

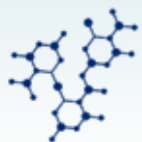


Allergy

CUE-221: Novel dual MOA anti-IgE

Differentiated Science Validated by Consistent Clinical Data

Provides CUE-221 Potential to Outperform Existing And Emerging Anti-IgE Therapies



Functionally distinct dual MOA

- Humanized anti-IgE IgG1 mAb **uniquely engineered by the innovators of omalizumab**
- Unique binding to IgE with distinct IgE conformation
- **Neutralizes free IgE with higher potency than omalizumab and unlike other anti-IgE's, down-regulates IgE synthesis**
- Robust preclinical safety profile

Consistent clinical data reinforce benchmark-setting potential in allergic disease

- **Rapid, durable IgE suppression** below limit of quantification, maintained beyond peak PK
- Supports anti-IgE pharmacodynamic activity in high-IgE patients
- Clinically meaningful improvements in CSU symptom scores
- PD supports dosing \geq QM
- Well tolerated and safe

Advancing toward late-stage value inflection

- **CUE-221 Phase 2 with placebo and active comparator; data readout expected in 2H 2026** (Ascendant CSU Study)
- Testing 3 SC QM doses vs placebo and omalizumab
- Primary Endpoint: HSS7 = 0 at week 12
- Will provide opportunity to further benchmark efficacy and refine dosing schedule

Driving next phase of growth through potential best-in class innovation

- Potential best-in class efficacy will be tested in global P2b study in food allergy
- **Food allergy is an IgE-driven disease, underscoring the potential of a therapy that neutralizes free IgE and blocks new IgE production**
- A second P2b/3 in CSU will follow based on readout from P2 CSU China study



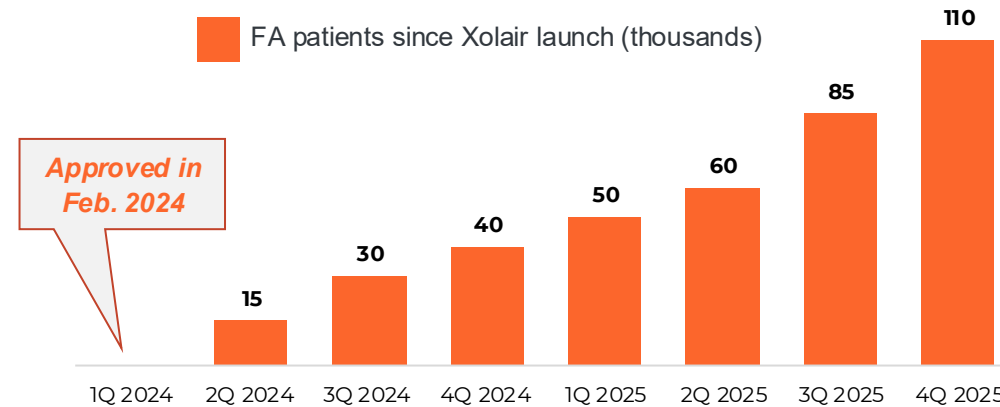
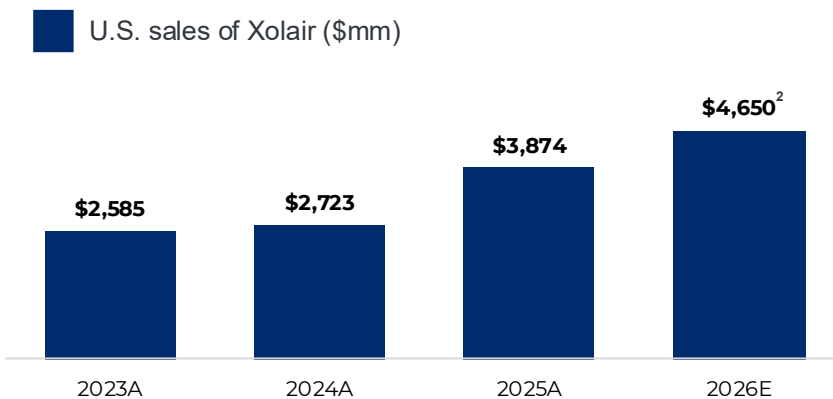
There is a Massive Unmet Need in Our Global Beachhead Indication: Food Allergy

377%

Claim lines with diagnoses of anaphylactic food reactions increased 377 percent between 2007 and 2016



AFTER FDA APPROVAL FOR FOOD ALLERGIES IN EARLY 2024, XOLAIR¹ SALES ARE INFLECTING



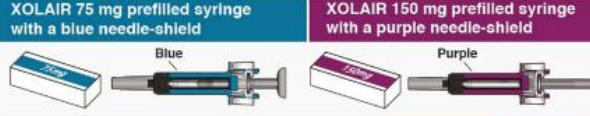
Source: Wall Street equity research; Food Allergy Research & Education (FARE), "Food Allergy Facts and Statistics," accessed 04/06/2026
 Note: FA = food allergy; ¹ Xolair is current standard of care. ² Roche provided verbal guidance of 20% growth for Xolair in 2026 during the FY-2025 earnings call














Omalizumab and an Omalizumab Biosimilar are the Only FDA-Approved Anti-IgE Therapies ^{1,2}

Significant Unmet Need Remains

- **Omalizumab works by neutralizing IgE¹**
 - In IgE mediated food allergy, patients are dosed every 2 to 4 weeks and may require up to 4 injections
 - Patients with high baseline IgE levels > 1850 IU/mL are not eligible for treatment¹
- **Other emerging anti-IgE therapies appear to only offer incremental improvements**
 - Longer half-life which may improve convenience, not efficacy³
 - Novel mechanisms of action that are still in early clinical development, with clinical validation yet to be established⁴

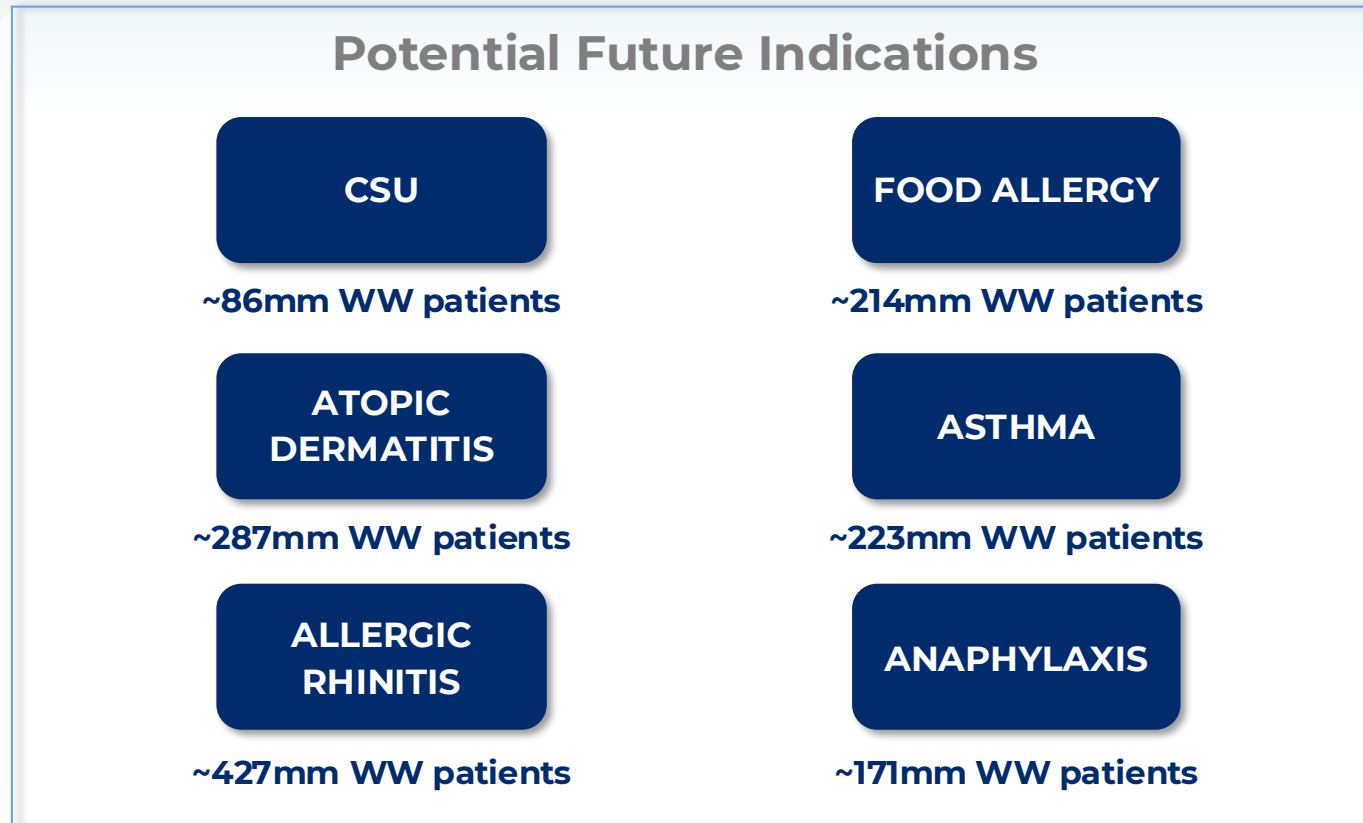
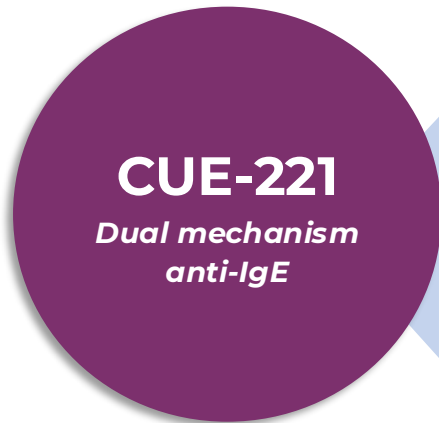


Dose	Syringes needed for the dose	75 mg	150 mg
75 mg	1 blue (75 mg)		
150 mg	1 purple (150 mg)		
225 mg	1 blue (75 mg) + 1 purple (150 mg)		
300 mg	2 purple (150 mg)		
375 mg	1 blue (75 mg) + 2 purple (150 mg)		
450 mg	3 purple (150 mg)		
525 mg	1 blue (75 mg) + 3 purple (150 mg)		
600 mg	4 purple (150 mg)		

Source: 1. Xolair (omalizumab) US Prescribing Information, 2. OMLYCLO (omalizumab-igec) US Prescribing Information 2025; 3. Shen et al Clinical and Translational Science, 2025; 4. Becklund et al. J Allergy Clin Immunol, 2022 and Brigger et al. J Allergy Clin Immunol 2025



CUE-221 is a Functionally Distinct Novel Dual MOA Anti-IgE Therapy, with Pipeline-in-a-Product Potential



Significant unmet need remains across the large opportunities in allergic disease

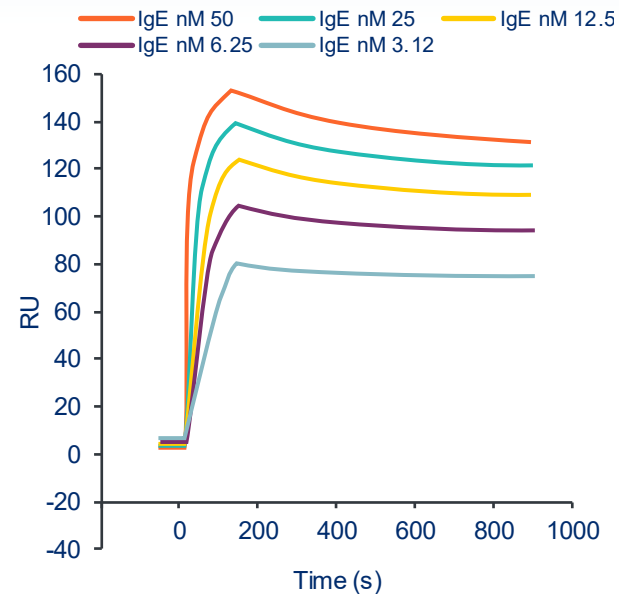
Source: Company filings, Wall Street equity research, World Health Organization, United Nations Department of Economic and Social Affairs, and academic research Note: Patient populations and TAM figures are WW estimates for 2030; IgE = Immunoglobulin E; CSU = chronic spontaneous urticaria



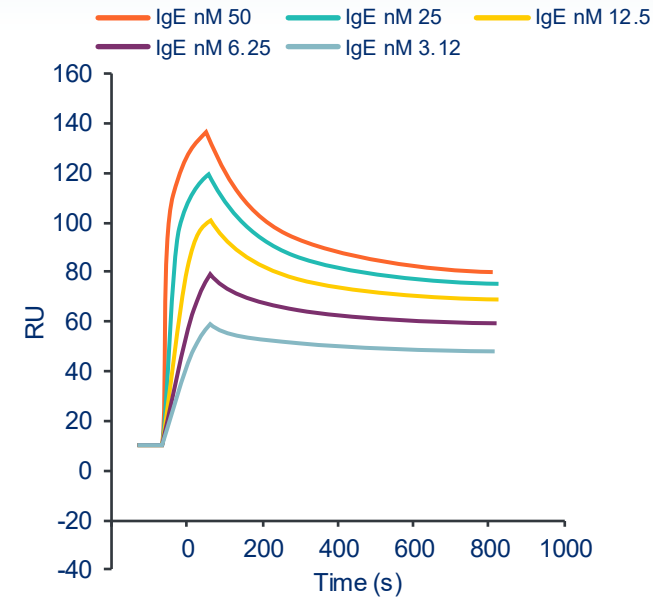
CUE-221: Designed for Deeper and Durable IgE Suppression

Higher binding affinity (picomolar range) and slower dissociation (off-rate) than omalizumab

CUE-221



OMALIZUMAB



- Humanized anti-IgG1 mAb engineered by the innovators of omalizumab
- Binds IgE at distinct epitopes, inducing a unique conformational change versus existing and emerging anti-IgE therapies
- Combines allosteric and orthosteric inhibition of FcεRI binding
- 4 to 8-fold higher IgE binding affinity and slower dissociation (off-rate) than omalizumab

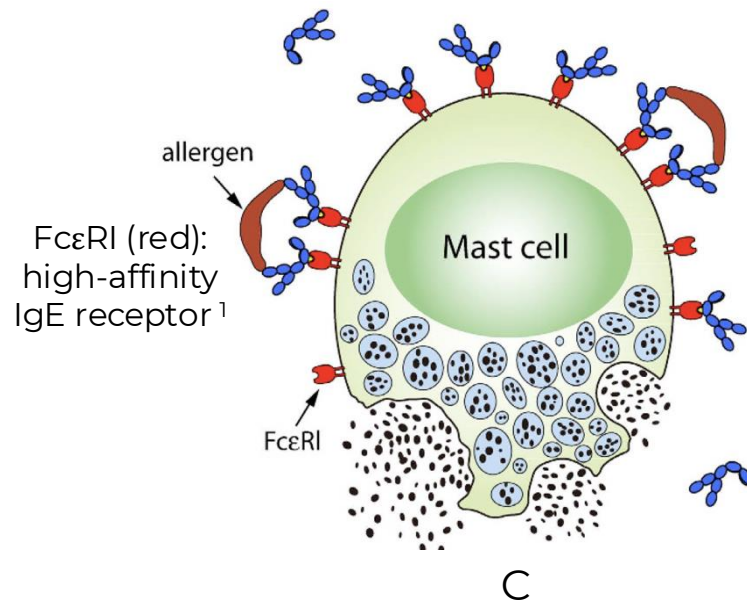


IgE Drives Allergic Disease by Engaging Two Critical Receptors

FcεRI and CD23 (FcεRII) regulate the allergic response

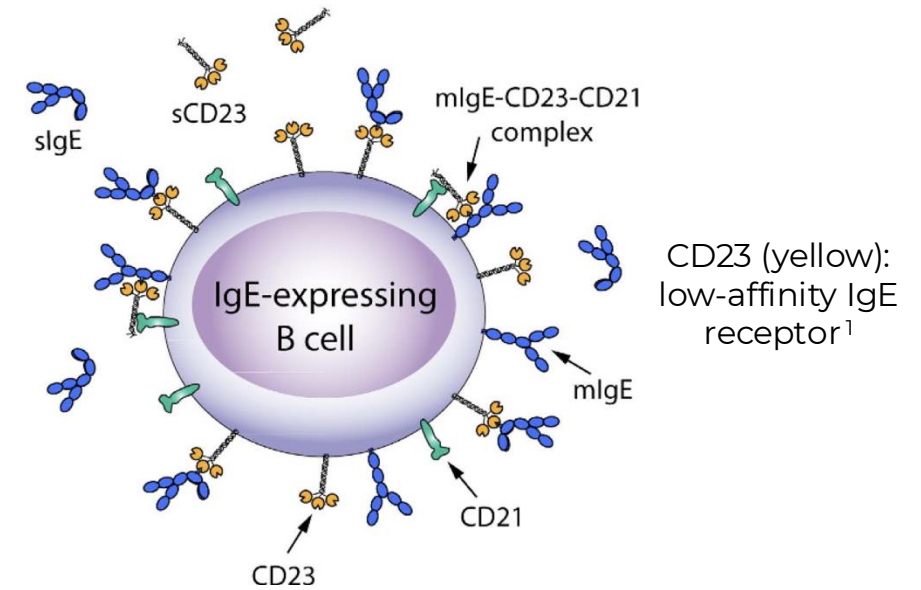
Cross-linking the FcεRI receptor leads to allergic reaction

IgE binds to FcεRI receptor on the surface of mast cells



Membrane bound CD23 suppresses IgE production when bound by IgE

CD23 receptor regulates IgE synthesis²



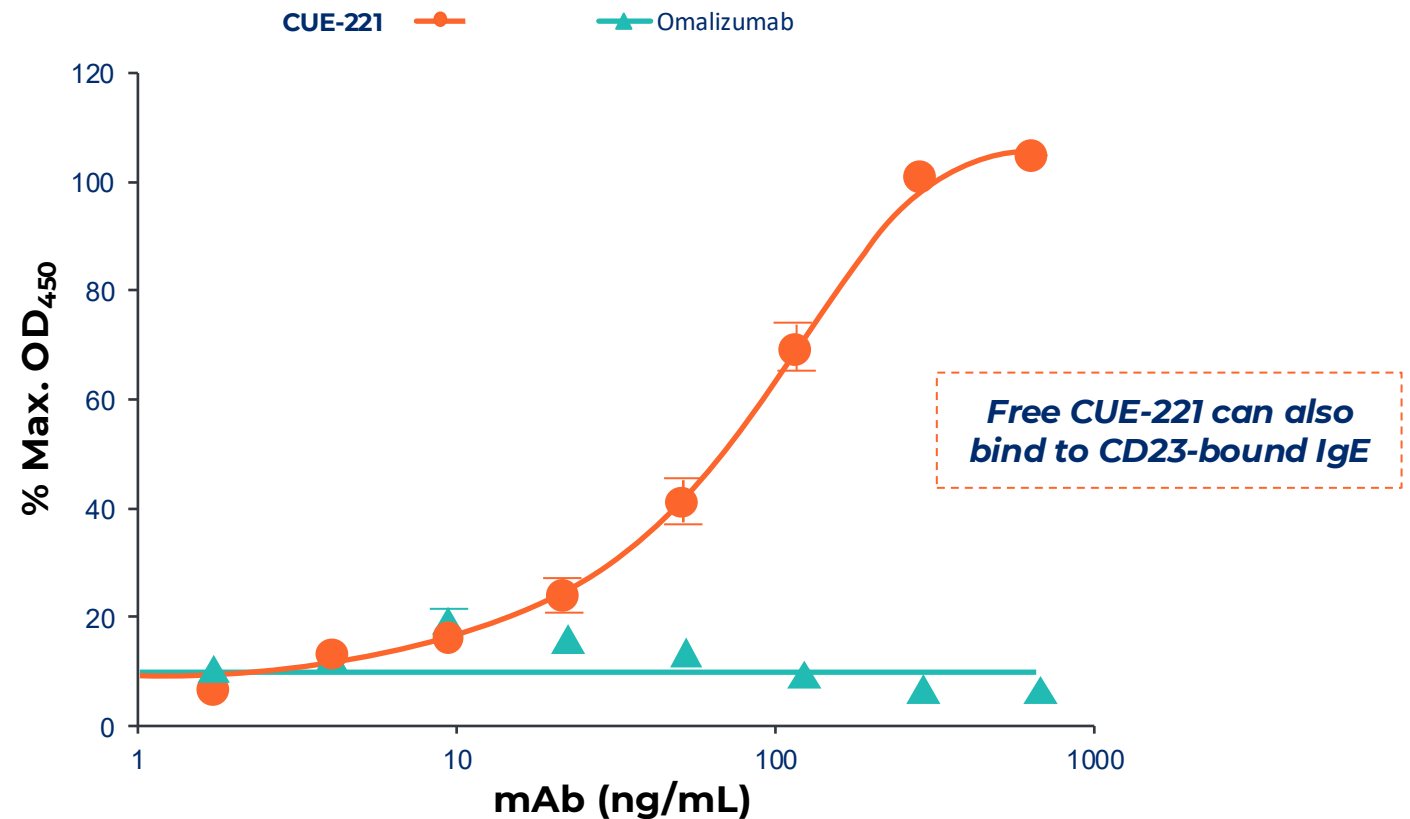
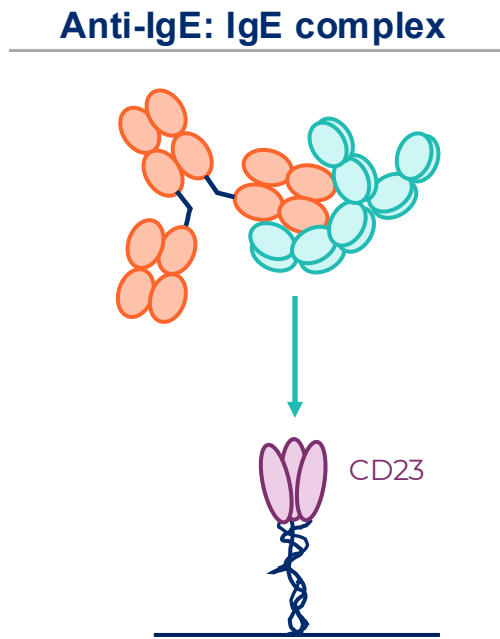
Soluble CD23 promotes IgE synthesis

Source: ¹Wright et al 2015 Nature Scientific Reports; ²Engeroff and Vogel 2020 Allergy



CUE-221's Differentiated IgE Binding Enables Distinct Functional Properties, Including Preserved Interaction with CD23-Bound IgE

CUE-221 does not block IgE binding to CD23, preserving CD23's natural regulatory function in high-IgE states



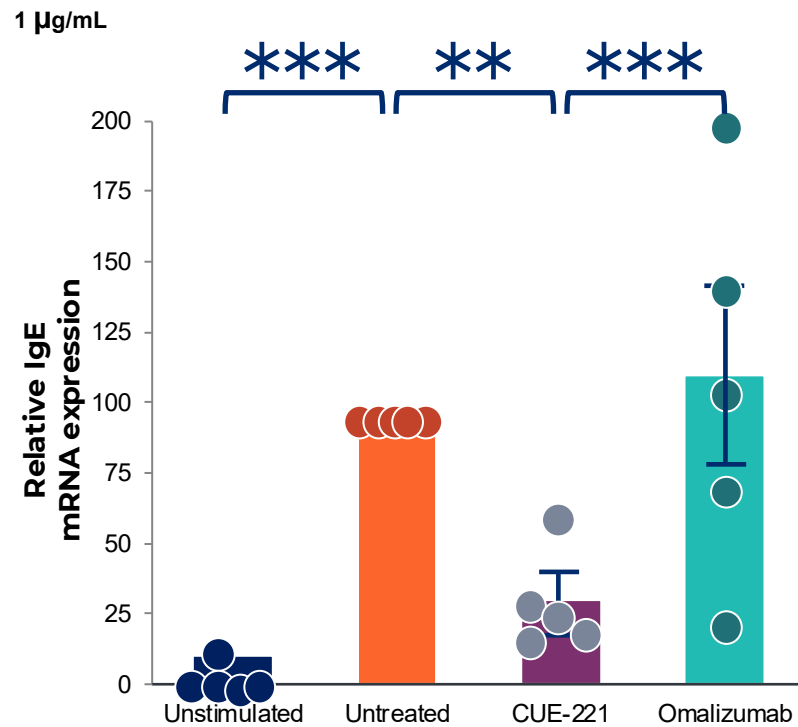
CD23-immobilized ELISA with IgE preloaded



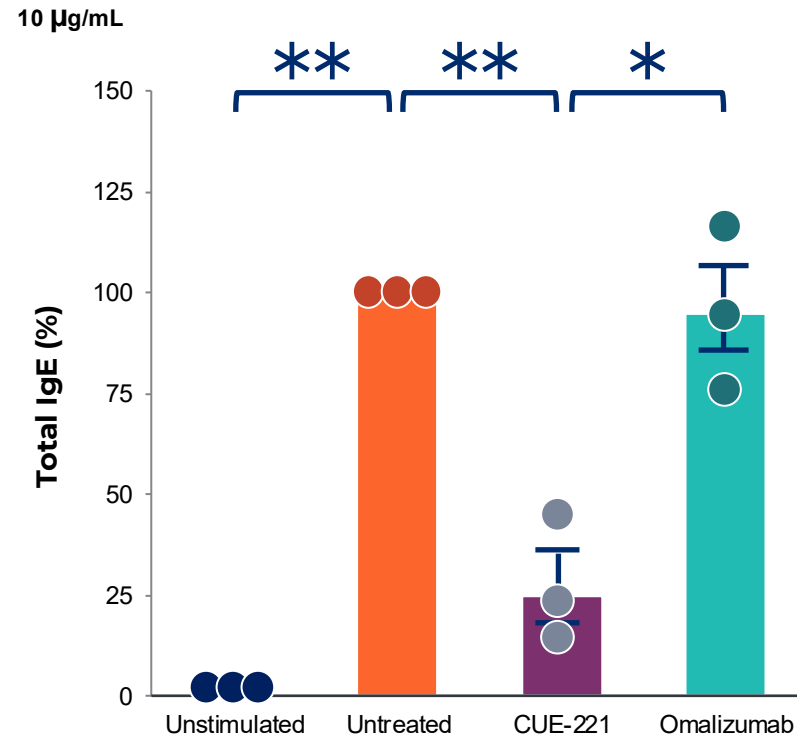
CD23 Acts as a Natural Brake on IgE Production in High IgE States

CUE-221 leverages the CD23-mediated IgE downregulation pathway, suppressing IgE mRNA and protein synthesis¹

IgE mRNA



IgE protein synthesis



Source: Adapted from Kou et al. J Clin Invest. 2022

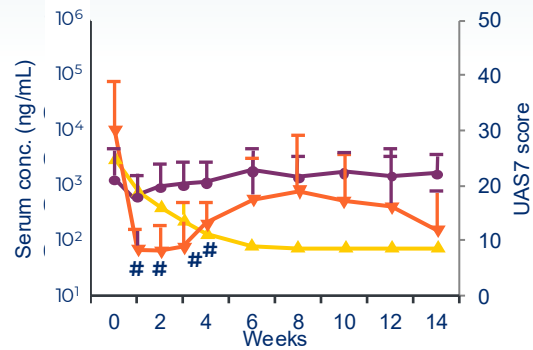
Note: ¹Human PBMCs were stimulated with IL-4 and anti-CD40 to induce IgE production, treated with mAbs for 11 days; Total IgE (by ELISA) and IgE mRNA levels (by real-time PCR; data not shown) measured relative to untreated controls; The unstimulated group corresponds PBMCs in which NO IL-4 and NO anti-CD40 was added, and thus is the background IgE (low/undetectable); The untreated group corresponds to a group that was stimulated with IL-4 + anti-CD40 but NO anti-IgE antibody added. This group produces a full IgE response and is used as the reference/baseline (100%).



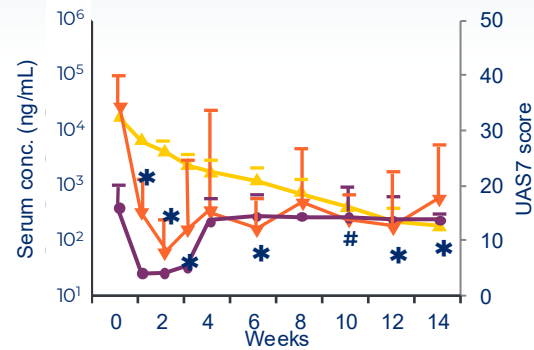
P1 SAD: Rapid and Prolonged Single-Dose PD Effect

Consistent with Dual IgE Neutralization and Down-regulation of Synthesis

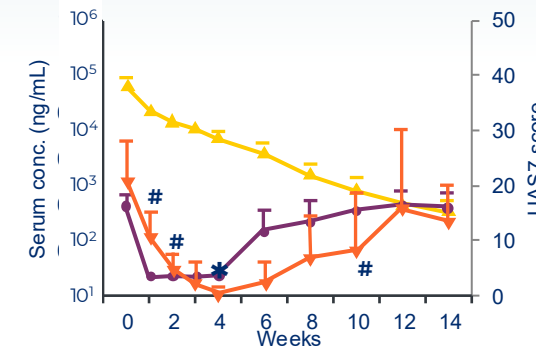
COHORT 1 (0.2 mg/kg)



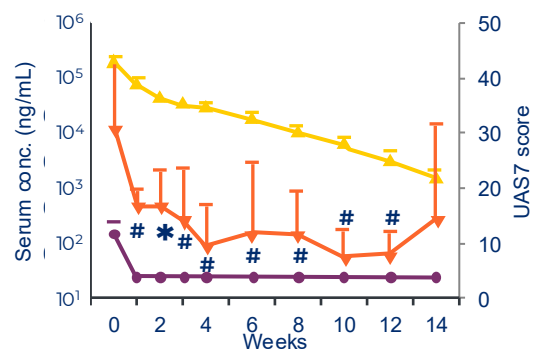
COHORT 2 (0.6 mg/kg)



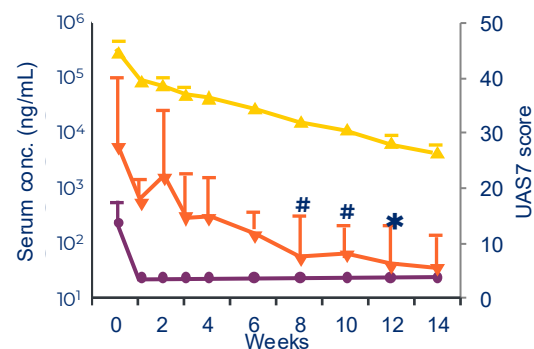
COHORT 3 (2 mg/kg)



COHORT 4 (6 mg/kg)



COHORT 5 (10 mg/kg)



▲ CUE-221 conc.
● Free IgE conc.
▼ UAS7 score

N = 15; 3 subjects per cohort

Durable reduction in free IgE beyond peak PK; improvement in symptom scores



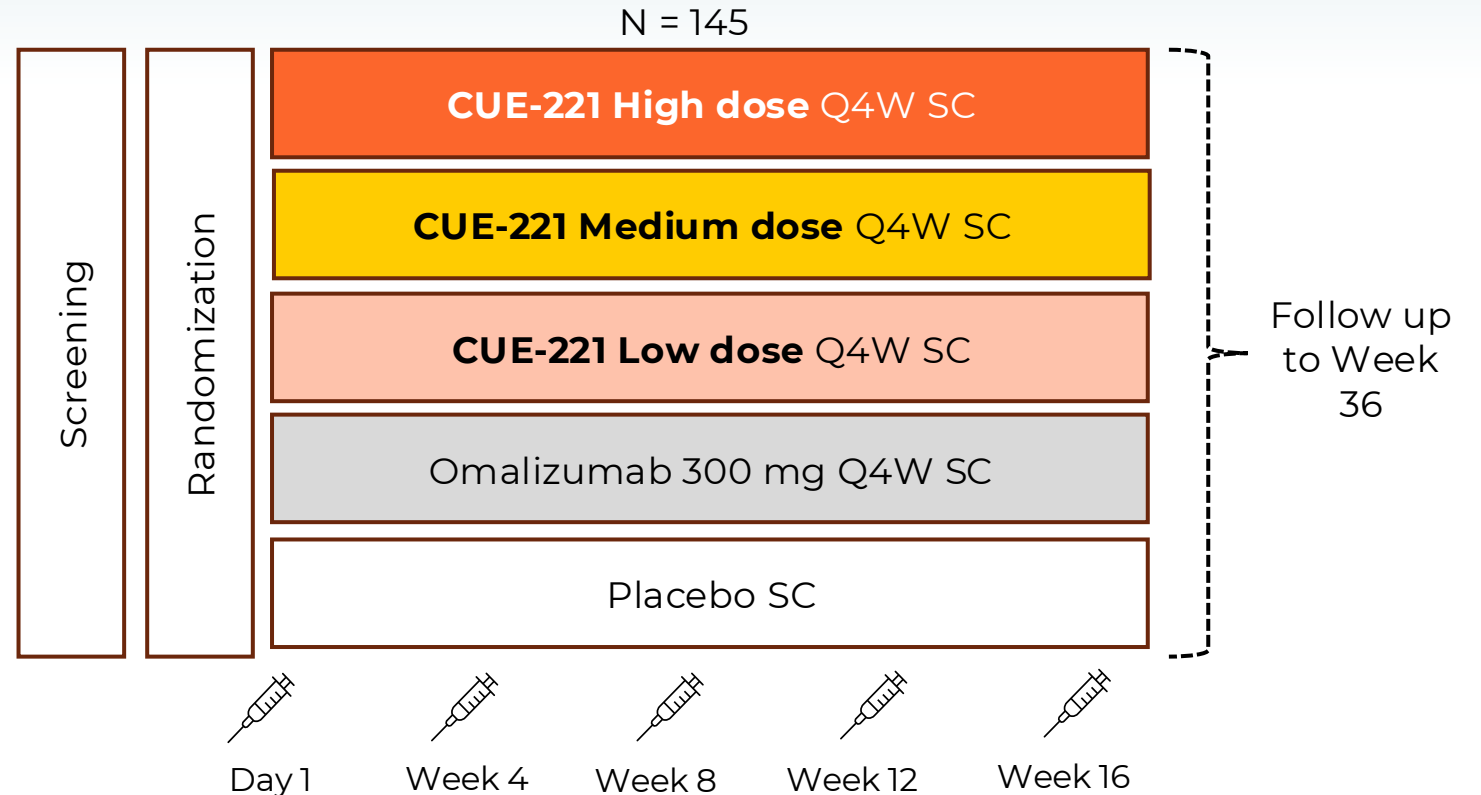
Potential for a Significant Value Inflection with Phase 2 CSU Study Results in 2H26

Phase 2 Study in China with placebo and active comparator (Ascendant CSU Study)

Will provide opportunity to further benchmark efficacy and refine dosing schedule for future global Phase 2 studies

Source: Data on File; CSU = chronic spontaneous urticaria; SC = subcutaneous; HSS7 = Hives severity score over 7 days

Phase 2 CSU Study Design



★ **Primary Endpoint:**
Percentage of patients with HSS7=0 at W12

