

Mining for Tomorrow's Cures

Licensing of CLN-418 (HBM7008) from Harbour Biomed

February 14th, 2023

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AGENDA

1. Introduction

Nadim Ahmed

2. CLN-418 Overview

Jeffrey Jones

3. Deal Terms & Financial Context

Corinne Savill

4. Strategic Perspective

Nadim Ahmed

5. Q&A

PRESENTERS

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Chief Medical Officer

Corinne Savill

Chief Business Officer

JOINING FOR Q&A





Patrick Baeuerle, Ph.D.

Chief Scientific Officer

Jeff Trigilio

Chief Financial Officer

Diversified pipeline leveraging novel technologies and differentiated mechanisms

Program Modality/MOA	IND-Enabling	Phase 1	Phase 2	Phase 3	Status	
Zipalertinib (CLN-081/TAS6417) EGFRex20ins inhibitor	NSCLC with EGFR exon 20 insertion mutations				BTD received; actively enrolling Phase 2b pivotal	 Co-commercialization and Co-development in US
CLN-049 FLT3 x CD3 bispecific	R/R AML, MDS				Actively enrolling	 or its Subsidiary retains Worldwide Rights
CLN-619 Anti-MICA/B IgG1	Pan-cancer				Actively enrolling	
CLN-418 B7H4x41BB bispecific immune activator	Multiple solid tumors				Actively enrolling	 ex-U.S. Rights retained by Harbour Biomed
CLN-978 CD19/CD3 T-cell antibody construct with HSA binding domain	B-cell NHL				FDA IND clearance received	 or its Subsidiary retains Worldwide Rights
CLN-617 Collagen-binding IL-12-IL2 fusion protein	Pan-cancer				IND-enabling studies	

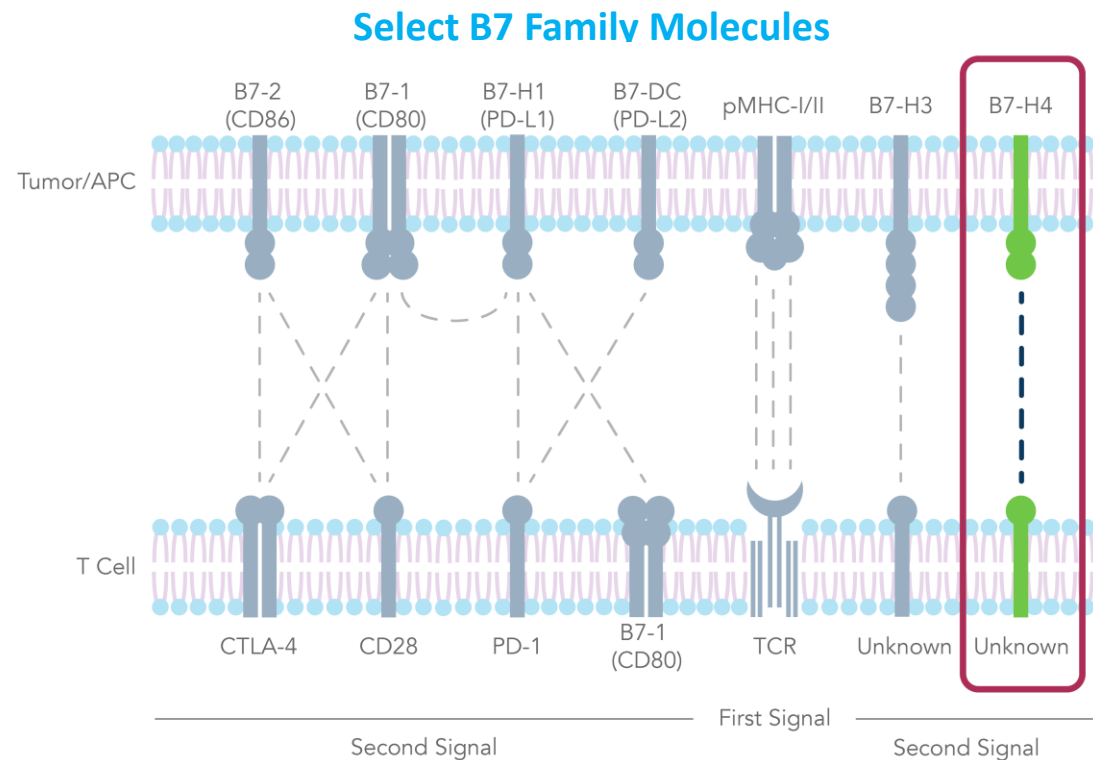
Potential for 6 clinical stage programs by YE23

B7H4: Tumor antigen that is member of key immune regulatory protein family

B7 family of ligands are part of the “master switches” of T-cell activation

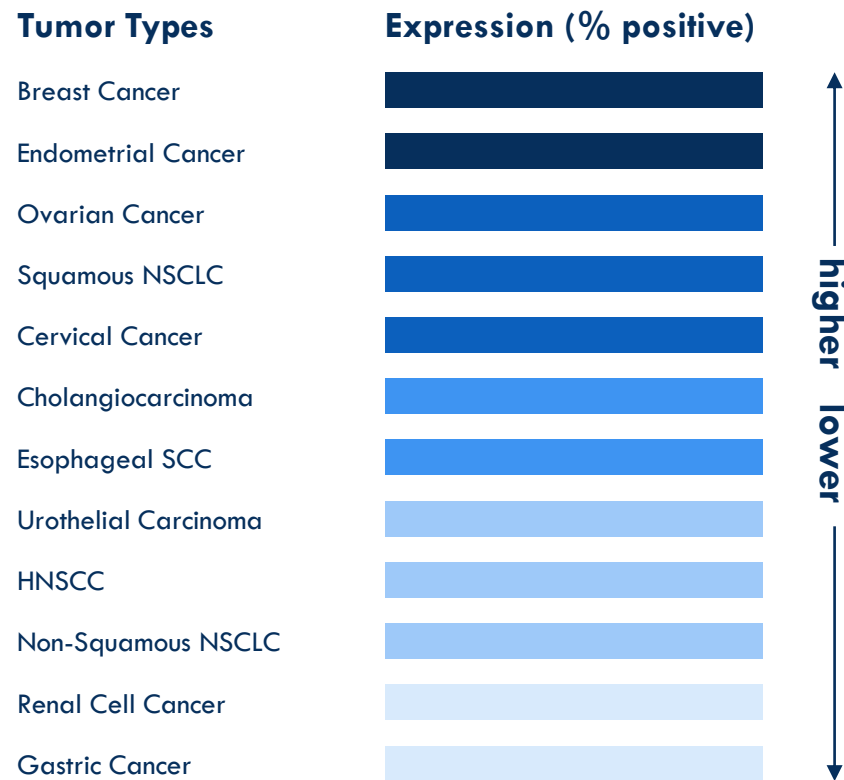
High surface expression of B7H4 on tumor cells, limited expression on normal cells

Inhibits T cell activation and correlates with poor prognosis in multiple tumor types

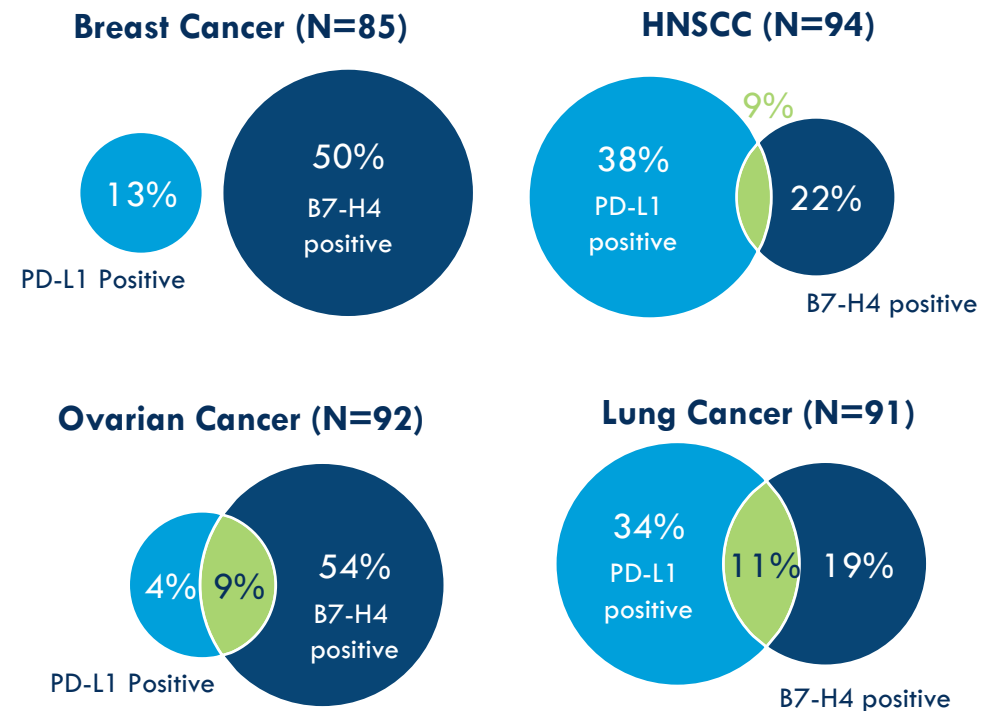


B7H4 is an attractive target for cancer immunotherapy

Expressed across range of solid tumors¹



Minimal overlap with PD-L1 expression²



- Highly expressed on multiple tumor types of high unmet need
- Expressed on tumors with low PD-L1 expression, potentially addressing tumor types for which immunotherapy exhibits limited efficacy ('cold tumors')

¹ In-house IHC data from Harbour BioMed

² Innovative B7H4 x CD3 & B7H4 x 4-1BB Bispecifics for Solid Tumor Therapies, Presented at 13th Annual Summit World Multispecifics, 2022

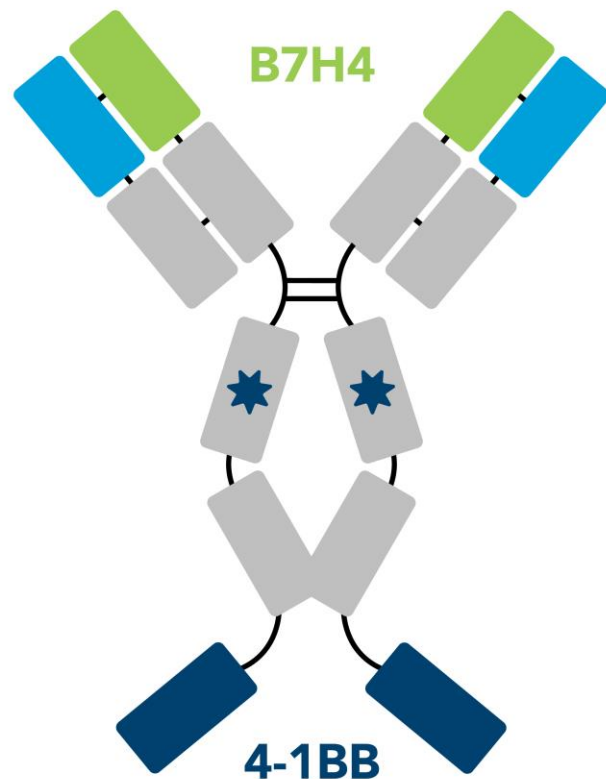
4-1BB agonism is an ideal way to harness the potential of B7H4

4-1BB: Key costimulatory molecule for T- and NK-cells

- 4-1BB activation drives proliferation and increased cytotoxicity of both T- and NK- cells
- 4-1BB monoclonal antibodies have demonstrated clinical activity, but have been limited by toxicity (e.g., urelumab)
- Contingent 4-1BB activation upon binding to B7H4 is designed to avoid these toxicities by localizing the T- and NK-cell activation to the tumor
- 4-1BB agonists have mechanistic rationale for high combination potential with other anti-cancer therapies

CLN-418: A potential first-in-class B7H4 x 4-1BB bispecific

Fully human bispecific antibody optimized using Harbour's Heavy Chain Antibody Based Immune Cell Engager (HBICE) Platform



CLN-418's Potential Differentiation

Expected wide therapeutic window

- Dual binding to tumor target B7H4 mediates avidity for stronger binding
- 4-1BB activation dependent on crosslinking of antibody avoids non-conditional immune agonism, based on preclinical data
- Silenced Fc domain avoids off target T-cell activation by Fc-gamma receptor positive cells

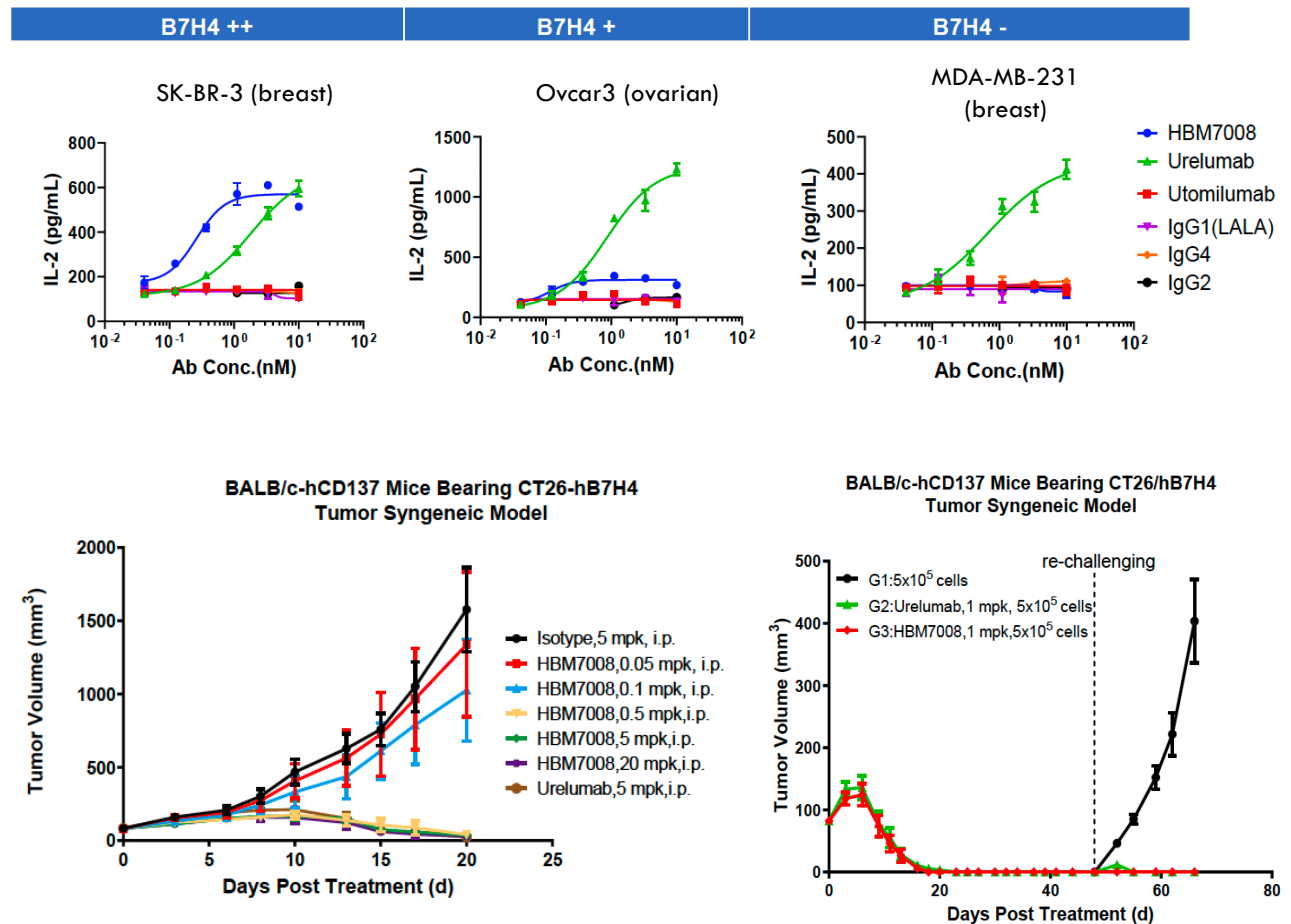
Convenient dosing with low immunogenicity risk

- Half-life extended via Fc domain
- Human B7H4 and 4-1BB binding domains

CLN-418 shows target-dependent, single-agent anti-tumor activity in preclinical studies

CLN-418 only elicits T-cell activation in the presence of B7H4

CLN-418 clears tumors in mice and drives a memory response that prevents tumor growth upon re-challenge



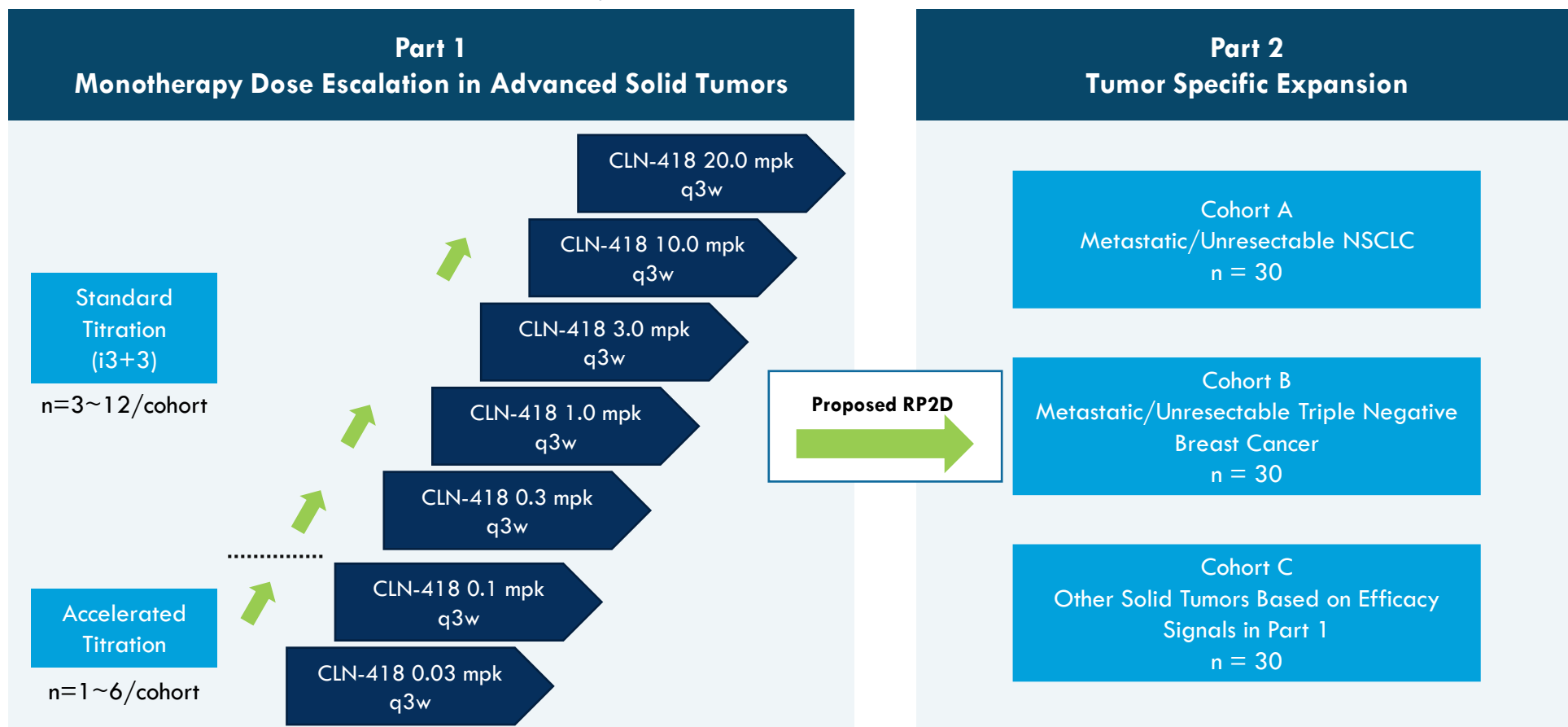
CLN-418 is positioned favorably relative to alternative approaches and is the only B7H4 x 4-1BB bispecific in the clinic



	Therapeutic Potential	CLN-418 B7H4 x 4-1BB	PDL1 x 4-1BB bsAb	B7H4 x CD3 T cell engagers	B7H4-ADC Ab-drug conjugate
41BB (effector moiety) driven	potential for SINGLE-AGENT ACTIVITY	✓	✓	✓	✓
	engages both T and NK CELLS	✓	✓	x	x
B7H4 (tumor target) driven	MULTIPLE INDICATION POTENTIAL (pan tumor target)	✓	✓	✓	✓
	TUMOR SELECTIVITY	higher	lower	higher	higher
B7H4 and 41BB driven	POTENTIAL THERAPEUTIC INDEX (favorable safety)	broader	broader	narrower	narrower
	potential for activity in PDL1 LOW TUMORS	higher	lower	higher	higher

CLN-418 Phase 1 study design (NCT05306444)

A first-in-human study to evaluate the safety and tolerability of the study drug CLN-418, and to determine the maximum tolerated dose and/or recommended Phase 2 study dose of CLN-418.



Parallel combination arms are being planned

INITIAL CLINICAL DATA EXPECTED 2024

Summary of terms



Upfront Payment

\$25 million for U.S. rights



Milestone Payments

Up to \$148 million in development-based milestones
Up to an additional \$415 million sales-based milestones



Royalties

Tiered royalties from low single digit to teens



Exclusive rights to U.S. market

Most attractive commercial territory for
new oncology products

Expands pipeline with another potential first-in-class, multi-indication molecule

Strong strategic fit within area of core expertise

- Places Cullinan at the forefront of bispecific antibody development in solid tumors, harnessing Cullinan's core expertise developing bispecific antibodies in hematology

B7H4 is a high-impact target

- B7H4 expressed widely across multiple tumors with low expression on normal tissue, potentially giving a wide therapeutic index
- Minimal overlap with PD-L1 creates opportunity where existing IO approaches have limited efficacy

4-1BB Bispecific is optimal approach to target B7H4

- 4-1BB is a key costimulatory molecule for T- and NK-cells
- Crosslinking activation of 4-1BB with B7H4 may preserve efficacy and limit toxicity
- CLN-418 optimized on Harbour's HBICE platform

Phase 1 study underway

- Monotherapy dose escalation ongoing; combination development planned
- Initial clinical data expected 2024

Attractive financial terms leave strong cash runway intact

- \$25 million upfront for exclusive U.S. development and commercialization rights
- Cullinan maintains \$525 million in *pro forma* cash post deal, sufficient to fund company into 2026, past multiple potential value creating milestones

Q&A

Nadim Ahmed

Chief Executive Officer

Jeffrey Jones, M.D., MPH, MBA

Chief Medical Officer

Corinne Savill

Chief Business Officer

Patrick Baeuerle, Ph.D.

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