Antibodies by Design[™]

XmAb[®] Antibody Therapeutics

Corporate Overview November 2019



Forward-Looking Statements

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Xencor: Engineering Antibody Immune Functions to Make Better Drugs

- XmAb[®] Fc domains: small changes, big functional impacts
 - Augments native immune functions, preserves half-life, stability and production
 - Over 750 issued patents and pending patents worldwide
- · Expansive bispecific antibody oncology pipeline advancing
 - 7 XmAb bispecific antibodies in Phase 1 clinical studies
 - 2 tumor microenvironment activators entered Phase 1 in Q2 2019
 - Novartis co-development and ex-U.S. license for XmAb14045 in Phase 1
 - Amgen's AMG 424 in Phase 1 study in myeloma; IND allowed for AMG 509 in prostate cancer
- Genentech co-development collaboration for novel IL15 cytokines
 - Wide-ranging combination strategy critical to advancing cytokines
 - Retained ability to perform clinical studies with broad spectrum of leading cancer therapies
 - IND submission for XmAb24306 in H2 2019
- XmAb late-stage clinical programs ongoing with partners

morphosys

- Tafasitamab/MOR208: Morphosys guided to BLA submission completed by year-end
- Ultomiris[®] (Alexion) approved in the U.S., Japan and EU for the treatment of adult patients with PNH; in the U.S. for patients with aHUS; additional indications in clinical testing













XmAb[®] Fc Domains

XmAb[®] Fc Domains Shift Focus of Antibody Drug Discovery by Creating New Axes for Differentiation



XmAb[®] Fc Domains Augment Natural Antibody Functions



Additional Fc domains: stability, complement activation

99.5% identical to natural antibody Plug-and-play substitution into any antibody



Development Pipeline Focused on Bispecific Fc Domains

Program (Targets)	Fc Domain	Primary Indication	Preclinical	Phase 1	Phase 2	Phase 3	Commercial Rights
obexelimab (XmAb5871) CD19	Immune Inhibitor	lgG4-RD SLE					☆xencor
XmAb7195 IgE	Immune Inhibitor	Asthma/ allergy					☆xencor
XmAb14045 CD123 x CD3	Bispecific	AML					Xencor Novartis *
XmAb13676 CD20 x CD3	Bispecific	B-cell malignancy					☆xencor
XmAb18087 SSTR2 x CD3	Bispecific	GEP-NET GIST					☆xencor
XmAb20717 PD-1 x CTLA-4	Bispecific Xtend	Oncology					☆xencor
XmAb22841 CTLA-4 x LAG-3	Bispecific Xtend	Oncology					⊘ xencor
XmAb23104 PD-1 x ICOS	Bispecific Xtend	Oncology					☆ xencor
XmAb24306 IL15Rβγ (IL15/IL15Rα-Fc)	Bispecific Xtend	Oncology					Genentech ** A Member of the Roche Group

* Novartis licensed ex-U.S. commercial rights, worldwide co-development

** Co-development with Genentech, Xencor option to co-promote in U.S.



XmAb[®] Fc Domains Have Created Numerous Differentiated Antibodies for Technology Partners

Program	Fc Domain	Indications	Preclinical	Phase 1	Phase 2	Phase 3	Approved	Commercial Rights
Ultomiris ® (ALXN1210)	Xtend™	PNH, aHUS						ALEXION
Tafasitamab (MOR208/XmAb5574)	Cytotoxic	NHL/CLL						morphosys
Talacotuzumab	Cytotoxic	Leukemia						CSL
VRC01LS	Xtend	HIV						
AMG 424 CD38 x CD3	Bispecific	Myeloma						AMGEN
AMG 509 STEAP1 x CD3	2+1 Bispecific	Prostate cancer						AMGEN
XmAb bispecific	Bispecific	Oncology						U NOVARTIS
XmAb bispecific	Bispecific	Oncology						Astellas

Technology licensing expands pipeline with very little opportunity cost



XmAb[®] Bispecific Fc Programs



XmAb[®] Bispecific Fc Domains Retain Beneficial Natural Antibody Properties



Stable cell line expression of XmAb[®]14045 Heterodimer Fc domain thermal stability 96.93% T_m = 72.1● ▲Cp (kcal/mol/●C) T_m = 77.5● A280 (mAU) 3.07% -5 n -10 Temperature (•C)

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Plug-and-play Fc Domain Enables Rapid Pipeline Generation and Prototyping



- Portfolio of CD3 bispecific molecules generated for development
 - Target T cells against tumors
- New oncology programs rapidly prototype different target combinations



Xencor Lead Bispecific Programs: T-Cell Engagement with Tuned Potency and mAb-like PK

XmAb[®]14045 (CD123 x CD3)



Cynomolgus monkey, single IV bolus Profound, sustained basophil depletion



XmAb®13676 (CD20 x CD3)



Cynomolgus monkey, single IV bolus Profound, sustained B-cell depletion



XmAb[®]18087 (SSTR2 x CD3)



huPBMC-SCID mouse xenograft Potent, dose-dependent tumor reduction





XmAb[®]14045: CD123 x CD3 Bispecific Antibody – A Full Length mAb to Be Dosed Intermittently



CD123 (IL-3 receptor α subunit)

- Found on early hematopoietic precursor cells and basophils
- Frequently expressed on hematologic malignancies

XmAb14045

- Stimulates targeted T cell-mediated killing of CD123expressing cells, regardless of T cell antigen specificity
- Ablation of Fc gamma receptor binding removes potential for receptor-mediated crosslinking and activation of T cells
- Fc preserves FcRn affinity for antibody-like half-life
- Does not require a continuous infusion
- Efficiently manufactured using standard antibody production methods

Collaboration with Novartis, U.S. Commercial Rights Retained

- Worldwide 50/50 cost share boosts development resources
- \$325M in milestones remaining, including \$90M in development milestones; low double-digit royalties on ex-U.S. sales



XmAb14045 Ongoing Phase 1 Study in Relapsed/Refractory AML: Initial Dose-escalation Data Presented at ASH 2018

Encouraging clinical activity

- 28% of evaluable patients achieved either complete remission (CR) or CR with incomplete hematologic recovery (CRi) at 2 highest initial doses (1.3 and 2.3 mcg/kg weekly)
- 2 patients with responses bridged to stem cell transplant; additional patient (transplantineligible) has remained in remission for 16+ weeks after discontinuation of therapy
- Dose escalation and dosing optimization continues





XmAb14045 Ongoing Phase 1 Study in Relapsed/Refractory AML: Initial Dose-escalation Data Presented at ASH 2018

Manageable cytokine release syndrome (CRS)

- Enrolled 66 heavily-pretreated patients
 - Median 3 prior therapies
 - 86% refractory to last therapy
 - 53% categorized as adverse risk (ELN 2017)
- CRS most common toxicity (55%)
 - CRS more severe on first dose
 - 6% experienced Grade 3 or 4 CRS
 - 29% experienced AEs within 24 hours consistent with CRS but not reported as such
- No clear evidence of drug-related myelosuppression

100% CRS Grade 1 2 3 4 3(12%) 80% 60% Percent of CRS 16(64%) 40% 6(30%) <mark>2(20%)</mark> 20% 2(11%) 1(9%) 5(25%) 3(25%) 2(11%) 3(12%) 2(20%) 2(16%) 2(11%) 1(9%) 1(5%) 1(4%) 0% C1D1 C1D15 C1D22 C2D1 C2D8 C2D22 C1D8 C2D15 n=25 n=18 n=20 n=18 n=12 n=12 n=11 n=10



CRS severity by infusion (Cohorts 9A-2B)

XmAb[®]13676: CD20 x CD3 Bispecific Antibody: Initial Dose Escalation Data to be Presented at ASH 2019

Encouraging early safety and efficacy data observed at active dose levels

- Phase 1 study initiated in February 2017; data cut off for ASH abstract on June 28, 2019
 - Part A to establish an initial priming dose; Part B to escalate dosing on subsequent doses
- In r/r NHL, 36 patients, median age of 61.5 years, median of 3.5 prior therapies
 - In the efficacy evaluable population and at doses ranging from 80 to 125 µg/kg, objective responses were observed in 33% of patients (n=6/18), including 42% of DLBCL patients (n=5/12, 2 CR)
 - The most common TR-TEAE was fever (n=18, 50%); others include chills (8, 22%), hypotension (7, 19%), neutropenia (7, 19%), thrombocytopenia (7, 19%), anemia (6, 16%)
 - CRS occurred in 42% of patients (n=15/36), and one patient receiving an initial dose of 125 µg/kg experienced Grade 4 CRS; all other CRS events were Grade 1 or 2
 - A priming dose of 45 µg/kg was chosen for Part B
- In r/r CLL, 8 patients, median age of 76 years, median of 4.5 prior therapies
 - 1 CR (Richter transformation) in 5 patients treated at 20 µg/kg, the highest dose administered so far in Part A
 - CRS occurred in 25% of patients (n=2/8), and one patient experienced Grade 3 CRS



Xencor's Dual Checkpoint/Co-Stim Bispecifics are **Designed to Promote Tumor-Selective T Cell Targeting**



Distinct and Novel Mechanisms-of-Action Define Xencor's Growing Oncology Pipeline



XmAb[®]20717

- PD-1 x CTLA-4 bispecific
- Two most validated checkpoint receptors
- Phase 1 study initiated July 2018

XmAb[®]22841

- CTLA-4 x LAG-3 bispecific
- Combinable with anti-PD1
- Triple checkpoint blockade
- Phase 1 study initiated May 2019

XmAb[®]23104

- PD-1 x ICOS bispecific
- Novel checkpoint x co-stim pairing
- Phase 1 study initiated May 2019



Genentech Collaboration Boosts Development Resources for Novel IL15 Cytokine Combinations for Oncology



XmAb24306

- IL15 is a highly active immune signaling protein that stimulates tumor killing NK cells and CD8+ T cells
- IL2 and IL15 share IL2Rβγ receptor interactions, but IL-15 avoids biased T reg activation
- Xencor's IL15 cytokines are built on a heterodimeric Fc domain and have potency tuned to improve therapeutic index, and incorporate Xtend[™] for longer half-life





Extensive clinical strategy to explore numerous combinations with Genentech's leading oncology portfolio

- Two-year research collaboration for IL15 programs
- Xencor retains ability to perform clinical studies, subject to requirements
- Xencor shares in 45% worldwide P&L and development costs; co-promotion option in U.S.
- Genentech receives worldwide commercial license to lead preclinical IL15 cytokine XmAb24306
- \$120M upfront and up to \$160M in XmAb24306 development milestone payments; up to \$180M for each new IL15 program
- IND application for XmAb24306 planned H2 2019







Obexelimab Inhibits Multiple Pathways of B Cells Without Killing B Cells

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Anti-CD19 with **Immune Inhibitor Fc Domain** obexelimab Ag BCR FcγRIIb **CD19 B** Cell

Fcγ**R**IIb binding up by ~400x

Potent suppression of B-cell responses without destroying B cells

- Phase 2 trials in IgG4-Related Disease and SLE
 - Final data on IgG4-RD presented at the American College of Rheumatology (ACR) Meeting in Nov. 2017
 - Topline data from SLE presented at ACR in Oct. 2018
- B-cell inhibition: proven for autoimmune disease
 - B-cell depletion (e.g., Rituxan) RA, MS, others off-label
 - B-cell growth inhibition (Benlysta) Lupus
- Current limitations of B-cell targeting antibodies
 - Tradeoff of potency against long-term B-cell ablation
 - No simple subcutaneous delivery

Obexelimab Target Product Profile

Monoclonal antibody that inhibits B-cell function to treat autoimmune diseases

No long-term immune suppression

Subcutaneous injection, every other week



Data from IgG4-RD Phase 2 Study Show Promising Activity

Obexelimab was well tolerated and showed promising activity in IgG4-RD

- Phase 2 primary objective: to evaluate the effect on the IgG4-RD Responder Index (RI) in patients with active IgG4-RD (proportion of patients with improvement in IgG4-RI at 6 mos
- Corticosteroids were tapered and discontinued in all five patients that were on corticosteroids at first obexelimab dose
- Most frequent AEs were GI infusion-related symptoms
- Plasmablasts decreased by about 70-80%, while B cells decreased by 40-55%, both within 2 weeks
- 14 of 15 patients achieved a decrease of ≥ 5 in the IgG4-RD RI. Initial response to therapy occurred quickly, most within two weeks of first dose
- 12 patients (80%) completed the study. All 12 achieved the primary endpoint of a decrease of IgG4-RD RI of ≥ 2 at Day 169
- Remission (IgG4-RD RI of 0 and no corticosteroids after month 2) was attained in 8 patients at Day 169; 4 others achieved an RI ≤ 4







ACR Meeting, November 2017

Obexelimab Lupus (SLE) Phase 2 Study Design



Primary objective: ability of obexelimab to maintain SLE disease activity improvement

Randomized, double-blinded, placebo-controlled

N = 104 patients, 20 U.S. sites, topline data at ACR meeting in October 2018



Obexelimab Shows Disease Modifying Activity in Randomized Phase 2 Study in SLE

- Positive trend in primary endpoint, proportion of patients without loss of improvement (LOI) at Day 225, not statistically significant
 - LOI: SLEDAI increase ≥4 points or new BILAG A or B score AND physician intent to treat with rescue medication
- Met pre-defined secondary endpoint, time to LOI, 76% improvement in median time to LOI (p=0.025)
- Obexelimab well-tolerated and safety profile is consistent with prior studies
 - Most common AEs were transient, infusion-related
 - No opportunistic infections or deaths reported
 - Low incidence of major organ flares





* Excludes 12 patient discontinuations for reasons other than LOI or toxicity



2019/2020 Milestones and Goals

Trial Initiations / IND Submissions

Initiate Phase 1 study of **XmAb23104** (PD-1 x ICOS) in advanced solid tumors



Initiate Phase 1 study of XmAb22841 (CTLA-4 x LAG-3) in advanced solid tumors

Support Genentech's IND for **XmAb24306** (IL15/IL15R α -Fc targeting IL15R $\beta\gamma$)

Initial Phase 1 Data Readouts

ASH) XmAb13676 (CD20 x CD3) in B cell malignancies at ASH 2019

XmAb18087 (SSTR2 x CD3) in NET and GIST (1H2020)

XmAb20717 (PD-1 x CTLA-4) in multiple solid tumor types (1H2020)

\$620.5 million in cash at September 30, 2019 Runway beyond 2024



Antibodies by Design™

XmAb[®] Antibody Therapeutics

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