



# Crescent Biopharma Overview

March 2025

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# Crescent Biopharma aims to advance the next wave of innovation in cancer therapy

Crescent's pipeline consists of **potentially best-in-class therapies for the treatment of solid tumors**

 ~\$200 million financing in October 2024 anticipated to fund operations through 2027 <sup>2</sup>	PROGRAM	MOA	DISCOVERY	IND-ENABLING	IND	POTENTIAL INDICATIONS
	<b>CR-001<sup>1</sup></b>	<b>PD-1 x VEGF</b> same cooperative MoA as ivonescimab				4Q25
<b>CR-002</b>	<b>Undisclosed</b> ADC #1 with Topol payload				Mid-2026	Solid tumors
<b>CR-003</b>	<b>Undisclosed</b> ADC #2 with Topol payload					Solid tumors

Crescent is the fifth company launched with assets discovered and developed in-house by Paragon Therapeutics, **an antibody discovery engine** founded by Fairmount Funds in 2021



Prior companies founded **using Paragon's engineered antibody technology** have collectively raised >\$2B and generated significant value



1. Anticipated expiration for filed provisional patent is 2045+

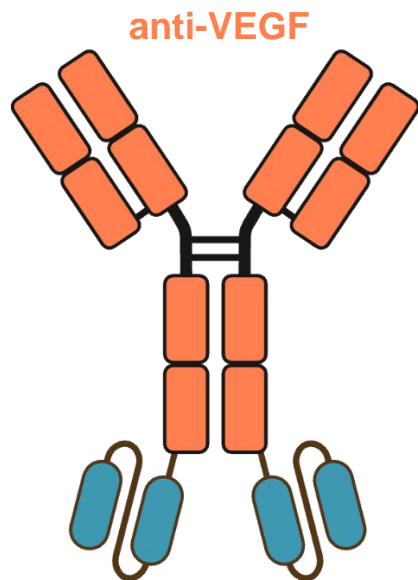
2. Financing scheduled to close immediately prior to the closing of the merger with GlycoMimetics

**Notes:** NSCLC: Non-small cell lung cancer. MoA: Mechanism of action. ADC: Antibody-drug conjugates. Topol: Topoisomerase

# Crescent is advancing three highly impactful oncology programs with best-in-class potential

## CR-001

PD-1 x VEGF cooperative tetravalent bsAb  
Same MoA as ivonescimab



anti-PD-1 scFvs

Designed to **reproduce ivonescimab's established pharmacology**

**Pipeline-in-a-program opportunity** across solid tumor indications

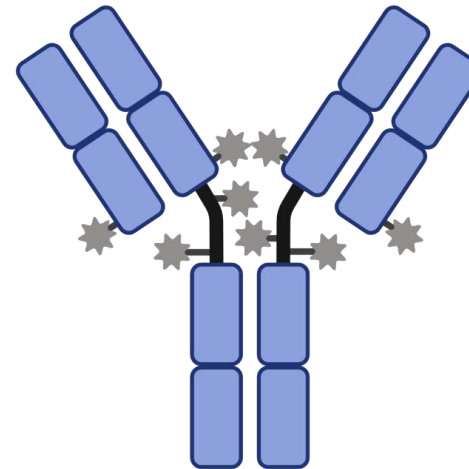
Potential to **move to frontline use in the \$50B+ PD-(L)1 immunotherapy market**

IND expected 4Q25

Interim PoC data expected 2H26

## CR-002 & CR-003

ADCs with topoisomerase inhibitor payloads  
Potentially best-in-class



Two unique, undisclosed targets with **significant potential across solid tumors** as single agents

Each has potential to **synergize with CR-001** in combination studies, further driving clinical efficacy

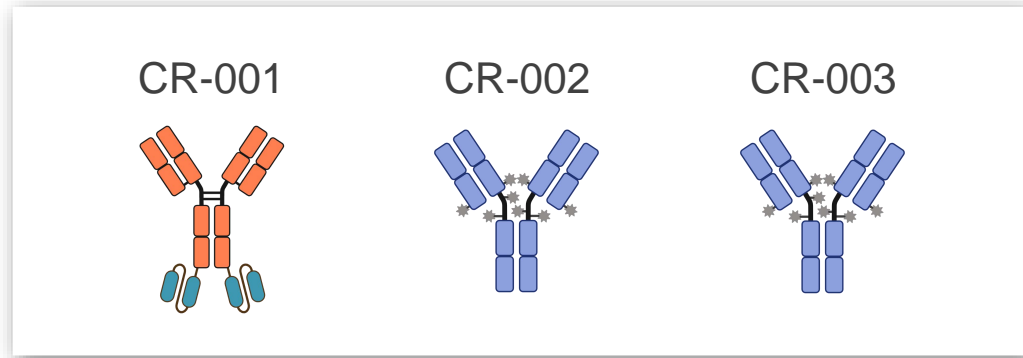
Both utilize the **best-in-modality cytotoxic payload**: topoisomerase inhibitor

CR-002 IND expected mid-26

Interim PoC data expected 2027

# Multiple ways to win: Crescent pipeline enables optionality with differentiating combination therapies

## Optimized Novel Monotherapies

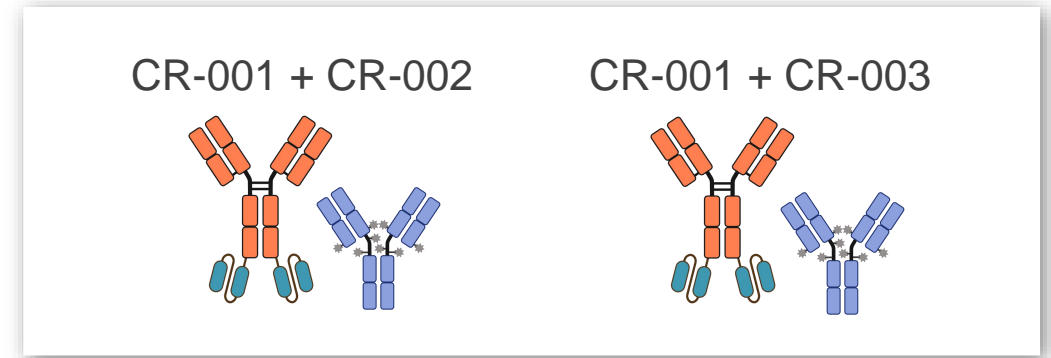


### Engineered for:

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Best-in-class efficacy	Safety
Efficacy across histologies	Stability
Pharmacokinetics	Developability

## Synergistic Combination Approaches



### Selected for:

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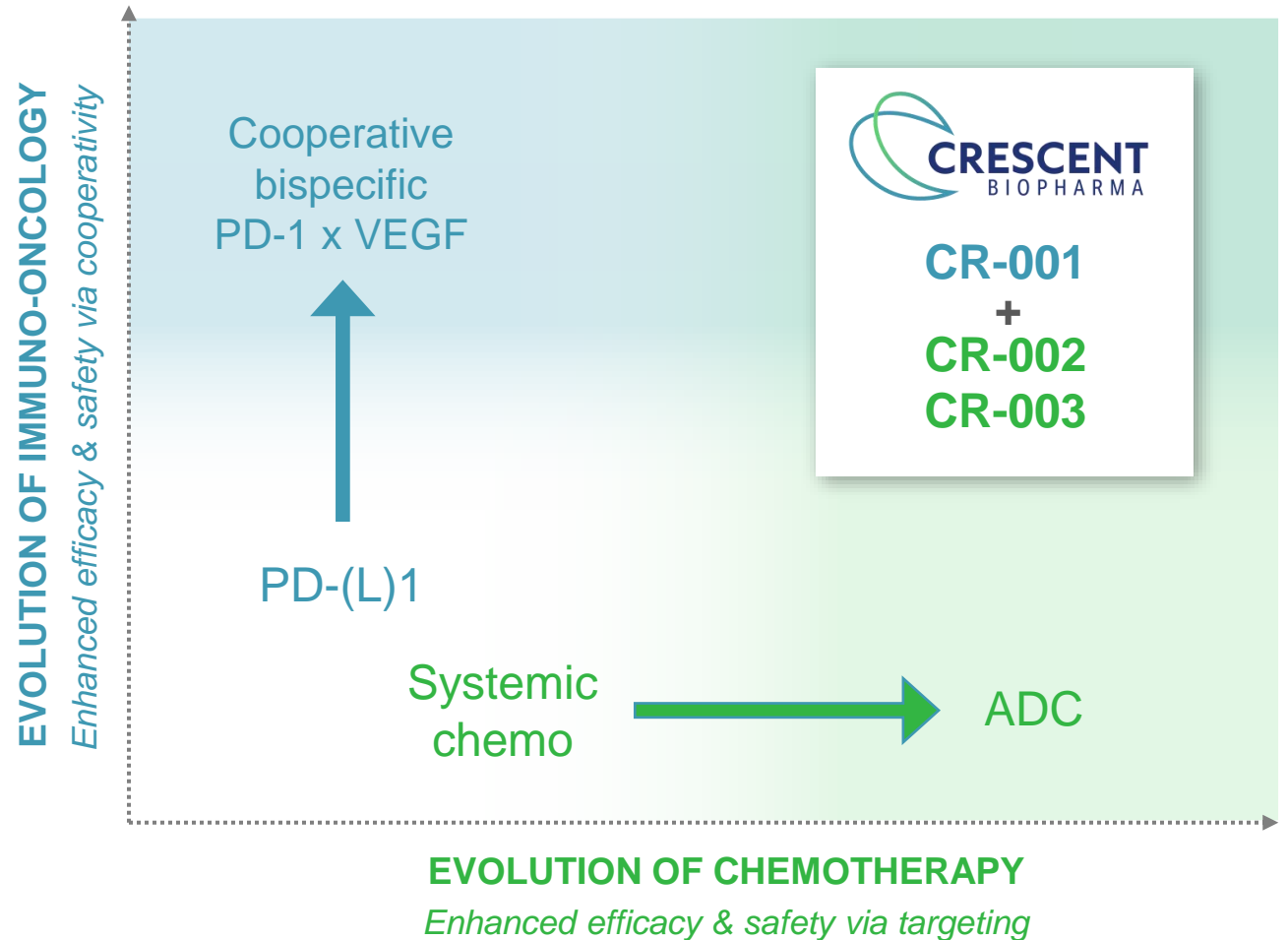
MoA synergy
Efficacy in overlapping histologies
Broad utility in solid tumors

# Crescent leverages three key advances in oncology for next-generation combinations within unique portfolio

## Three revolutions underway in oncology:

- Immuno-oncology is potentially moving from PD-(L)1 to cooperative PD-1 x VEGF
- Chemo is moving from systemic toxins to tumor-targeted toxins via improved ADCs
- Monotherapies are being replaced by synergistic combinations of targeted therapies

**Crescent is developing leading assets in both categories, designed to combine for maximum efficacy in priority indications**



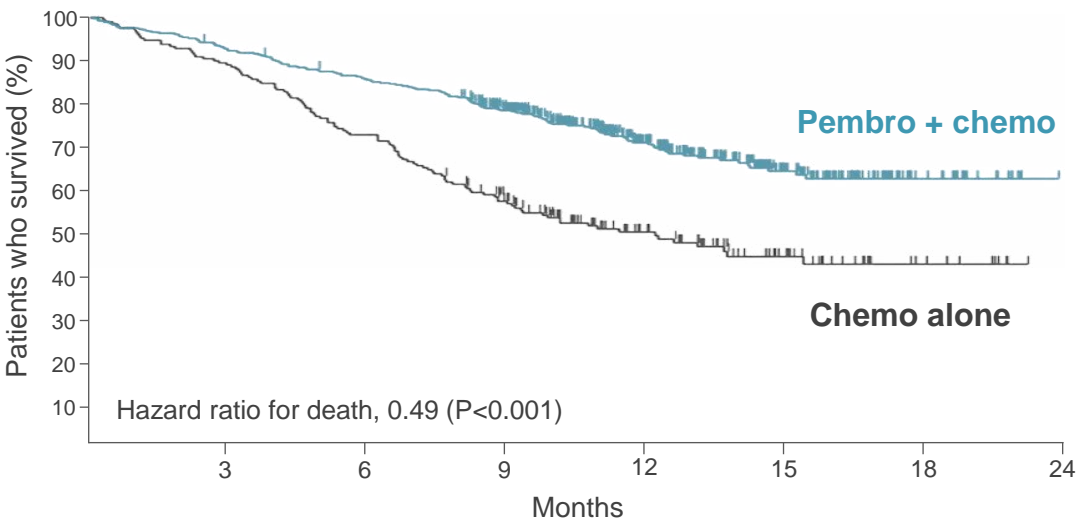
# PD-(L)1-targeted therapies, annualizing \$50B+, have transformed oncology – with Keytruda now the best-selling drug in the world

PD-(L)1 inhibitors have significantly prolonged cancer survival, shifting 1L treatment to immunotherapy

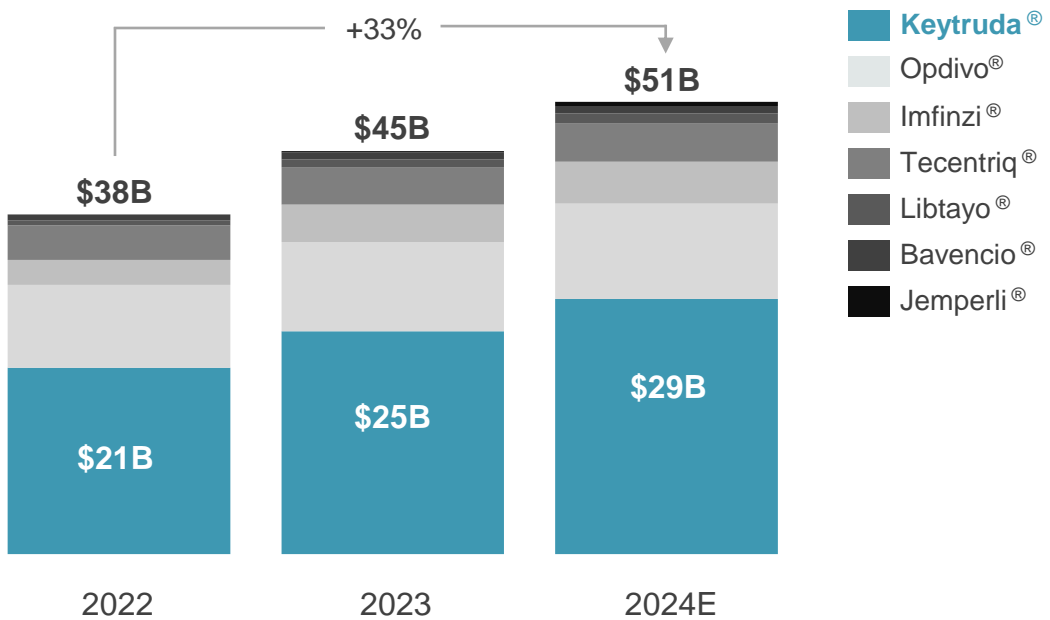
PD-(L)1-targeted therapy is one of the largest drug classes, with Keytruda (pembrolizumab) the dominant player

## Example

In 1L NSQ NSCLC, addition of pembrolizumab to chemo significantly improved mOS (NR vs 11.3 months<sup>1</sup> with HR 0.49)



## Anti-PD-(L)1 global sales



Keytruda alone is approved in 20+ oncology indications with expected revenue of ~\$30B in 2024

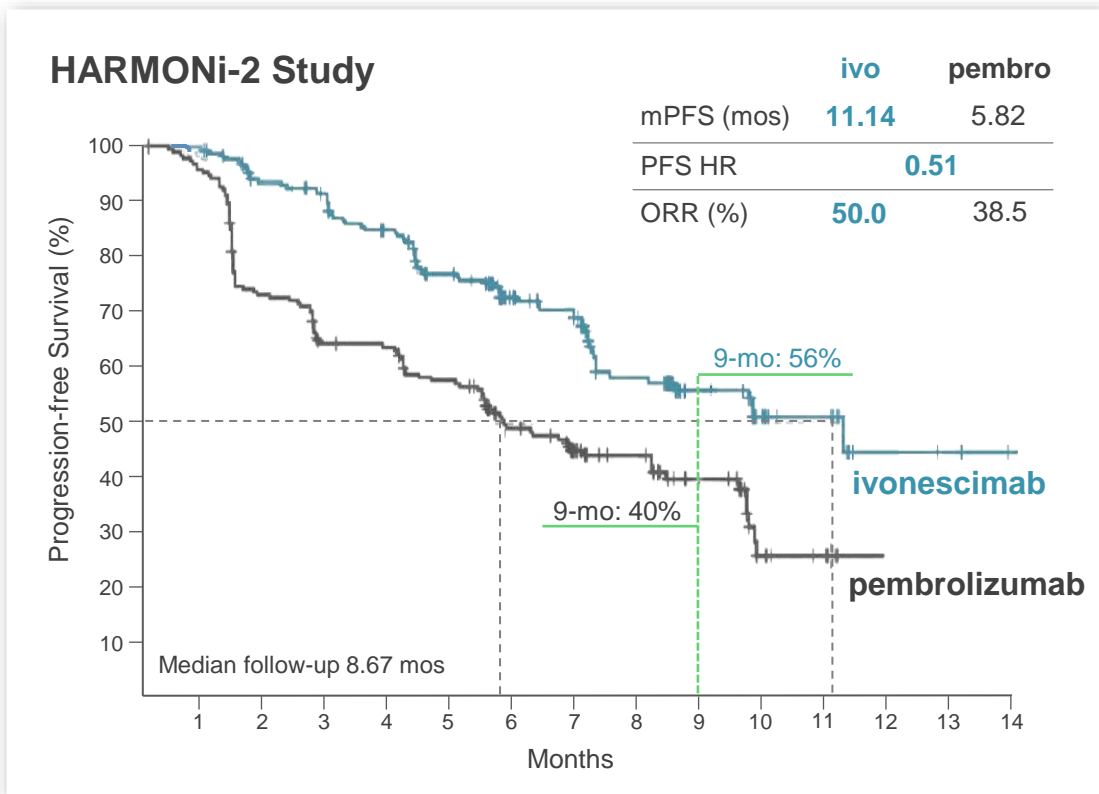


1. 5-year follow-up demonstrated mOS of 22.0 vs 10.6 months.  
 Notes: 1L: First-line. NSQ: Non-squamous. NSCLC: Non-small cell lung cancer. mOS: median overall survival. NR: Not reached. HR: Hazard ratio.  
 Sources: 2018 Gandhi (NEJM); 2023 Garassino (J Clin Oncol); GlobalData; FactSet; Pembrolizumab FDA Label

# Ivonescimab, a cooperative PD-1 x VEGF bispecific, doubled progression-free survival vs. Keytruda in a P3 NSCLC trial

Ivonescimab is the first drug to demonstrate superiority in PFS over pembrolizumab in a randomized Phase 3

Ivonescimab's novel mechanism of action raises the bar on efficacy and safety



## ✓ Broader Efficacy

Ivonescimab demonstrates benefit in patients where anti-PD-(L)1 efficacy has historically been modest (e.g., squamous, PD-(L)1<sup>low</sup>)

	PD-L1 <sup>low</sup> (TPS 1-49%)	PD-L1 <sup>high</sup> (TPS ≥50%)	Non-squamous	Squamous
HR	0.54	0.46	0.54	0.48

## 🛡️ Promising Safety

Ivonescimab had **lower AEs than expected** vs. anti-VEGF monotherapy. This suggests a **differentiated profile** due to cooperativity-driven tissue targeting

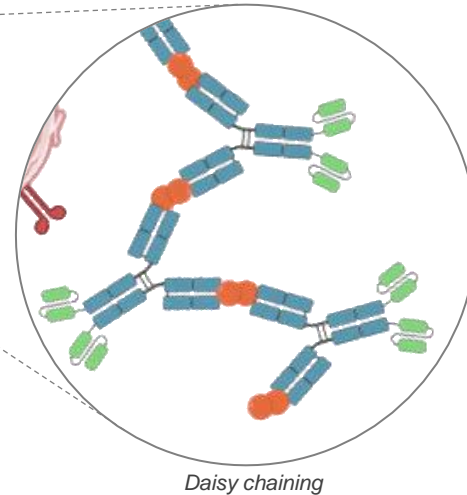
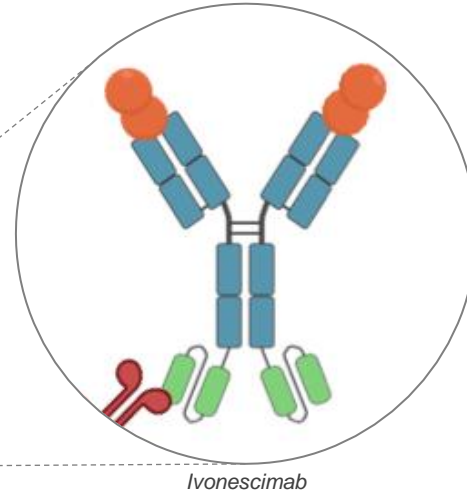
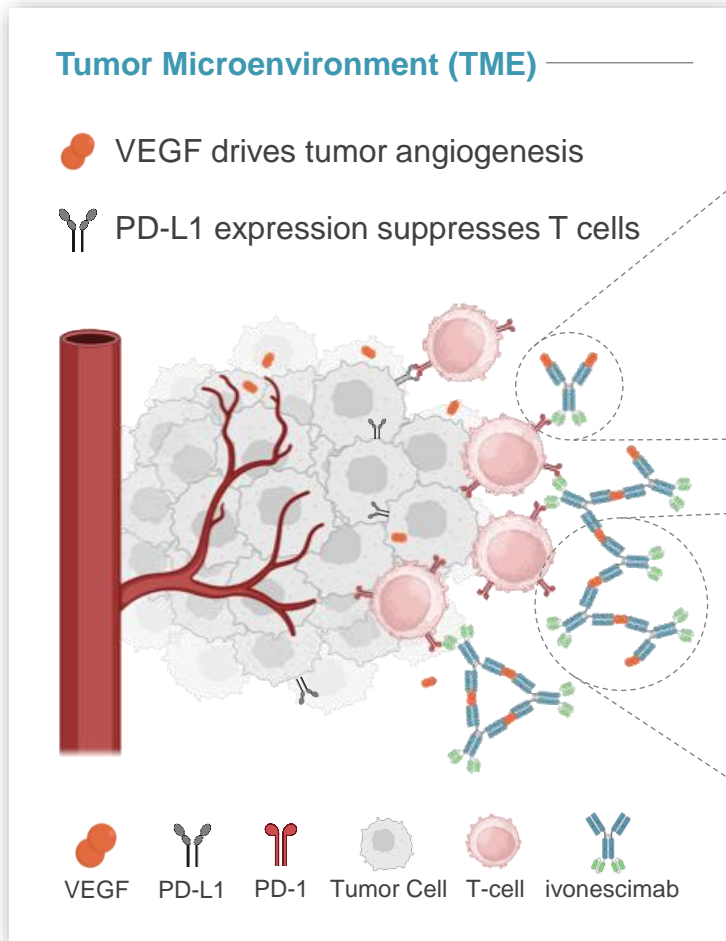
Dual blockade of PD-1 and VEGF through a cooperative bispecific antibody has led to unprecedented clinical results, demonstrating superiority to pembrolizumab and a **\$15B+ market cap for ivonescimab's ex-China sponsor, Summit Therapeutics**



# CR-001

*Cooperative, tetravalent  
PD-1 x VEGF bispecific antibody*

# Ivonescimab's novel, cooperative MoA is hypothesized to drive enhanced anti-tumor activity while maintaining tolerability



## Tumor Targeting

Dual blockade of PD-1 and VEGF through a novel tetraivalent bispecific format with cooperative binding effects has led to **unprecedented clinical results** in third party trials

PD-1 arm concentrates VEGF inhibition in the TME, potentially **sparing healthy tissue and reducing AEs**

## Cooperativity

Ivonescimab's **cooperative binding** blocks PD-1 / PD-L1 interactions *and* inhibits VEGF

VEGF binding to ivonescimab increases affinity to PD-1 and vice versa, enhancing both T-cell activation and VEGF-signaling blockade. This helps explain the **cross-trial outperformance** of ivonescimab vs. an anti-PD-L1 + anti-VEGF combination

PD-1 binding strength (affinity) is increased by >18x in the presence of VEGF

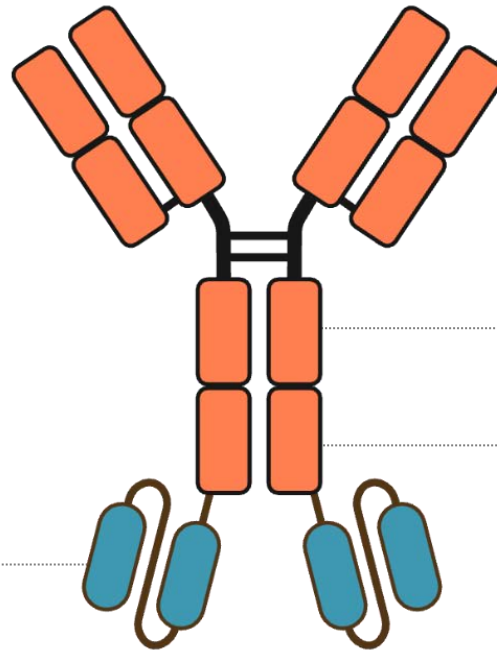
# CR-001 is a highly potent PD-1 x VEGF bsAb designed to recapitulate ivonescimab's cooperative pharmacology

## Same design as ivonescimab

Pairs anti-VEGF IgG & anti-PD-1 scFvs  
Avoids risks of alternative, clinically unprecedented constructs (e.g., VEGF trap, anti-PD-L1 IgG, ADCC)

## Highly potent & stable scFvs

Designed to be the best possible anti-PD-1 epitope / binding domain  
Anti-PD-1s have historically outperformed anti-PD-L1s in meta-analyses of solid tumor studies  
Contains proprietary engineering to enable functional and stable scFvs



## Potential for reduced AEs

Cooperative binding increases anti-VEGF activity in TME, reducing AE risks in healthy tissue  
Identical VEGF potency to preserve safety

## Effector-null human IgG Fc

Equivalent to ivonescimab  
ADCC carries additional AE risk

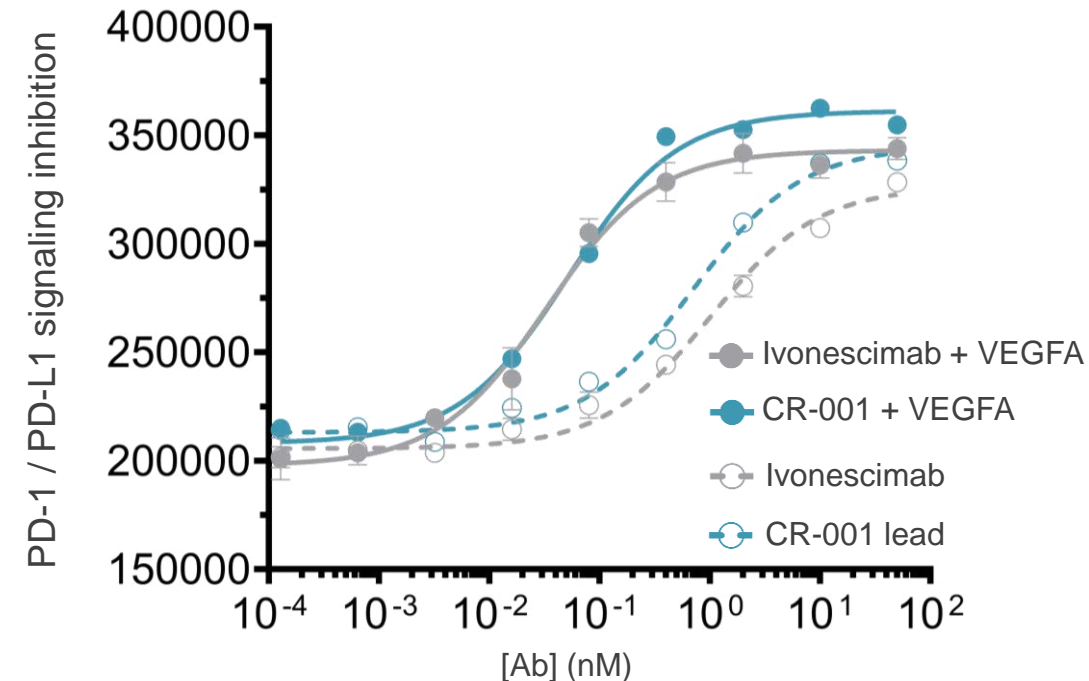
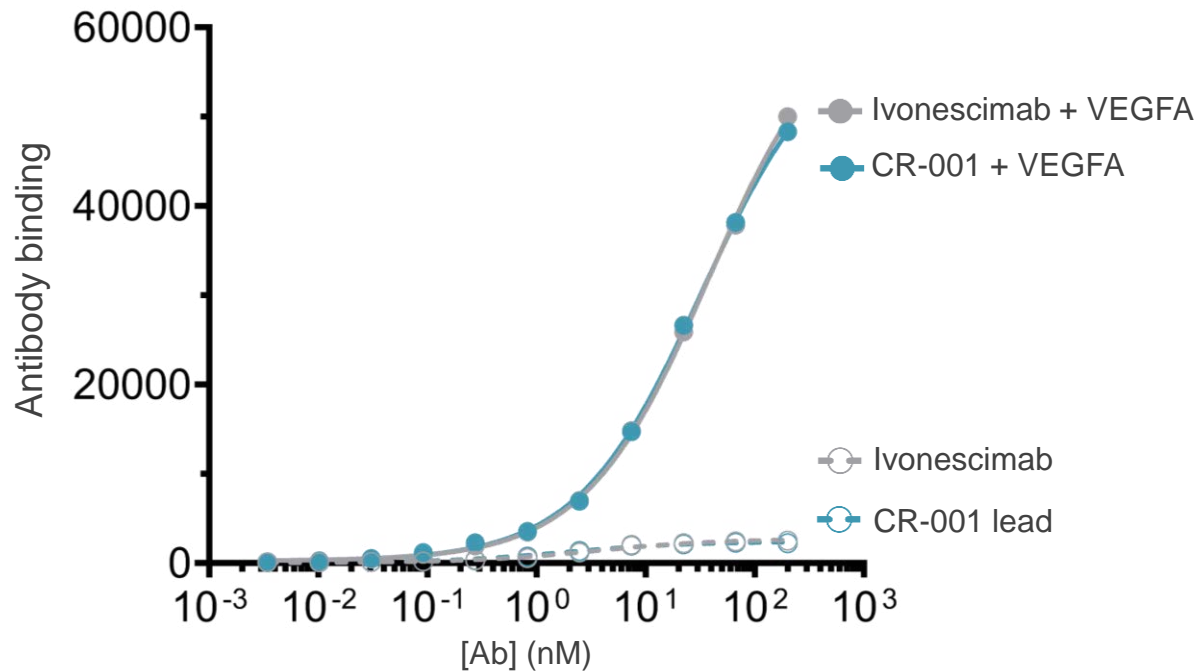
## Designed to match ivonescimab PK

Native FcRn binding to match distribution and elimination of ivonescimab

# CR-001 replicates ivonescimab's cooperative binding effect which leads to cooperative inhibition of PD-1 signaling in presence of VEGF

CR-001, like ivonescimab, **increases PD-1 binding** on PD-1+ Jurkat cells **in the presence of VEGF**...

...leading to **higher potency** in an NFAT reporter assay in **the presence of VEGF**.



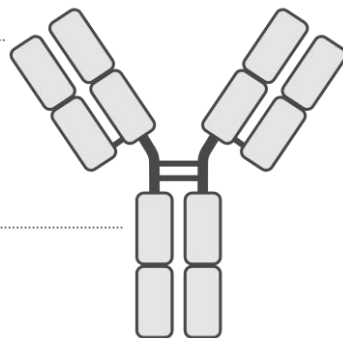
**CR-001 lead demonstrates same cooperative effect as ivonescimab across multiple assays**

# CR-001 engineering replicates ivonescimab function with biophysical properties that maximize flexibility in development

Standard mAbs can be improved with established protein engineering approaches...

CDRs improved via diversification and/or affinity maturation to maximize potency

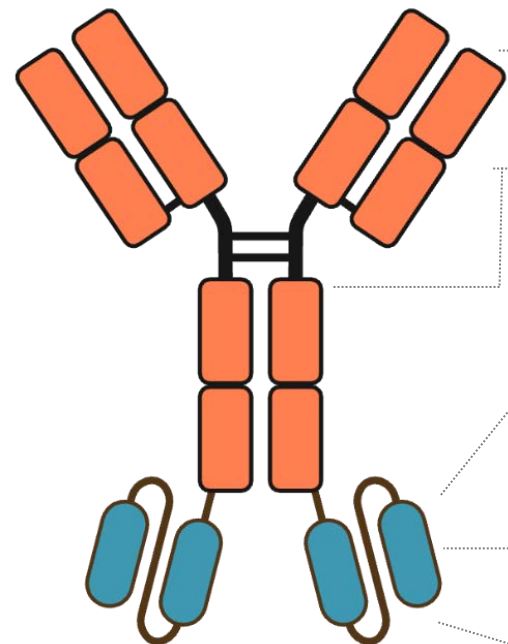
Fc engineering tunes ADCC, CDC, half-life, etc.



...but ensuring cooperative effect, stability, and developability of a tetravalent PD-(L)1 x VEGF bispecific antibody is more difficult

## Ivonescimab's unique structure and geometry – and resulting cooperative function – is challenging to replicate

Alternative constructs risk not reproducing ivonescimab's superior efficacy and safety in clinical practice



IgG format bound to VEGF dimer **required to daisy chain**; different potency may alter chaining kinetics and VEGF trap geometry does *not* work

**Fc silencing** helps reduce risk of AEs

Leading anti-PD-1s are unstable and aggregate in scFv format, requiring significant engineering; **CR-001 maintains >95% monomer at 150mg/mL**

Bispecific antibodies often cannot achieve high concentrations with low enough viscosity to maximize development optionality; **CR-001 is low viscosity (<16 cP) up to 150mg/mL**

**CR-001 has novel composition of matter IP** related to proprietary, stabilized scFvs

# CR-001 has potential to transform SoC across a multitude of oncology indications, with numerous first-in-class opportunities



## SOFT TISSUE

- Alveolar soft part sarcoma
- Soft tissue sarcoma



## HEMATOLOGICAL

- Classical Hodgkin lymphoma
- Primary mediastinal large B-cell lymphoma (PMBCL)



## REPRODUCTIVE

- Cervical
- Endometrial
- Fallopian tube
- Ovarian (epithelial)
- ★ ● Triple negative breast cancer (TNBC)
- Urothelial



## BRAIN

- Glioblastoma



## HEAD & NECK

- Head & neck squamous cell carcinoma (HNSCC)
- Nasopharyngeal
- Thyroid



## LIVER & BILIARY

- Biliary tract
- Hepatocellular carcinoma (HCC)



## KIDNEY

- Renal cell carcinoma (RCC)



## CHEST/THORACIC

- Esophageal
- ★ ● EGFRm non-small cell lung cancer (NSCLC)
- ★ ● Non-squamous NSCLC
- ★ ● Squamous NSCLC
- ★ ● Small cell lung cancer (SCLC)
- Pleural mesothelioma



## GASTROINTESTINAL

- Colorectal (all comers)
- Colorectal (MSI-H / dMMR)
- Gastric / Gastroesophageal junction (GEJ)
- Primary peritoneal

- Anti-VEGF approvals
- Anti-PD-(L)1 approvals
- Anti-VEGF and anti-PD(L)-1 approvals
- ★ Ongoing / announced global study from Summit or BioNTech



## TISSUE-AGNOSTIC

- High microsatellite instability (MSI-H) / deficient DNA mismatch repair (dMMR)
- High tumor mutational burden (TMB-H)



## SKIN

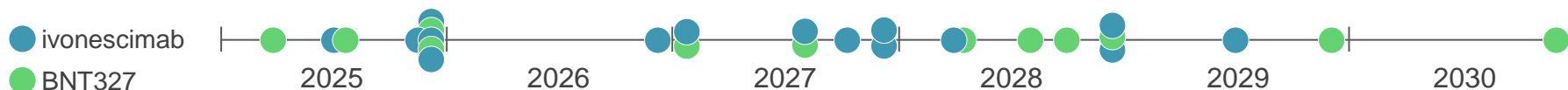
- Basal cell carcinoma
- Cutaneous squamous cell carcinoma
- Melanoma
- Merkel cell carcinoma



# Development programs across key late-stage competitors include numerous P3s with PFS & OS readouts, paving the way for CR-001

Leading PD-(L)1 x VEGF programs, with **similar expected cooperativity to CR-001**, will generate **Phase 3 PFS & OS catalysts for years to come**

Pivotal readouts expected:



Program	Company	Study	Phase	Indication	Population	Combo	Comparator	Data Expected
ivo PD-1 x VEGF	Akesobio	HARMONi-A	3 (China)	mNSCLC	2L EGFRm NSQ	chemo	chemo	OS late 2025
		HARMONi-2	3 (China)	mNSCLC	1L PDL1+ TPS ≥ 1%	none	anti-PD-1	OS mid 2025
		AK112-306	3 (China)	mNSCLC	1L squamous	chemo	anti-PD-1 + chemo	OS late 2025
		AK117-302	3 (China)	HNSCC	1L R/M PD-L1+ CPS ≥ 1	anti-CD47	anti-PD-1	PFS & OS 2027
		AK112-308	3 (China)	TNBC	1L mTNBC	chemo	chemo	PFS 2026, OS 2028
		AK112-309	3 (China)	BTC	1L A/M BTC ECOG 0-1	chemo	anti-PD-L1 + chemo	PFS & OS 2027
BNT327 PD-L1 x VEGF	BIONTECH	HARMONi	3 (global)	mNSCLC	2L EGFRm NSQ	chemo	chemo	PFS & OS 2025
		HARMONi-3	3 (global)	mNSCLC	1L SQ & NSQ	chemo	anti-PD-1 + chemo	PFS & OS 2027-8
		HARMONi-7	3 (global)	mNSCLC	1L PD-L1+ TPS > 50%	none	anti-PD-1	PFS & OS 2028-9
		PM8002-BC010C	2/3 (China)	mNSCLC	2L EGFRm NSQ	chemo	chemo	PFS & OS 2025
		BNT327-06*	2/3 (global)	mNSCLC	1L	chemo	anti-PD-1 + chemo	PFS & OS 2029-30
		[announced]*	2/3 (TBA)	NSCLC	1L	[TBA]	[TBA]	[TBA]
		PM8002-C013C	3 (China)	TNBC	1L	chemo	chemo	PFS & OS 2027-8
BNT327-03	3 (global)	SCLC	1L ES-SCLC	chemo	anti-PD-L1 + chemo	PFS & OS 2028		
PM8002-BC011C	2/3 (China)	SCLC	1L ES-SCLC	chemo	PD-L1 + chemo	PFS & OS late 2025		
PM8002-C014C	3 (China)	SCLC	2L	chemo	chemo	PFS & OS 2027-8		

# Parallel clinical development paths offer potential for both first-in-class and lower risk opportunities for CR-001

## First-in-class opportunities

Focus on potential first-in-class opportunities with **rapid path to market** (i.e., efficient development strategy, **anticipated high likelihood of PFS and OS success**)

Numerous indications with **clinically meaningful anti-PD-(L)1 +/- VEGF efficacy** and potential to combine with chemo / orthogonal MoAs



*Illustrative*

## Fast-follower in clinically validated indications

Plan to **rapidly follow ivonescimab** in indications where clinical validation vs. anti-PD-(L)1 is highly differentiating

High conviction **CR-001 can replicate ivonescimab's efficacy** given similar construct and equivalent MoA



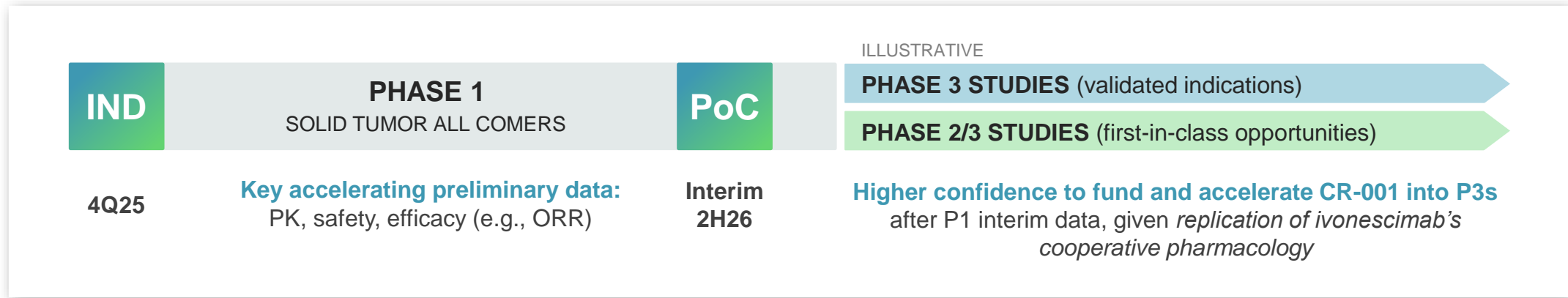
*Potential indications based on ongoing Phase 3 trials*

TWO PARALLEL DEVELOPMENT PATHS FOR CR-001



# CR-001 Phase 1 data offer potential for early acceleration – a rarity for a solid tumor oncology program

Phase 1 interim proof-of-concept readout is a potentially significant value-generating event for CR-001



Preliminary data from early Phase 1 cohorts provides substantial **validation of program** because CR-001's structural design and preclinical data are similar to those of ivonescimab

Early Phase 1 data, as single agent and in combination with SoC, **enables rapid late-stage development** in multiple solid tumor types, unlocking broad first-in-class and fast-follower opportunities

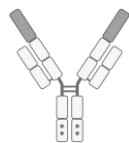
CR-001 is markedly **differentiated from novel constructs disconnected from ivonescimab's MoA**; alternative formats may require significantly more patients worth of safety and efficacy data in tumor-specific expansion cohorts and/or Phase 2s to establish conviction before initiating Phase 3s

High conviction in CR-001's clinical profile can be reached in **~9-12 months** from Phase 1 initiation, offering potential for **significant early value inflection**

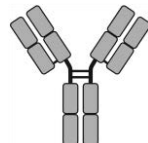
# Only five known constructs with potential to exhibit ivonescimab-like cooperative pharmacology

	Anti-PD-1 scFv-based			Anti-PD-1 VHH-based	Anti-PD-L1 VHH-based
<b>Program</b>	CR-001	ivonescimab	CTX-10726	LM-299	BNT327 / PM8002
<b>Company</b>					
<b>Stage</b>	Preclinical	Phase 3 (Global)	Preclinical	Phase 1/2 (China)	Phase 3 (Global)
<b>Anti-VEGF IgG</b>	Bevacizumab	Bevacizumab	Bevacizumab	Bevacizumab	Bevacizumab
<b>Anti-PD-(L)1</b>	Anti-PD-1 scFvs	Penpulimab scFvs	Anti-PD-1 scFvs	Novel anti-PD-1 VHHs	Novel anti-PD-L1 VHHs
<b>Fc function</b>	Fc null, to avoid potential AEs	Fc null, to avoid potential AEs	Silenced Fc-γR binding	Fc null, to avoid potential AEs	Fc null, to avoid potential AEs
<b>Cooperative pharmacology</b>	✓	✓	No preclinical data disclosed	Expected (not disclosed); unclear impact of VHH structure	Expected (not disclosed); unclear impact of PD-L1 VHH

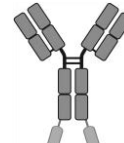
## Examples of alternative constructs



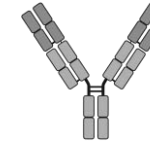
- Anti-PD-L1 IgG with enhanced ADCC
- VEGF trap



- Anti-PD-1 mAb with off-target VEGFR2 binding through same variable domains



- Anti-PD-1 IgG
- Novel anti-VEGF VHHs
- Inverted format



- Bevacizumab
- Anti-PD-1 Fabs
- PD-1 domains attached to IgG N-terminus instead of C-terminus

# CR-001 preclinical data reproduce ivonescimab's breakthrough pharmacology & are rapidly advancing to generate significant value



**Unprecedented third-party data validate PD-1 x VEGF cooperativity**

*Ivonescimab significantly improved PFS versus pembrolizumab in Phase 3 in 1L NSCLC – the first therapy to do so head-to-head*



**Transformative MoA for \$50B+ market**

*Poised to transform NSCLC standard of care, with broad application across \$50B+ anti-PD-(L)1 market*



**CR-001's proprietary engineering is designed to replicate ivonescimab**

*CR-001 is a highly potent PD-1 x VEGF bsAb reproducing cooperative binding qualities critical to ivonescimab*



**Promising pipeline of next-gen ADCs**

*CR-002 and CR-003 offer complementary development opportunities for CR-001*

# CR-002 & CR-003

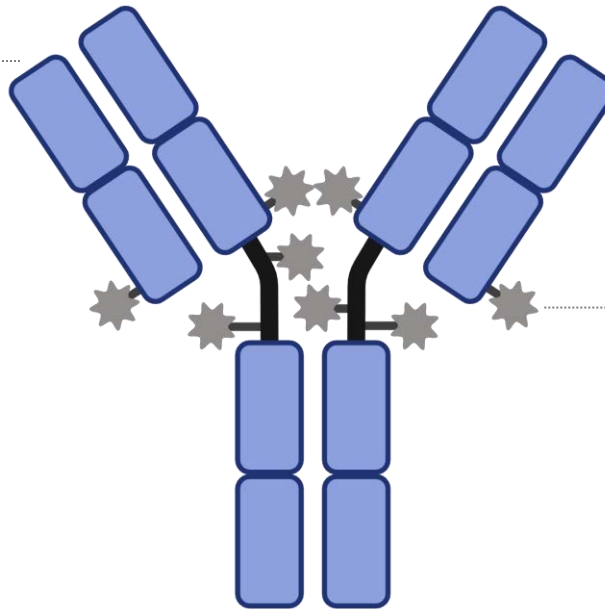
*Topoisomerase inhibitor ADCs  
against validated targets*

# CR-002 and CR-003 are potentially best-in-class topoisomerase inhibitor ADCs, with applicability across solid tumors

## Validated, undisclosed solid tumor ADC targets

Targets for CR-002 and CR-003 to be disclosed as programs approach IND

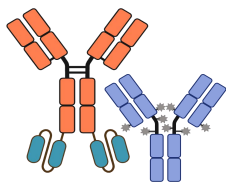
Each unique target has **potential in multiple solid tumor indications**



## Best-in-modality topoisomerase inhibitor payloads

Topoisomerase inhibitor payloads have consistently demonstrated **superior efficacy and safety** over microtubule inhibitor payloads

Each ADC is expected to have **bystander-killing effect**



## Potential to synergize with CR-001 and other immunotherapies

Each ADC can be leveraged in **combination studies** in solid tumors

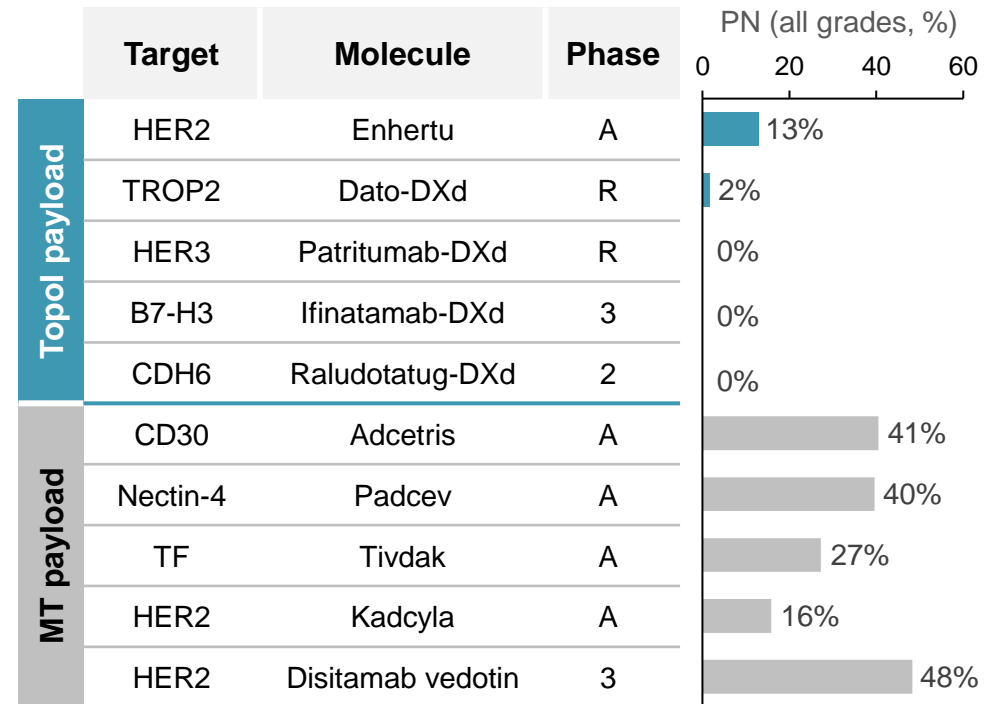
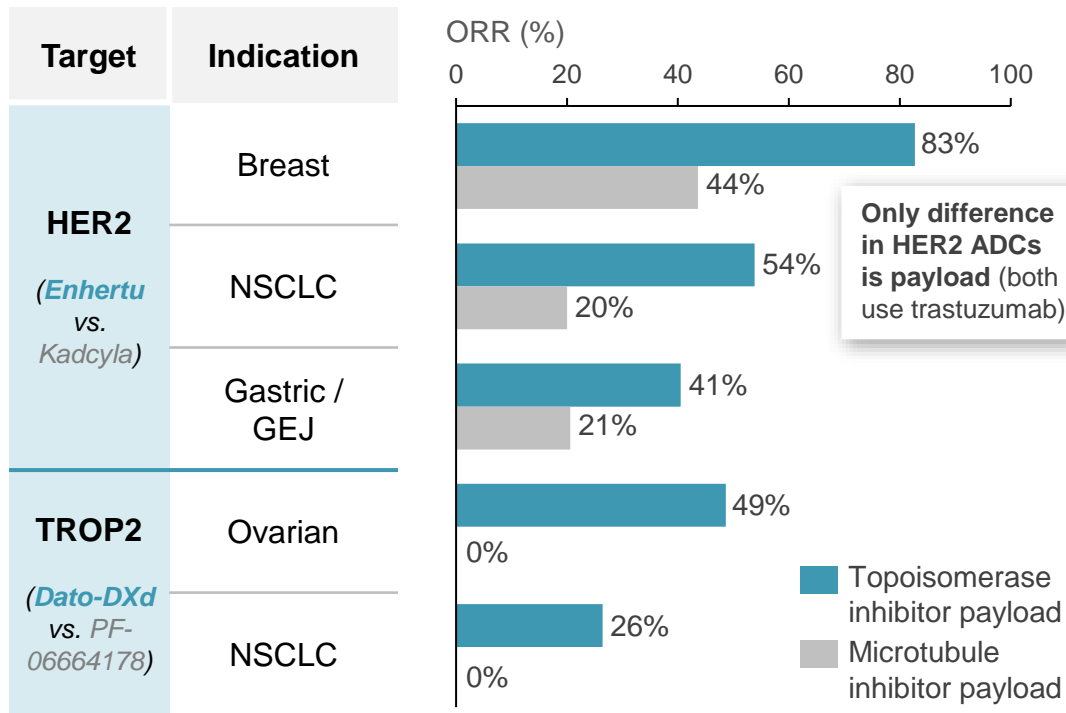
Multiple indications with ongoing PD-(L)1 x VEGF bispecific development and *separate* development of ADCs **accelerate clinical path for combinations**

# ADCs with topoisomerase inhibitor payloads have demonstrated best-in-modality efficacy and safety

CROSS-TRIAL COMPARISONS

Topol payload-based ADCs have **demonstrated superior ORR** vs. microtubule inhibitor-based ADCs in cross-trial comparisons...

... and have shown much **lower rates of peripheral neuropathy**, a critical AE that can **drive dose reductions & discontinuations**



CR-002 and CR-003 utilize the **best-in-ADC payload in their potentially best-in-class profiles**

**Corporate**

# Rapidly growing leadership team with deep experience building the next generation of biotechnology companies

## Executive Team



**Jonathan Violin, PhD**  
Interim CEO



**Chris Doughty**  
Chief Business Officer



**Ryan Lynch**  
Chief Accounting Officer



**Barbara Bispham**  
General Counsel



**Wenjie Cheng, PhD**  
SVP Technical Operations

## Board of Directors



**Peter Harwin**  
Chair



**Alex Balcom**



**Susan Moran, MD**

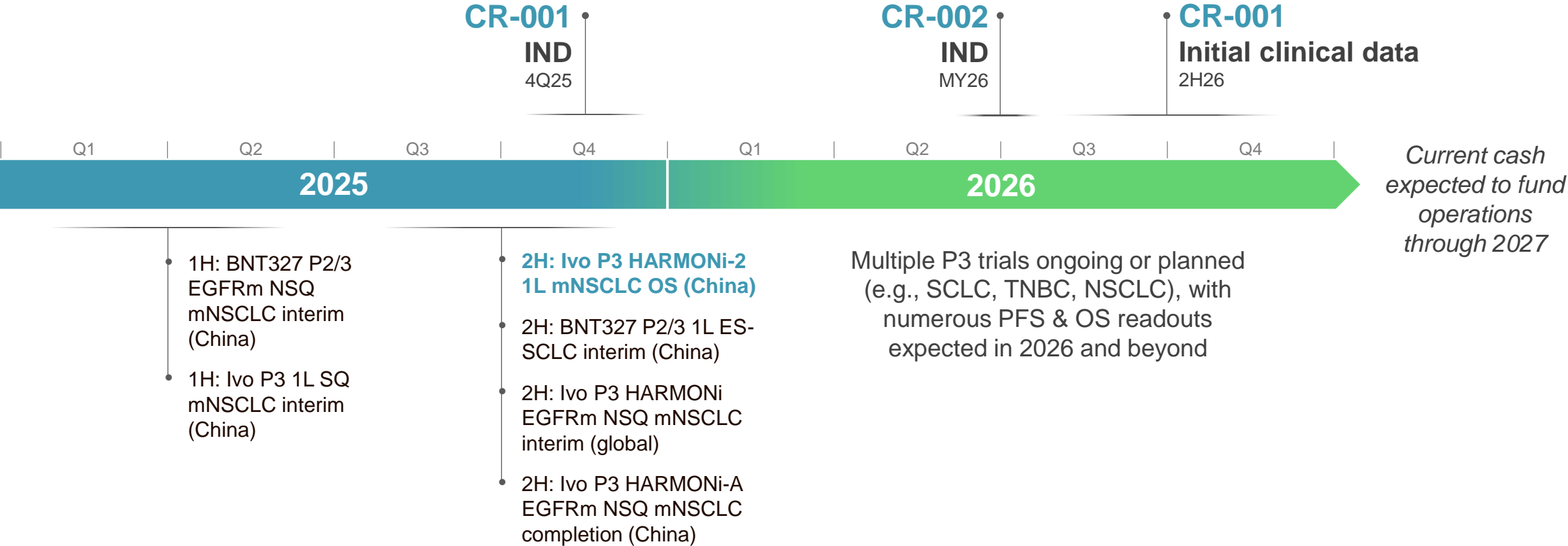


**Jonathan Violin, PhD**







# Financing expected to fund Crescent programs through key anticipated value-generating catalysts



# Estimated capitalization following close of transactions

		Shares on an as-converted basis	Expected ownership of the combined company
	Shares of common stock outstanding	64,532,953	<b>3.1%</b>
	Shares of common stock outstanding	105,137,814	<b>96.9%</b>
	Series A shares	298,298,000	
Pre-closing financing	Shares of common stock	1,339,680,730	
	Pre-funded warrants	273,643,080	
Estimated total shares of common stock of the combined company post-closing		2,081,292,577	



**Thank you**