic treatment strategy: sequential combination with CPIs (2041)

#### **EXPERIENCED MANAGEMENT TEAM**





#### Vlad Vitoc, MD, MBA

Founder, Chairman, and **Chief Executive Officer** 

- 22+ years in Oncology Pharma/ Biotech: Commercial, Medical
- 12 compounds launched across 20+ tumor types
- Leadership roles at Bayer (Nexavar), Astellas (Tarceva, Xtandi), Cephalon (Treanda), Novartis (Zometa), Incyte (Jakafi)







#### Sergei Gryaznov, PhD

**Chief Scientific Officer** 

- 25+ years as Scientist
- Expert Drug Discovery and Development, Oncology with 120+ publications
- Head of the J&J Oligonucleotide Center of Excellence Worldwide
- Expert of telomeres and telomerase in cancer. coinventor of THIO







**Jeffrey Himmelreich, MBA** 









**Head of Finance** 

 Active CPA licensed in the state of Pennsylvania and is a Chartered Global Management Accountant

• 20+ years of financial expertise











#### SIGNIFICANT MARKET OPPORTUNITY





## Developing agents for the top tumor types markets globally

#### NSCLC (#1 WW)

Mortality: 1.7M Sales: \$34B

#### **CRC (#2 WW)**

Mortality: 1.0M Sales: \$ 20B



#### \$42B CPIs Group

- 5 CPIs approved for NSCLC sold \$12B
- >30% of NSCLC drug sales
- >40% of total CPI sales
- Keytruda®: \$7.5B in NSCLC of \$21B total



#### Partnership with Regeneron (Libtayo®)

- Profile similar to Keytruda®
- Libtayo® is entrant #5 in CPIs
- Needs superior efficacy to Keytruda®
- Sequential combination with THIO is key

#### **Checkpoint Inhibitors Market**













#### **COMPARABLE COMPANIES**



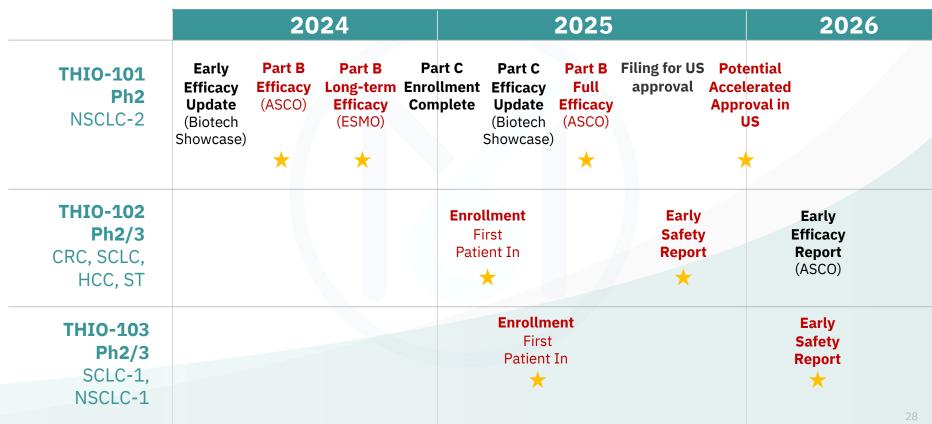


- On June 3, 2022, Bristol Myers Squibb (BMS) announced the acquisition of Turning Point Therapeutics in an all-cash transaction for **\$4.1B** in equity value.
- On October 9, 2023, BMS acquired Mirati for <u>\$4.8B</u> in cash, plus up to \$1B in contingent value right.

#### **MULTIPLE VALUE-DRIVING MILESTONES**



#### ★ Major inflection points





### **APPENDIX**

#### CAPITALIZATION TABLE & CASH BALANCE



<b>Capitalization</b>	<b>Table</b> (as of 9/30/2023)
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Common stock	13,761,123
Options (WAEP: \$2.62) <sup>1</sup>	7,860,736
Warrants (WAEP: \$5.59)	924,760
Fully Diluted Shares Outstanding	22,546,619

Cash Balance of \$6.10<sup>2</sup> million (as of 9/30/2023)

Note: Directors and officers, and their affiliates, own 39.2% of the 22,546,619 fully diluted shares outstanding

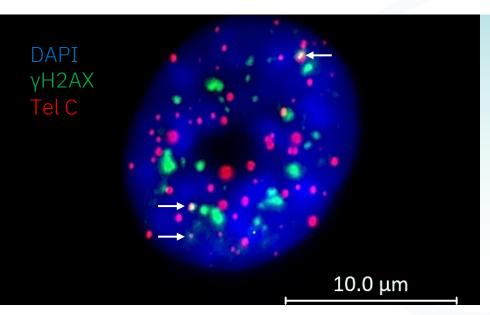
<sup>&</sup>lt;sup>1</sup> 5,252,847 options held by directors and officers

<sup>&</sup>lt;sup>2</sup> Includes the net proceeds from a public offering of common stock in April 2023

#### BIOMARKER – TIFS (TELOMERE DYSFUNCTION INDUCED FOCI)

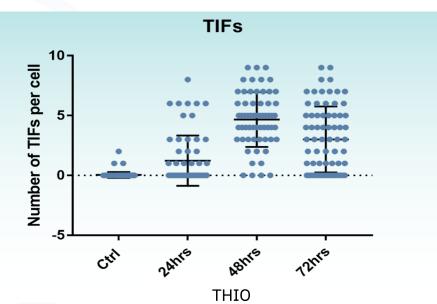


## Confocal microscopy image of LLC cell nucleus after treatment with THIO



- Yellow dots indicated TIFs by THIO
- Green dots yH2AX
- Red dots telomeres

## Quantification of TIFs induced in LLC cell by 3 µM of THIO



- TIFs induction reached max after ~ 48h
- Formation of TIFs indicated on-target MOA of THIO