Proteins by Design[®]

XmAb[®] Antibody Therapeutics

Corporate Overview October 2024



Forward-Looking Statements

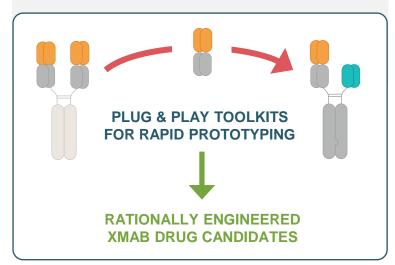
Certain statements contained in this presentation, other than statements of historical fact, may constitute forwardlooking statements within the meaning of the Private Securities Litigation Reform Act of 1995. Such statements include, but are not limited to, statements regarding Xencor's development plans and timelines; potential regulatory actions; expected use of cash resources; the timing and results of clinical trials; the plans and objectives of management for future operations; and the potential markets for Xencor's product and development candidates. Forward-looking statements are based on the current expectations of management and upon what management believes to be reasonable assumptions based on information currently available to it, and involve numerous risks and uncertainties, many of which are beyond Xencor's control. These risks and uncertainties could cause future results, performance or achievements to differ significantly from the results, performance or achievements expressed or implied by such forward-looking statements. Such risks include, but are not limited to, potential delays in development timelines or negative preclinical or clinical trial results, reliance on third parties for development efforts and changes in the competitive landscape including changes in the standard of care, as well as other risks described in Xencor's filings with the Securities and Exchange Commission (SEC). Xencor expressly disclaims any duty, obligation or undertaking to update or revise any forward-looking statements contained herein to reflect any change in Xencor's expectations with regard thereto of any subsequent change in events, conditions or circumstances on which any such statements are based, except in accordance with applicable securities laws. For all forward-looking statements, we claim the protection of the safe harbor for forward looking statements contained in the Private Securities Litigation Reform Act of 1995.



Proven Power of XmAb® Engineering: Proteins By Design®

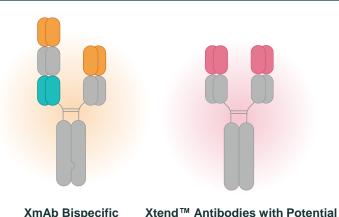
Small changes, big functional impacts

- XmAb Fc Domains augment native immune functions in molecules and/or control their structure, while preserving desired attributes
- XmAb engineered antibodies are designed to solve complex biologic problems
- Strong patent portfolio with over 1,600 patents issued and pending worldwide



Advancing an optimized portfolio of XmAb drug candidates

- Oncology: 3 novel TCEs advancing in Phase 1 studies; narrow focus for vudalimab in mCRPC and 1L NSCLC
- Autoimmune: Upcoming study initiation plans
 - 4Q'24: XmAb942 (Xtend™ TL1A)
 - 1H'25: Plamotamab (CD20xCD3) in RA
 - 2H'25: XmAb657 (CD19xCD3)



T-cell Engagers (TCEs) Best-in-class Half-life Extension

Partnerships leverage modular XmAb technology

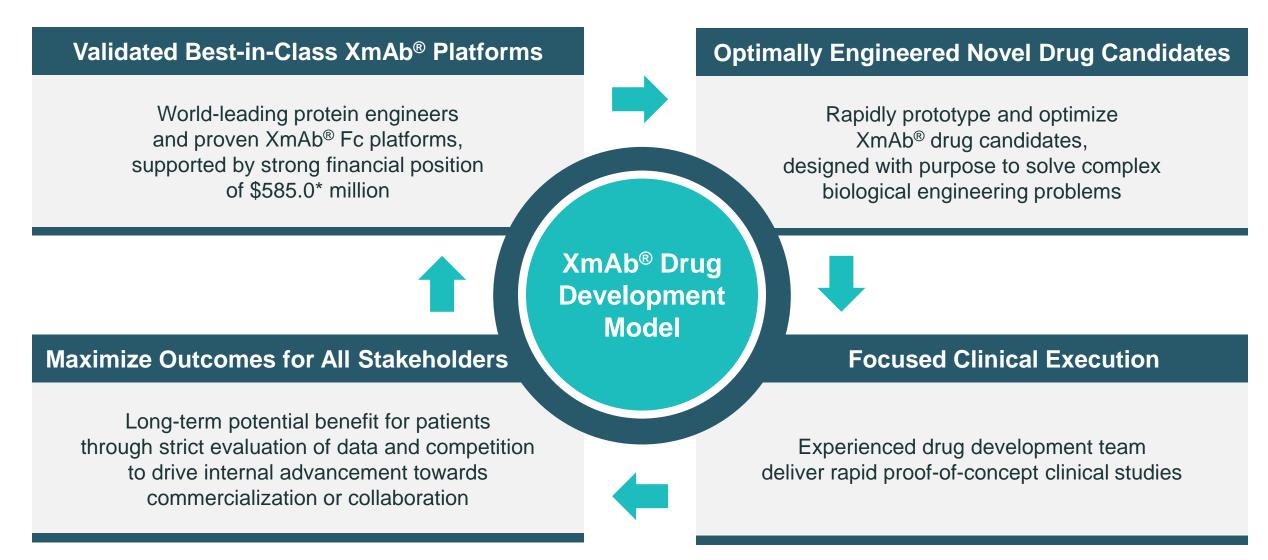
- More than 15 technology license partnerships greatly broadens scope with little-to-no effort
- Multiple commercialized XmAb antibodies

ULTOMIRIS[®] MONJUVI[®]/MINJUVI[®]





Xencor's Disciplined Drug Development Strategy



Axencor

* As of 6/30/2024. Includes cash, cash equivalents & marketable debt. Updated 8/5/2024.

Next-Gen XmAb® Drug Design in Oncology & Autoimmune Diseases

Pipeline focus on T-cell engagers and bispecific mechanisms

Program	Targets	XmAb [®] Platforms	Indications	Discovery	Preclinical	Phase 1	Phase 2	Phase 3
Solid Tumor C	ncology: T-cell Eng	gagers (CD3 & CD28)						
XmAb819	ENPP3 x CD3	2+1 Bispecific	ccRCC					
XmAb808	B7-H3 x CD28	2+1 Bispecific, Xtend™	Prostate cancer, oncology	+ pembrolizumab				
XmAb541	CLDN6 x CD3	2+1 Bispecific, Xtend	Ovarian cancer, oncology					
XmAb Program	Undisclosed TCE	Bispecific, Xtend	Solid tumor oncology					

Solid Tumor Oncology: T-cell Selective, Dual Checkpoint Inhibitor

Vudalimab PD-1 x CTLA-4 Bispecific, Xtend mCRPC +/- chemotherapy
1L NSCLC + chemotherapy

Immunology Programs

Plamotamab	CD20 x CD3	Bispecific	Rheumatoid Arthritis			1H'25
XmAb942	TL1A	Xtend, FcKO	Inflammatory Bowel Diseases (IBD)		4Q'24	
XmAb657	CD19 x CD3	2+1 Bispecific, Xtend	Autoimmune Diseases		2H'25	
XmAb Program	TL1A x IL23	Bispecific, Xtend	Autoimmune Diseases			

ccRCC clear cell renal cell carcinoma NSCLC non-small cell lung cancer mCRPC metastatic castration-resistant prostate cancer FcKO Fc knock out

Immunology



XmAb® Technologies Create Numerous Differentiated Antibodies for Technology Partners

Technology licensing expands pipeline with very little opportunity cost

Selected Programs	Fc Domain	Indications	Preclinical	Phase 1	Phase 2	Phase 3	Marketed	Commercial Rights
Ultomiris®	Xtend™	PNH, aHUS, gMG, NMOSD						AstraZeneca Rare Disease
Monjuvi®	Cytotoxic	DLBCL						Incyte
Xevudy®	Xtend	COVID-19				NOT CURRENTL	Y AUTHORIZED IN U.S.	NIR
Obexelimab	Immune Inhibitor	IgG4-RD, wAIHA						Zenas### BioPharma
Tobevibart (VIR-3434)	Cytotoxic Xtend	Hepatitis B Hepatitis D						NIR
Xaluritamig STEAP1 x CD3	2+1 Bispecific	Prostate cancer						AMGEN
Efbalropendekin alfa IL15/IL15Rα-Fc	Bispecific Xtend	r/r multiple myeloma Oncology	+ daratumumab + cevostamab + atezolizumab					Genentech A Member of the Roche Group
ASP2138 CLDN18.2 x CD3	2+1 Bispecific	Oncology						Astellas
JNJ-9401 PSMA x CD28	Bispecific	Prostate cancer						Johnson&Johnson Innovative Medicine
JNJ-1493 CD20 x CD28	Bispecific	Heme-Onc						Johnson&Johnson Innovative Medicine

Registered trademarks: Ultomiris[®] (Alexion Pharmaceuticals, Inc.), Monjuvi[®] (Incyte Holdings Corp.) Xevudy[®] (Glaxo Group Limited)

Key



XmAb® Bispecific T Cell Engagers

XmAb **2+1** Bispecific Antibody Format

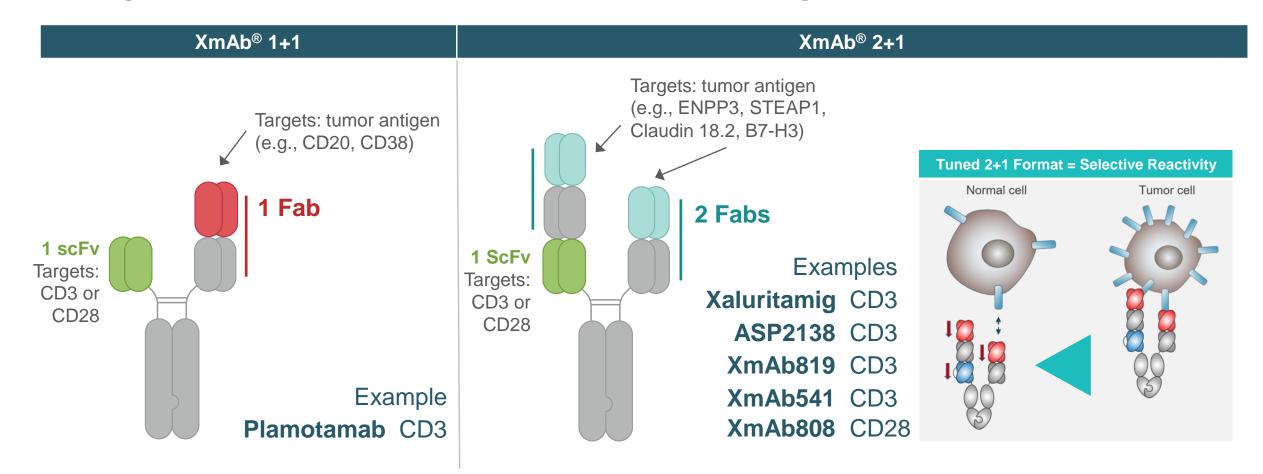
XmAb819 (ENPP3 x CD3)

XmAb541 (CLDN6 x CD3)

XmAb808 (B7-H3 x CD28)



XmAb[®] T Cell Engagers Use Multiple Formats and Affinity Designs to Customize for Each Tumor Target

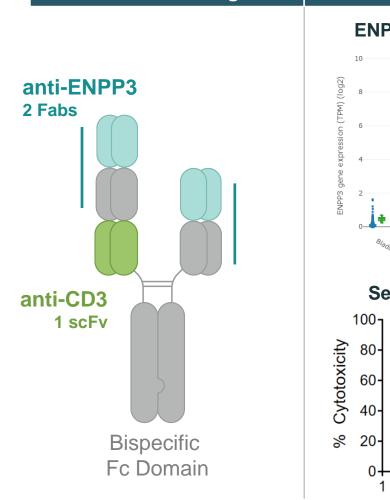


CD3 affinity tuned for reduction of cytokine release syndrome and off-tumor cell killing Tumor antigen binding affinity tuned for tumor expression density and to match format



XmAb[®]819: CD3 T-cell Engager for Renal Cell Carcinoma in Phase 1

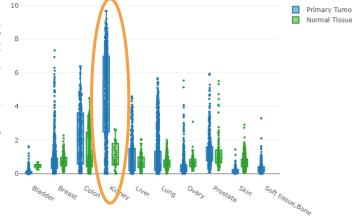
Encouraging Initial Data in Ongoing Dose Escalation in ccRCC



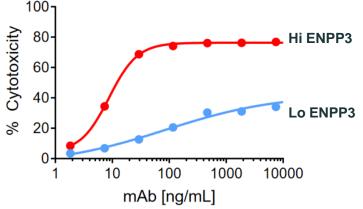
XmAb 2+1 Design

XmAb819 (ENPP3 x CD3)

ENPP3: high RCC; low healthy tissues



Selective T cell directed cytotoxicity



Phase 1 Dose Escalation Study

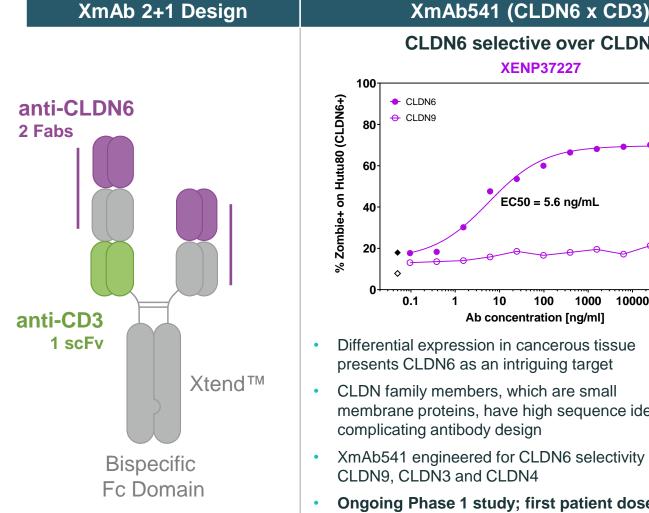
- Dose escalation ongoing in relapsed/refractory clear cell RCC (ccRCC - nearly uniformly high ENPP3 expression
- Administration: IV and subcutaneous cohorts dosing weekly, with priming and step-up doses
- Following determination of RP2D, expansion cohorts planned in ccRCC and other histologies with high ENPP3 expression
- Companion diagnostic under development for potential patient selection in other histologies

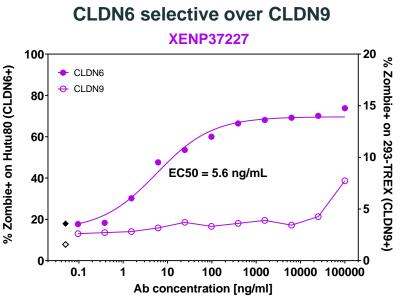
Observed in escalation (update 09-Sep-2024)

- Clear initial evidence of anti-tumor activity, including RECIST responses, in recent cohorts
- Duration of treatment for several patients in earlier dose cohorts has extended beyond one year
- Cytokine release syndrome (CRS) manageable
- No MTD reached; tolerability from recent dose cohorts continues to support dose escalation
- Investigators remain highly engaged, and enrollment into new dose cohorts has been rapid
- Intravenous and subcutaneous cohorts continue dose
 escalation in parallel
- Evaluation of expansion into additional tumor types is ongoing
- Clinical update and first dose expansion cohort expected to start during 1H'25
 NCT05433142

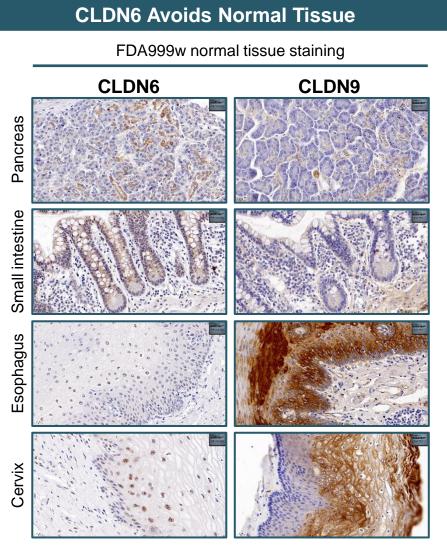


XmAb[®]541: CD3 T-cell Engager for Ovarian Cancer & Solid Tumors





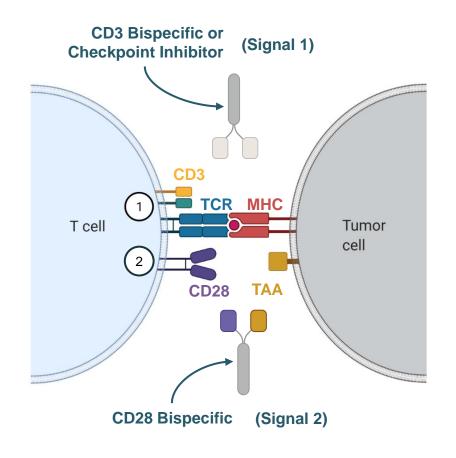
- Differential expression in cancerous tissue presents CLDN6 as an intriguing target
- CLDN family members, which are small membrane proteins, have high sequence identity, complicating antibody design
- XmAb541 engineered for CLDN6 selectivity over similar CLDN9, CLDN3 and CLDN4
- Ongoing Phase 1 study; first patient dosed 1H24 NCT05433142





CD28 Bispecific Antibodies Provide a Boost to T Cell Activation

CD28 provides "Signal 2" activation



XmAb[®] CD28 T cell engagers feature low affinity, monovalent binding

- Avoid historic CD28 safety concerns (superagonism)
- Well behaved: stable, silent FcγR interactions, with Xtend[™] Fc technology
- Potential to combine with anti-PD1 and/or CD3 T cell engagers

Multiple wholly owned early-stage and actively advancing programs

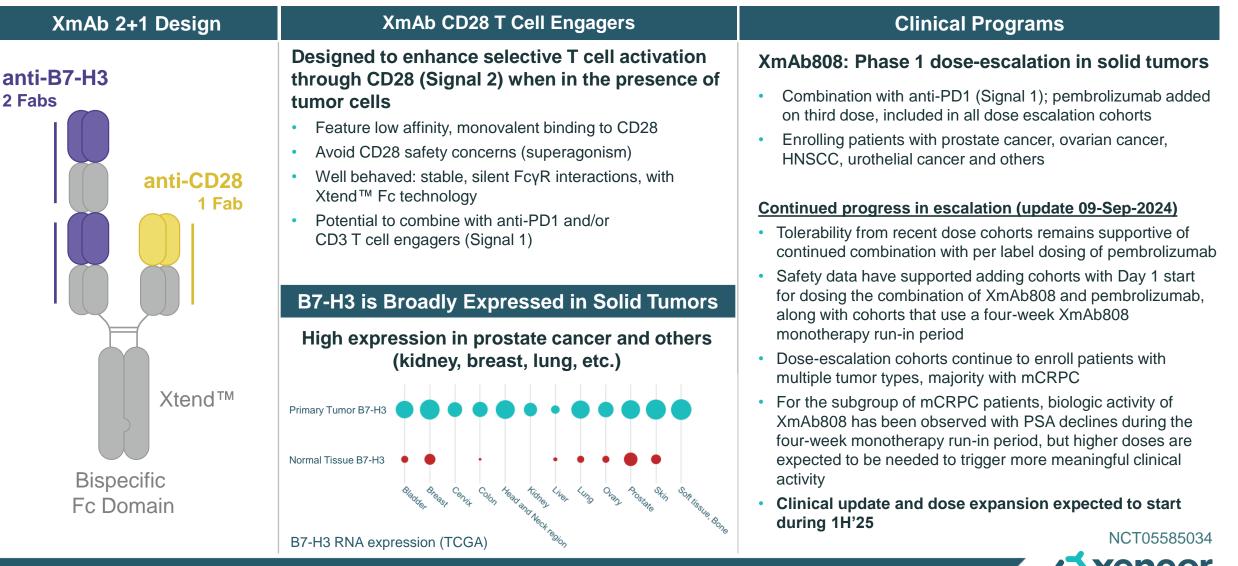
- Ongoing Phase 1 study of XmAb808 (B7-H3 x CD28) in combination with pembrolizumab in solid tumors
- Presented preclinical data from multiple research-stage programs targeting CEACAM5, STEAP1, ENPP3, Trop2 and MSLN

Two narrow, target-limited collaborations with J&J for CD28 bispecifics

- JNJ-9401 (PSMA x CD28; Phase 1) for combination with J&J CD3 bispecifics; collaboration includes access to J&J prostate-cancer franchise for clinical combinations across Xencor's portfolio
- JNJ-1493 (CD20 x CD28; Phase 1) for J&J's use in combination with agents, such as CD3 bispecifics



XmAb808: Tumor-specific CD28 T-cell Engager Targeted to Broadly Expressed Tumor Antigen B7-H3 in Phase 1



New Pipeline Programs: B-cell Depleting T-cell Engagers

Plamotamab (CD20 x CD3)

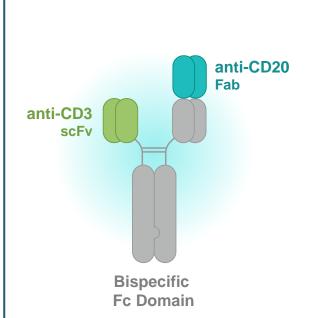
XmAb657 (CD19 x CD3)



Plamotamab Phase 2 Ready, Subcutaneous CD20 x CD3 BsAb

Planned proof-of-concept for the T-cell engager class in autoimmune and inflammatory disease

XmAb[®] CD20 x CD3 Bispecific Design



- Plamotamab designed in a 1+1 format and selected for extended activity and favorable tolerability observed in NHPs
- Human half-life ~18 days;
 estimated 80% SC
 bioavailability
- Robust manufacturing process with high yield and excellent formulation stability data

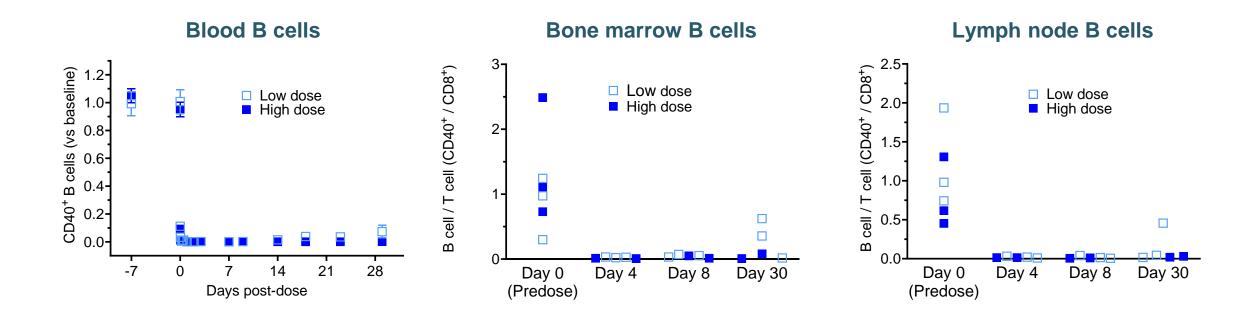
Positioned for Success

- N=154 from dose escalation and expansion cohorts with both IV and SC formulations in B-cell cancers
- Comparable preliminary efficacy data to leading commercial CD20 x CD3 in patients with prior CAR-T
- IV & SC dosing regimens with improved CRS data vs. leading commercial CD20 x CD3¹
- Existing inventories of drug product and drug substance for seamless integration into the next phase of clinical development

BsAb bispecific antibody IV intravenous, SC subcutaneous NHP non-human primate CRS cytokine release syndrome 1 No head-to-head trial has been conducted evaluating plamotamab against other data included herein. Differences exist between clinical trial design, patient populations and the product candidates themselves, and caution should be exercised when comparing data across trials.



Single Dose of Plamotamab in NHPs Durable B-cell Depletion Observed in Blood and Lymphoid Organs

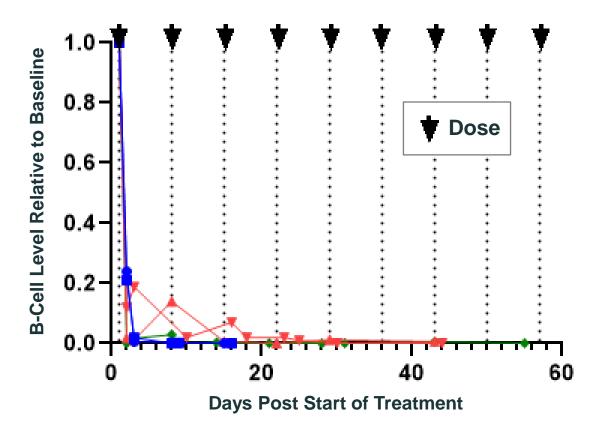




>95-99% Peripheral B-cell Depletion Observed in Lymphoma Patients with IV & SC Plamotamab in Phase 1 Monotherapy Study

- Patients were identified (N=5) that had baseline absolute B-cell count
 > 30 cells/µL in the blood
- >90% reductions in B-cell count also observed at lower doses

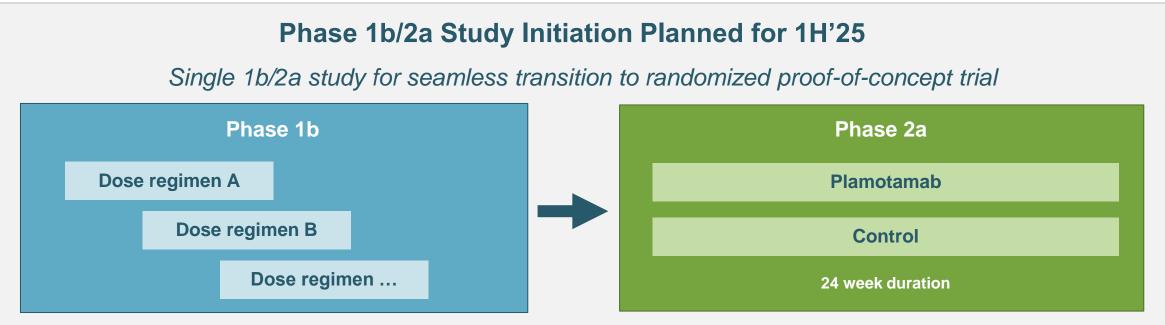
Relative Change in B-cell Count





Plamotamab: Plan for Phase 1b/2a Rheumatoid Arthritis Study Start

Maximal efficiency to clinical proof of concept in multi-drug resistant rheumatoid arthritis (MDR-RA)



- Quickly refine priming/step-up dosing regimens used in lymphoma studies
- Assess SC and IV routes, and pre-medication regimen including corticosteroids, to be run in parallel on a staggered start
- Assess safety, biomarkers, initial efficacy in RA patients

- Advance selected dosing regimen into placebo-controlled trial in MDR-RA patients
- Single-cycle dosing in line with other B-cell depleting agents
- 24-week efficacy endpoint with interim efficacy analysis at week 12 with paired biomarker assessment



XmAb657 CD19 x CD3 Optimized for Autoimmune Disease

Rational XmAb[®] Design

- High affinity and stability anti-CD19 binder
 - Bivalent to efficiently target B cells expressing very low levels of CD19 (e.g., plasma cells and plasmablasts)
- Affinity-tuned and highly stable anti-CD3 binder
- Uses Xencor's clinically validated 2+1 format
- Heterodimeric Fc domain engineered to abrogate effector function and improve half-life
- Xtend[™] Fc for long half life

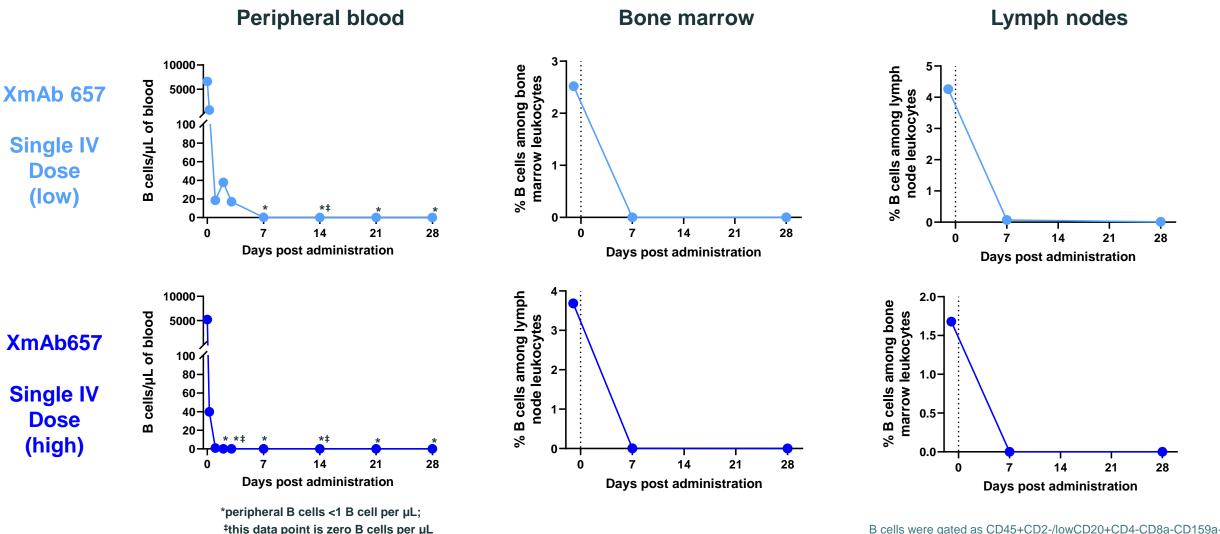
Positioned for Success

Ongoing NHP studies have shown effective B-cell depletion with single dose

- Broad opportunity set of disease indications supports multiple development pathways for success
- EULAR 2024 and subsequent updates of CD19 CAR-T clinical data highlighted potential issues with CAR-T approach on efficacy and safety
- Rational design of XmAb657 supports best-in-class potential for clinical outcomes
- Current timeline to FIH study in 2H'25 puts Xencor on-track to be a leading CD19 x CD3 program within autoimmune disease



Single Dose of XmAb657 in NHPs **Deep B-cell Depletion Sustained for at Least 28 Days**

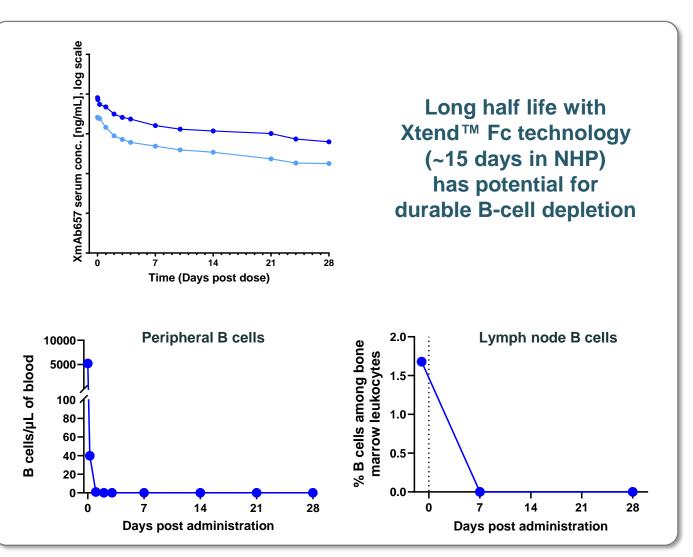


[‡]this data point is zero B cells per µL

(encor

XmAb657: Rationally Designed for Autoimmune Disease FIH Planned 2H'25

- Has been observed to demonstrate deep and durable B-cell depletion in NHPs, enabled by potentially best-in-class pharmacokinetics
- Has been well tolerated in NHP with no clinical signs of CRS
- GMP production campaign initiated
- Further plans to investigate subcutaneous dosing and priming
- First-in-human study planned to initiate in 2H'25





New Pipeline Programs: TL1A Portfolio

XmAb942 (Xtend[™] TL1A)

XmAb TL1A x IL-23



Inflammatory Bowel Disease (IBD) is a Devastating Disease with Significant Unmet Medical Need

~3m Estimated diagnoses in the US¹

Two common forms: Crohn's disease Ulcerative colitis

Economic burden estimated at \$5.4B in 2023²

1 Clarivate 2 GlobalData 3 Prescient whitepaper

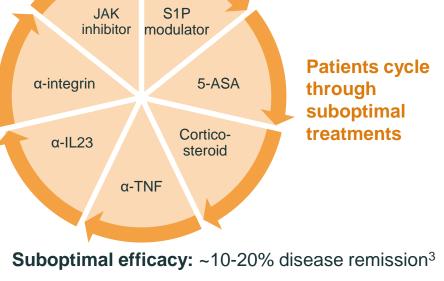
Significant Health Burden

- Impaired quality of life
- Lower life expectancy
- Surgeries, hospitalization
- Increased risk for intestinal resection
- Increased risk for colorectal cancer

Severe Symptoms of IBD

- Fatigue
- Fever
- Reduced appetite
- Mental health

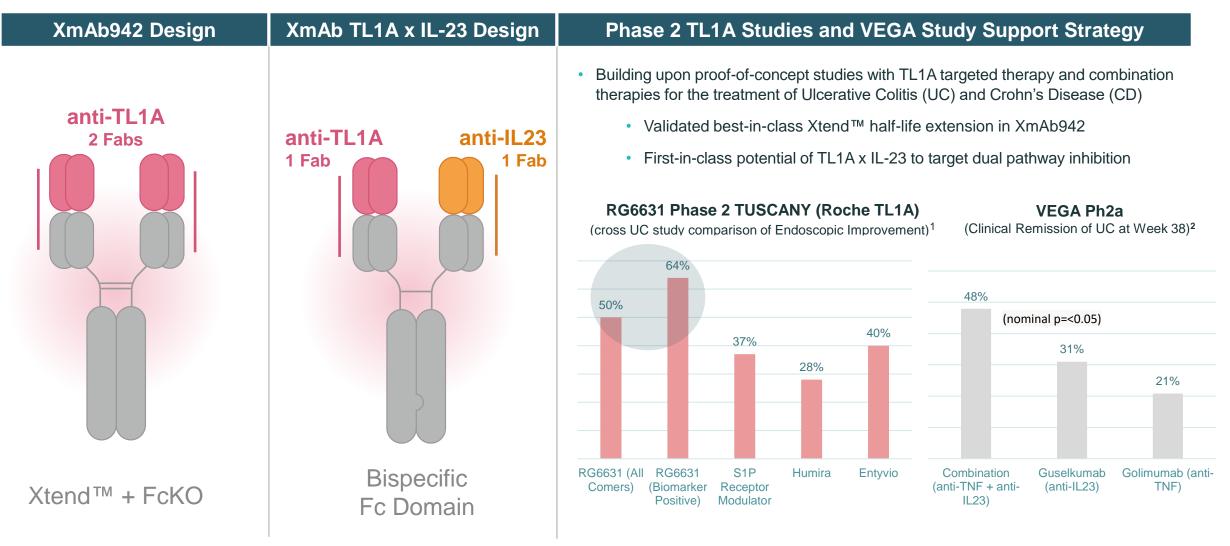




- Adverse events: Infection, malignancy, thromboembolism, cardiac
- Burdensome regimens: poor patient compliance



Development of XmAb942 and XmAb TL1A x IL-23 for IBD



Sourced from Roivant presentation of TUSCANY, Entyvio (anti-integrin) and Humira (anti-TNF) data from VARSITY P3 study, S1P receptor modulator data from ELEVATE 52 P3 study
 Guselkumab plus golimumab combination therapy versus guselkumab or golimumab monotherapy in patients with ulcerative colitis (VEGA); Feagan and Shao et al.; The Lancet G&H; Feb 2023

Xtend[™] Fc: Validated Half-Life Extension (HLE) Technology Enabling Potential Best-in-Class Anti-TL1A

Clinically validated with significantly improved half-life and dose frequency

- Ultomiris half-life extended >4x as compared to Soliris; maintenance dose frequency reduced by 4X¹
- VRC01LS half-life extended >4X as compared to parental (71 days vs 15 days)²

Similar safety and immunogenicity risk as parental antibodies in studied antibodies using Xtend Fc domains^{3,4,5}

Antibody thermostability maintained in studied antibodies using Xtend Fc domains^{6,7}

Superior or comparable to other HLE technologies (e.g., YTE) across multiple studies and parameters^{6,7,8}

Typical HLE scaling from cyno to human is ~3.5x⁹

Clinical Half-Life and Maintenance Dosing Ultomiris vs. Soliris¹⁰

Product	Half-life (days) ¹¹	Dosing Interval ^{1,12}	
Ultomiris (with Xtend™)	49.7-64.3	Q8W	
Soliris	11.33-12.1	Q2W	

Proprietary Xtend[™] Fc Domain has been incorporated into ≥ 35 molecules that have been tested in clinical studies

Xtend is commonly referred to as 'LS' in academic literature

¹ Ultomiris & Soliris drug labels 2 Ledgerwood Clin Exp Imm 2015 3 Lee et al. Blood 2019 4 Gaudinski et al. PLOS Med 2018 5 Vu et al. J Neurol 2023 6 Ko et al. Exp Mol Med 2022 7 Internal Data 8 Ko et al. Nature Letter 2014 9 Haraya & Tachibana. BioDrugs (2023) 37:99–108 10 Data adapted from FDA and EMA drug labels 11 Reported Half-life across approved indications 12 Maintenance dosing interval in adults



XmAb942: Novel High-Affinity Anti-TL1A mAb Designed for Extended Half-Life, Under Development for the Treatment of IBD

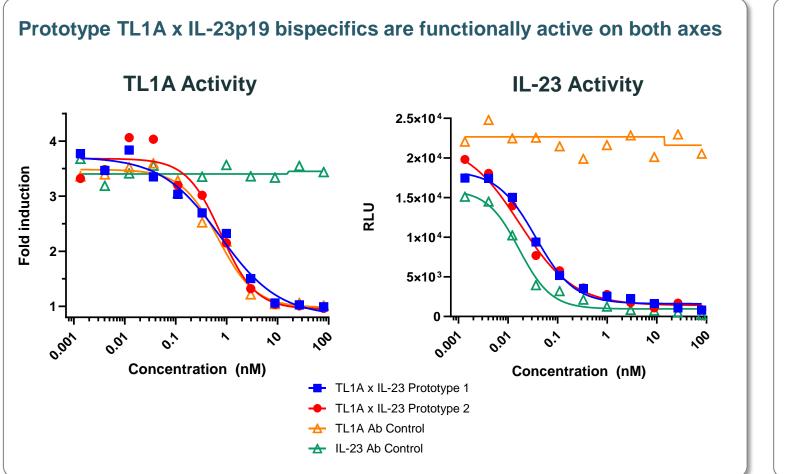
- XmAb942 utilizes Xtend[™] Fc domain technology with potentially class-leading potency
- Half-life in non-human primate studies, 23 days, supports Q8W to Q12W dosing in humans
- High concentration formulation for subcutaneous dosing
- Preclinical discovery and characterization presented during UEG Week 2024
- First-in-human clinical studies to begin 4Q'24

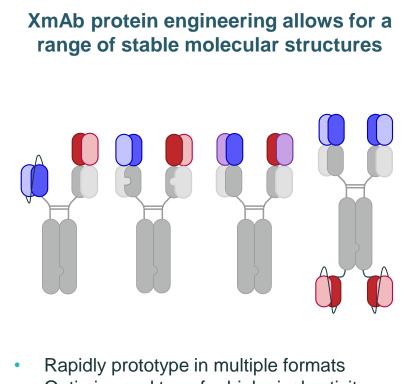
Company	Program ¹	Potent	SC Dosing	Q8-12W Dosing	Half-life extension	Low Immunogenicity
Xencor	XmAb942	\bigcirc	\bigcirc			Predicted
Merck (Prometheus) ^{2,3}	MK-7240	\bigotimes	\bigcirc	\bigotimes	\bigotimes	\bigcirc
Roche (Roivant) ^{4,5}	RG-6631	\bigcirc	\bigcirc	\bigotimes	\bigotimes	\bigotimes
Sanofi (Teva) ⁶	TEV-48574	\bigcirc	\bigcirc	\bigotimes	\bigotimes	ТВД

1 No head-to-head trial has been conducted evaluating XmAb942 against other data included herein. Differences exist between clinical trial design, patient populations and the product candidates themselves, and caution should be exercised when comparing data across trials 2 PRA023 Progress Update (Prometheus presentation) 3 Feagan et al. The Anti-TL1A Antibody PRA023 Demonstrated Proof-of-Concept in Crohn's Disease: Phase 2a APOLLO-CD Study Results (DOP87) Abstract citation ID: jjac190.0127 4 Banfield et al. Br J Clin Pharmacol. 2020;86:812–824 5 Clarke et al. mAbs. 2018;10:4, 664-677 6 Danese et al. Clin Gastroenterology and Hepatology. 2021;19:11, 2324-32.e6



XmAb[®] TL1A x IL-23 to Have First-in-Class Potential First-in-Human Study Planned in 2026





Optimize and tune for biological activity



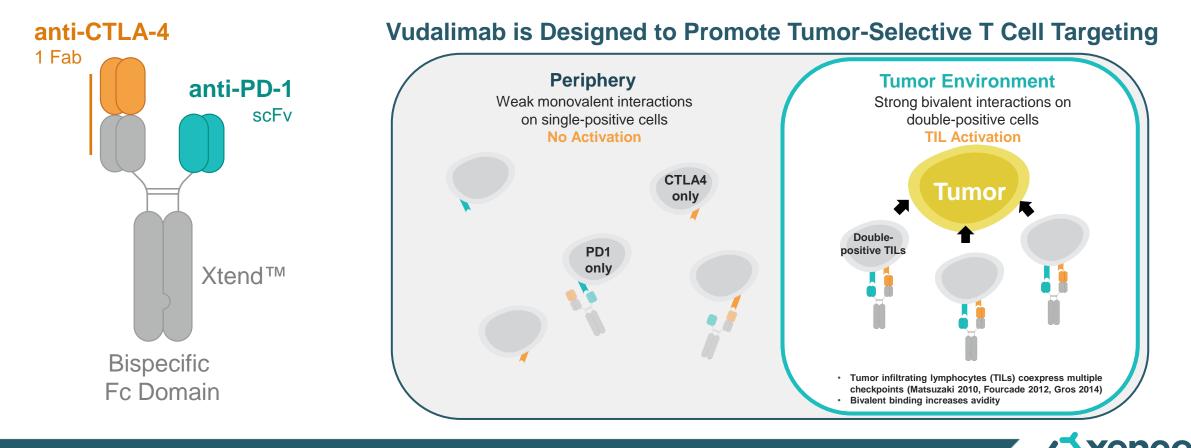
T Cell Selective, Dual Checkpoint Inhibitor

Vudalimab (PD-1 x CTLA-4)

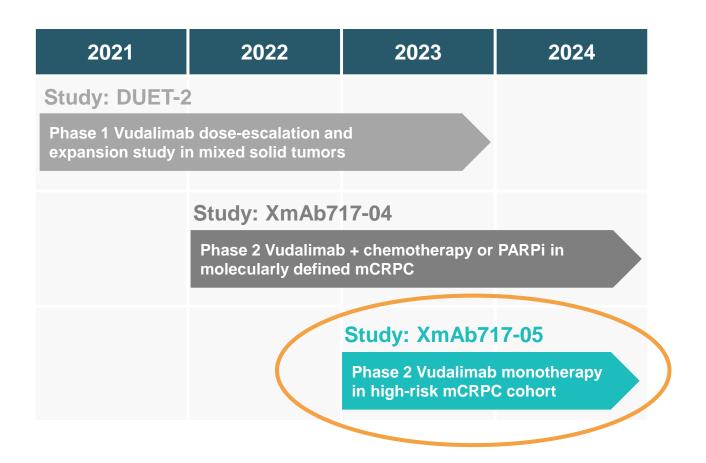


Vudalimab: Selective PD-1 x CTLA-4 Bispecific Antibody

- Monotherapy generally well tolerated, with a potentially differentiated profile from two-antibody combination therapy (PD-1 + CTLA-4)
- Xencor's initial focus on tumors with poor PD-1 inhibitor activity: e.g., prostate cancer



Evolution of Vudalimab Development Program for mCRPC



mCRPC Cohort in Study XmAb717-05

- Visceral, soft tissue, or lymph node metastases ("high-risk" mCRPC)
- Measurable disease by RECIST 1.1
- Progressed after all approved, medically appropriate therapies
- < 2 prior chemotherapy regimens
- No prior treatment with anti-CTLA-4 or PD-1

Study status (February 7, 2024)

• 14/30 mCRPC patients enrolled

Vudalimab Q3W flat dose schedule: 1000 (< 80 kg) or 1200 mg (≥ 80 kg) IV



Preliminary Data for Vudalimab Monotherapy in mCRPC

Vudalimab monotherapy has been associated with clinical response in 5 out of 12 evaluable mCRPC patients with visceral or lymph node metastases

 Findings consistent with observations of durable responses (41 and 27 weeks) in Phase 1 mCRPC cohort in 2 patients with visceral and lymph node (extrapelvic and/or intrapelvic) metastases

Vudalimab monotherapy safety profile has been consistent with other checkpoint inhibitor therapies

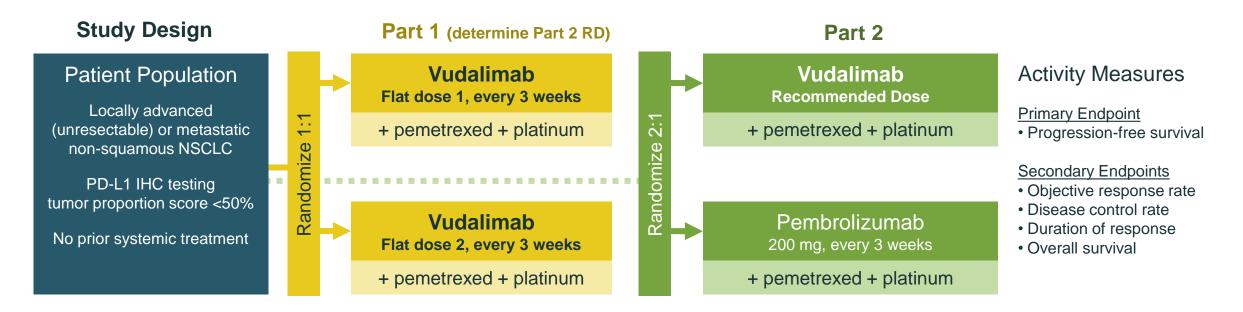
• Low rate of discontinuation of treatment due to adverse events

Additional data will inform future development efforts in monotherapy, and/or contribution of monotherapy to combination treatment



Phase 1b/2 Study in 1L NSCLC in combination with chemotherapy

Part 1 dose comparison, Part 2 randomized vs. pembro; First patient dosed in Q4 2023



Encouraging proof-of-concept data in NSCLC supports evaluation of vudalimab in 1L

- Study of volrustomig (PD-1 x CTLA-4) in checkpoint-naïve NSCLC shows superior PFS over pembrolizumab
 - Volrustomig + chemo vs. pembrolizumab + chemo
- Vudalimab Phase 1 Cohort C (20 patients with NSCLC) activity in 3-4L patients
 - Heavily-pretreated population: 95% checkpoint experienced, 40% ≥ 2 prior checkpoints, median 3 prior therapies
 - 14% objective response rate (2 partial responses of 14 evaluable); 50% disease control rate (7/14)

NCT06173505



Guidance for Progress Across XmAb® Portfolio Programs in 2024

XmAb Drug Candidate2024 Priority

Solid Tumors: T-Cell Engagers (CD3 & CD28)

XmAb819	ENPP3 x CD3	Advance dose escalation toward target dose levels in 2024
XmAb808	B7-H3 x CD28	Advance dose escalation toward target dose levels in 2024
XmAb541	CLDN6 x CD3	Dose first patient during 1H 2024, enroll Phase 1 study

Immunology

XmAb942		Present preclinical data during UEG Week 2024	\checkmark
	Xtend™ TL1A	Initiate first-in-human Phase 1 study in Q4 2024	
Plamotamab CD20 x CD3		Define clinical development plan	\checkmark
XmAb657	CD19 x CD3	GMP campaign and IND preparation	\checkmark



Potential Inflection Points for Xencor's Clinical Portfolio in 2025

XmAb Drug Candidate		Indication	1H'25	2H'25
Oncology Portfo	olio			
XmAb819	ENPP3 x CD3	ccRCC	Initiation of dose expansion	
XmAb808	B7-H3 x CD28	Solid tumor	Initiation of dose expansion	
XmAb541	CLDN6 x CD3	Ovarian+		Advance toward target dose levels
Vudalimab	PD-1 x CTLA-4	mCRPC	Mono & combo cohort expansion readout	
		NSCLC	Evaluate chemo combination safety	
Immunology Por	rtfolio			
XmAb942	Xtend™ TL1A	IBD+	SAD readout	MAD readout and Phase 2 start
Plamotamab	CD20 x CD3	Rheumatoid arthritis	Initiate Phase 1/2 study	
XmAb657	CD19 x CD3	Autoimmune		Initiate FIH study

As presented 09-Sep-2024 SAD Single ascending dose MAD multiple ascending dose FIH first-in-human



Proteins by Design[®]

XmAb[®] Antibody Therapeutics

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