

UNITED STATES
SECURITIES AND EXCHANGE COMMISSION
WASHINGTON, D.C. 20549

FORM 8-K

CURRENT REPORT
Pursuant to Section 13 or 15(d)
of The Securities Exchange Act of 1934

Date of Report (Date of earliest event reported): September 9, 2024

XENCOR, INC.

(Exact name of registrant as specified in its charter)

Delaware
(State or Other Jurisdiction of
Incorporation)

465 North Halstead Street, Suite 200
Pasadena, California
(Address of Principal Executive Offices)

001-36182
(Commission
File Number)

20-1622502
(IRS Employer
Identification Number)

91107

(Zip Code)

(626) 305-5900

(Registrant's telephone number, including area code)
(Former name or former address, if changed since last report.)

Check the appropriate box below if the Form 8-K filing is intended to simultaneously satisfy the filing obligation of the registrant under any of the following provisions:

- Written communications pursuant to Rule 425 under the Securities Act (17 CFR 230.425)
- Soliciting material pursuant to Rule 14a-12 under the Exchange Act (17 CFR 240.14a-12)
- Pre-commencement communications pursuant to Rule 14d-2(b) under the Exchange Act (17 CFR 240.14d-2(b))
- Pre-commencement communications pursuant to Rule 13e-4(c) under the Exchange Act (17 CFR 240.13e-4(c))

Securities registered pursuant to Section 12(b) of the Act:

Title of Each Class	Trading Symbol(s)	Name of Each Exchange on Which Registered
Common Stock, par value \$0.01 per share	XNCR	Nasdaq Global Market

Indicate by check mark whether the registrant is an emerging growth company as defined in Rule 405 of the Securities Act of 1933 (§230.405 of this chapter) or Rule 12b-2 of the Securities Exchange Act of 1934 (§240.12b-2 of this chapter).

Emerging growth company

If an emerging growth company, indicate by check mark if the registrant has elected not to use the extended transition period for complying with any new or revised financial accounting standards provided pursuant to Section 13(a) of the Exchange Act.

Item 7.01. Regulation FD Disclosure.

On September 9, 2024, Xencor, Inc. (the “Company”) posted a presentation on the “Investors” section of the Company’s website (www.xencor.com), which includes descriptions of initial data from the ongoing Phase 1 dose-escalation study of XmAb819 (ENPP3 x CD3), descriptions of initial data from the ongoing Phase 1 dose-escalation study of XmAb808 (B7-H3 x CD28), and presentations of new programs to be developed for patients with autoimmune diseases and associated near-term clinical timelines. The information contained in, or that can be accessed through, the Company’s website is not a part of this filing. A copy of the presentation is furnished as Exhibit 99.1 to this Current Report on Form 8-K and incorporated herein by reference. The presentation is current as of September 9, 2024 and the Company disclaims any obligation to update this material.

The information furnished under this Item 7.01 of this Current Report on Form 8-K, including Exhibit 99.1 hereto, shall not be deemed “filed” for purposes of Section 18 of the Securities Exchange Act of 1934, as amended (the “Exchange Act”), or incorporated by reference in any filing under the Securities Act of 1933, as amended, or the Exchange Act, except as shall be expressly set forth by specific reference in such filing.

Item 8.01. Other Events.

On September 9, 2024, the Company issued a press release announcing clinical progress updates in early-stage oncology programs, XmAb819 (ENPP3 x CD3) and XmAb808 (B7-H3 x CD28), and XmAb drug candidates to be evaluated for the treatment of patients with autoimmune and inflammatory diseases, including plamotamab (CD20 x CD3), XmAb657 (CD19 x CD3), XmAb942 (Xtend™ TL1A) and the XmAb TL1A x IL-23 program. A copy of the press release is filed as Exhibit 99.2 to this Current Report on Form 8-K and incorporated herein by reference.

Item 9.01. Financial Statements and Exhibits.

(d) Exhibits.

Exhibit No.	Description
99.1	Presentation dated September 9, 2024.
99.2	Press release issued by Xencor, Inc. on September 9, 2024.
104	Cover Page Interactive Data File, formatted in Inline Extensible Business Reporting Language (iXBRL).

SIGNATURES

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned hereunto duly authorized.

Date: September 9, 2024

XENCOR, INC.

By: /s/ Celia Eckert
Celia Eckert
General Counsel & Corporate Secretary

Xencor R&D Strategy Call

September 9, 2024



Today's Agenda

- Overview
- Rationale for bispecific antibodies in autoimmune & inflammatory diseases
- New pipeline programs: B-cell depleting T-cell engagers
 - Plamotamab (CD20 x CD3)
 - XmAb657 (CD19 x CD3)
- New pipeline programs: TL1A portfolio
 - XmAb942 (Xtend™ TL1A)
 - XmAb TL1A x IL-23
- Potential first-in-class T-Cell engagers in solid tumor oncology
 - XmAb819 (ENPP3 x CD3)
 - XmAb808 (B7-H3 x CD28)

Forward Looking Statements

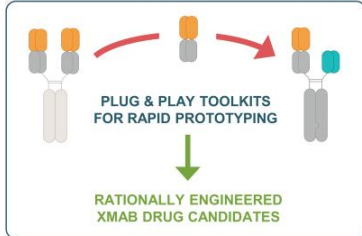
Certain statements contained in this presentation, other than statements of historical fact, may constitute forward-looking statements within the meaning of the Private Securities Litigation Reform Act of 1995. Such statements include, but are not limited to, statements regarding Xencor's development 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. 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.

Disclaimer: We use our trademarks and our logo in this presentation. This presentation also includes trademarks, tradenames and service marks that are the property of other organizations. Solely for convenience, trademarks and tradenames referred to in this presentation appear without the "®" and "™" symbols, but those references are not intended to indicate, in any way, that we will not assert, to the fullest extent under applicable law, our rights or that the applicable owner will not assert its rights, to these trademarks and tradenames.

Proven Power of XmAb® Engineering: Proteins By Design®

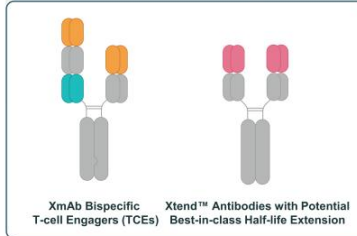
Small changes, big functional impacts

- 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)



Partnerships leverage modular XmAb technology

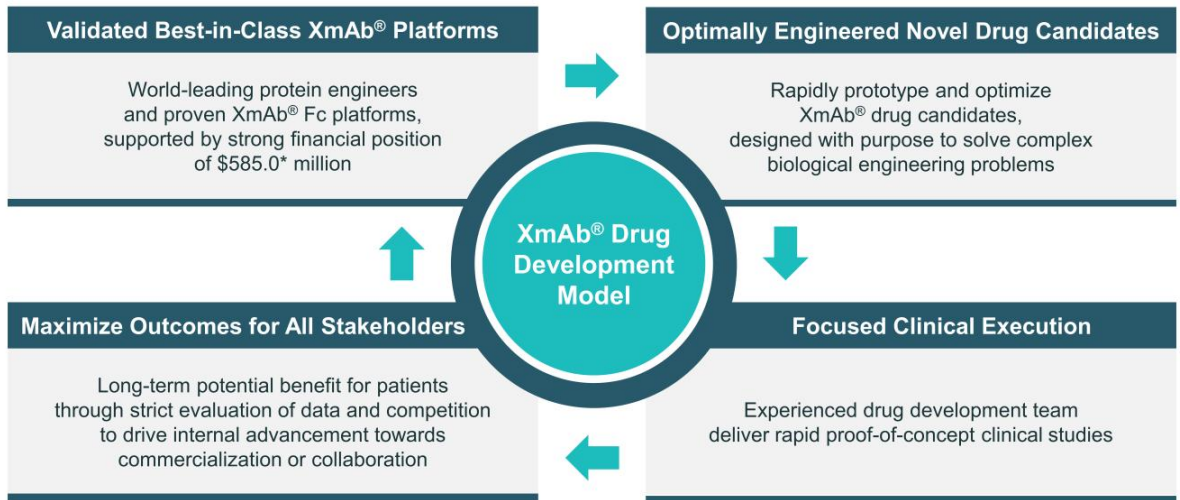
- More than 15 technology license partnerships greatly broadens scope with little-to-no effort
- Multiple commercialized XmAb antibodies

ULTOMIRIS® MONJUVI®/MINJUVI®

COLLABORATION PORTFOLIO INCLUDES



Xencor's Disciplined Drug Development Strategy



* 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
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Solid Tumor Oncology: T-cell Engagers (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

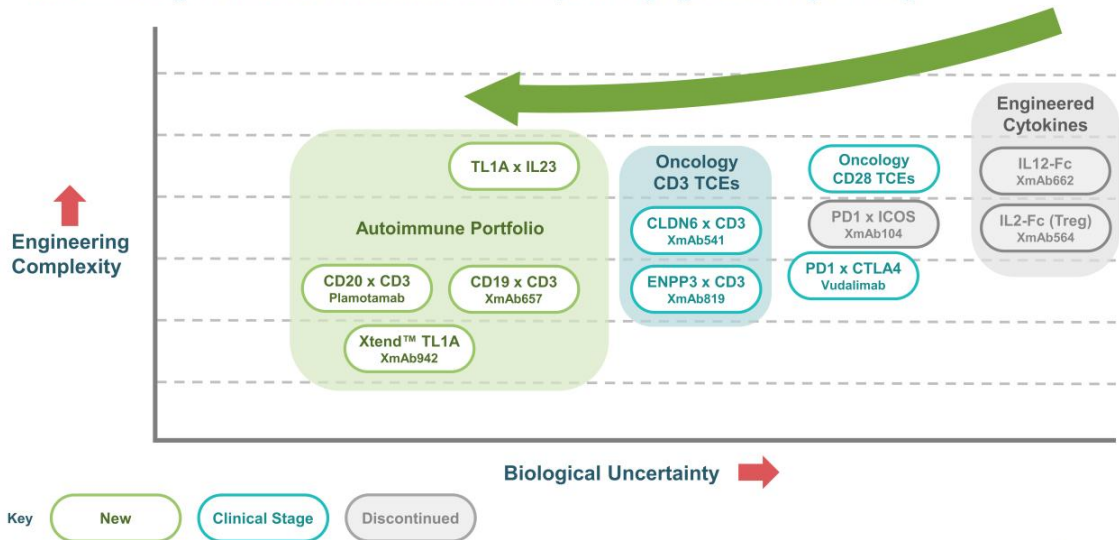
Piamotamab	CD20 x CD3	Bispecific	Rheumatoid Arthritis				1H'25	
XmAb942	TL1A	Xtend, FcKO	Inflammatory Bowel Diseases (IBD)			4Q'24		
XmAb657	CD19 x CD3	2+1 Bispecific, Xtend	Autoimmune Diseases			2H'25		
XmAb Program	TL1A x IL23	Bispecific, Xtend	Autoimmune Diseases					

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

Key Solid tumors Immunology Planned Study Initiation

Rebalanced Portfolio Optimized for XmAb® Drug Development

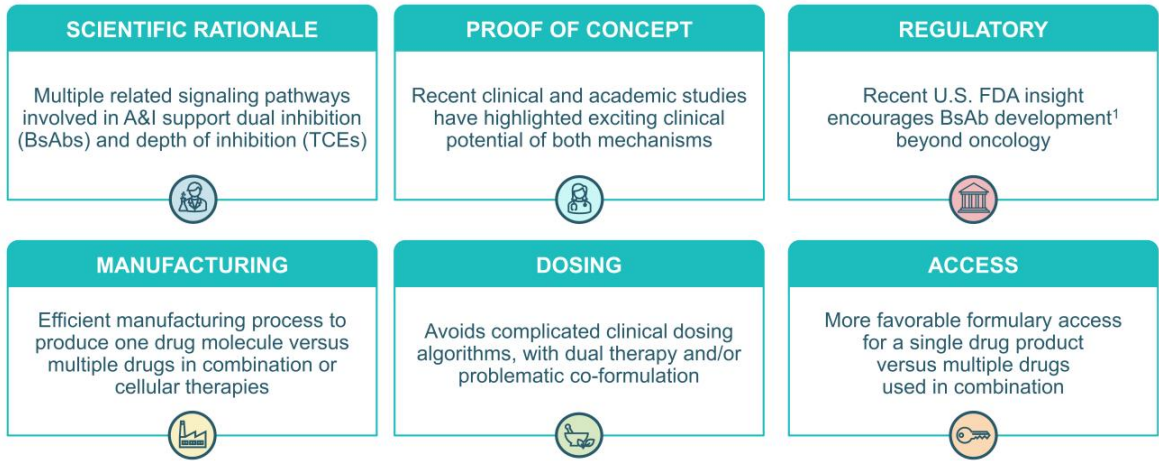
Validated targets across autoimmune disease, leveraging XmAb engineering



**Rationale for bispecific
antibodies in autoimmune and
inflammatory (A&I) diseases**



New Era Emerging for Bispecific Antibody Drug Development in Autoimmune and Inflammatory Diseases



¹BsAb bispecific antibody TCE T-cell engager 1 "The agency has been encouraging drug development in this area. In 2021, FDA finalized a guidance on BsAb development programs." (U.S. FDA, 2024)

Well Validated Targets and Bispecific Antibody Formats Could Enable New Biology to Create Breakthrough Medicines

Newly published data shows potential for multiple types of bispecific antibodies in autoimmune disease

Highly potent B-cell depletion demonstrated promise for patients with severe rheumatic and inflammatory autoimmune disorders in small academic studies, and depth of B-cell depletion has been linked to better clinical outcomes in larger randomized controlled trials

nature medicine

Article <https://doi.org/10.1038/s41591-024-02064-5>

Bispecific T cell engager therapy for refractory rheumatoid arthritis

EULAR 2024

Received: 13 December 2023 | Laura Bucci¹✉, Medańska-Haga, Filippo Pappalardo², Carlo Tu^{3,4}, Arjan Weidman⁵, Inês Phillips, Markus Eckstein⁶, Stefanie Stefan Uderjohr⁷, Alina Re, George Saketani⁸, et al.

Check for updates

Nat Med. 2024

NOBILITY: Exploration of PD-Efficacy Relationship

Sustained B-cell depletion, undetectable CD19, and LLCO by HSP-C1 at Week 24 and Week 52

Undetectable B-cell depletion, detectable CD19, and LLCO by HSP-C1 at Week 24 or Week 52

Patients with sustained B-cell depletion, as measured by HSP-C, were 72% more likely to achieve CRP at Week 76 (12 months after last golimumab dose)

Group	n	Percentage of patients achieving CRP at Week 76
Sustained B-cell depletion	50	~50%
Undetectable B-cell depletion	29	~30%
Patients with B-cell depletion	18	18%

CRP at Week 76 $P < 0.01$

Combination therapy using two approved antibodies showed additive efficacy in Phase 2 in colitis (Janssen) and new real-world multicenter studies

The Lancet Gastroenterology & Hepatology

Volume 9, Issue 4, April 2023, Pages 307-320

Articles

Guselkumab plus golimumab combination therapy versus guselkumab or golimumab monotherapy in patients with ulcerative colitis (VEGA): a randomised, double-blind, phase 2, proof-of-concept trial

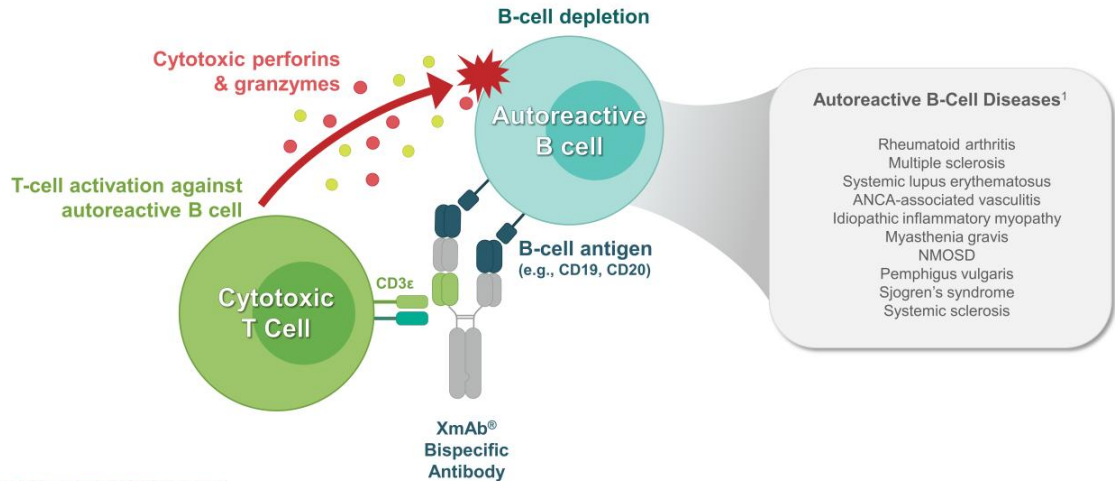
Front. Immunol. 2023

Lancet Gastroenterol Hepatol. 2023

Dual targeted therapy in patients with psoriatic arthritis and spondyloarthritis: a real-world multicenter experience from Spain

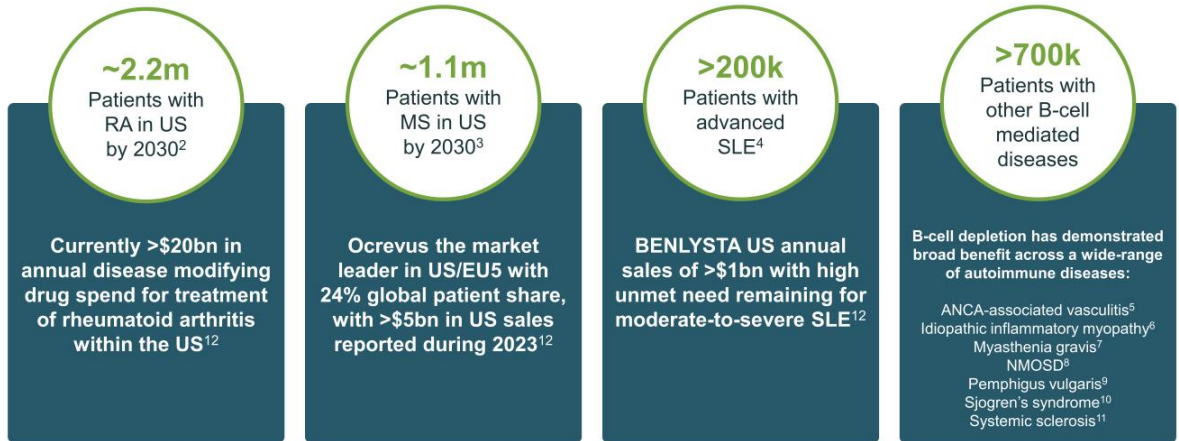
Cristina Valero-Martínez¹, Judit Font Urgelles², Meritxell Sallés³, Beatriz E. Joven-Ibáñez⁴, Alexia de Juanes⁴, Julio Ramírez⁵, Xavier Juanola⁶, Raquel Almodóvar⁷, Ana Laiz⁸, Mireia Moreno⁹, Manel Pujol¹⁰, Emma Beltrán¹¹, José Antonio Pinto-Tasende¹², Laura Crespi¹³, Luis Sala-Icardo¹⁴, Santos Castañeda¹⁵ and Rosario García-Vicuña¹⁶✉

Deep B-Cell Depletion with T-cell Engagers Could Help “Reset” the Immune System for Patients with Autoimmune Disease



¹ Schett G, et al. Ann Rheum Dis 2024;0:1-12.

XmAb® CD20 & CD19 TCEs Can Address Significant Unmet Needs for Autoimmune Disease Responsive to Targeted B-Cell Depletion¹



¹ Based on randomized controlled trials with positive primary endpoints (Schett G, et al. Ann Rheum Dis 2024;0:1-12. ² J Manag Care Spec Pharm. 2018; 24(10):1010-1017. ³ JAMA Neurol. 2023; 80(7):693-701. ⁴ Arthritis Rheumatol. 2021 Jun; 73(6): 991-996. ⁵ J Clin Med. 2022;11(9):2573. ⁶ BMC Musculoskelet Disord. 2012; 13: 103. ⁷ Front Neurol. 2024; 15:1339167. ⁸ Mult Scler. 2024; 13524585231224683. ⁹ JAMA Dermatol. 2019; 155(5): 627-629. ¹⁰ Arthritis Care Res (Hoboken). 2017; 69(10):1612-1616. ¹¹ J Manag Care Spec Pharm. 2020 Dec;26(12):1539-1547. ¹² GlobalData.

**Rheumatoid Arthritis:
Where are we and where do
we need to go?**

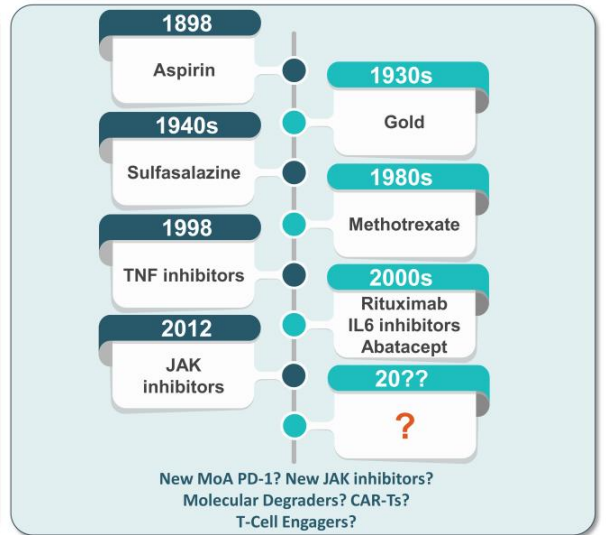
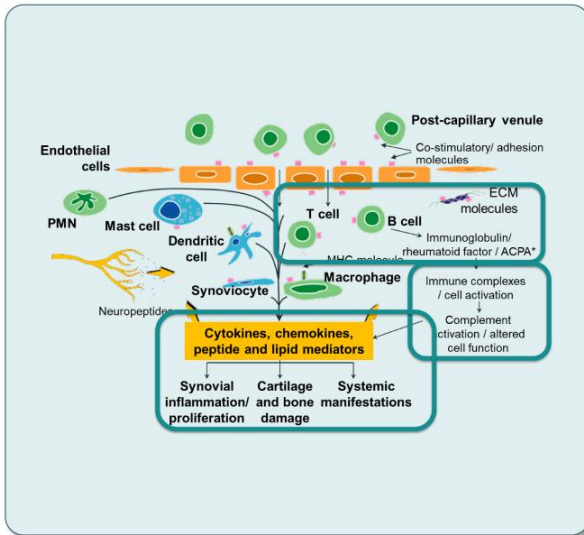
Roy Fleischmann, MD MACR

Adjunct Professor of Medicine
University of Texas Southwestern Medical Center

Medical Director, Metroplex Clinical Research Center
Dallas, Texas



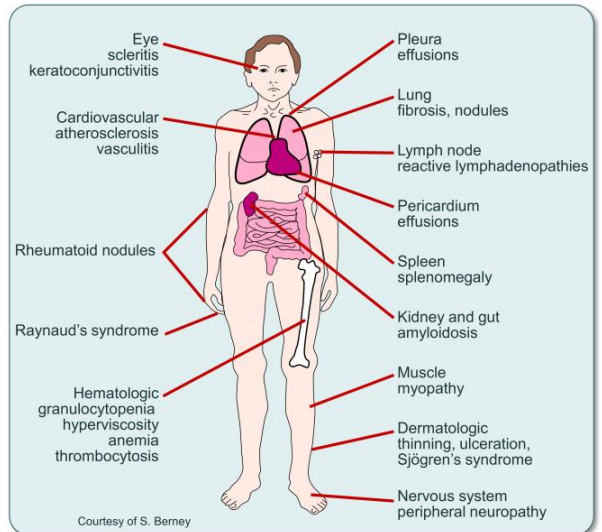
Immunopathogenesis of Rheumatoid Arthritis



*ACPA = anti-citrullinated protein antibody. Detected as anti-CCP Ab. Citrulline is a post-translational modification of arginine, e.g. at inflammatory sites

Rheumatoid Arthritis Manifestations

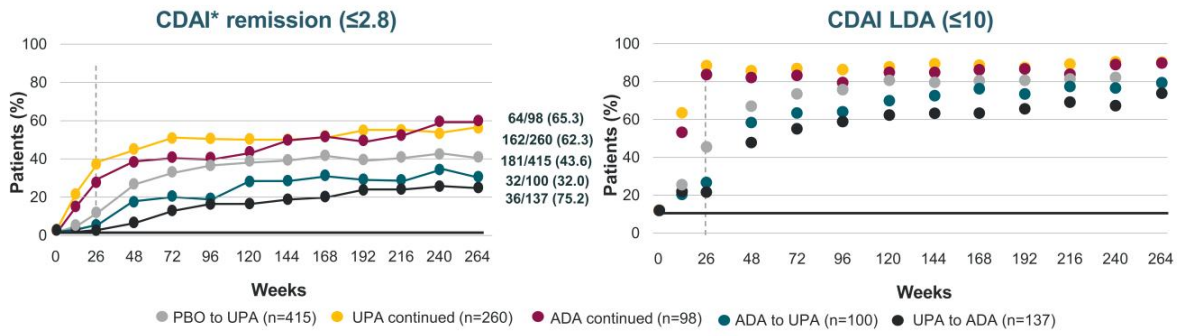
- Systemic, inflammatory polyarthritis that leads to joint destruction, deformity, and loss of function.
- Pathology involves synovial membranes and peri-articular structures of joints, typically resulting in uncontrolled inflammation with pannus formation and clinical symptoms of pain, swelling and stiffness which may lead to irreversible damage and deformity with functional limitation.



Courtesy of S. Berney

ACR Subcommittee on RA Guidelines. Arthritis Rheum. 2002;46:328; Goronzy JJ, Weyand CM. In: Klippel JH, et al, eds. Primer on the Rheumatic Diseases 2001; 12th ed. Atlanta, GA: Arthritis Foundation; 2001. p 209-16. Anderson RJ. *ibid.* 218; Arnett FC, et al. Arthritis Rheum. 1988;31:315

Even with the Most Effective Medications, 40% of RA Patients Do Not Reach Remission Even If They Continue with the Medication (AO)



What degree of disease activity can be reached with effective medications for RA?

SELECT-COMPARE: UPA + MTX vs ADA + MTX. Vertical line at Week 26 indicates the end of the PBO-controlled period. AO As Observed PBO placebo UPA Upadacitinib ADA adalimumab CDAI Clinical Disease Activity Index LDA low disease activity * CDAI remission is a stricter clinical metric than DAS28. Fleischmann R et al. EULAR 2023. Poster POS0849.

Consequences of Inadequately Treated RA

Cardiovascular Disease (CVD)

RA patients have an increased risk of CV events^{1,2}. Risk of CVD death 50% higher vs. general population, which correlates with CV risk factors and inflammation^{3,4}. DMARDs reduce CV event risk^{5,6} if disease activity reduced⁷.

Serious Infection (SIE)

RA is associated with a 2-fold increased risk of SIE, thought to be due to defective immune system and comorbidities such as diabetes, pulmonary or renal disease and functional disability⁸. TNF α inhibitors increase the risk 2-fold and glucocorticoids 4-fold.

Venous and Pulmonary Thrombosis


Active RA is associated with a > 2-fold increase in the development of deep venous thrombosis and pulmonary embolus compared to the general population⁹.

Lymphoma

Severe disease activity in RA patients is correlated with a 70-fold increased risk of developing malignant lymphomas, particularly diffuse large B cell lymphoma¹⁰.

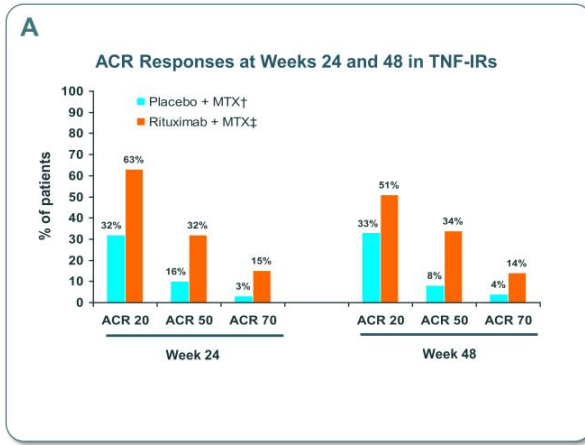
¹ Avina-Zubieta JA, et al. Risk of cardiovascular mortality in patients with rheumatoid arthritis: a meta-analysis of observational studies. *Arthritis Rheum* 2008;59:1690-7. ² Solomon DH, et al. Cardiovascular morbidity and mortality in women diagnosed with rheumatoid arthritis. *Circulation* 2003; 107:1303-7. ³ Dei Rincon I, et al. Escalante A. Association between carotid atherosclerosis and markers of inflammation in rheumatoid arthritis patients and healthy subjects. *Arthritis Rheum* 2002;48:1833-40. ⁴ Myasoedova E, et al. Lipid paradox in rheumatoid arthritis: the impact of serum lipid measures and systemic inflammation on the risk of cardiovascular disease. *Ann Rheum Dis* 2011;70:482-7. ⁵ Micha R, et al. Systematic review and metaanalysis of methotrexate use and risk of cardiovascular disease. *Am J Cardiol* 2011;108:1362-70. ⁶ Barnabe C, et al. Systematic review and metaanalysis: anti-tumor necrosis factor a therapy and cardiovascular events in rheumatoid arthritis. *Arthritis Care Res (Hoboken)* 2011;63:522-9. ⁷ Solomon DH, et al. Disease Activity in Rheumatoid Arthritis and the Risk of Cardiovascular Events. *Arthritis and Rheumatology*. Vol. 67, No. 6, June 2015, pp 1449-55. ⁸ Listing J, et al. *Rheumatology*, 2013 52(1): 53-61. ⁹ Choi HK, et al. *Ann Rheum Dis* 2013;72:1182-87. ¹⁰ Saecklund E, et al. *Arthritis Rheum* 2006;54:692-701.

Unmet Needs in the Treatment of RA

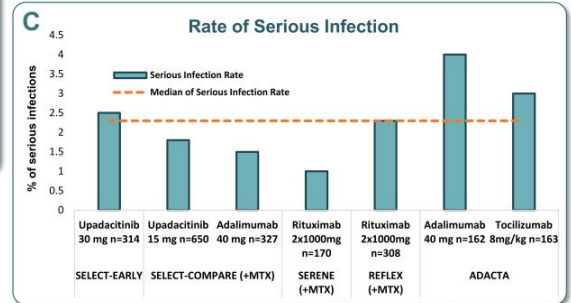
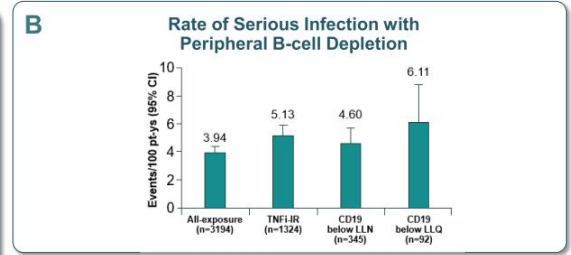
Current Landscape	Trends
<ul style="list-style-type: none">• ~ 1,300,000 patients with RA in the US²  <p data-bbox="448 376 959 443">~15-18% of patients do not respond adequately or are unable to tolerate the currently approved medications¹</p> <ul style="list-style-type: none">• This means that ~ 200,000 - 230,000 patients in the U.S. alone require new therapeutic options• We do not have the necessary tools to predetermine which patient will have a complete clinical response without adverse events to a specific mechanism or action or specific molecule.	<p data-bbox="1050 293 1394 320">Survey of 25 rheumatologists¹:</p> <p data-bbox="1050 322 1394 400">What do they suspect will be the significant changes over the next 5 years in the treatment of RA?</p> <ul style="list-style-type: none">■ Convinced that bDMARDs and Jakinibs should be initiated earlier■ Emerging novel MoA offering improved efficacy, safety and tolerability■ Novel B-cell depleting therapies, CAR-T cell therapy, combination biologics, and more targeted, effective treatment options

¹ Xencor survey of investigators ² Helmick and Lawrence et al: Estimates of the prevalence of arthritis and other rheumatic conditions in the United States, Part I. Arthritis Rheum. 2008;58(1):15-25

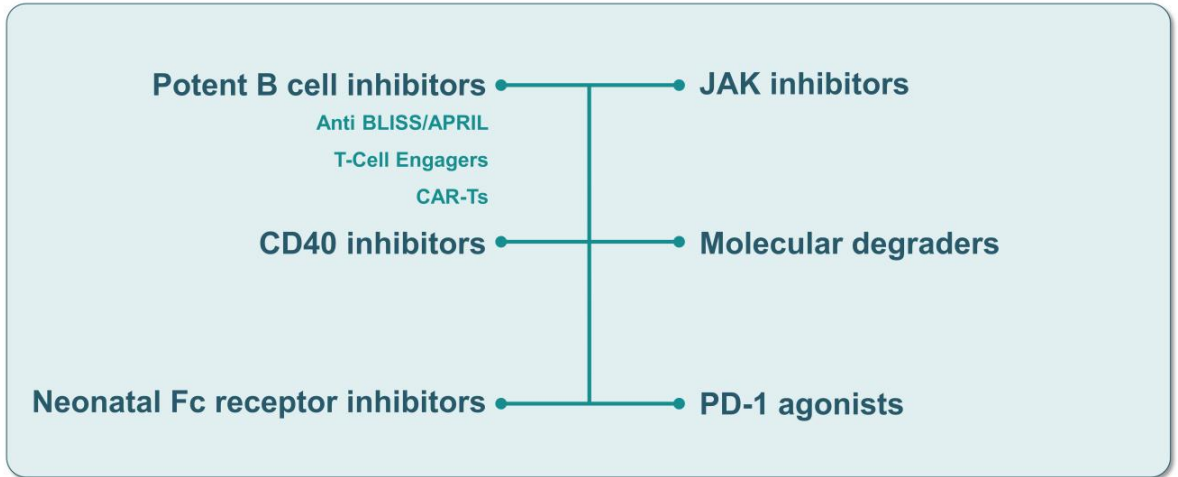
Clinical Experience of Rituximab in RA



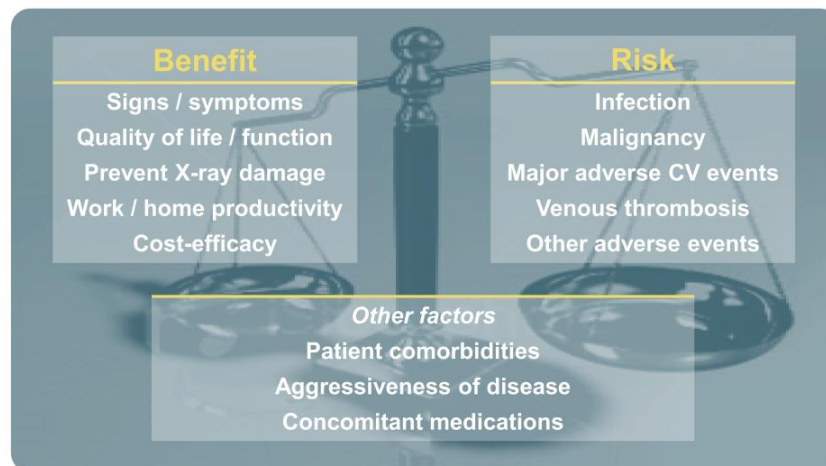
A REFLEX Study; Cohen S, et al. 2006 Arthritis Rheum 54(9): 2793-2806B TNF-IR: TNF inhibitors Inadequate Responders; B Patients with up to 9.5 years of follow-up analyzed 22 years after any RTX treatment for limited return (LLN; <80 cells/ μ L) of CD19 B cells and CD19 cell counts below lower limit of quantification (LLQ; <20 cells/ μ L). CD19 cell counts were measured from peripheral blood; No measurements from other tissue compartments reported; van Vollenhoven, R. et al. 2015 J. Rheum. 42(10):1761-1766 C Cross-trial comparison of serious infection rates (24-week endpoint except for SELECT-COMPARE: 26 weeks).



New Therapies on the Horizon in RA



A Highly Effective Medication with a Risk Profile That Can Be Mitigated, Has a Very Favorable Benefit/Risk Profile for Patients with Rheumatoid Arthritis



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

Plamotamab (CD20 x CD3)

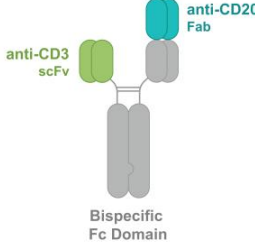
XmAb657 (CD19 x CD3)



Plamotamab

Phase 2 Ready, Subcutaneous CD20 x CD3 BsAb

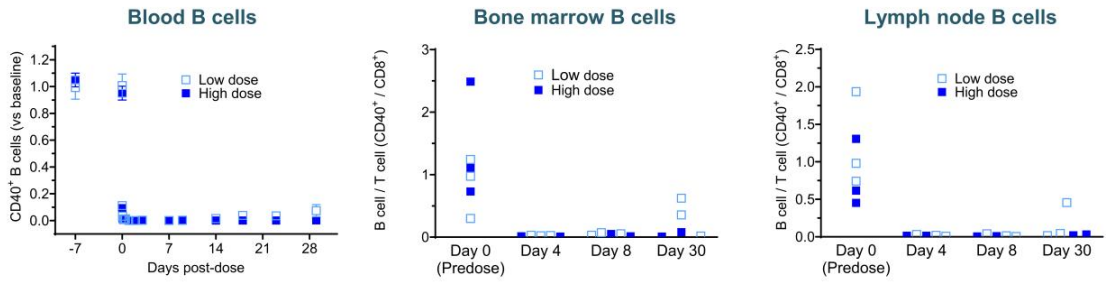
Planned proof-of-concept for the T-cell engager class in autoimmune and inflammatory disease

XmAb [®] CD20 x CD3 Bispecific Design	Positioned for Success
 <ul style="list-style-type: none">• 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	<ul style="list-style-type: none">• 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.

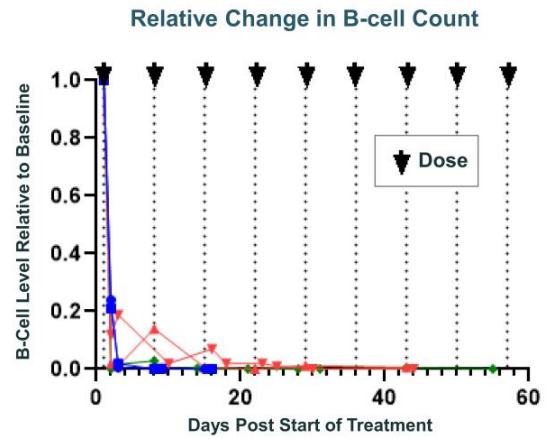
Single Dose of Plamotamab in NHPs

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/ μ L in the blood
- >90% reductions in B-cell count also observed at lower doses



Phase 1 Monotherapy Study of Plamotamab

Heavily Pre-Treated Population with High Rates of Prior CAR-T

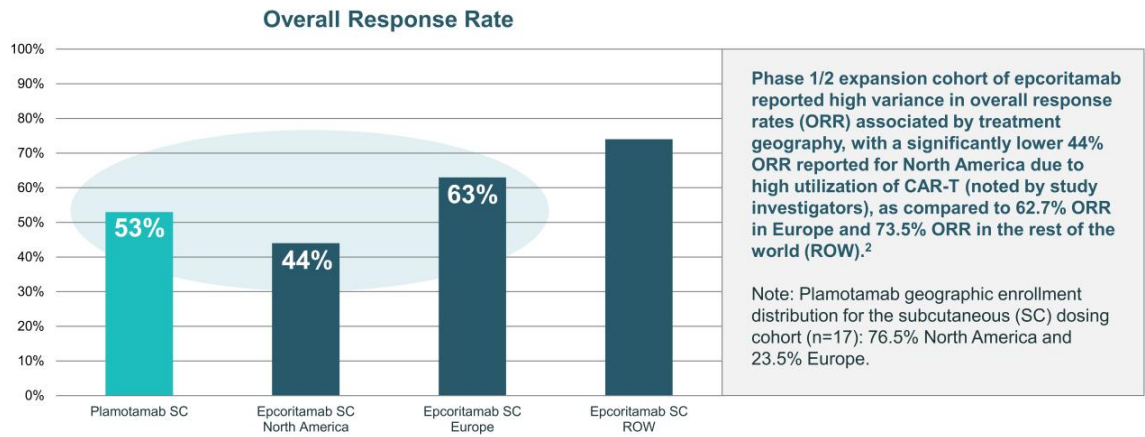
DLBCL + HGBCL Patient Characteristics

Characteristics	RD (IV) (n = 35)	SC all cohorts (n = 20)
Age, median (range)	69 (36-86)	67 (27-90)
Baseline ECOG		
0, n (%)	13 (37.1)	7 (35.0)
1	19 (54.3)	13 (65.0)
2	3 (8.6)	0
Bulky disease at study entry, n (%)		
> 6 cm	10 (28.6)	5 (25.0)
> 10 cm	5 (14.3)	0
Median number of prior systemic therapies	4.0 (2-11)	4.0 (2-10)
Refractory to last therapy, n (%)	25 (71.4)	10 (50.0)
Prior transplantation, n (%)	3 (8.6)	4 (20.0)
Prior CAR-T, n (%)	21 (60.0)	17 (85.0)

RD (IV) recommended IV dose DLBCL diffuse large B-cell lymphoma HGBCL high-grade B-cell lymphoma ECOG Eastern Cooperative Oncology Group

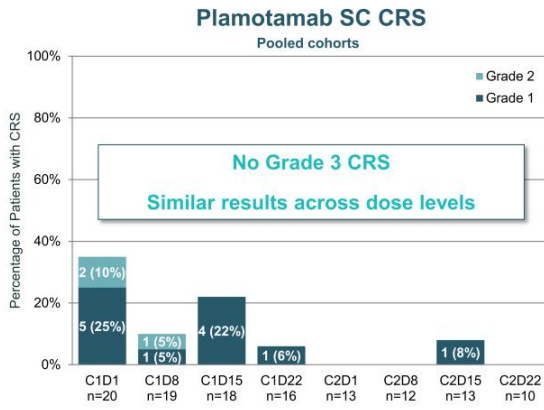
Plamotamab ORR Compared to Commercial CD20 x CD3¹

Regional differences in lymphoma prior therapy markedly impact outcomes

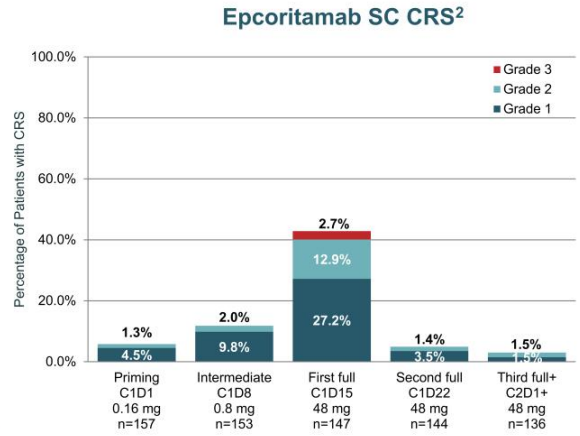


¹ No head-to-head clinical trial has been conducted evaluating plamotamab against epcoritamab. Differences exist between trial design and patient populations, and caution should be exercised when comparing data across unrelated trials. ² Thieblemont and Lugtenburg et al.; J Clin Oncol 41:2238-2247.

No Grade 3 CRS and Lower Grade CRS Observed, Compared to Commercial CD20 x CD3¹



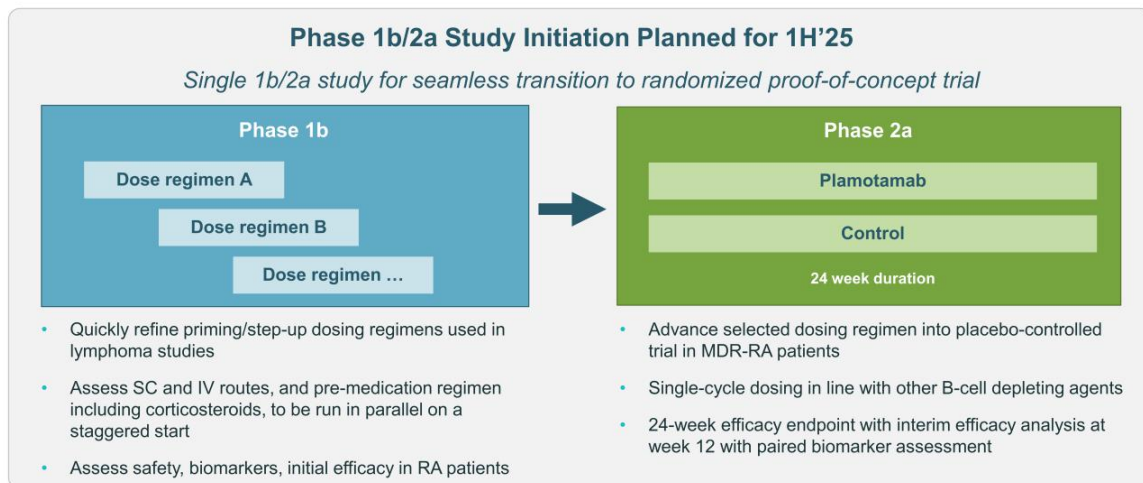
**Summary of CRS at Recommended IV Dose regimen:
< 50% incidence overall, no Grade 3, Cycle 1 limited**



¹ No head-to-head clinical trial has been conducted evaluating plamotamab against epcoritamab. Differences exist between trial design and patient populations, and caution should be exercised when comparing data across unrelated trials. ² Epcoritamab, a Novel, Subcutaneous CD3xCD20 Bispecific T-Cell-Engaging Antibody, in Relapsed or Refractory Large B-Cell Lymphoma: Dose Expansion in a Phase I/II Trial; Thiebtemont and Lugtenburg et al.; J Clin Oncol 41:2238-2247. Data cutoff: January 31, 2022.

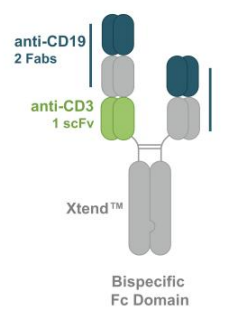
Plamotamab: Plan for Phase 1b/2a Rheumatoid Arthritis Study Start

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



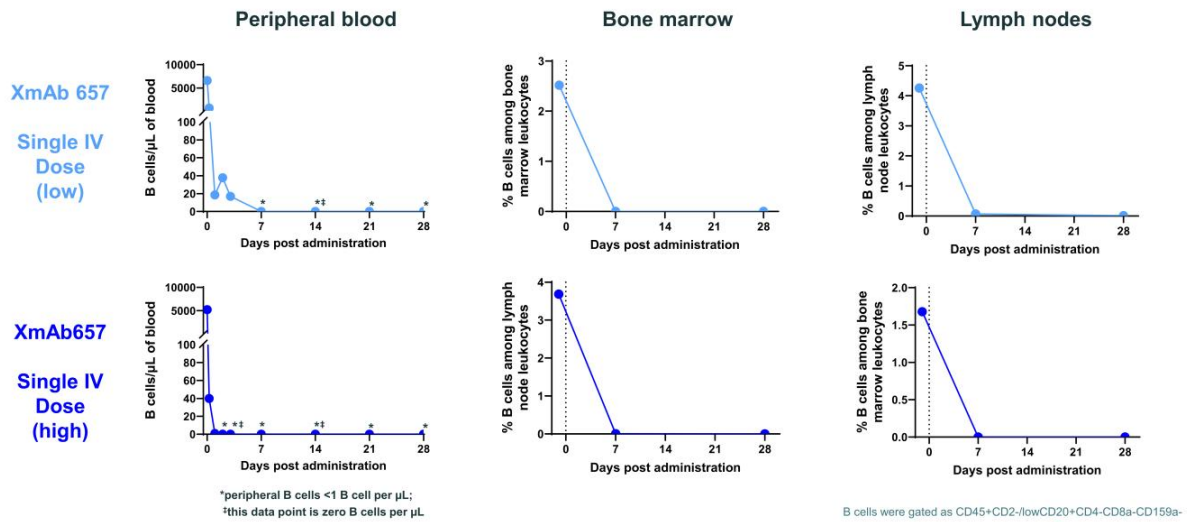
XmAb657

CD19 x CD3 Optimized for Autoimmune Disease

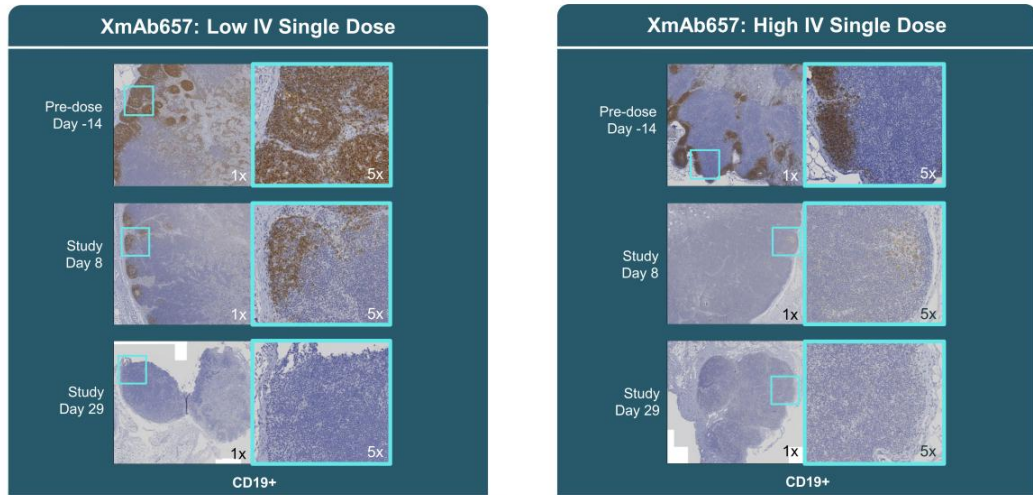
Rational XmAb® Design	Positioned for Success
 <p>The diagram illustrates the structure of XmAb657. It features two anti-CD19 Fab fragments (dark blue) and one anti-CD3 scFv fragment (green) attached to a bispecific Fc domain (grey). The Fc domain is labeled as Xtend™ Bispecific Fc Domain.</p> <ul style="list-style-type: none">• 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	<p>Ongoing NHP studies have shown effective B-cell depletion with single dose</p> <ul style="list-style-type: none">• 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



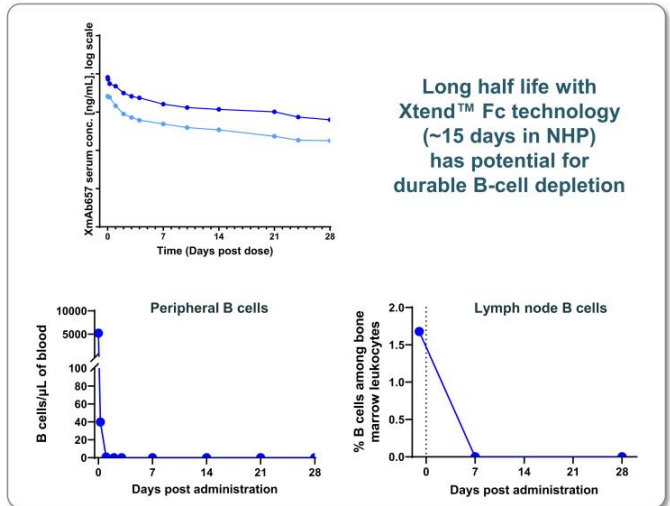
Deep B-Cell Depletion in Lymph Nodes in NHPs Confirmed by CD19+ IHC



CD19 IHC reagent antibody non-interfering with XmAb657

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 TLA1 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

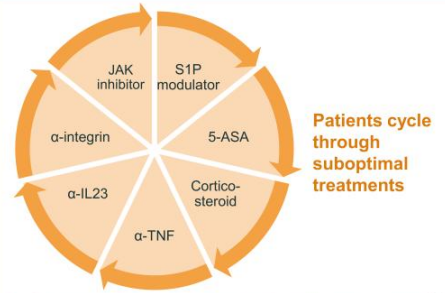
Significant Health Burden

- 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

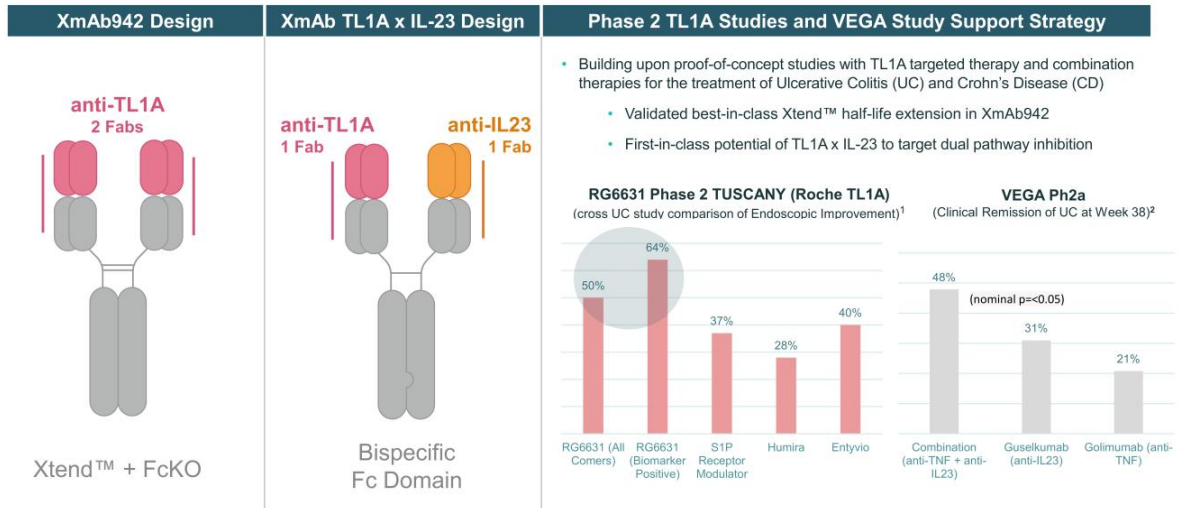
Current Standards of Care are Lacking



- **Suboptimal efficacy:** ~10-20% disease remission³
- **Adverse events:** Infection, malignancy, thromboembolism, cardiac
- **Burdensome regimens:** poor patient compliance

¹ Clarivate ² GlobalData ³ Prescient whitepaper

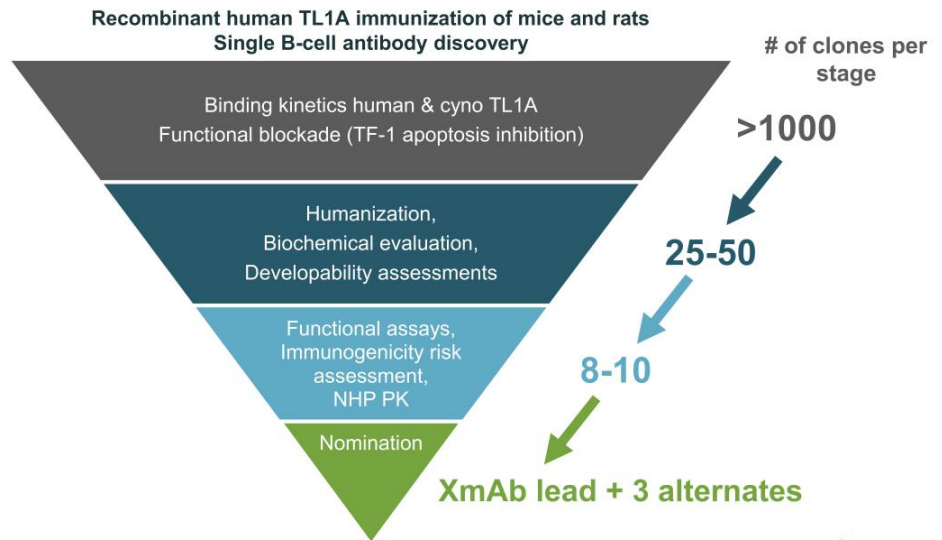
Development of XmAb942 and XmAb TL1A x IL-23 for IBD



¹ 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
² 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

Discovery Campaign for Anti-TL1A Generation

Design of lead and backups in less than 6 months



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 ≥ 21 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 >22 days supports Q8W to Q12W dosing in humans
- High concentration formulation for subcutaneous dosing
- **First-in-human clinical studies to begin 4Q'24**

Discovery & characterization of XmAb942 accepted for presentation during UEG Week on Tues., Oct. 15

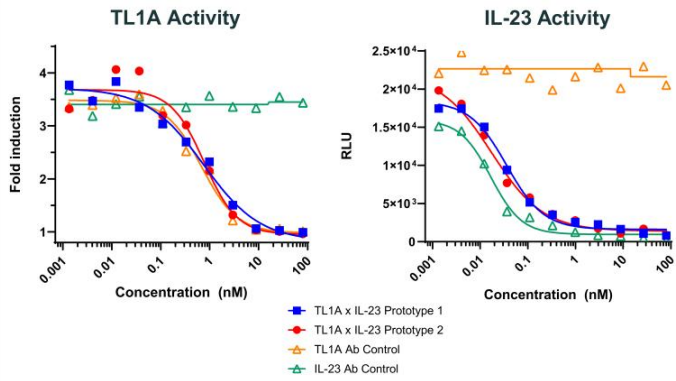
Company	Program ¹	Potent	SC Dosing	Q8-12W Dosing	Half-life extension	Low Immunogenicity
Xencor	XmAb942	✓	✓	✓	✓	✓ Predicted
Merck (Prometheus) ^{2,3}	MK-7240	✗	✓	✗	✗	✓
Roche (Roivant) ^{4,5}	RG-6631	✓	✓	✗	✗	✗
Sanofi (Teva) ⁶	TEV-48574	✓	✓	✗	✗	TBD

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

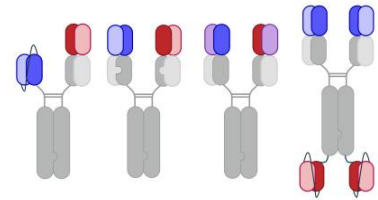
XmAb® TL1A x IL-23 to Have First-in-Class Potential

First-in-Human Study Planned in 2026

Prototype TL1A x IL-23p19 bispecifics are functionally active on both axes



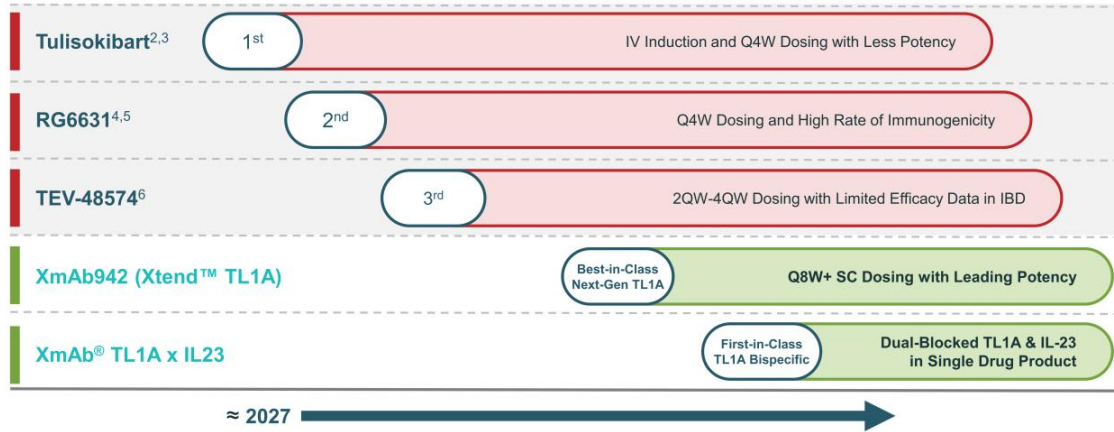
XmAb protein engineering allows for a range of stable molecular structures



- Rapidly prototype in multiple formats
- Optimize and tune for biological activity

Xencor Positioned for Best-in-Class TL1A Portfolio in \$23bn+ Global IBD Market¹

Potential Commercializations for First-Gen Programs and XmAb® TL1A Portfolio



Timelines are illustrative only and subject to FDA approvals ¹ Estimate of US, UK, Spain, Japan, Italy, Germany, France and Canada market size in 2030 (GlobalData) 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: jacc190.0127 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

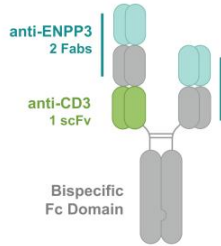
**Potential First-in-Class
T-Cell Engagers
in Solid Tumor Oncology**



XmAb® T-Cell Engager Programs Designed to Address Unmet Need with Potential Across Multiple Tumor Types

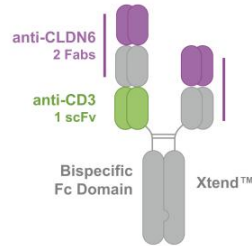
XmAb819 (ENPP3 x CD3)

- Engineered for greater selectivity for ENPP3-expressing tumor cells compared to normal cells, which also express ENPP3 at lower levels
- In development for patients with relapsed/refractory clear cell RCC (ccRCC), which has nearly uniformly high ENPP3 expression
- Dose-escalation ongoing



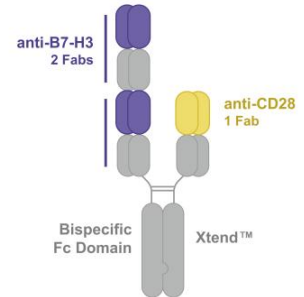
XmAb541 (CLDN6 x CD3)

- Engineered for CLDN6 selectivity over similar CLDN9, CLDN3 and CLDN4
- In development for patients with CLDN6+ tumors, including ovarian cancer
- Dose-escalation ongoing



XmAb808 (B7-H3 x CD28)

- Engineered to provide tumor-selective co-stimulation only when bound to tumor cells
- Combination with anti-PD1 (pembrolizumab)
- In development for patients with solid tumors
- Dose-escalation ongoing



XmAb819 Status Update¹: Encouraging Initial Data in Ongoing Dose Escalation in ccRCC

>\$2bn

peak sales
potential in
ccRCC²

XmAb819

Potential first-in-class
ENPP3 x CD3

Dose escalation on-track
with RECIST responses in
recent dose cohorts

XmAb819 remains on-track to reach target dose levels by year-end

Observed in escalation:

- 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**

¹ Update provided 09-Sep-2024, based on 30-Aug-2024 data cutoff ² Based upon internal Xencor projections of non-risk adjusted peak sales ccRCC clear cell renal cell carcinoma MTD maximum tolerated dose

XmAb808 Status Update¹: Continued Progress in Dose Escalation

>\$3bn
peak sales
potential²

XmAb808

Potential first-in-class
B7-H3 x CD28

Dose escalation on-track with
PSA reductions observed for
patients with mCRPC during
monotherapy run-in period

XmAb808 remains on-track to reach target dose levels by year-end

Observed in escalation:

- Tolerability from recent dose cohorts remains supportive of continued combination with per label dosing of pembrolizumab
- Safety data have supported adding cohorts with Day 1 start for dosing the combination of XmAb808 and pembrolizumab, along with cohorts that use a four-week XmAb808 monotherapy run-in period
- Dose-escalation cohorts continue to enroll patients with multiple tumor types, majority with mCRPC
- For the subgroup of mCRPC patients, biologic activity of XmAb808 has been observed with PSA declines during the four-week monotherapy run-in period, but higher doses are expected to be needed to trigger more meaningful clinical activity
- **Clinical update and dose expansion expected to start during 1H'25**

¹ Update provided 09-Sep-2024, based on 16-Aug-2024 data cutoff ² Based upon internal Xencor projections of non-risk adjusted peak sales mCRPC metastatic castration-resistant prostate cancer

Guidance for Progress Across XmAb® Portfolio Programs in 2024

XmAb Drug Candidate **2024 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	Xtend™ TL1A	Present preclinical data during UEG Week 2024 on October 15	
		Initiate first-in-human Phase 1 study in Q4 2024	
Plamotamab	CD20 x CD3	Define clinical development plan	✓
XmAb657	CD19 x CD3	GMP campaign and IND preparation	✓

Potential Inflection Points for Xencor's Clinical Portfolio in 2025

XmAb Drug Candidate		Indication	1H'25	2H'25
Oncology Portfolio				
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 Portfolio				
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

SAD Single ascending dose MAD multiple ascending dose FIH first-in-human

Xencor R&D Strategy Call

September 9, 2024





Xencor Announces XmAb Drug Candidates in Autoimmune Disease with Near-Term Clinical Plans and Shares Clinical Progress in Early-Stage Oncology Programs

-- Phase 1 healthy volunteer study of half-life extended anti-TL1A antibody XmAb942 to dose first subject in Q4 2024, with data anticipated in the first half of 2025 --

-- XmAb® T-cell engagers plamotamab (CD20 x CD3) and XmAb657 (CD19 x CD3) to be evaluated in autoimmune diseases, with respective Phase 1b/2a and Phase 1 studies to initiate in 2025 --

-- Ongoing Phase 1 dose escalation of XmAb819 (ENPP3 x CD3) in advanced clear cell renal cell carcinoma shows initial encouraging clinical activity including RECIST responses --

-- Management hosting webcast and conference call at 8:00 a.m. ET / 5:00 a.m. PT Today --

PASADENA, Calif.-- Sep. 9, 2024-- Xencor, Inc. (NASDAQ: XNCR), a clinical-stage biopharmaceutical company developing engineered antibodies for the treatment of cancer and other serious diseases, today announced four new XmAb® programs in development for the treatment of patients with autoimmune diseases and provided updates from dose-escalation studies evaluating its first-in-class oncology programs, including XmAb819 (ENPP3 x CD3) in patients with advanced clear cell renal cell carcinoma and XmAb808 (B7-H3 x CD28) in patients with advanced solid tumors.

"Xencor's clinical pipeline of XmAb bispecific T-cell engagers and newly announced autoimmune programs have multiple near-term milestones and offer a balance of opportunities to deliver novel treatment options that could potentially make a real difference in patients' lives. The foundation of our portfolio is world-class protein engineering, using our XmAb platforms to potentially solve complex engineering problems and rationally build drug candidates that address specific clinical opportunities," said Bassil Dahiyat, Ph.D., president and chief executive officer at Xencor. "Our goal is clear—fully leverage our protein engineering strengths and reduce exposure to biological uncertainties to increase our overall opportunities for clinical success."

Clinical Progress Updates in Early-Stage Oncology Programs: XmAb819 (ENPP3 x CD3) and XmAb808 (B7-H3 x CD28)

XmAb819: ENPP3 x CD3 bispecific T-cell engager in Phase 1 dose escalation for patients with advanced clear-cell renal cell carcinoma (ccRCC)

XmAb819 is designed to engage the immune system, activating T cells for highly potent and targeted killing of tumor cells expressing ENPP3, an antigen highly expressed on kidney cancers. Xencor's XmAb 2+1 multivalent format used in XmAb819 enables greater selectivity of ENPP3-expressing tumor cells compared to normal cells, which express lower levels of ENPP3.

- **Clinical update:** Initial evidence of anti-tumor activity has been observed in recent dose-escalation cohorts in the ongoing Phase 1 study, including RECIST responses, and the duration of treatment for several patients in earlier dose cohorts has extended beyond one year. Cytokine release syndrome remains manageable, and the tolerability profile from recent dose cohorts, including no maximum tolerated dose being reached, supports continued dose escalation toward target dose levels.
- **Guidance:** The Company continues to anticipate reaching target dose levels by year end and plans to provide a clinical update around initiation of the first dose expansion cohort during the first half of 2025.

XmAb808: B7-H3 x CD28 bispecific T-cell engager in Phase 1 dose escalation in advanced solid tumors

XmAb808 is a tumor-selective, co-stimulatory CD28 bispecific antibody that binds to the broadly expressed tumor antigen B7-H3 and is constructed with the XmAb 2+1 format. Co-stimulation is required for T cells to achieve full activation, and targeted CD28 bispecific antibodies may provide conditional co-stimulation of T cells when the antibodies are bound to tumor cells.

- **Clinical update:** The majority of patients enrolled into the ongoing Phase 1 dose-escalation study are men with metastatic castration-resistant prostate cancer (mCRPC). In this group of patients, prostate specific antigen (PSA) declines have been observed during the four-week monotherapy safety run-in period. Tolerability from recent dose cohorts remains supportive of continued dose escalation in combination with pembrolizumab.
- **Guidance:** The Company continues to anticipate reaching target dose levels by year end and plans to provide a clinical update around initiation of dose expansion cohorts during the first half of 2025.

XmAb Drug Candidates for the Treatment of Patients with Autoimmune and Inflammatory Diseases and Planned Clinical Studies: Plamotamab (CD20 x CD3), XmAb657 (CD19 x CD3), XmAb942 (Xtend™ TL1A) and the XmAb TL1A x IL-23 Program

Plamotamab: CD20 x CD3 bispecific T-cell engager to be evaluated in patients with multi-drug resistant rheumatoid arthritis (MDR-RA), with Phase 1b/2a study anticipated to initiate in the first half of 2025

Xencor plans to initiate a Phase 1b/2a proof-of-concept study for plamotamab in MDR-RA in the first half of 2025. The Phase 1b portion of the study will select a priming and step-up dose regimen based on the regimen established in oncology, and will assess the initial safety, efficacy, and biomarkers of plamotamab in patients with MDR-RA. The selected dose regimen will then be evaluated in the randomized Phase 2a portion, with efficacy determined at week 24.

Xencor previously completed a Phase 1 clinical study of plamotamab in hematologic cancers, completing enrollment in late 2023. Results from the study showed favorable tolerability and comparable preliminary efficacy data, when cross compared to results from studies of a competitor molecule within the class, with similar patient baseline characteristics. Based on these clinical outcomes, significant B-cell depletion observed in preclinical studies, and the emergent biology supportive of B-cell targeted T cell engagers for the treatment of patients with autoimmune diseases, Xencor plans to evaluate plamotamab in MDR-RA, in which patients progressed through two prior lines of therapy.

XmAb657: Rationally designed CD19 x CD3 bispecific T-cell engager for patients with autoimmune diseases, with first-in-human Phase 1 study anticipated to initiate in the second half of 2025

Xencor has leveraged its XmAb protein engineering platforms to create XmAb657, a potent, potentially long-acting CD19 x CD3 bispecific antibody, utilizing the XmAb 2+1 bispecific antibody format and Xtend™ Fc technology. In non-human primate studies, a single dose of XmAb657 deeply reduced B cells by over 99.98% in the peripheral compartment, bone marrow and lymph nodes, which was sustained for at least 28 days. Half-life was estimated to be 15 days, which indicates a potential for durable B-cell depletion in clinical studies. XmAb657 was well tolerated preclinically, with no clinical signs of cytokine release syndrome. Xencor plans to initiate a first-in-human study during the second half of 2025.

XmAb942: A novel high-affinity anti-TL1A antibody designed for extended half-life, under development for the treatment of inflammatory bowel diseases (IBD), with first-in-human Phase 1 study anticipated to initiate in the fourth quarter 2024

XmAb942 is a monospecific anti-TL1A antibody, utilizing Xencor's Xtend Fc domain and proprietary Fc silencing technology, with potentially class-leading potency, and is under development for patients with IBD. The two most common forms of IBD are Crohn's disease and ulcerative colitis. Half-life preclinically was greater than 22 days, potentially supporting an 8- to 12-week dosing regimen in humans. An abstract with preclinical characterization was accepted for presentation at the United Europe Gastroenterology Week (UEGW) in Vienna, Austria on Tuesday, October 15. Xencor anticipates dosing the first subject in a first-in-human, single-ascending dose study of XmAb942 in the fourth quarter of 2024, with interim data during the first half of 2025.

XmAb TL1A x IL-23 Program: Potential first-in-class bispecific antibody to combine two validated biological pathways of interest into one drug candidate for the treatment of IBD, leveraging Xencor's world-class protein engineering

An expertly engineered XmAb TL1A x IL-23p19 bispecific antibody could potentially provide dual targeting of important inflammatory pathways for autoimmune and inflammatory disease, while avoiding the complexities of dosing and formulary access for two separate TL1A and IL23 targeted drugs. Xencor anticipates initiating first-in-human studies during 2026.

Conference Call and Webcast

Xencor will host a conference call and webcast today at 8:00 a.m. ET (5:00 a.m. PT) to review the topics outlined in this news release.

The live webcast may be accessed through "Events & Presentations" in the Investors section of the Company's website, located at investors.xencor.com. Telephone participants may register to receive a dial-in number and unique passcode that can be used to access the conference call. A recording will be available for at least 30 days.

About Xencor

Xencor is a clinical-stage biopharmaceutical company developing engineered antibodies for the treatment of patients with cancer and other serious diseases. More than 20 candidates engineered with Xencor's XmAb® technology are in clinical development, and multiple XmAb medicines are marketed by partners. Xencor's XmAb engineering technology enables small changes to a proteins structure that result in new mechanisms of therapeutic action. For more information, please visit www.xencor.com.

Forward-Looking Statements

Certain statements contained in this press release may constitute forward-looking statements within the meaning of applicable securities laws. Forward-looking statements include statements that are not purely statements of historical fact, and can generally be identified by the use of words such as "potential," "can," "will," "plan," "may," "could," "would," "expect," "anticipate," "seek," "look forward," "believe," "committed," "investigational," and similar terms, or by express or implied discussions relating to Xencor's business, including, but not limited to, statements regarding expectations for clinical progress, planned presentations of clinical data, new XmAb candidates and programs, planned and in process clinical trials, the quotations from Xencor's president and chief executive officer, and other statements that are not purely statements of historical fact. Such statements are made on the basis of the current beliefs, expectations, and assumptions of the management of Xencor and are subject to significant known and unknown risks, uncertainties and other factors that may cause actual results, performance or achievements and the timing of events to be materially different from those implied by such statements, and therefore these statements should not be read as guarantees of future performance or results. Such risks include, without limitation, the risks associated with the process of discovering, developing, manufacturing and commercializing drugs that are safe and effective for use as human therapeutics and other risks, including the ability of publicly disclosed preliminary clinical trial data to support continued clinical development and regulatory approval for specific treatments, in each case as described in

Xencor's public securities filings. For a discussion of these and other factors, please refer to Xencor's annual report on Form 10-K for the year ended December 31, 2023 as well as Xencor's subsequent filings with the Securities and Exchange Commission. You are cautioned not to place undue reliance on these forward-looking statements, which speak only as of the date hereof. This caution is made under the safe harbor provisions of the Private Securities Litigation Reform Act of 1995, as amended to date. All forward-looking statements are qualified in their entirety by this cautionary statement and Xencor undertakes no obligation to revise or update this press release to reflect events or circumstances after the date hereof, except as required by law.

Contacts

For Investors:

Charles Liles

cliles@xencor.com

(626) 737-8118

For Media:

Cassidy McClain

Inizio Evoke

cassidy.mcclain@inizioevoke.com

(619) 694-6291