



# Crescent Biopharma Overview

October 2024

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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.

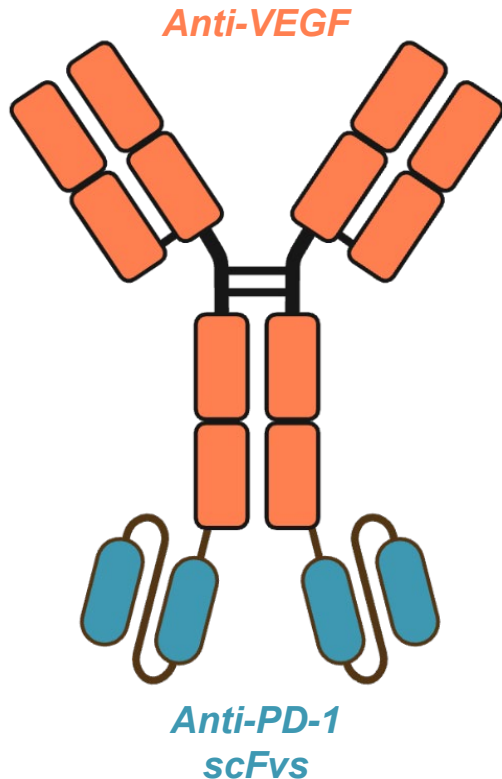
- Crescent is the **fifth company** launched with assets **discovered and in-development in-house** by Paragon Therapeutics, a leading biotech incubator founded by Fairmount Funds in 2021.
- Prior companies founded with Paragon assets have **collectively raised >\$2B and generated significant value.**

Program	MoA	Stage			Potential Indications
		Discovery	IND-enabling	Clinical	
CR-001 <sup>1</sup>	PD-1 x VEGF <i>(same cooperative MoA as ivonescimab)</i>				YE25 / 1Q26 NSCLC, other solid tumors
CR-002	Undisclosed #1 <i>(ADC, Topol payload)</i>				Mid-26 Solid tumors
CR-003	Undisclosed #2 <i>(ADC, Topol payload)</i>				Solid tumors

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

## CR-001

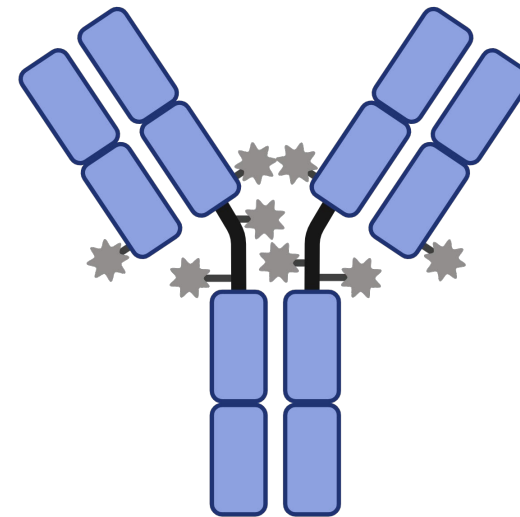
*PD-1 x VEGF cooperative tetravalent bsAb;  
same MoA as ivonescimab*



- Designed to **reproduce ivonescimab's established pharmacology**.
- **Pipeline in a program opportunity** across solid tumor indications, with potential to **move to frontline use in the \$50B+ PD-(L)1 immunotherapy market**.
- **IND expected YE25 / 1Q26.**

## CR-002 & CR-003

*ADCs with topoisomerase inhibitor payloads;  
potentially best-in-class*



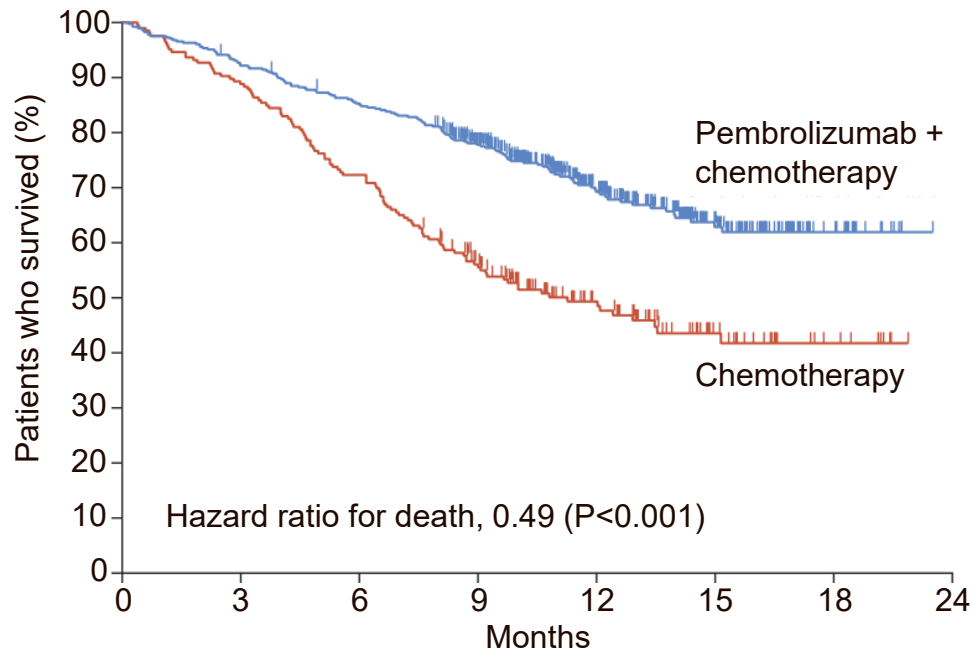
- Two unique, undisclosed targets with **significant potential across solid tumors** as single agents.
- Each has potential to **synergize with CR-001** in combination studies, further driving clinical efficacy.
- Both utilize the **best-in-modality cytotoxic payload: topoisomerase inhibitor**.
- **CR-002 IND expected mid-26.**

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

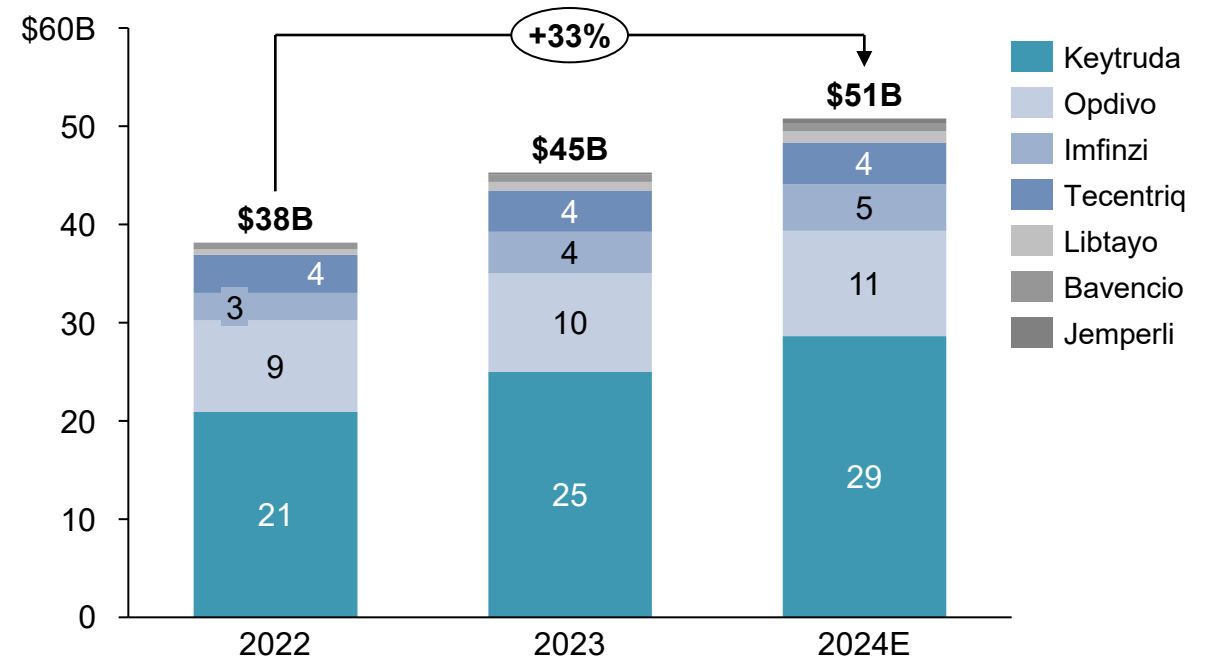
PD-(L)1 inhibitors have **significantly prolonged survival, shifting 1L treatment to immunotherapy**

PD-(L)1-targeted therapies are **one of the largest drug classes, with Keytruda (pembrolizumab) the dominant player**

- For example, in 1L NSQ NSCLC, addition of pembrolizumab to chemo **significantly improved mOS (NR vs 11.3 months<sup>1</sup> with a HR of 0.49).**



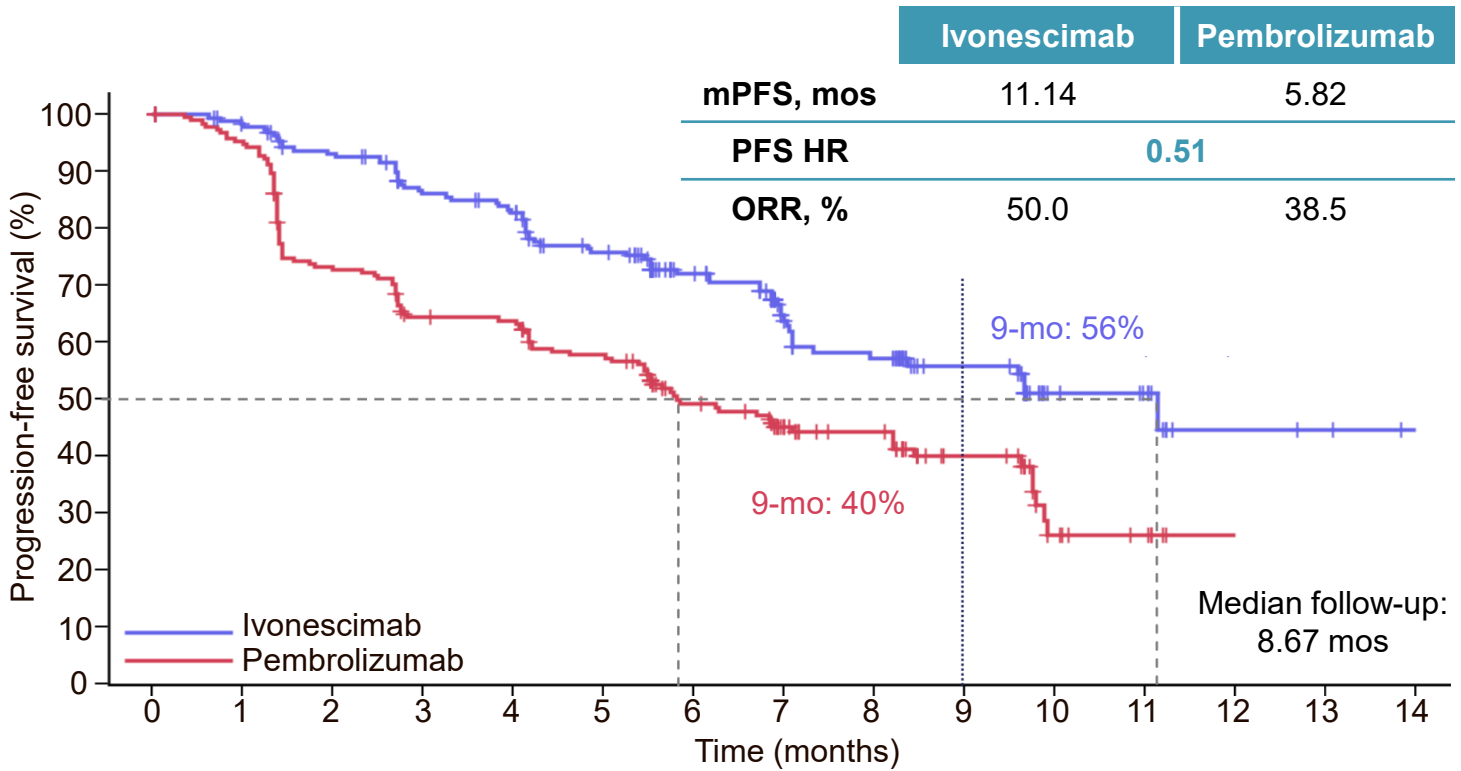
anti-PD-(L)1 global sales



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

# 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 MoA **raises the bar on efficacy and safety**

**1 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

**2 Promising safety:** Ivonescimab had **lower AEs than expected** versus anti-VEGF monotherapy. This suggests a **differentiated profile** driven by 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 ivo's ex-China sponsor, Summit Therapeutics.**



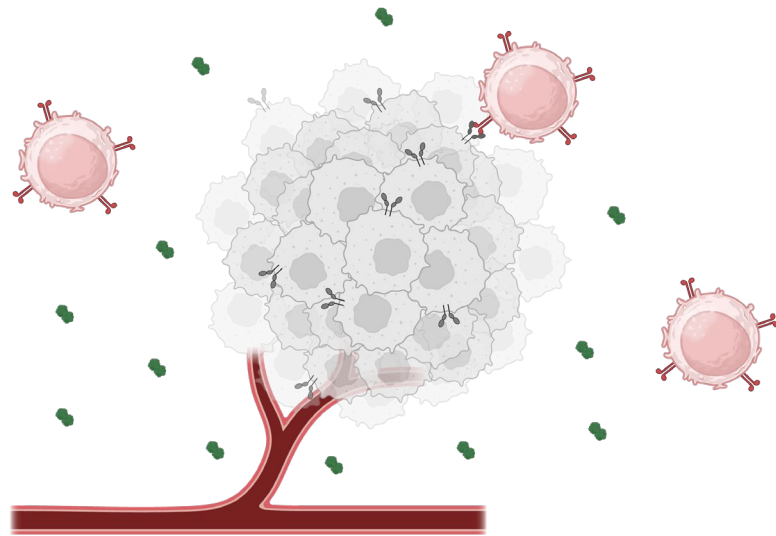
Notes: HR: hazard ratio. PFS: progression-free survival. AE: adverse event. NSQ: Non-squamous SQ: Squamous. Akeso has licensed ivonescimab to Summit in North America, South America, Europe, Africa, Middle East, and Japan. Akeso maintains rights in Asia (ex-Japan / Middle East) and in Oceania. Sources: 2024 Zhou (WCLC Presentation on HARMONI-2); Summit Therapeutics; 2018 Paz-Area (NEJM); 2019 Mok (Lancet); 2022 De Castro Jr (J Clin Oncol); Avastin Label

# CR-001

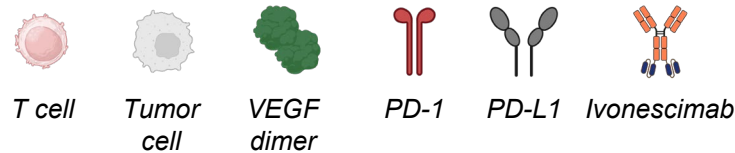
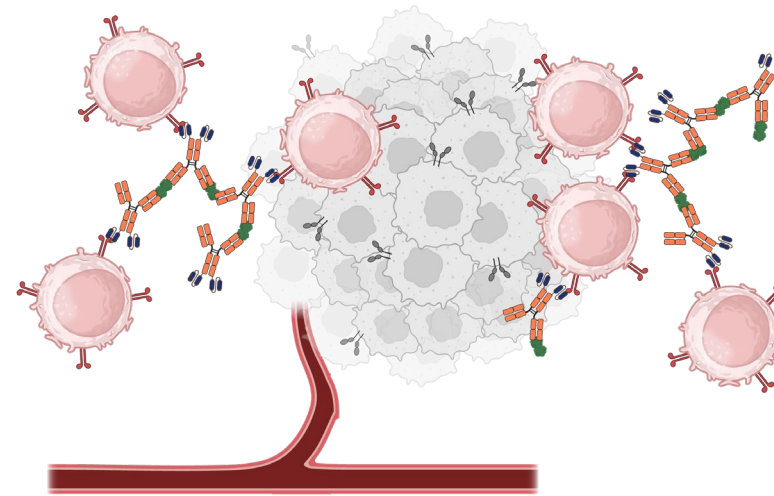
*Cooperative, tetravalent  
PD-1 x VEGF bispecific antibody*

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

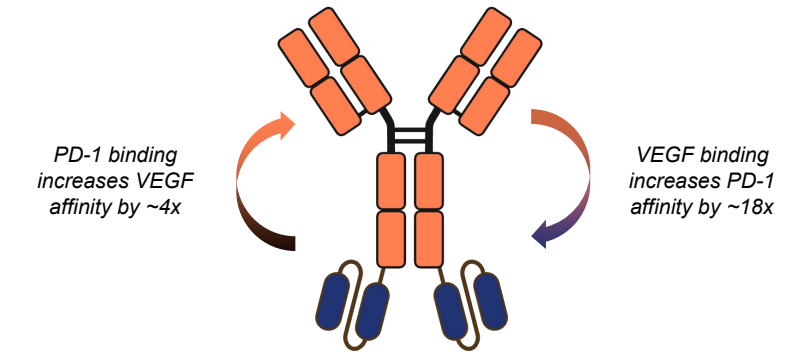
**A** VEGF drives **tumor angiogenesis** and PD-L1 expression **suppresses T cells**



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



✓ **Cooperativity:** 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.



✓ **Tumor targeting:** PD-1 arm concentrates VEGF inhibition in the TME, **potentially sparing healthy tissue** and reducing AEs.

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



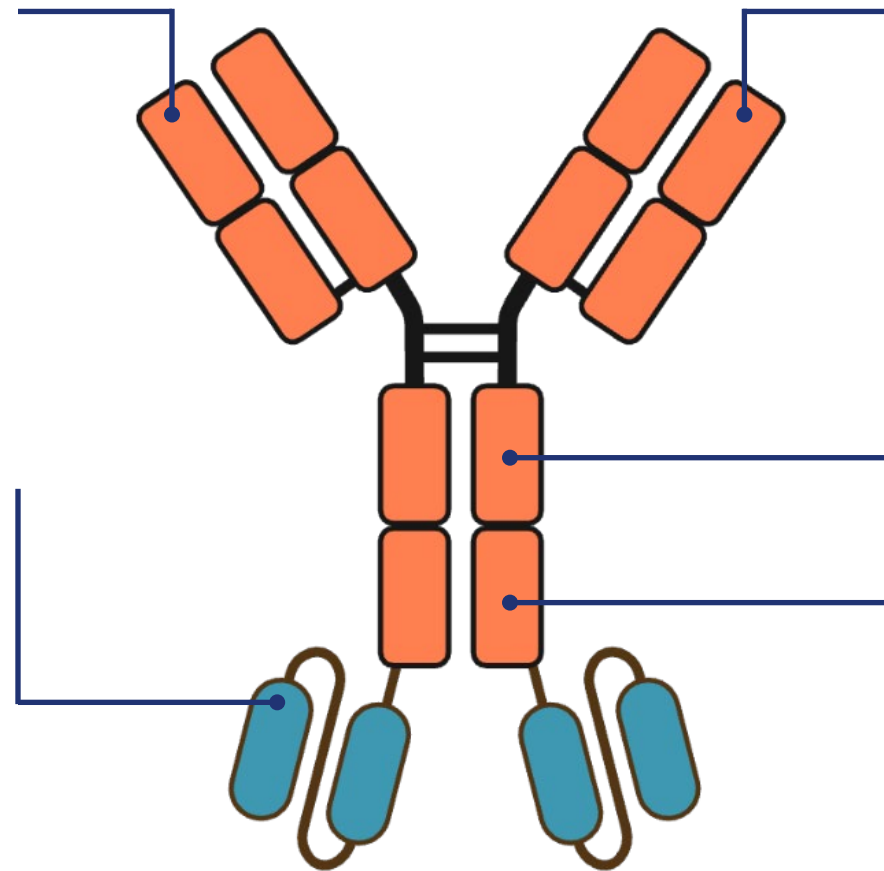
# CR-001 is a highly potent PD-1 x VEGF bsAb designed to recapitulate ivonescimab's cooperative pharmacology

## Same design as ivonescimab

- Pairs anti-VEGF IgG & anti-PD-1 scFvs
- **Avoids risk 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

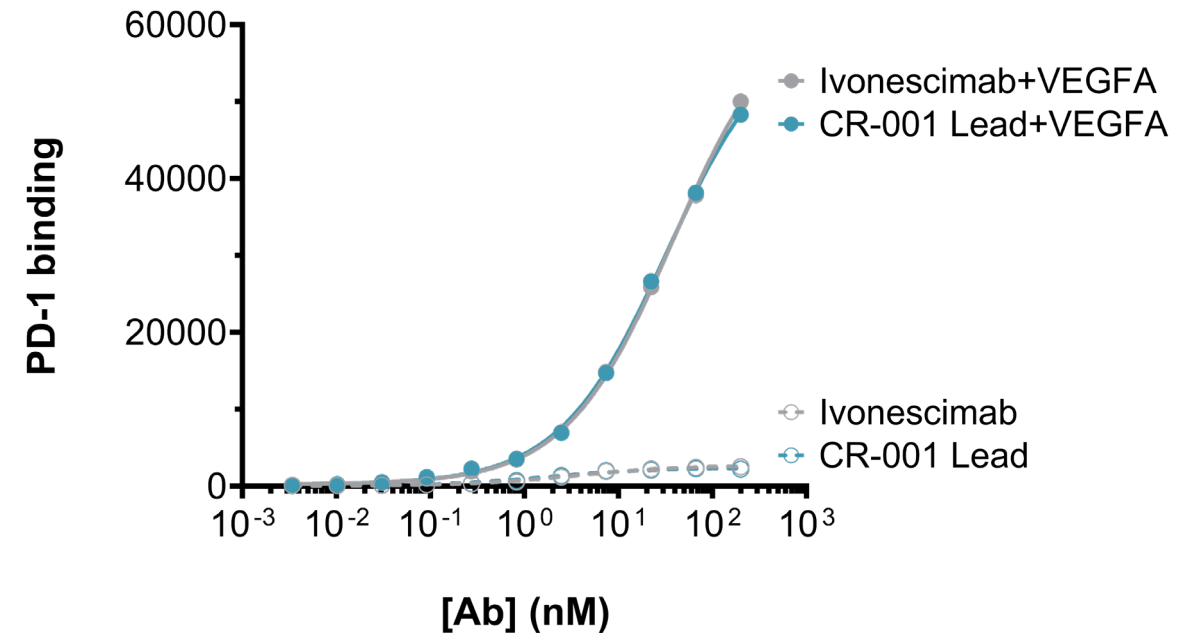
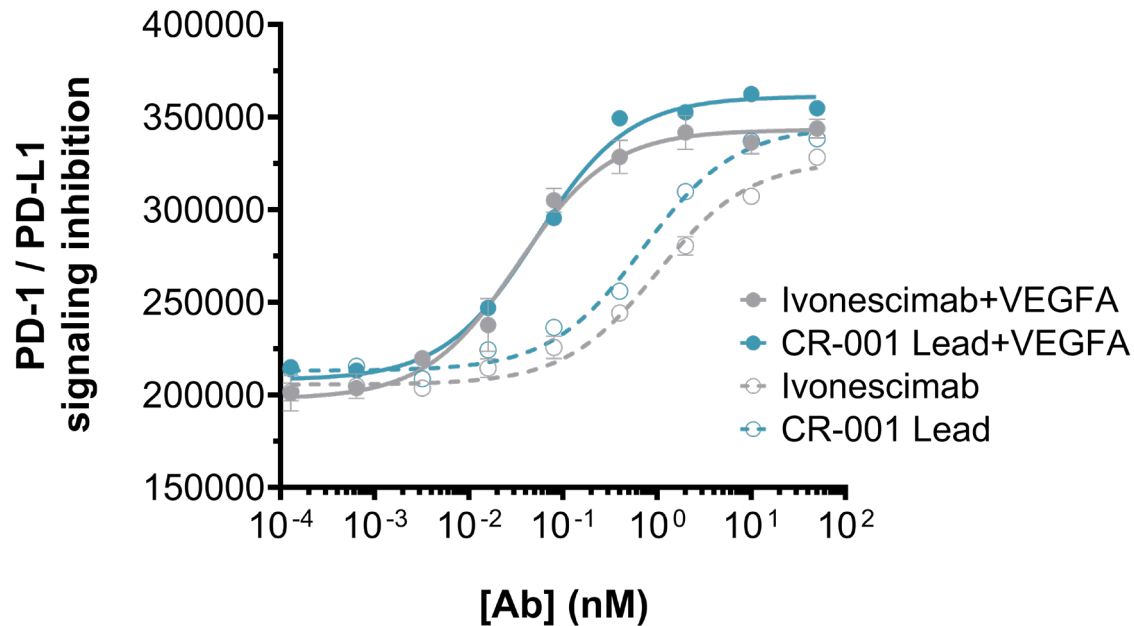
## Designed to match ivonescimab PK

- Native FcRn binding to **match distribution and elimination** of ivonescimab

# CR-001 replicates ivonescimab's cooperative effect, with greater binding to and inhibition of PD-1 signaling in presence of VEGF

CR-001 lead, like ivonescimab, is **more potent** in an NFAT reporter assay **in the presence of VEGF**...

... and also **increases PD-1 binding** on PD-1+ Jurkat cells **in the presence of VEGF**.

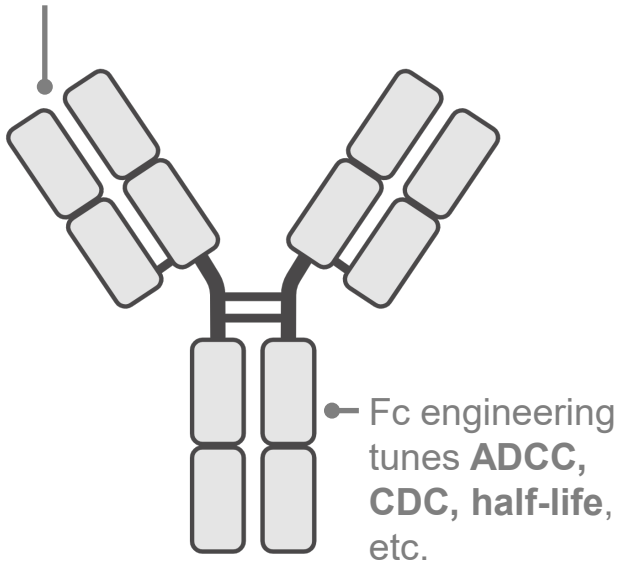


CR-001 lead **demonstrates same cooperative effect** as ivonescimab across multiple assays.

# Replicating ivonescimab's tetravalent format and cooperativity, with stable scFvs, requires complex protein engineering

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

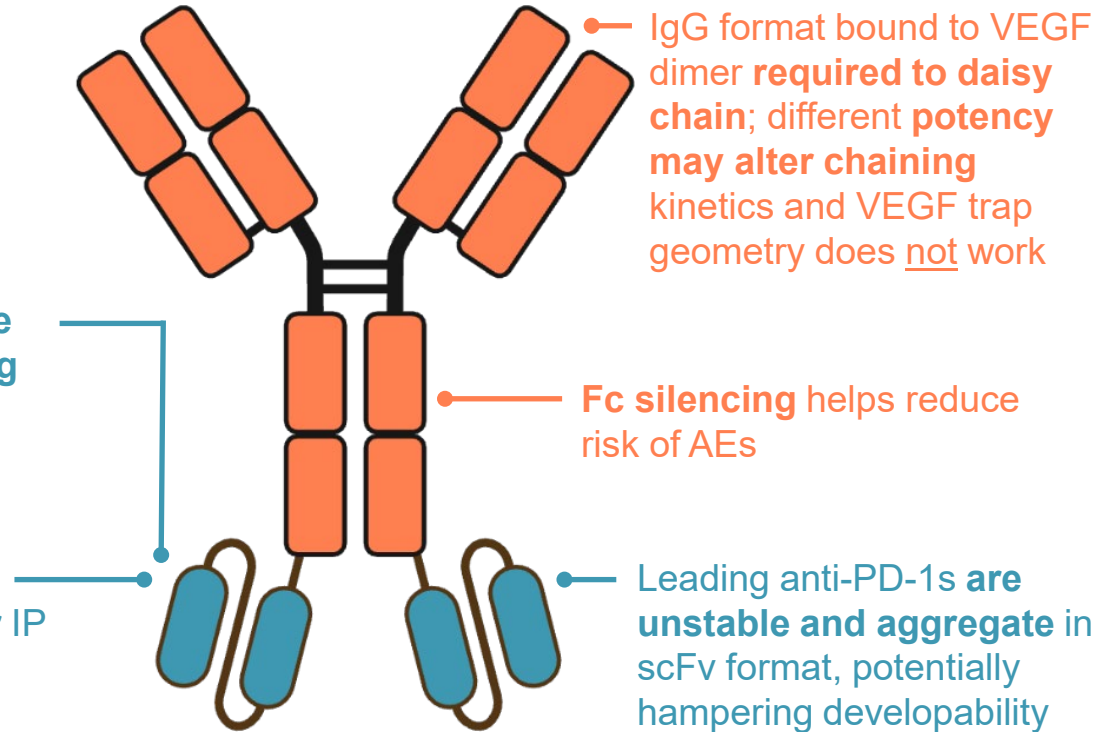
CDRs improved via diversification and/or affinity maturation to maximize potency



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

scFv format can require significant engineering to ensure stability

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



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.

# 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

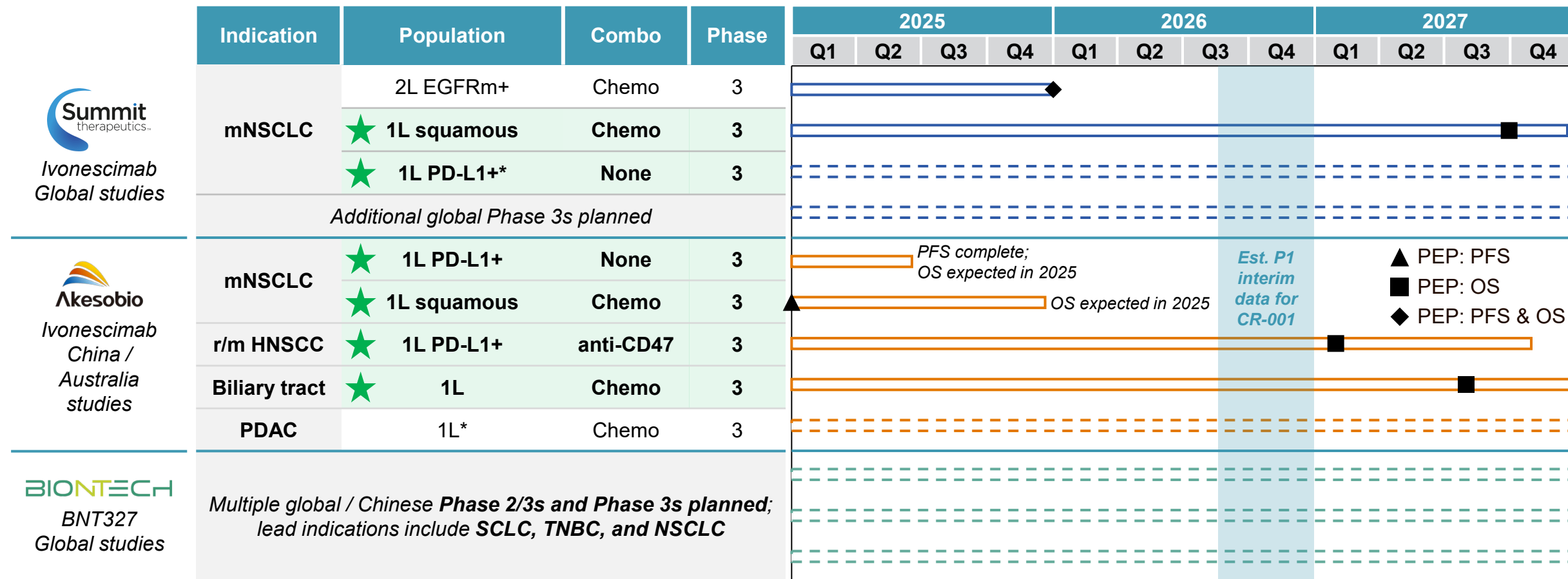
- High microsatellite instability (MSI-H) / deficient DNA mismatch repair (dMMR)
- High tumor mutational burden (TMB-H)



## 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

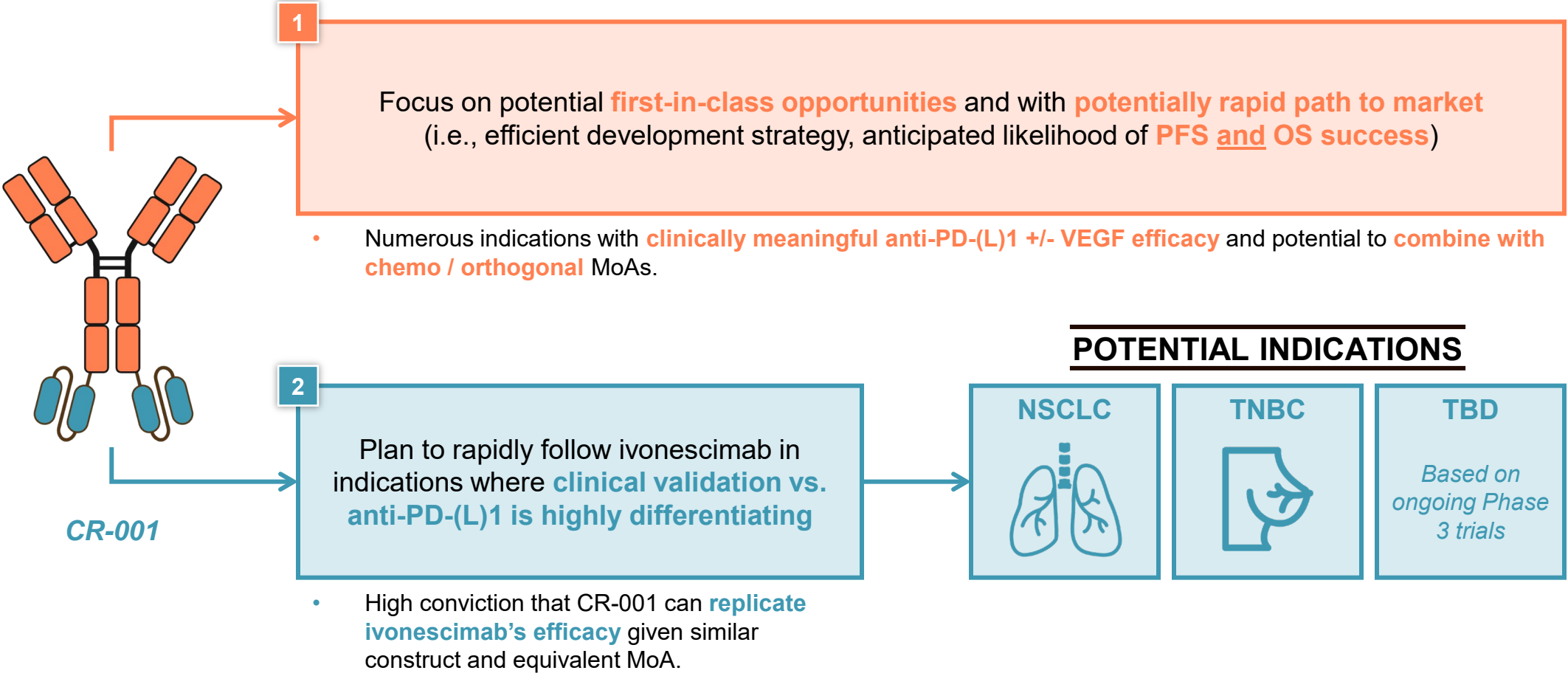


★ vs. PD-(L)1 comparator

Active and contemplated global & Chinese / Australian Phase 3s – across tumor types, lines of therapy, and combinations – will help guide clinical development for CR-001.

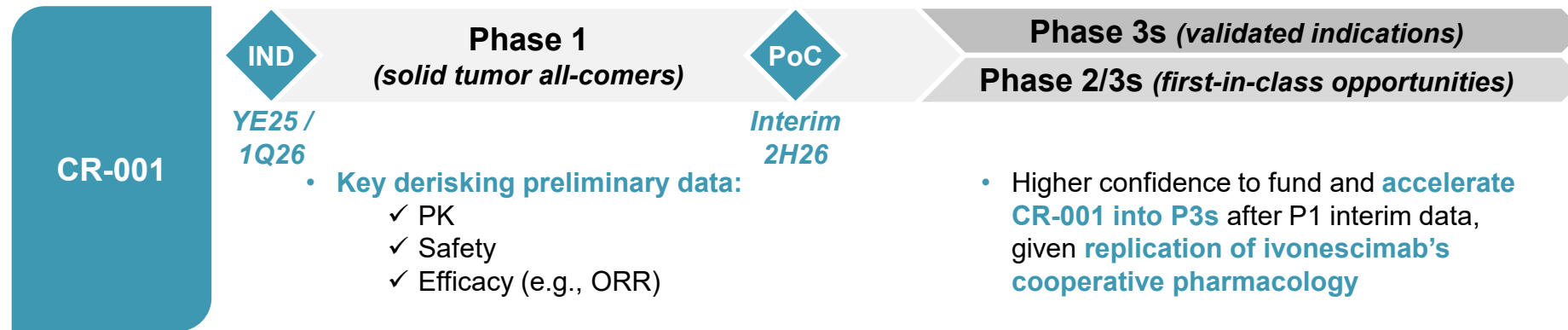
\*Summit has announced P3 in 1L PD-L1+ NSCLC, monotherapy vs. pembro, but has not released trial details. Akeso has announced P3 in 1L pancreatic ductal adenocarcinoma (PDAC), combined with chemo, but has not released trial details. Notes: List of trials is not exhaustive. All confirmed trials have been initiated prior to 2025. NSCLC = non-small cell lung cancer; TNBC = triple negative breast cancer; SCLC = small cell lung cancer; HNSCC = head & neck squamous cell carcinoma. PFS and OS readouts estimated based on PEP (primary endpoints) and completion dates listed on ClinicalTrials.gov. Sources: ClinicalTrials.gov; Company websites; Company presentations

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



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

ILLUSTRATIVE



Phase 1 interim proof-of-concept data are a potentially **significant value-generating event** for CR-001.

- Preliminary data from early Phase 1 cohorts **provides substantial validation of program** because CR-001 **structural design and preclinical data are similar to ivonescimab**.
- Early Phase 1 data, as single agent and in combination with SoC, rapidly enables 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, which 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**, offering potential for significant **early value inflection**.

# CR-001 preclinical data reproduces ivonescimab's breakthrough pharmacology and is rapidly advancing to generate significant value



**Unprecedented third-party data validates 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*



**Built by the proven Paragon team**

*Paragon has a demonstrated track record discovering and developing best-in-class molecules*



# CR-002 & CR-003

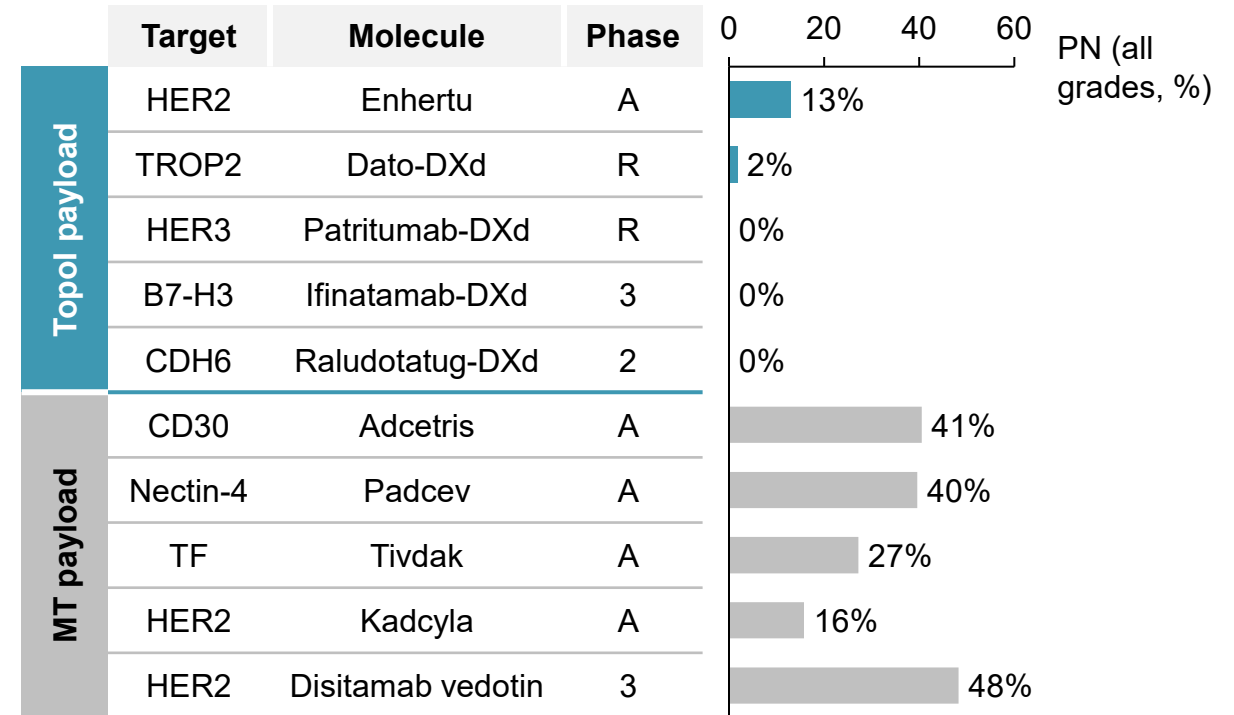
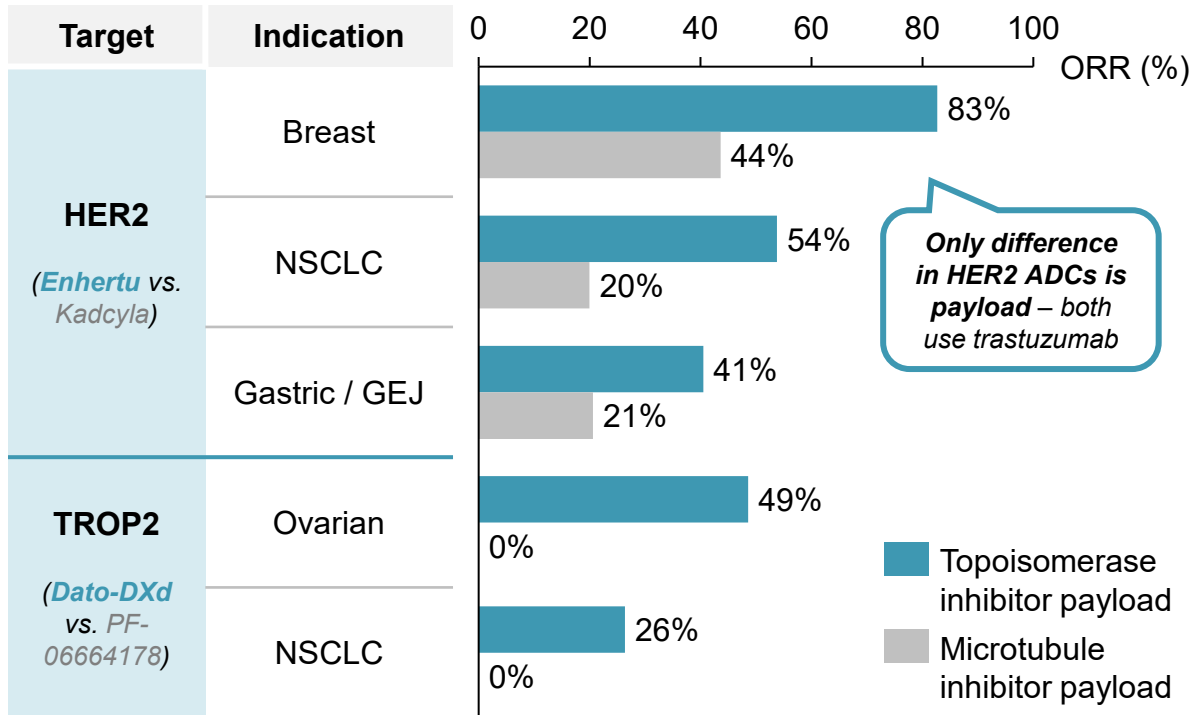
*Topoisomerase inhibitor ADCs  
against validated targets*

# 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**



# Leadership with deep experience building leading biotechnology companies

Partnership with Paragon Therapeutics provides proven expertise in antibody engineering and development



**Jonathan Violin**  
Interim CEO



**Chris Doughty**  
CBO



**Peter Harwin**  
Board of Directors



**Evan Thompson**  
COO



**Hussam Shaheen**  
Head of Research



**Keri Lantz**  
Head of Finance



**Damon Banks**  
Head of Legal



**Neta Batscha**  
SVP, Strategy & Operations



**Mike Meehl**  
SVP, Biologics Research



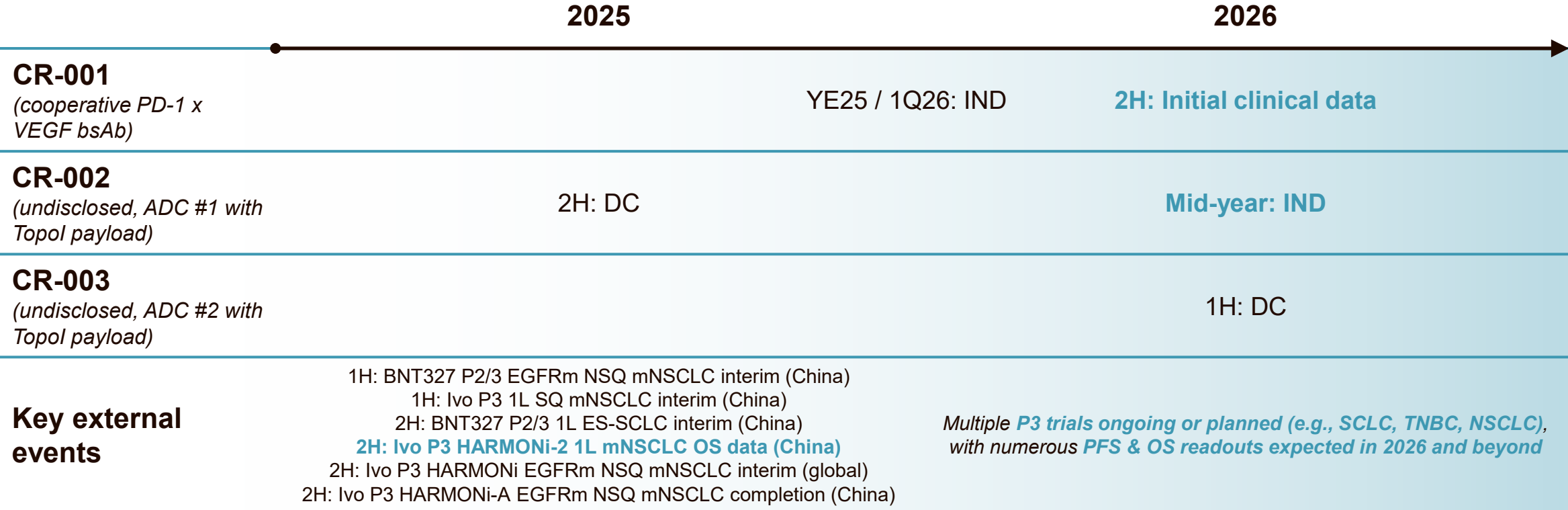
**Jason Oh**  
SVP, Biology



**Shawn Russell**  
SVP, CMC



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



Notes: mNSCLC = metastatic non-small cell lung cancer; TNBC = triple negative breast cancer; SCLC = small cell lung cancer; ES = extensive stage. NSQ = non-squamous; SQ = squamous; EGFRm = mutant EGFR.  
Sources: ClinicalTrials.gov; Company websites

# Estimated capitalization following close of transactions

		Shares on an as-converted basis	Expected ownership of the combined company
<b>GlycoMimetics</b> <ul style="list-style-type: none"> <li>Shares of common stock outstanding</li> </ul>		64,532,953	3.1%
<b>Crescent Biopharma</b> <ul style="list-style-type: none"> <li>Shares of common stock outstanding</li> <li>Series A shares</li> </ul>		105,137,814	
			298,298,000
<b>Pre-closing financing</b> <ul style="list-style-type: none"> <li>Shares of common stock</li> <li>Pre-funded warrants</li> </ul>		1,339,680,730	96.9%
<b>Estimated total shares of common stock of the combined company post-closing</b>		2,081,292,577	



**Thank you**