



HOOKIPA Pharma

Next-generation Immunotherapies for the
Treatment of Cancer and Other Serious Diseases

January 2025

NASDAQ: HOOK



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Merger Transaction Overview

Transaction Structure

- Proposed all stock transaction where HOOKIPA Pharma Inc. (“HOOKIPA” or “HOOK”) acquires Poolbeg Pharma plc (“Poolbeg” or “POLB”)
- HOOKIPA remains TopCo with existing Nasdaq listing
- Concurrent Financing: Private placement of approximately \$30M+ to be funded into HOOKIPA immediately following transaction close
- Implied ownership split pre-merger without PIPE on a fully-diluted basis: POLB shareholders 55.0% / HOOKIPA shareholders 45.0%
- In addition, CVR's for HB-200, HB-400 & HB-500 Programs (for pre-PIPE HOOKIPA shareholders)
- POLB expected to apply for cancellation of its shares on AIM Market at transaction close and become a private sub of HOOKIPA

Capitalization and Use of Proceeds

- Combined company expected to have sufficient capital to realize meaningful value inflection points
 - HB-700: Phase 1 interim data expected in H1 2026
 - POLB 001: Phase 2a topline data expected in H2 2026
 - HB-500: Phase 1b Primary completion expected in H2 2025
- Expected to provide cash runway through year-end 2026*

Transaction Timeline

- Possible offer announcement in line with Rule 2.4 of the UK City Code on Takeovers and Mergers
- HOOKIPA to either announce a firm intention to make an offer for POLB, or not, under Rule 2.7 of the UK City Code on Takeovers and Mergers
- Concurrent financing contingent on transaction close

Post-Closing



- Combined company anticipates benefiting from a strong international leadership team comprised of individuals with both significant industry experience and a track record of success





Investment Highlights

- Next-generation cancer immunotherapy portfolio driven by lead assets each with \$10B+ market opportunities with potential to gain significant market share ¹⁻²
 - Phase 1-ready multi-KRAS-targeting HB-700 (derisked by clinical POC with platform asset eseba-vec (HB-200))
 - Phase 2-ready small molecule POLB 001
- Strategic partnership with Gilead on HBV and HIV programs in Phase 1b with potential of up to \$417.5M potential future opt-in, development and commercial milestones and significant sales royalties
- Multiple near-term clinical data catalysts in multiple programs over next 24 months in large therapeutic areas with unmet medical needs
- Clinically validated antigen-specific T cell activation platform designed to produce durable, robust anti-tumor activity underpins HB-700
- Proven leadership team with extensive experience in successfully developing and commercializing medicines with a track record of delivering shareholder returns

HOOKIPA Has a Diversified Core Immunotherapy Pipeline

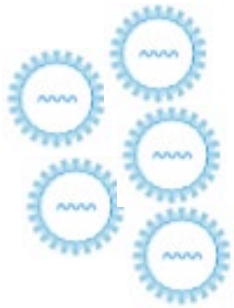
Product	Modality	Indication	Preclinical	Phase 1	Phase 2	Phase 3	Key Milestone
Oncology Programs							
HB-700	Next-generation immunotherapy	KRAS Mutated Tumors	Phase 1-ready				FPD in Phase 1 trial expected mid-2025
POLB 001	p38 MAPK inhibitor	Immunotherapy-induced CRS	Phase 2-ready				FPD in Phase 2 trial expected H2 2025
Partnered Programs in Infectious Disease							
HB-400¹	Next-generation immunotherapy	HBV	Gilead-led Phase 1 ongoing				Primary completion expected H1 2025
HB-500²	Next-generation immunotherapy	HIV	HOOKIPA-led Phase 1 ongoing				Primary completion expected H2 2025

HOOKIPA Pipeline has Additional Partnership Opportunities

Product	Modality	Indication	Preclinical	Phase 1	Phase 2	Phase 3	Key Milestone
Additional Programs							
Eseba-vec (HB-200)	Next-generation immunotherapy	HPV16+ HNSCC	Mature Phase 2 data with POC in combo with CPI				Final Phase 2 data expected H2 2025
GLP-1 Program	GLP-1R agonist	Obesity and diabetes					Topline POC data expected H1 2026
AI Programs	Novel target discovery	Influenza					Potential partnership
		RSV					Potential partnership

HOOKIPA's Next-Gen Vaccine Platform Designed to Supercharge Immunity

T cell activation platform based on work of Nobel laureate and HOOKIPA co-founder Rolf Zinkernagel⁵



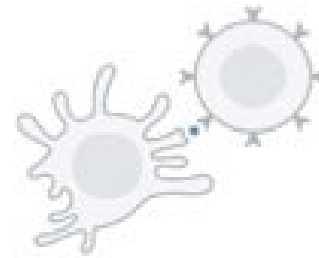
Drug Design

Heterologous and alternating 'prime-boost' arenavirus vectors with target antigens



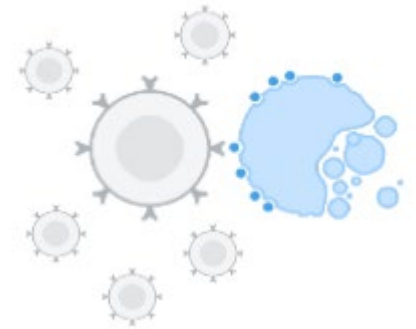
Infection of APCs

Dendritic cells or macrophages



Activation of T cells

Tumor-specific T cell expansion and activation



Tumor Cell Killing

Robust anti-tumor activity

Unprecedented levels of cancer-specific T cells with polyfunctionality & durability with continued treatment¹⁻⁴



HB-700

Next-generation multi-KRAS targeting cancer immunotherapy with strong commercial potential

KRAS is the Most Prevalent Oncogenic Driver

~1.5M people worldwide

are diagnosed annually with KRAS-mutated NSCLC, CRC, or PDAC

~20%

All NSCLC

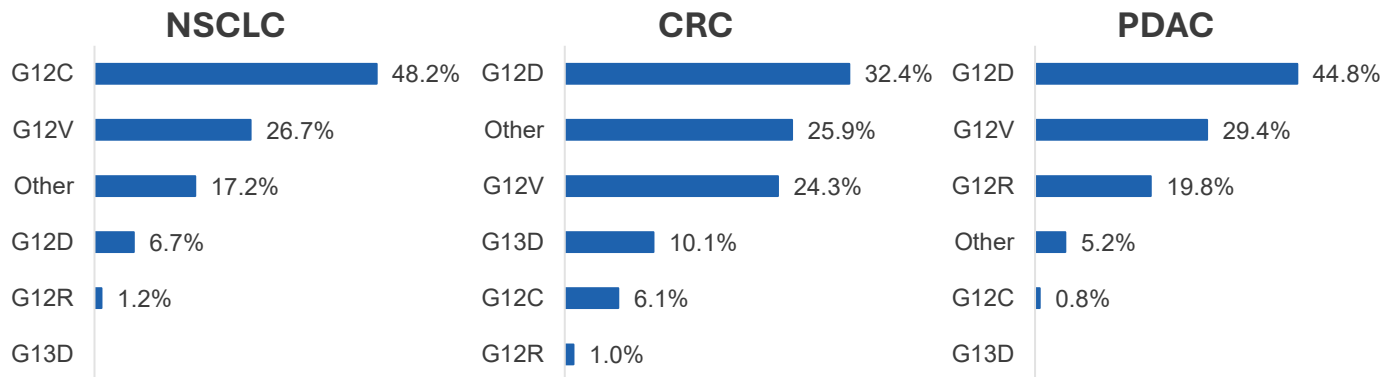
~30-40%

All CRC

≥88%

All PDACs

Prevalence of the top 5 KRAS mutations by tumor type



KRAS Market Opportunity

Targeting KRAS has been challenging due to lack of activity, poor selectivity or treatment resistance

Approved standards of care are small molecules targeting a single mutation

~\$5-6B

2038 global KRAS market size estimate

Unmet needs remain for a therapy that can

- Address multiple mutation subtypes
- Drive deep and durable responses
- Achieve immunogenic tumor cell death



A Novel Multi-KRAS Mutant Targeting Cancer Immunotherapy

Designed for uniquely strong anti-tumor T cell activation

Targets 5 of the most prevalent KRAS mutations¹

G13D, G12V, G12C, G12D & G12R

Strong preclinical proof-of-concept package²

KRAS mutation specific T cell activation in humanized mice; preclinical profile supports diverse combinations

Derisked by clinical POC with platform asset eseba-vec

Dose selection and treatment schedule based on alternating 2-vector therapy in eseba-vec program - demonstrated clinically durable and robust anti-tumor activity

Phase 1-ready

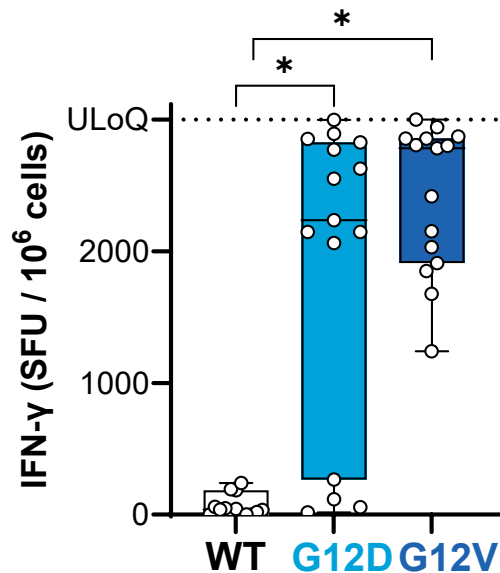
Nonclinical development and clinical trial material manufacturing completed

Strong commercial potential

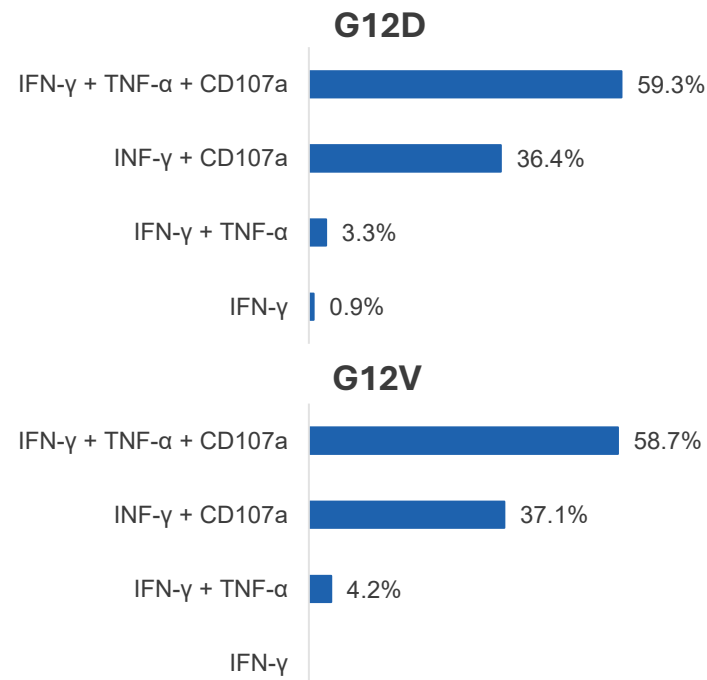
Large addressable populations in NSCLC, CRC and PDAC

Preclinical Proof of Concept: Highly Immunogenic with Potent Target Cell Killing in Humanized Mice

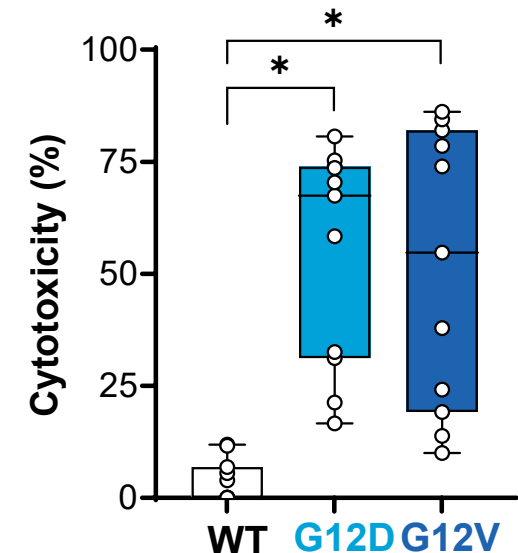
High frequencies of circulating KRAS mutation specific CD8⁺ T cells



KRAS^{mut} specific T cells exhibit a polyfunctional profile



KRAS^{mut} specific T cells are functional and specifically kill target cells



Proposed Open-Label Phase 1/2 in Metastatic KRAS-Mutated NSCLC

Clinical POC in NSCLC followed by expansion in related indications

Eligibility Metastatic/advanced NSCLC & CRC
KRAS mutated and HLA selected
≥1 prior line SoC

Eligibility Metastatic/ advanced NSCLC
PD-L1 scores of TPS ≥50%
KRAS-mutated and HLA selected
No prior line of therapy

Phase 1 Dose De-Escalation

2L+ NSCLC & CRC

HB-700
Intended dose; n = 3 + 3

Phase 1 Dose Expansion

2L+ NSCLC

NSCLC
n = 6

Primary Endpoint:

RP2D

Secondary Endpoint:

Safety, T cell response,
initial antitumor activity

Phase 2 Combination

1L NSCLC

HB-700 + Pembrolizumab
n = 30

Primary Endpoint:

ORR by RECIST v1.1

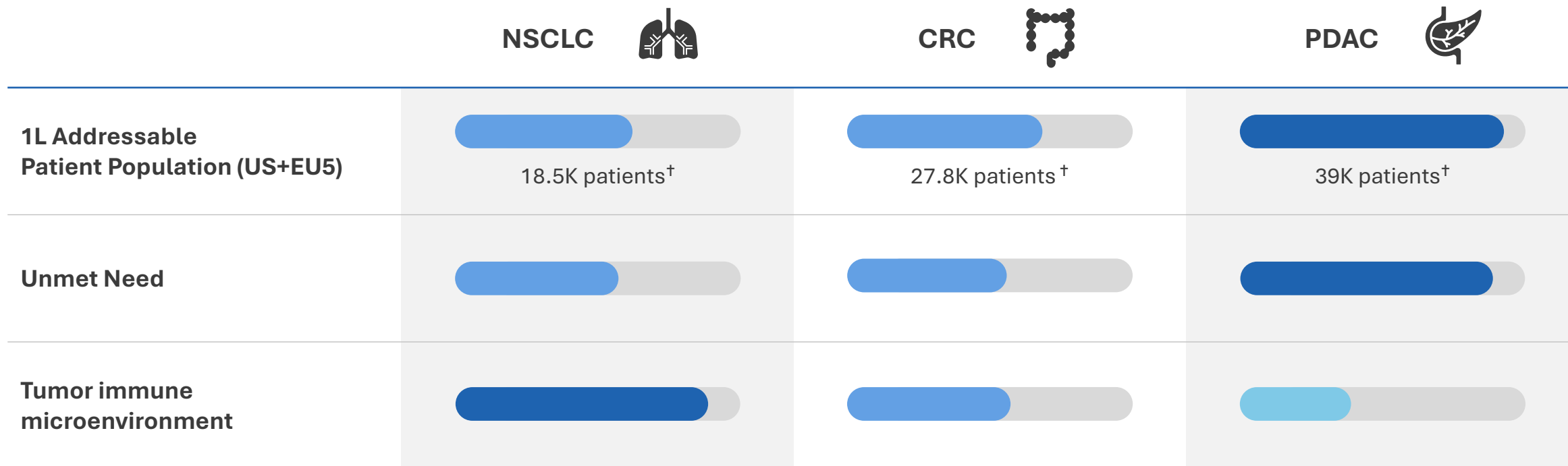
Secondary Endpoint:

Safety, DCR, DoR, PFS, OS, T cell
response and tumor infiltration

Expect FPD in Phase 1 trial mid-2025 with interim Phase 1 data in KRAS mutant tumors in H1 2026

HB-700 KRAS Immunotherapy has Potential for \$1.5B Peak Worldwide Net Sales* Across 1L NSCLC, CRC and PDAC, if approved

Additional expansion opportunities in 2L and neoadjuvant/adjutant for locally advanced disease

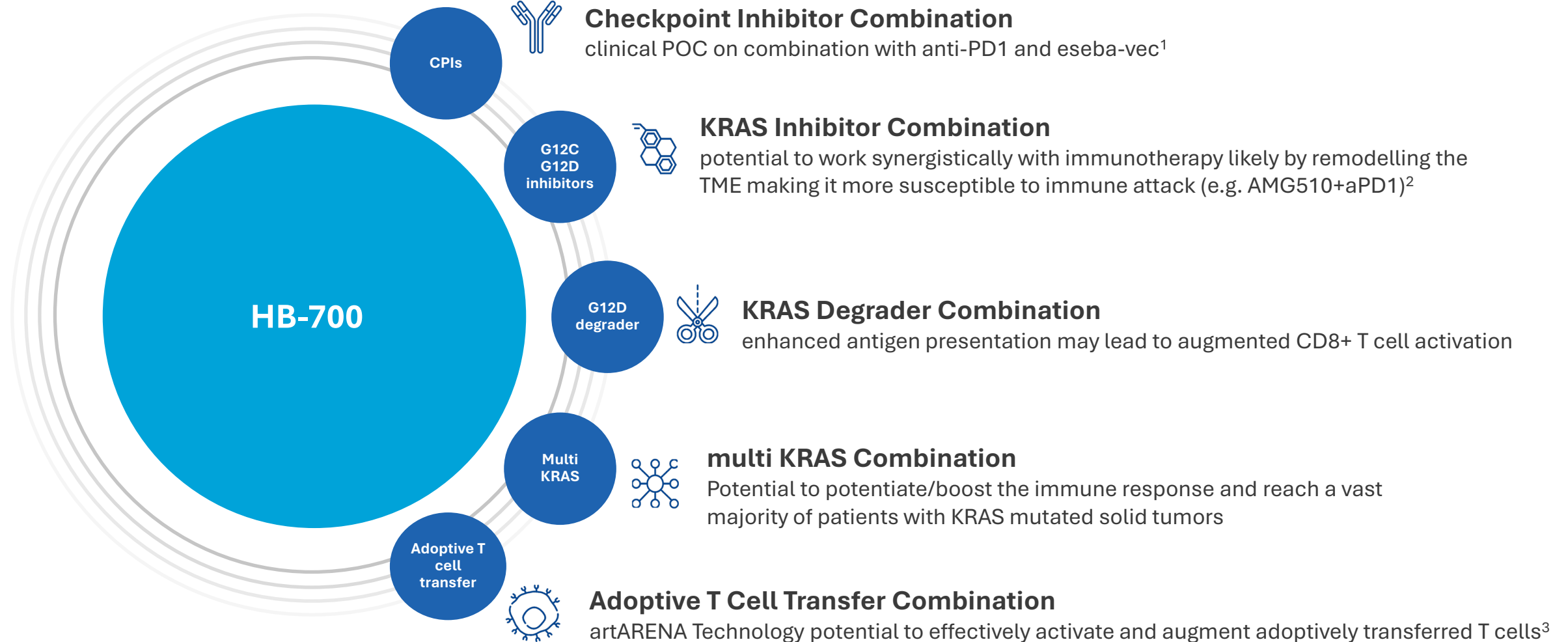


Relative Opportunity for HB-700



*Assumptions: HLA restriction is included (54% NSCLC, 61% PDAC, 48% CRC). US launch: 2032, 3YTP; EU5 launch: 2033, 5YTP. Settings 1L NSCLC, CRC and PDAC with 10-18% market share
[†] Source: Trinity Life Sciences

HB-700 Potential to Combine with Diverse Approved and Emerging Therapies





POLB 001

Potentially breakthrough orally delivered p38 MAPK inhibitor to prevent cancer immunotherapy-induced CRS



CRS Associated with Immunotherapies is a High Unmet Need

Effective prophylaxis represents a >\$10B market opportunity⁴

Cytokine Release Syndrome (CRS)

A severe, potentially life-threatening side effect of cancer immunotherapies

No approved therapies for prevention

Approved options for CRS management (tocilizumab) **have not adequately*** prevented Grade 2+ CRS in clinical trials

>70%¹ of patients experience CRS

on certain CAR T / bispecific antibody therapies and are restricted to specialist cancer centers

Estimated \$5B annually²⁻³

in direct costs to US health systems by 2030
CRS of all grades can require hospitalization

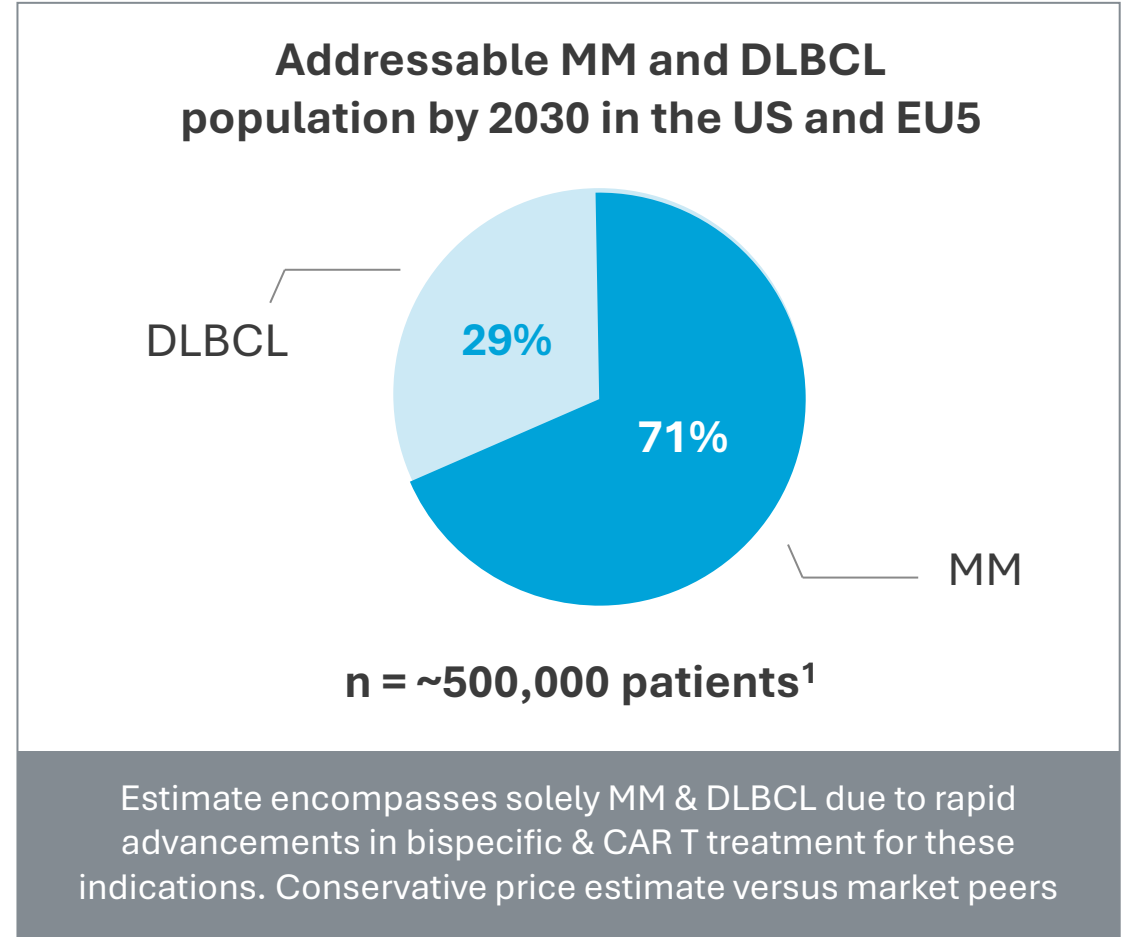
CRS Preventative Therapy: >\$10B US Market Opportunity

A significant opportunity exists for POLB 001 as an adjunct therapy to bispecific and CAR T treatment³

1st, 2nd and 3rd line+ MM and DLBCL patients in the US and EU5, receive CAR T cell and bispecific antibody therapy¹

An effective primary prophylactic for CRS could **enable outpatient administration and broader uptake** of immunotherapies²

Potential across additional hematological malignancies, solid tumors and new areas like severe influenza





Potential to Make Immunotherapies Safer & More Accessible

Selective p38 MAPK Inhibitor

- Selectively prevent excessive inflammation without immunosuppression
- Oral agent
- Strong patent portfolio

Strong Preclinical & Clinical Data

- Phase 2-ready
- Favorable safety and tolerability profile
- Potent TNF- α inhibition shown in two Phase 1 trials
- Potent inhibition of IL-6 and other key inflammatory markers in clinical & preclinical models

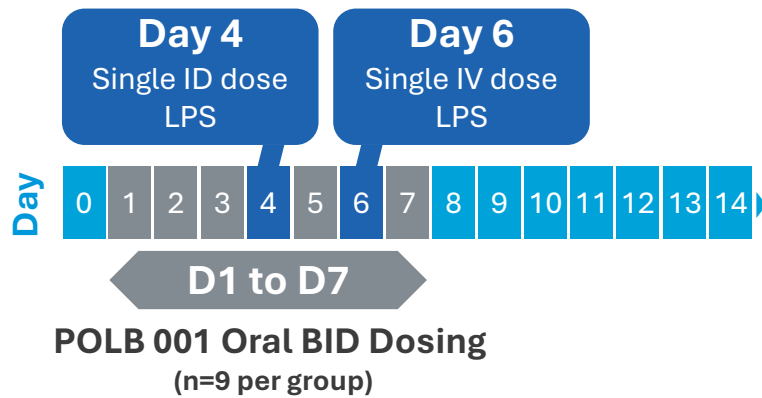
Significant Market Opportunity

- >\$10B market opportunity
- Potential for Orphan Drug Designation
- No approved therapy for CRS prevention

LPS Human Challenge: Potent Inhibition of Excessive Inflammation

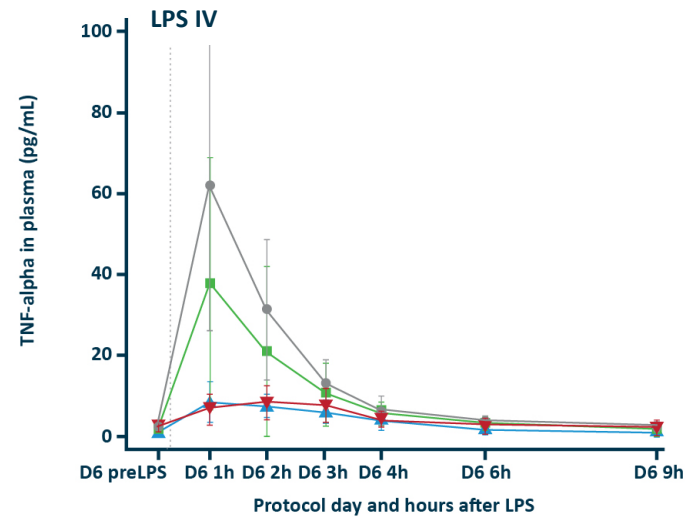
Supportive of potential of POLB 001 as a prophylactic for cancer immunotherapy-induced CRS

Trial design



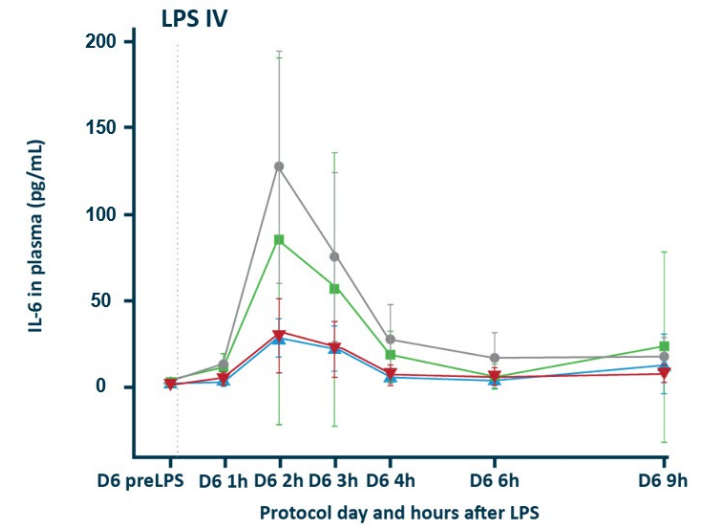
Potential to effectively prevent CRS while preserving key immune system functionality

TNF-α



73.5% and 56.2% maximal reduction compared to placebo for 70 mg and 150 mg doses respectively ($p = 0.0003$)

IL-6



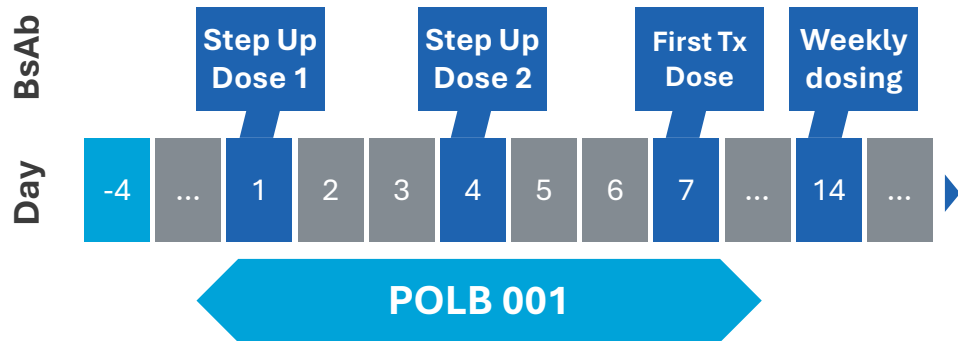
57.4% and 63.5% maximal reduction for 70 mg and 150 mg doses respectively ($p = 0.0002$)

● Placebo ■ 30 mg POLB 001 ▲ 70 mg POLB 001 ▼ 150 mg POLB 001

Planned POLB 001 Phase 2a Investigator Initiated Trial

Prevention of CRS in R/R Multiple Myeloma Patients Receiving Bispecific Ab

Trial design



Phase 2a IIT

Single Arm

N = 30 - 40

BID Oral POLB 001

Single dose level

SoC BsAb*

Key Objectives/Endpoints

Incidence of Grade 2+ CRS

Incidence of CRS all grades

Confirm safety and pharmacokinetics

Exploratory biomarker analysis

Expect FPD in Phase 2 trial H2 2025 with topline data expected in H2 2026

*Clinical trial collaboration and supply agreements with a large pharma company expected for approved BsAb



Gilead Strategic Partnerships

HB-400 and HB-500



Aim to Develop Functional Cures for HBV and HIV

HB-400

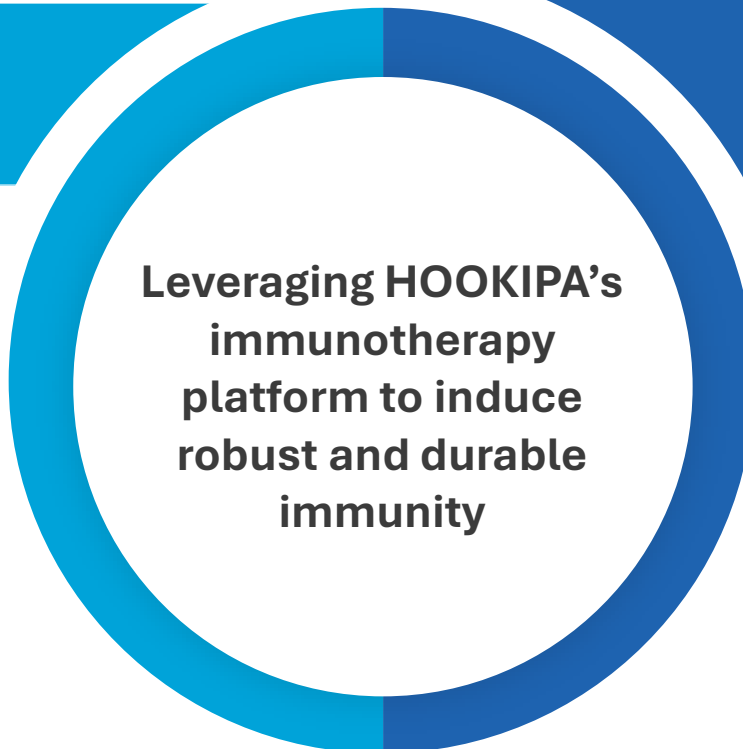
For the treatment of HBV

Alternating, two-vector non-replicating arenaviral HBV immunotherapy

High Potential candidate in Gilead's efforts to develop a curative regimen of treatments

Phase 1 enrollment completed

**Primary completion
expected H1 2025**



**Leveraging HOOKIPA's
immunotherapy
platform to induce
robust and durable
immunity**

HB-500

For the treatment of HIV

Alternating, two-vector replicating arenaviral HIV immunotherapy

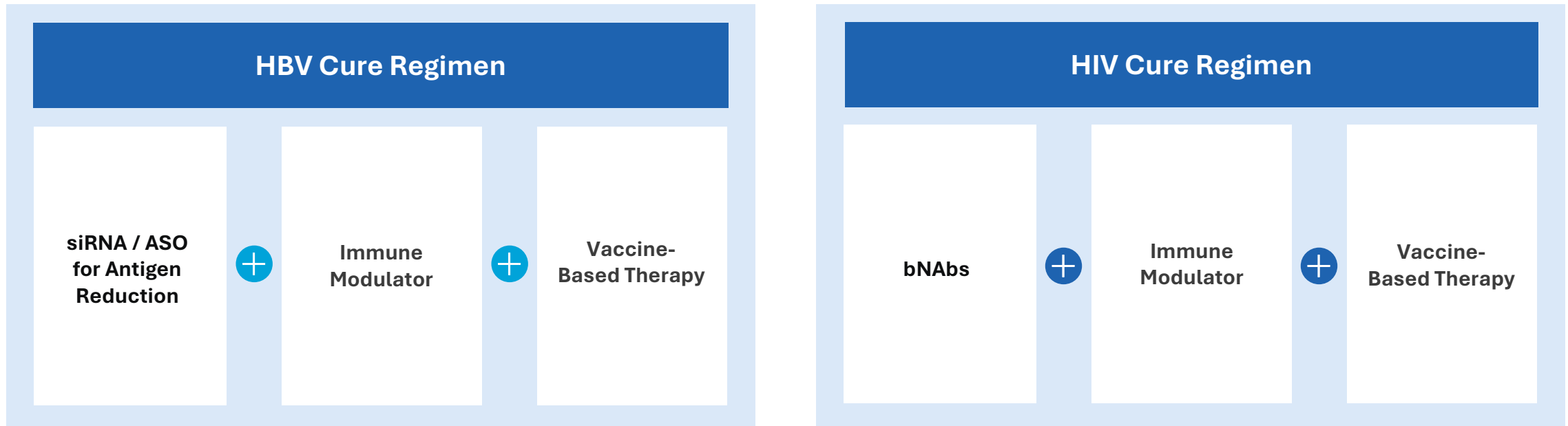
Ongoing Phase 1b study

FPD July 1, 2024, with enrollment expected to complete by Jan 2025

**Primary completion
expected H2 2025**

HOOKIPA's Immunotherapies Are Prominently Featured in Gilead's Pipeline

Using combination strategies and novel mechanisms with the goal to drive viral suppression and durable immunity¹⁻⁴



HOOKIPA's immunotherapies have potential to build long-term immune responses



Partnership Offers Significant Revenue Potential

HB-400 for HBV

Gilead responsible for clinical development

Next potential milestone payment: start of Phase 2

HOOKIPA Responsibilities

- Vector design
- Manufacturing and supply of clinical material

Terms

- \$185M potential future development + commercialization milestones
- High-single digit to mid-teen % royalties
- All costs borne by Gilead, including HOOKIPA spend

HB-500 for HIV

Gilead retains exclusive option post Phase 1

HOOKIPA Responsibilities

- Vector design
- Conducting Phase 1b clinical trial

Terms

- \$232.5M potential future opt-in, development + commercial milestones
- Mid-single digit to low double-digit % royalties



Additional Programs

Eseba-vec (HB-200), Oral GLP-1, AI Programs



Eseba-vec (HB-200) in Recurrent/Metastatic HPV16+ HNSCC

Strong scientific thesis with mature Phase 2 data and POC in combination with checkpoint inhibitors

Leading Phase 2 Data in HPV16+ R/M OPC CPS \geq 20

- 52% response rate in CPS \geq 20, best among vaccine approaches²
- 16% complete response in CPS \geq 20 HNSCC patients²
- Durable responses leading to progression free survival of 16.3 months²
- Favorable safety profile and well tolerated

Clearly Defined Registrational Path

- FDA-endorsed strategy for potential **accelerated approval**
- **EMA PRIME designation received**
- Robust clinical and preclinical data package

Large Addressable Market with Expansion Opportunities

- Initial opportunity: ~1,500-3,000 patients with 1L HPV16+ R/M OPC (CPS \geq 20)¹
- Potential expansion across HPV16+ OPC continuum
- Other HPV16+ cancers (non-OPC HNSCC, anal, cervical, penile, vulvar, vaginal)
- Additional ~17,500-20,000 patient opportunity¹



Additional Pipeline Programs in Large Market Opportunities

GLP-1 Program – *Oral GLP-1R agonist*

Obesity and Diabetes Treatment

Phase 1 ready asset

Proprietary Delivery Technology

Potential to overcome oral delivery challenges of peptide-based biologicals¹

Proof-of-concept trial initiation expected H1 2025

AI Programs – *Drug Discovery*

RSV and Influenza

Preclinical assets

AI Platform Programs, Drug Discovery & Repurposing Opportunities²

Integrated proprietary multi-parametric clinical data to identify novel host response targets

A number of potential targets and clinical-stage repurposing candidates have been identified
Discussions ongoing in respect to collaborations



Combined Company and Financial Overview



Global Company with Strong Patents and Cash Runway

HOOK (NASDAQ)

Combined company expected to have operations in EU, UK & US

Robust patent portfolio covering:

- Platform patents
- Product-specific patents
- Oncology platform patents

Expected to be Debt Free with Cash Runway Through YE 2026*

Offering Size

Approximately
\$30M+

Expected to Fund Key Catalysts

HB-700

Phase 1 interim data expected H1 2026

POLB 001

Phase 2a topline data expected H2 2026

HB-500

Phase 1b primary completion expected H2 2025

Clinical Milestones in High Interest Areas Over the Next 24 Months

Cash runway expected to be extended through YE2026* including HB-700, POLB 001 & HB-500 milestones

Program	Indication	2025		2026		2027	
		H1	H2	H1	H2	H1	H2
HB-700 ¹	KRAS Mutated Tumors	Phase 1/2 FPI		Phase 1 interim data		Phase 1/2 full readout	
POLB 001 ¹	Immunotherapy-induced CRS		Phase 2 FPI		Phase 2 topline data		
HB-400 ²	HBV	Phase 1b primary completion					
HB-500 ³	HIV		Phase 1b primary completion				
Eseba-vec ¹	HPV16+ HNSCC		Final Phase 2 readout				
GLP-1 Program ¹	Oral GLP-1	POC trial FPI		POC trial topline data			

Proposed Team with Proven Execution and Operational Leadership



Malte Peters, MD, PhD
Chief Executive Officer



Mark Winderlich, PhD
Chief Development Officer



Cathal Friel
Executive Chairman, Poolbeg Co-Founder



Ian O'Connell
Chief Financial Officer, Poolbeg Co-Founder



David Allmond
Chief Business Officer



John McEvoy
Chief Legal Officer





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- Multiple near-term clinical data catalysts in multiple programs over next 24 months in large therapeutic areas with unmet medical needs
- Clinically validated antigen-specific T cell activation platform designed to produce durable, robust anti-tumor activity underpins HB-700
- Proven leadership team with extensive experience in successfully developing and commercializing medicines with a track record of delivering shareholder returns



Appendix

HOOKIPA-Poolbeg Merger Brings a New Management Team with a Track Record of Delivering Shareholder Returns



- World class rare and orphan focused biopharma co-founded and comprising of Poolbeg senior management team
- Listed 2016 c.\$50M and acquired in 2023 for \$1.48B
- Restructured and fixed underperforming assets, driving development & commercial success across multiple markets
- Made strategic choices to rapidly generate substantial value for shareholders, including:
 1. Approval of Filisuvez for EB and market launch
 2. In-licensing of Lomitapide
 3. Acquisition of Aegerion Pharmaceuticals
 4. Acquisition of Chiasma Inc



- Poolbeg's co-founders Cathal Friel and Ian O'Connell took control of distressed hVIVO via a vehicle they co-founded called Open Orphan (later renamed hVIVO).
- Grew sales revenue from c.\$30M in 2019 to an expected c.\$77M for 2024, and market cap from c.\$15M to c.\$175M
- Restructured to refocus operations on core strengths and implement efficiencies to drive revenue growth
- Poolbeg Pharma spun-out of Open Orphan, bringing virology expertise



- Mark Winderlich and Malte Peters co-led development and approval strategy of tafasitamab in combination with lenalidomide in 2L+ DLBCL in US, EU, and other countries using real-world data
- Assembled an experienced development team, that successfully led 3 large Phase 3 studies in 1L DLBCL, 2L FL, 1L MF, leading to the acquisition of Morphosys by Incyte and Novartis
- Led felzartamab clinical development in oncology & AI, which was acquired by hBIO, later Biogen.



- Malte Peters was responsible for the development and approvals of a PI3K inhibitor, BRAF/MEK inhibitor, CDK4 inhibitor, c-MET inhibitor, and other molecules
- Introduced the concept of patient selection based on molecular profiles, leading to proof of concept and accelerated approvals from Phase 1 and 2 trials
- Led the clinical team at Sandoz, Novartis' generic division, to achieve approval of rituximab and Enbrel biosimilars



POLB 001: An Oral p38 MAPK Inhibitor That Selectively Targets Key Inflammatory Pathways Without Broad Immunosuppression

Phase 2 ready asset with a comprehensive pre-clinical and clinical data package

Favorable Safety and Tolerability Profile



97 subjects dosed during Phase I FIH and LPS Challenge studies



No SAEs or discontinuations due to AEs, all were of mild intensity



No clinically meaningful findings in clinical laboratory test results, vital signs or ECG



Favorable safety & tolerability profile

Designed to Prevent Immunotherapy-Induced CRS



Suitable for at-home dosing (used in LPS Challenge Study)



Hepatic metabolism and biliary excretion profile favorable for multiple myeloma and renally impaired populations



BID oral regimen designed to provide targeted protection during CRS risk period

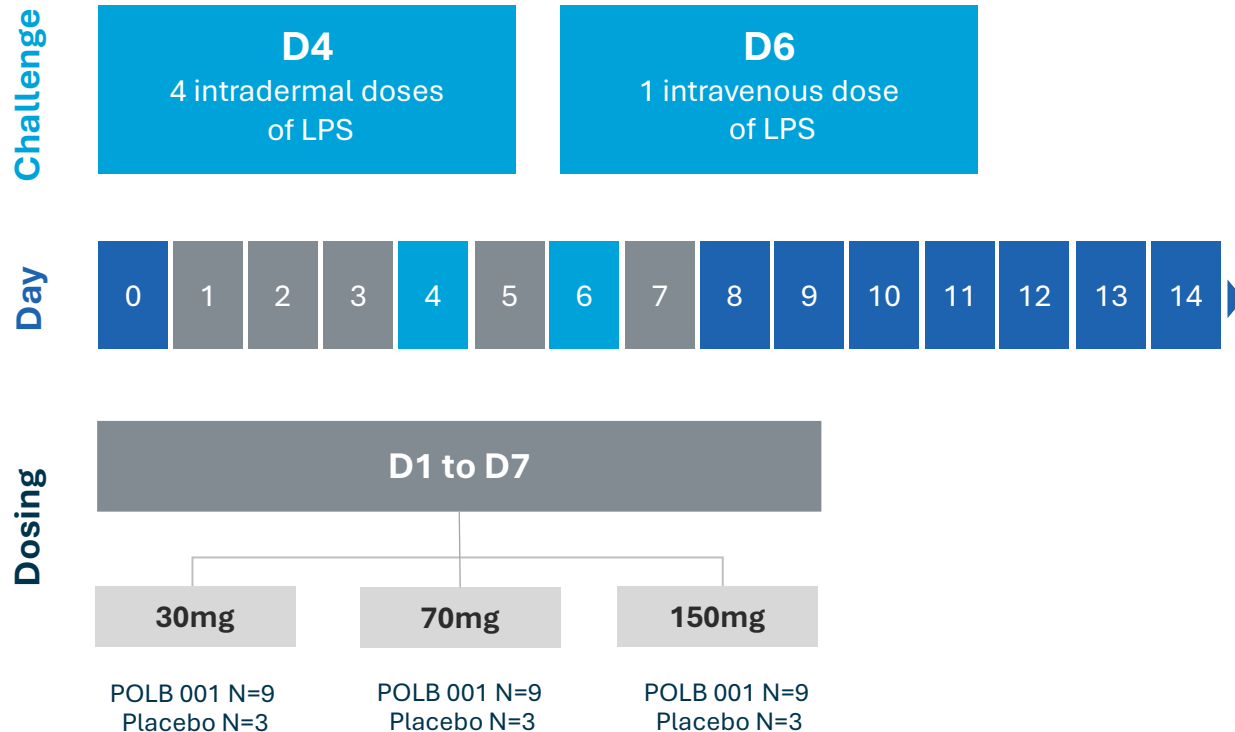


Half-life of 7-14 hours provides adequate exposure and avoids excessive exposure beyond periods of CRS risk

POLB 001: Benefit in Treatment of LPS-Induced Inflammation

Randomized, double-blind, placebo-controlled, inflammatory challenge trial in healthy volunteers

Trial design



Endpoints

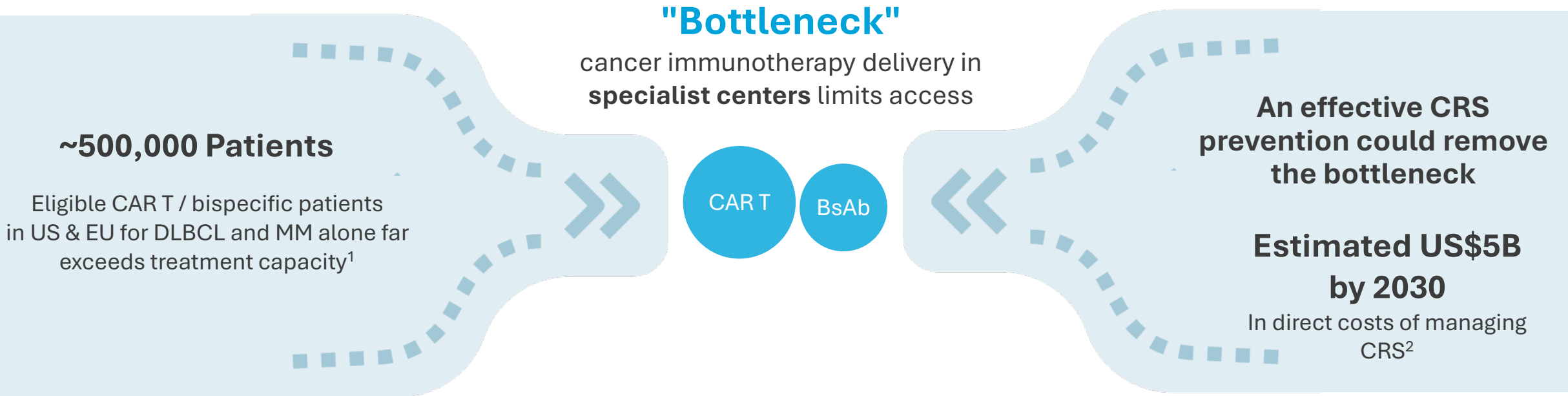
Intravenous LPS challenge

- Bloods (cytokines, vascular markers, CRP)
- *Ex-vivo* LPS response
- Safety & tolerability (inc. vital signs, AE's, ECG, Hematology)

Local inflammatory responses were also measured via intradermal LPS challenge on day 4

POLB 001 is Designed to Address a High Unmet Medical Need

Effective prevention of CRS by POLB 001 may enable broader access to cancer immunotherapies



“ Bispecific antibodies will only be delivered in specialist cancer centers until there is a way to make them safer. POLB 001 could make treatment safe enough to extend them to a much wider patient population.

Professor Gareth Morgan, US

“ The development of an oral CRS preventive therapy will mean no or shorter hospital stays.

French KOL

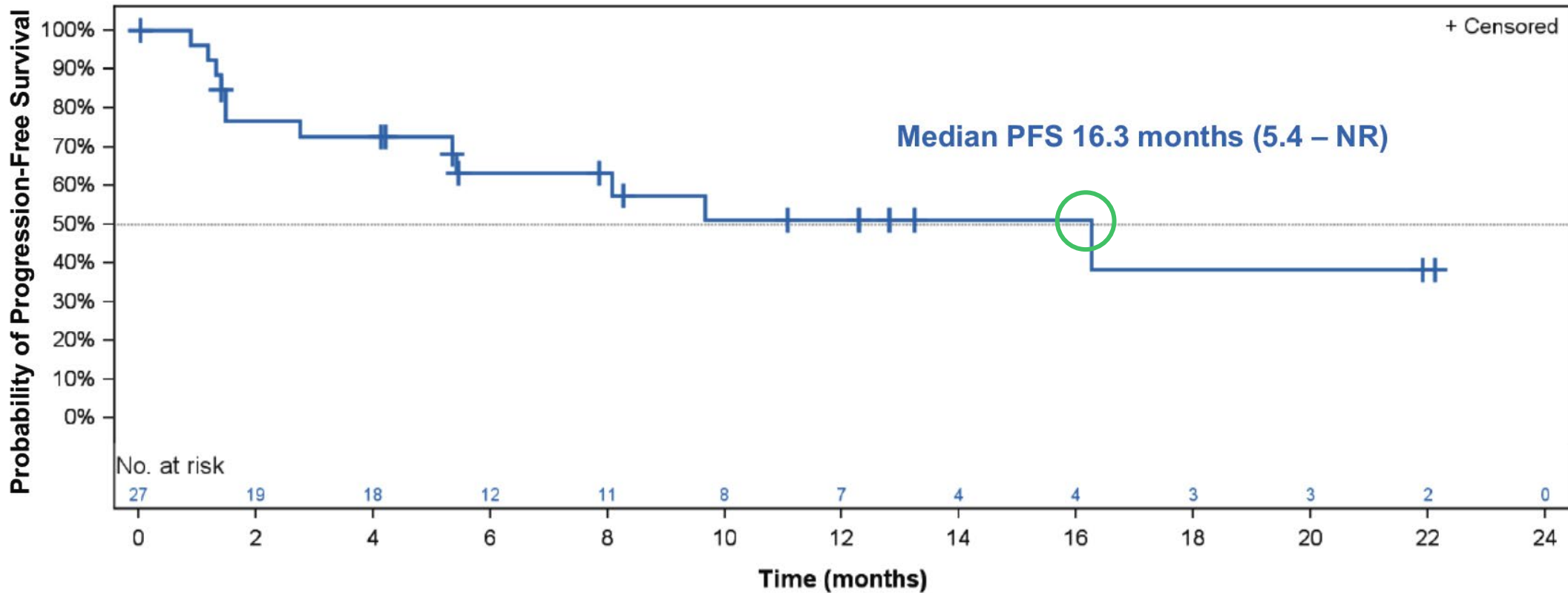
Eseba-vec: Synergistic Activity in Combination with Pembrolizumab

Eseba-vec (HB-200) is a next-generation immunotherapy targeting HPV+ cancers

	Eseba-vec monotherapy in 2L+ CPS ≥ 1 N = 27	Pembro monotherapy in 1L CPS ≥ 20 N = 133)	Eseba-vec+ Pembro in 1L CPS ≥ 20 N = 25 ¹
Overall Response Rate	4%	23%	52%
Complete Response Rate	No CR	8%	16%
% Tumor Shrinkage	33%	Not reported	84%
Disease Control Rate	44%	53%	80%
Median Progression Free Survival	~3.0 mos	3.4 mos	16.3 mos

Eseba-vec Exhibits Promising Preliminary PFS

In 1L CPS \geq 20 HPV+ R/M H&NSCC



Eseba-vec Has Broad Potential Across Multiple HPV16+ Cancers

Up to ~20,000 US patients and ~39,000 patients globally

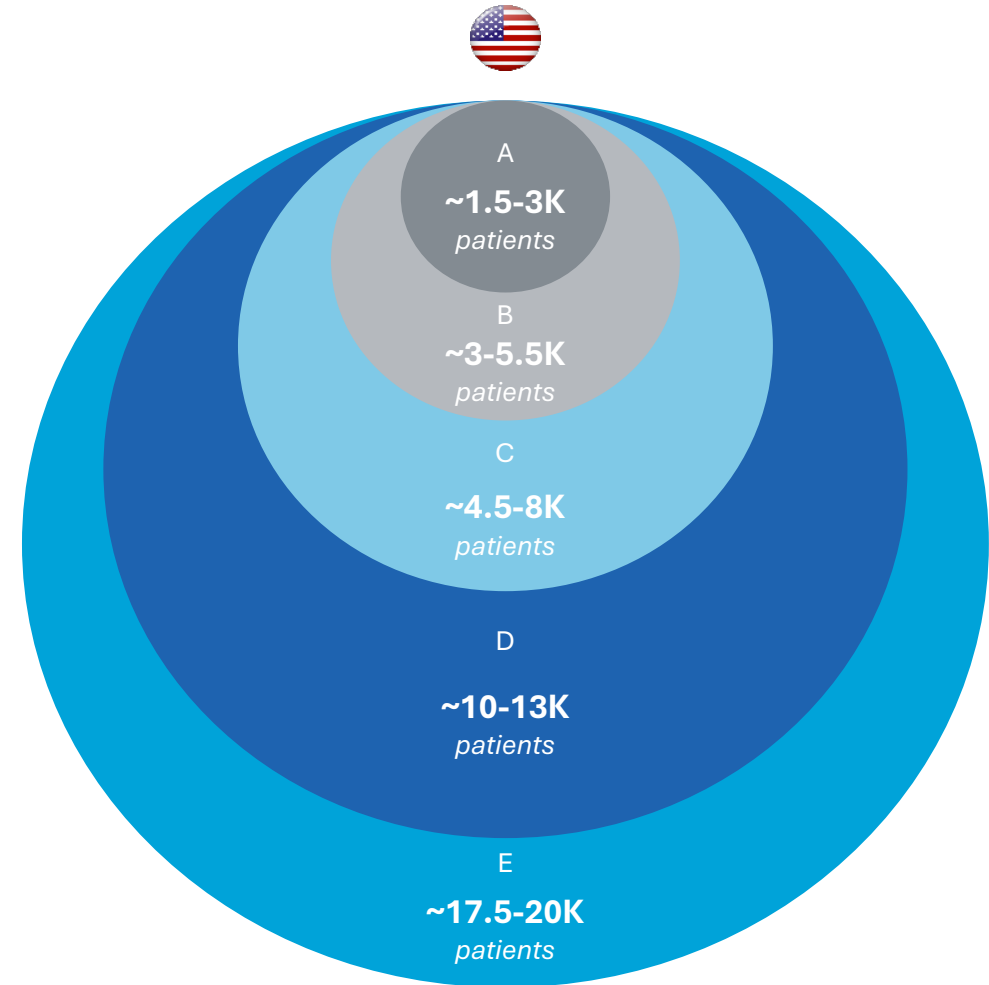
OPC Opportunities

- A** **Immediate OPC Opportunity**
1L HPV16+, CPS \geq 20 R/M OPC
- B** **Expand to CPS 1-19**
1L HPV16+, CPS >1 R/M OPC
- C** **Expand to Neoadjuvant/Adjuvant & 2L+ HPV16+ OPC**

Beyond OPC

- D** **Expand to Recurrent/Metastatic Non-OPC HPV16+ HNSCC**
- E** **Additional 2L+ Anogenital Opportunity HPV16+ Solid Tumors**

*Assuming full expansion beyond initial label (HPV16+ HNSCC and anogenital tumors)



Cumulative Patient Numbers



Special Note Regarding Forward-Looking Statements

This presentation contains “forward-looking statements” within the meaning of the Private Securities Litigation Reform Act of 1995, as amended, including statements regarding HOOKIPA's expectations regarding the terms, benefits, impacts and timing of the proposed combination (the "Proposed Combination") between HOOKIPA and Poolbeg Pharma plc ("Poolbeg") and the proposed private placement, as well as statements regarding any or all of the following (assuming completion of the Proposed Combination and proposed private placement, as applicable): the success, cost and timing of HOOKIPA's product development activities and clinical trials; the timing, scope or likelihood of regulatory filings and approvals, including accelerated approval of HB-200 by the U.S. Food and Drug Administration (“FDA”), and final FDA, European Medicines Agency or other foreign regulatory authority approval of HOOKIPA's current and future product candidates; key milestones for HOOKIPA's product candidates; HOOKIPA's ability to develop and advance its current and future product candidates and programs into, and successfully complete, clinical trials, including for HB-700, POLB-001, HB-500 and HB-200; the potential of HOOKIPA's arenavirus platform to treat additional HPV16+ tumors and its applicability to additional antigens; the expected timing of patient enrollment and dosing in clinical trials, completing clinical trials, and the availability of data from clinical trials; expected revenue from clinical, regulatory and commercial milestones for HOOKIPA's partnered programs, including HB-500 and HB-400; the market opportunity for HOOKIPA's product candidates, if approved, in the indications they seek to treat, including the blockbuster potential of HB-700 and the market opportunity for POLB 001; the potential of POLB 001 to receive orphan drug designation; the potential of POLB 001 as an adjunct therapy for to bispecific and CAR T treatment; HOOKIPA's expected capital needs, sufficiency of resources to achieve anticipated milestones, and cash runway; the potential to develop product candidates in partnerships with third parties; and other statements that are not historical fact. 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