

R&D Spotlight: CLYM116 Initial Phase 1 Results

September 3, 2026



Forward Looking Statements

This presentation contains “forward-looking statements”, including without limitation statements regarding: future expectations, plans and prospects for Climb Bio; expectations regarding the therapeutic benefits, clinical potential and clinical development of CLYM116; the anticipated timelines for reporting clinical data from Climb Bio’s ongoing and planned clinical trials of CLYM116; the potential commercial opportunity and limited competitive landscape and CLYM116 in IgA nephropathy (IgAN); projections and estimates derived from pharmacokinetic and pharmacodynamic modeling; the sufficiency of Climb Bio’s cash resources for the period anticipated; and other statements containing the words “anticipate,” “believe,” “continue,” “could,” “estimate,” “expect,” “intend,” “may,” “plan,” “potential,” “predict,” “project,” “should,” “target,” “would,” “will,” “working” and similar expressions. Forward-looking statements are based on management’s current expectations of future events and are subject to a number of risks and uncertainties that could cause actual results to differ materially and adversely from those set forth in, or implied by, such forward-looking statements. Climb Bio may not actually achieve the plans, intentions or expectations disclosed in these forward-looking statements, and you should not place undue reliance on these forward-looking statements. These risks and uncertainties include, but are not limited to, important risks and uncertainties associated with: the ability of Climb Bio to timely and successfully achieve or recognize the anticipated benefits of its technology transfer and exclusive license agreement with Beijing Mabworks Biotech Co., Ltd.; changes in applicable laws or regulations; the possibility that Climb Bio may be adversely affected by other economic, business and/or competitive factors; Climb Bio’s ability to advance budoprutug and CLYM116 on the timelines expected or at all and to obtain and maintain necessary approvals from the U.S. Food and Drug Administration and other regulatory authorities; expectations regarding formulation and device readiness; obtaining and maintaining the necessary approvals from institutional review boards at clinical trial sites and independent data safety monitoring boards; replicating in clinical trials positive results found in early-stage clinical trials or nonclinical studies; top-line data may change as more patient data become available and are subject to audit and verification procedures; competing successfully with other companies that are seeking to develop treatments for IgAN and other immune-mediated diseases; maintaining or protecting intellectual property rights related to CLYM116 and/or its other product candidates; the outcome of any legal proceedings or other disputes; managing expenses; the possibility that models used in clinical development may be inaccurate; and raising the substantial additional capital needed, on the timeline necessary, to continue development of budoprutug, CLYM116 and any other product candidates Climb Bio may develop. For a further discussion of other risks and uncertainties, and other important factors, any of which could cause Climb Bio’s actual results to differ materially from those contained in the forward-looking statements, see the “Risk Factors” section, as well as discussions of potential risks, uncertainties and other important factors, in Climb Bio’s most recent filings with the U.S. Securities and Exchange Commission. In addition, the forward-looking statements included in this presentation represent Climb Bio’s views as of the date hereof and should not be relied upon as representing Climb Bio’s views as of any date subsequent to the date hereof. Climb Bio anticipates that subsequent events and developments will cause Climb Bio’s views to change. However, while Climb Bio may elect to update these forward-looking statements at some point in the future, Climb Bio specifically disclaims any obligation to do so, except as required by law.

This presentation also contains estimates and other statistical data made by independent parties and by us relating to market size and other data about our industry. These data involve a number of assumptions and limitations, and you are cautioned not to give undue weight to such estimates. In addition, projections, assumptions and estimates of our future performance and the future performance of the markets in which we operate are necessarily subject to a high degree of uncertainty and risk.

Webcast Agenda



CLYM116 Opportunity

Aoife Brennan, M.B., Ch.B.
President and CEO, Climb Bio



CLYM116: Initial Phase 1 Data in Healthy Volunteers

Edgar Charles, M.D.
Chief Medical Officer, Climb Bio



Closing Remarks and Q&A session

including Climb Bio management team

CLYM116 Topline Phase 1 Data Exceeded Target; Phase 2 Underway

Best-in-class profile, with long half-life, potent and prolonged PD effects, and Q12W dosing

Phase 1 Healthy Volunteer Data

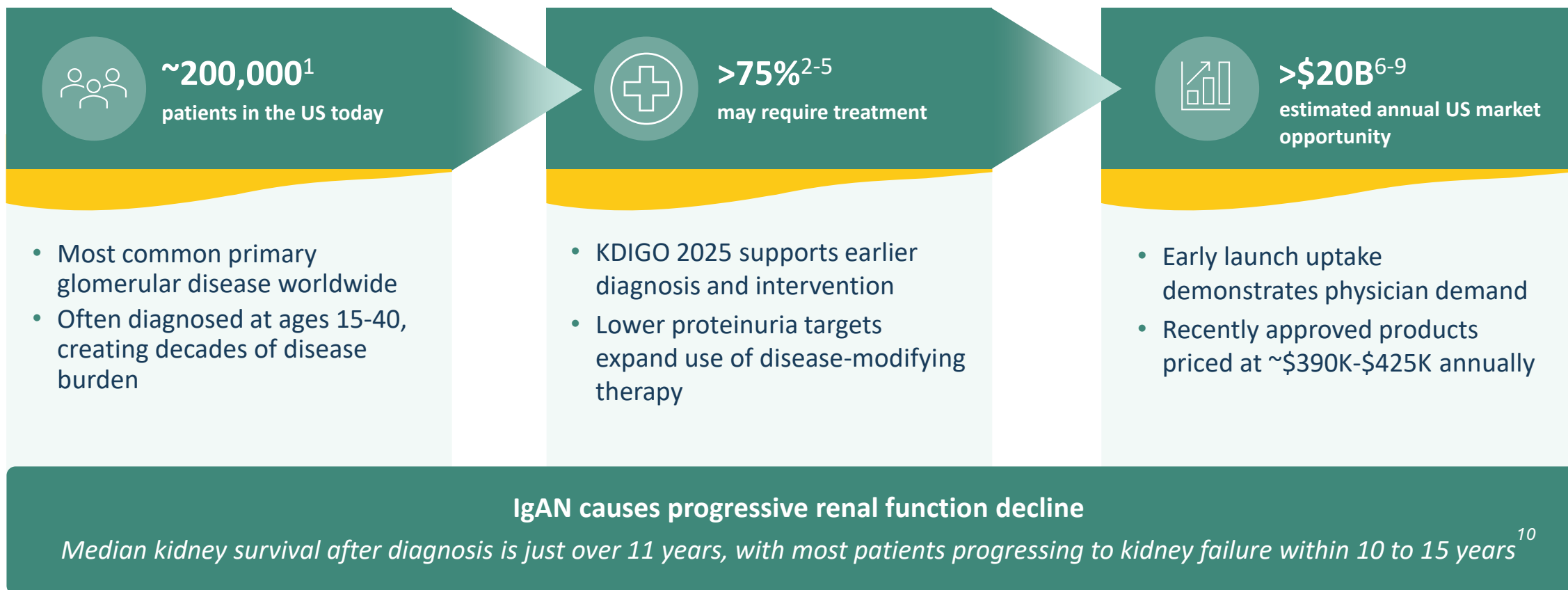
- **~29-day half-life**, 3x longer than sibeprenlimab, with linear clearance throughout the effective dose range
- **>90% free APRIL suppression** following a single, SC injection of 320 mg, with potential for sustained APRIL suppression with a Q12W dosing regimen
- **~60-75% suppression of IgA, Gd-IgA1, and IgM** following a single, SC injection of 320 mg, with suppression maintained through 12 weeks
- **Well-tolerated; no hypogammaglobulinemia**

Development Strategy Summary

- **NAVIGATE-2, a Phase 2 study in IgAN, enrolling with initial data anticipated in 1H 2027**
- **Target dosing regimen defined:** 800 mg loading dose followed by 400 mg Q12W, supported by integrated PK/PD modeling
- **High concentration (200 mg/mL) formulation complete**, enabling delivery of 400 mg in a single SC injection
- **Pre-filled syringe development complete, autoinjector compatible drug product**
- Anticipate advancing a **single regimen to Phase 3 in 2027**, subject to regulatory feedback

IgAN: A Lifelong Disease Requiring Durable Disease Modification

Earlier and expanding diagnosed population support a significant market opportunity



APRIL is a Validated Target in IgAN with Further Potential to Unlock

Phase 3 results and recent approvals establish the APRIL class and create an opportunity for differentiated next-generation therapies

First Generation: Where We Are Now

APPROVED	Sibeprenlimab ^{1,2} APRIL	51.2% placebo-adjusted UPCR reduction	Stabilization of eGFR through 2 years in Phase 3	Q4W
APPROVED	Atacicept ^{3,4} BAFF/APRIL	41.8% placebo-adjusted UPCR reduction	Stabilization of eGFR through 36 weeks in Phase 2*	Q1W
BLA FILED	Povetacept ^{5,6} BAFF/APRIL	49.8% placebo-adjusted UPCR reduction	Stabilization of eGFR through 48 weeks in Phase 1/2*	Q4W

Above reflects cross-trial comparisons and not data from head-to-head studies; differences exist between trial designs and participant characteristics, and caution should be exercised when comparing data across trials.

Next Generation: Where We're Headed

Depth and Durability of Suppression

Near-complete APRIL suppression

Robust IgA Response

Reduced pathogenic forms of IgA and IgA immune complex formation

Convenient Dosing Regimen

Substantive decrease from 12 to 4 injections annually

*Phase 3 eGFR data not yet available.

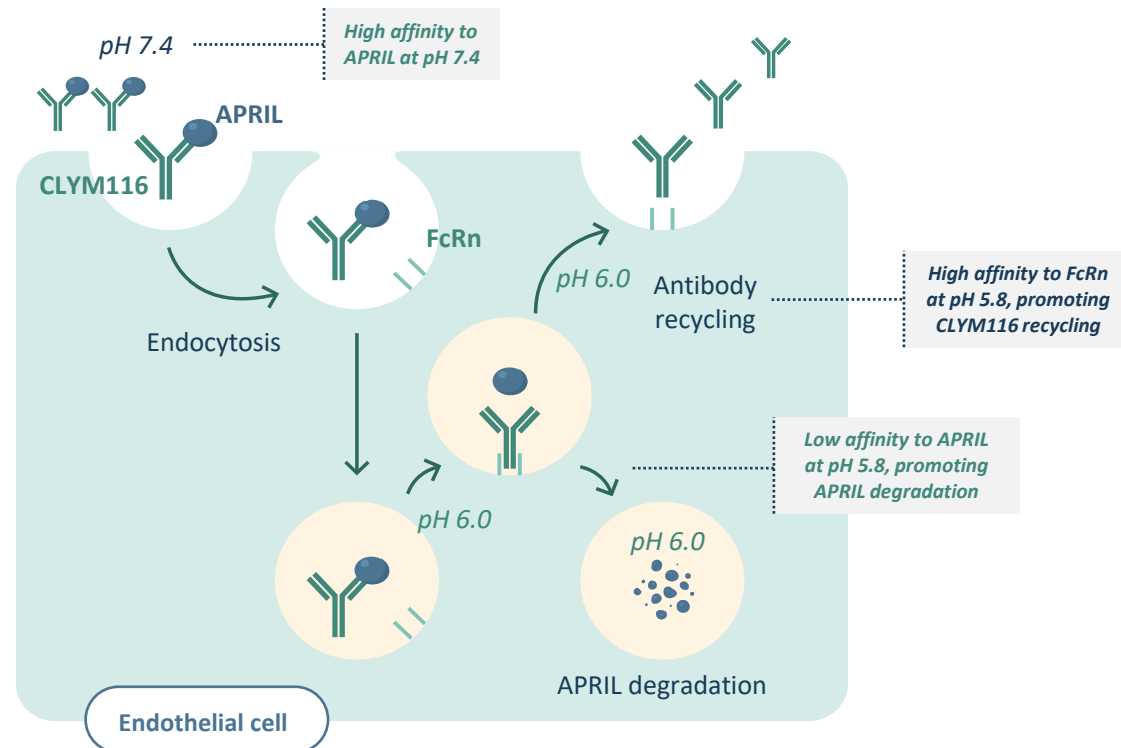
APRIL = a proliferation-inducing ligand, BAFF = B-cell activating factor, eGFR = estimated glomerular filtration rate, IgAN = IgA nephropathy, UPCR = urine protein creatinine ratio

¹ Perkovic NEJM 2025, ² Rizk GlomCon 2026, ³ Lafayette NEJM 2025, ⁴ Lafayette Kid Intl 2024, ⁵ Vertex Press Release, March 9, 2026, ⁶ Madan KI Reports 2025

CLYM116 “Sweeper” Designed for a Differentiated Clinical Profile

Anti-APRIL mAb, combining the utility of a sweeper with Fc mutations that increase serum half-life to deliver improved clinical activity

pH-dependent binding and FcRn recycling allow CLYM116 to release captured APRIL for degradation and re-enter circulation to bind again



Enhanced clearance beyond conventional neutralization:

- ✓ **Deeper target suppression**
Repeated capture and clearance of APRIL
- ✓ **Sustained activity**
Antibody recycling enables continued target engagement
- ✓ **Less frequent dosing**
Prolonged pharmacology may support an extended maintenance interval beyond half-life extension alone

Phase 1 Data Support Best-in-Class Potential of CLYM116

Data from healthy volunteer study demonstrate differentiation across key clinical attributes



EFFICACY

Robust APRIL & IgA Suppression

- **>90% free APRIL suppression** with a single 320 mg SC dose
- **~60-75% suppression of IgA, Gd-IgA1, and IgM** following a single 320 mg SC dose, with suppression maintained through 12 weeks



CONVENIENCE

Potential for Every 12-Week Dosing

- **Half-life of ~29 days**
- **Low injection burden with target regimen; 5 injections in year 1, with 4 injections annually thereafter**



SAFETY

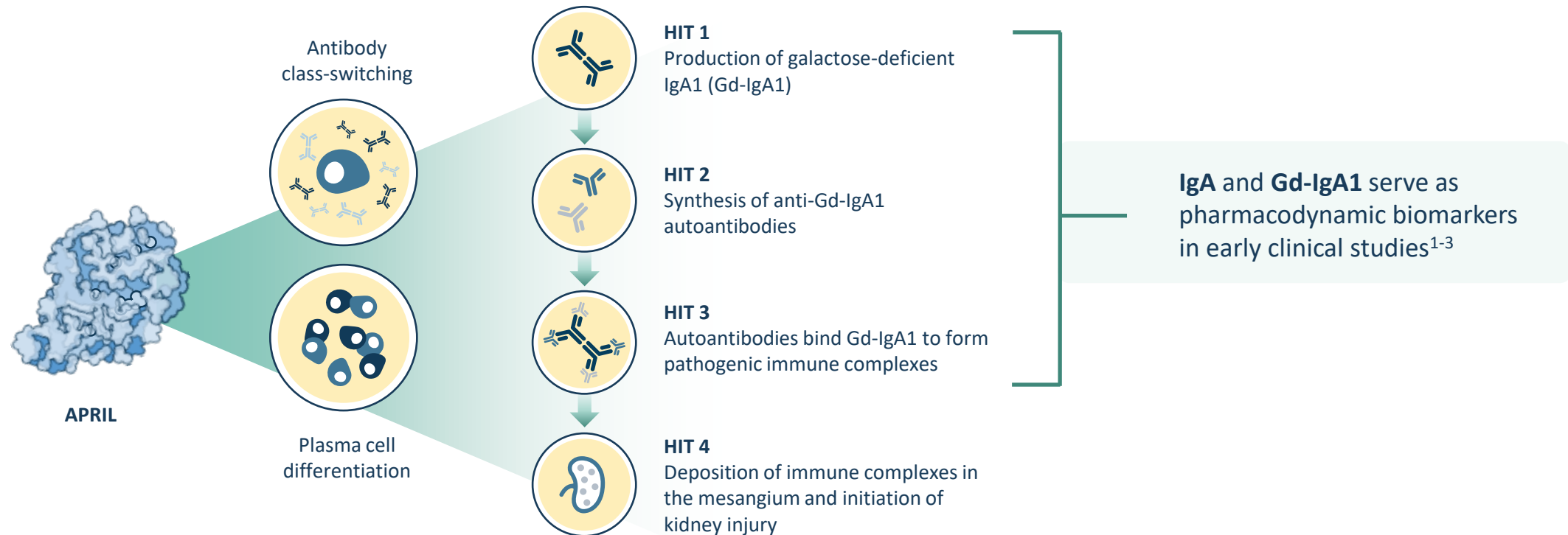
Favorable Safety Profile

- **Generally well-tolerated**
- **No SAEs/DLTs**
- **No hypogammaglobulinemia**

CLYM116

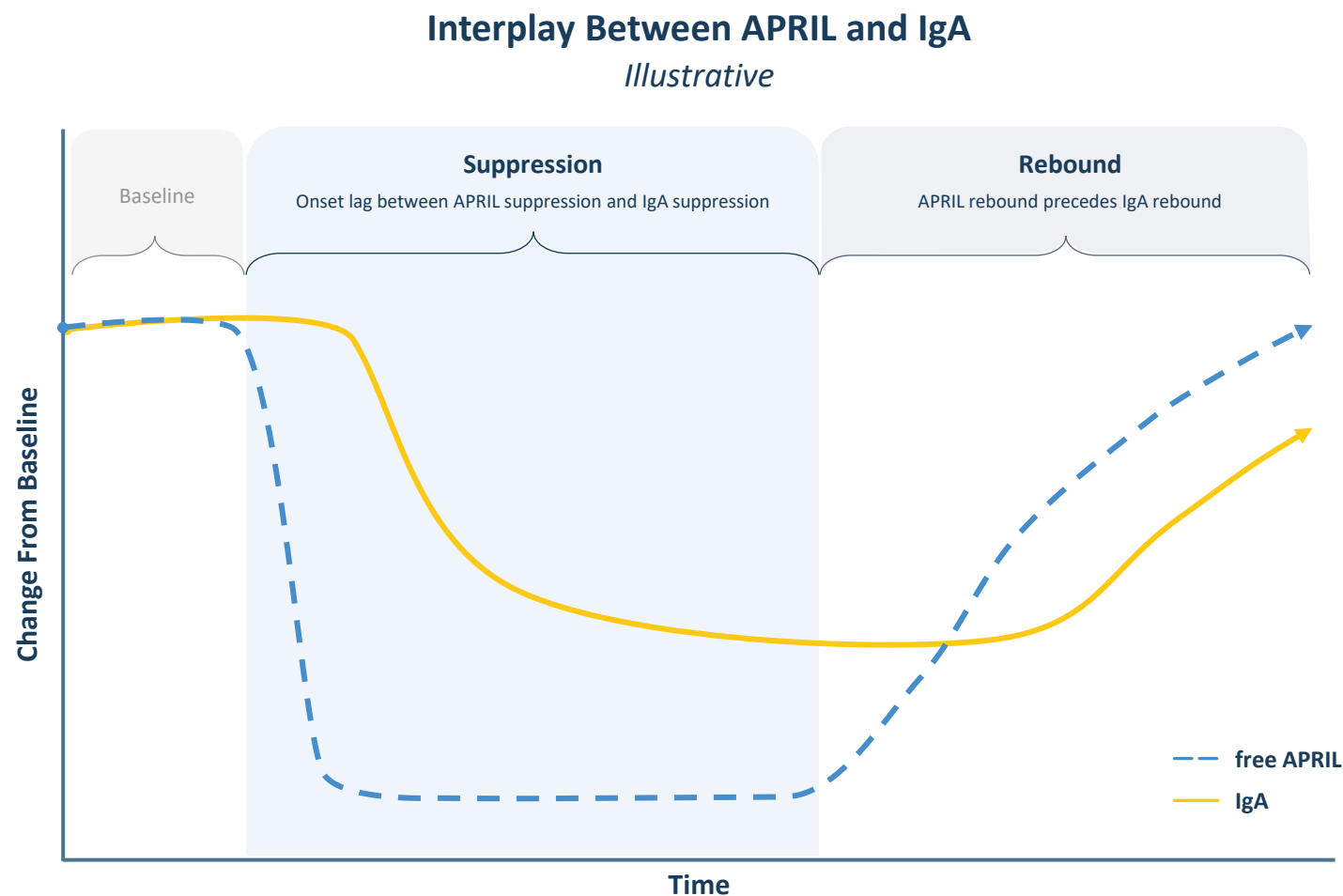
APRIL: A Central Driver of IgAN Pathogenesis

In IgAN, APRIL inhibition has been demonstrated to prevent the production of pathogenic IgA and the consequent immune complex formation that leads to kidney damage



Defining a Best-in-Class Q12W anti-APRIL Profile

Near-complete APRIL suppression throughout the dosing interval to drive optimal PD effects



CLYM116 Australia Phase 1 Design & Objectives

Randomized, placebo-controlled ascending-dose study in healthy volunteers

Robust evaluation around the 400 mg RP2D to support dose selection and exposure

Study Objectives:

- Safety and tolerability
- Pharmacokinetics
- Pharmacodynamics, including effects on serum APRIL and immunoglobulins (IgA, IgM, IgG, Gd-IgA1)

ASCENDING DOSE COHORTS, N = 46

Subcutaneous administration
~8 subjects per cohort (6 CLYM116: 2 placebo)



Dose & Formulation for Future Studies:

400 mg dose enabled by a 200 mg/mL formulation in a single 2 mL injection

Demographics and Baseline Characteristics

Cohorts were generally well balanced, with participant characteristics consistent with a healthy volunteer population

	25 mg	80 mg	160 mg	320 mg	480 mg	320 mg MAD	POOLED CLYM116	POOLED PLACEBO
N	6	6	5	6	6	6	35	11
Age, years	30.3 (7.4)	33.2 (14.3)	34.6 (10.1)	30.3 (4.6)	43.7 (14.6)	41.2 (15.8)	35.6 (12.2)	35.6 (11.8)
Female (n, %)	1 (17%)	5 (83%)	4 (80%)	3 (50%)	3 (50%)	4 (67%)	20 (57%)	8 (73%)
Race (n, %)								
White	2 (33%)	3 (50%)	4 (80%)	5 (83%)	4 (67%)	5 (83%)	23 (66%)	5 (45%)
Asian	4 (67%)	1 (17%)	1 (20%)	0	2 (33%)	1 (17%)	9 (26%)	5 (45%)
Other	0	2 (33%)	0	1 (17%)	0	0	3 (9%)	1 (9%)
Body Weight, kg	76.0 (17.5)	60.3 (10.6)	68.0 (8.3)	75.5 (11.7)	70.8 (10.4)	78.0 (16.9)	71.3 (13.7)	62.8 (11.6)
BMI, kg/m ²	24.9 (3.4)	21.6 (2.1)	25.4 (2.7)	25.6 (3.8)	24.4 (2.3)	27.1 (4.5)	24.8 (3.4)	22.8 (2.8)
Serum IgA, mg/dL	2.95 (1.20)	1.87 (0.63)	2.04 (1.03)	1.35 (0.42)	2.37 (0.52)	1.73 (0.31)	2.05 (0.86)	2.36 (0.72)

Mean (SD) used for age, body weight, body mass index (BMI), and serum IgA

CLYM116 Demonstrated Favorable Safety and Tolerability in HV

Generally well-tolerated, with no serious adverse events or hypogammaglobulinemia

Safety

CLYM116 was generally well-tolerated in the ongoing Phase 1 trial

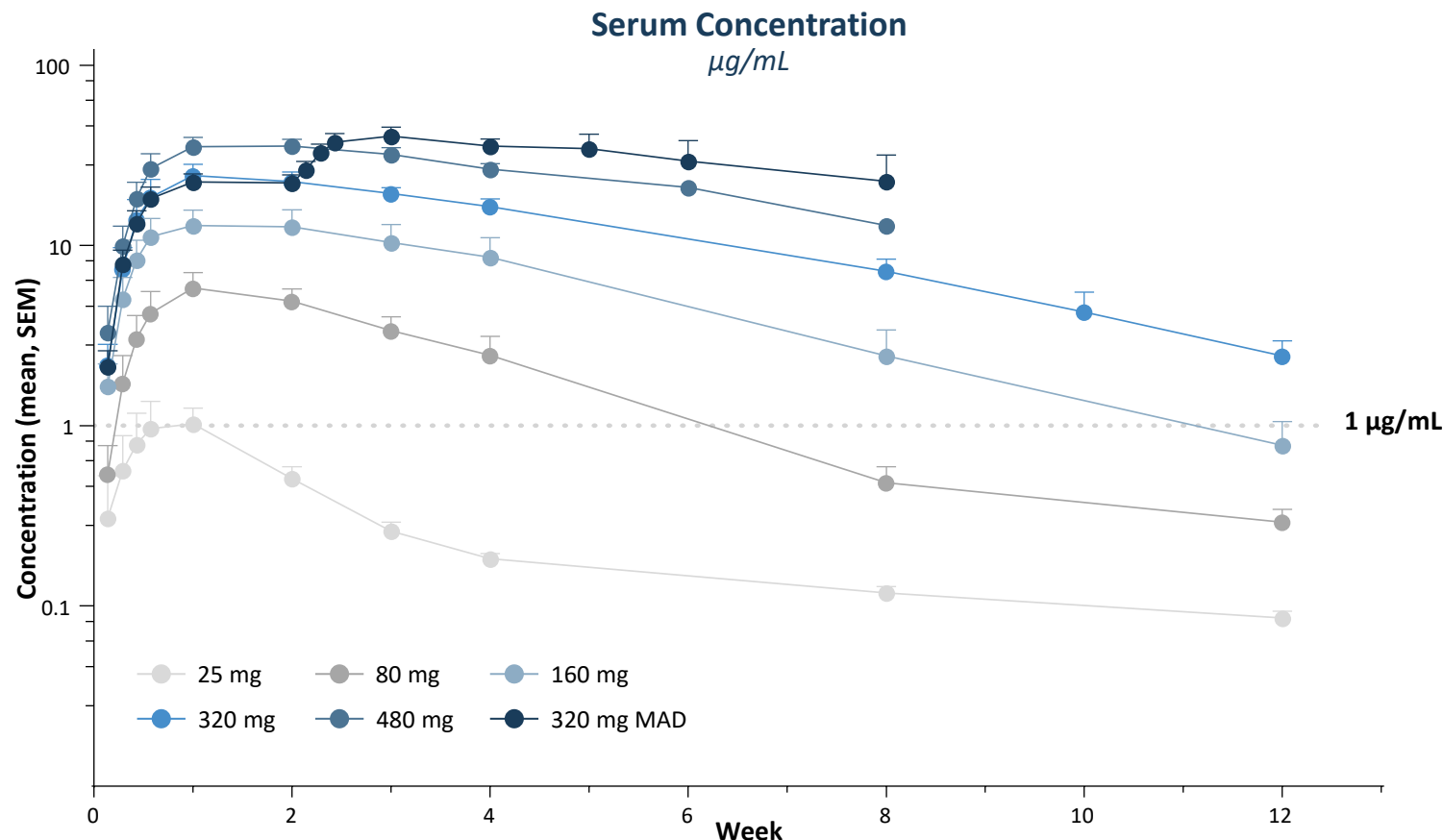
- No dose-limiting toxicities, SAEs, Grade ≥ 3 AEs, or AE-related discontinuations
- All AEs mild to moderate (Grades 1-2), transient, and self-resolving
- No hypogammaglobulinemia
- Injection site reactions observed in 5 subjects, all Grade 1 and resolved without intervention

	25 mg	80 mg	160 mg	320 mg	480 mg	320 mg MAD	POOLED CLYM116	POOLED PLACEBO
N	6	6	5	6	6	6	35	11
≥ 1 TEAE, n (%)	4 (67%)	5 (83%)	3 (60%)	3 (50%)	5 (83%)	5 (83%)	25 (71%)	5 (45%)
≥ 1 TRAE, n (%)	0	1 (17%)	3 (60%)	1 (17%)	3 (50%)	0	8 (23%)	N/A
≥ 1 severe TEAE, n	0	0	0	0	0	0	0	0
Discontinued due to AE, n	0	0	0	0	0	0	0	0

TEAEs occurring in >2 subjects in the CLYM116 cohorts included headache (20%), upper respiratory tract infection (14%), and injection-site reactions (14%).

CLYM116 Exposure Profile Supports Long Dosing Interval

~29-day half-life, 3x longer than sibeprenlimab, with linear clearance throughout the effective dose range



Pharmacokinetic Profile

- Dose-proportional exposure
- **Half-life of ~29 days**, compared to 9.3 days for sibeprenlimab^{1,2}
- Linear clearance observed at concentrations above 1 $\mu\text{g/mL}$
- **No TMDD observed** during the 12-week follow-up **at doses of 160 mg and above**, likely reflective of APRIL degradation due to sweeper mechanism

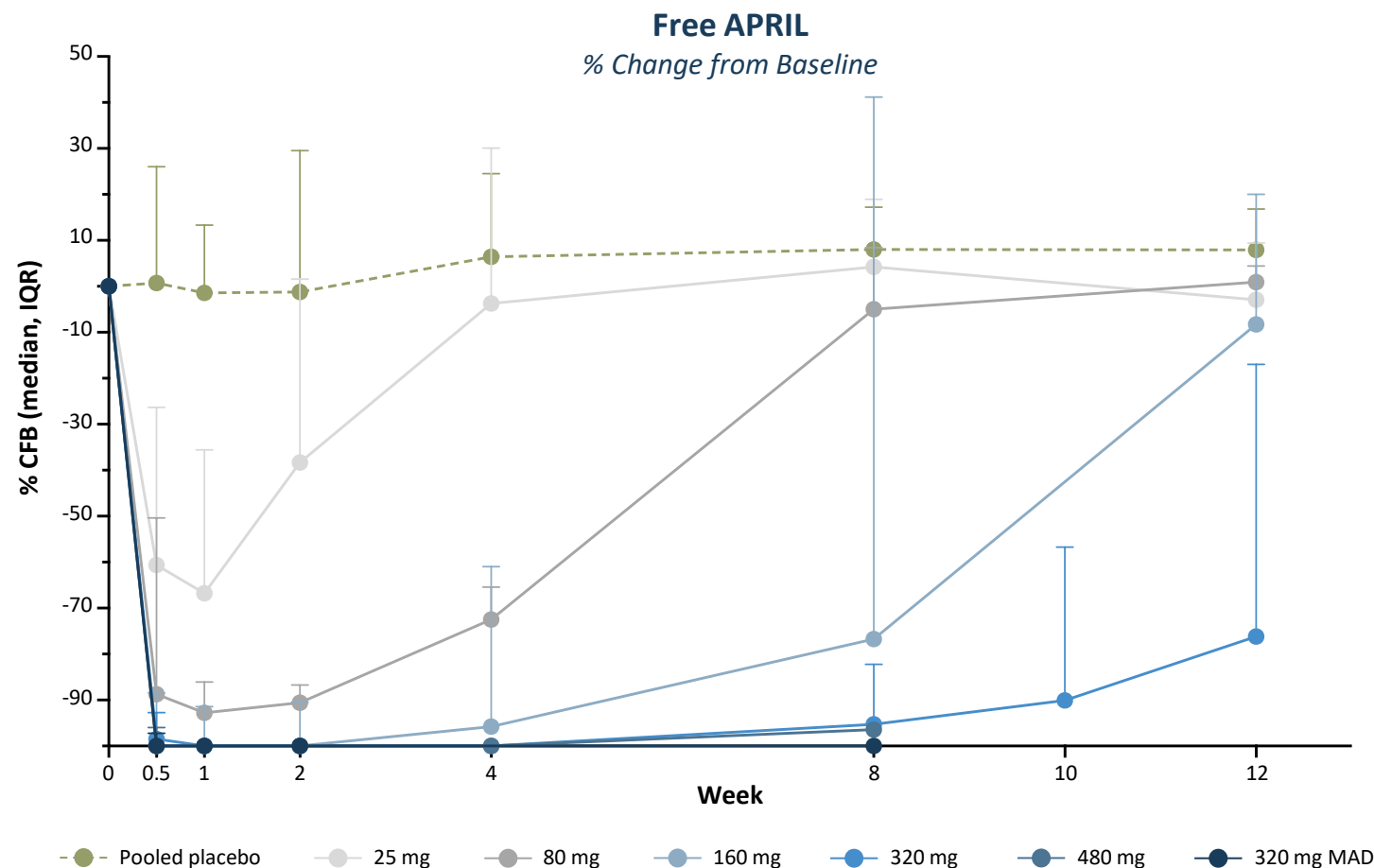
Data as of 26 August 2026; 12-week data available for all subjects in the 25, 80, 160, and 320 mg SAD cohorts. Follow-up ongoing in the 480 mg SAD and 320 mg MAD cohorts (data incomplete for weeks 5-8). 320 mg MAD cohort doses administered on Day 1 and Day 15. Half-life dependent on dose and dose interval. Projected CLYM116 half-life based on 320 mg MAD.

APRIL = a proliferation-inducing ligand, MAD = multiple ascending dose, SAD = single ascending dose, TMDD = target mediated drug disposition

¹ Zhang Clin Pharm Drug Dev 2023, ² Sibeprenlimab US Prescribing Information

CLYM116 Achieved Rapid and Sustained Free APRIL Suppression

Near-complete APRIL suppression after a single 320 mg SC dose, demonstrating robust target engagement and prolonged pharmacodynamic effect

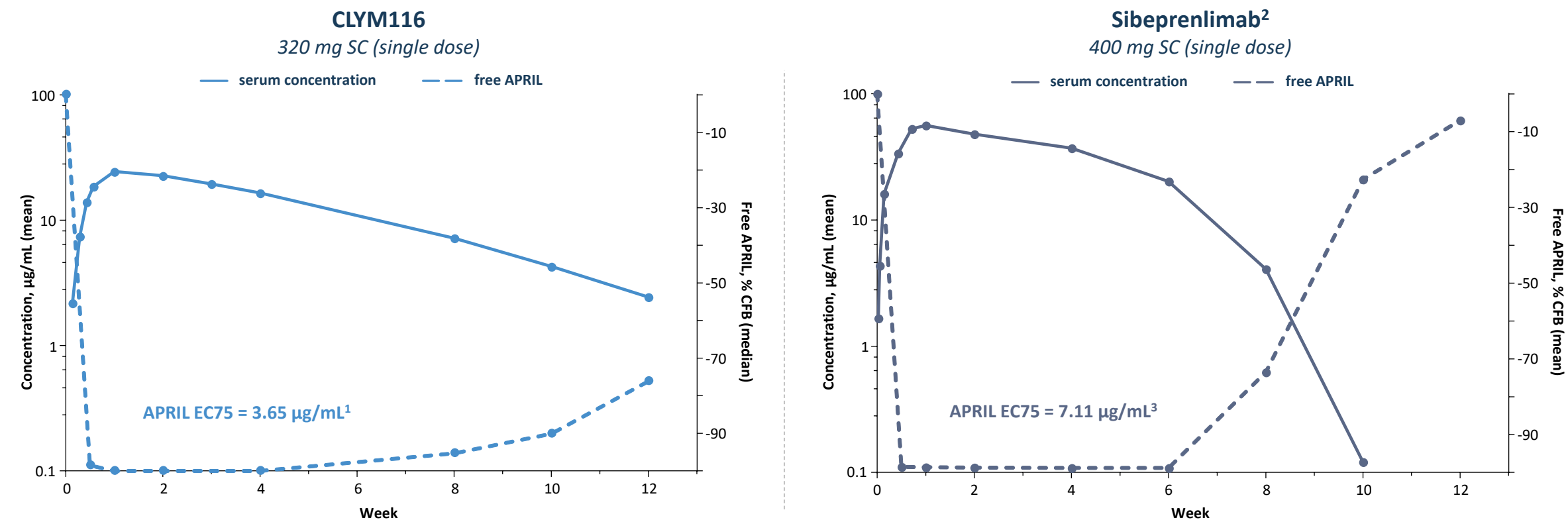


APRIL Suppression

- **>99% suppression** by Day 4 with doses ≥ 160 mg
- **>90% suppression** achieved through **Week 10** at the 320 mg dose, with **>75% suppression** maintained through **Week 12**
- Follow-up ongoing for the 480 mg SAD and 320 mg MAD doses

CLYM116 Demonstrated PK/PD Advantages vs. First-Gen Anti-APRIL

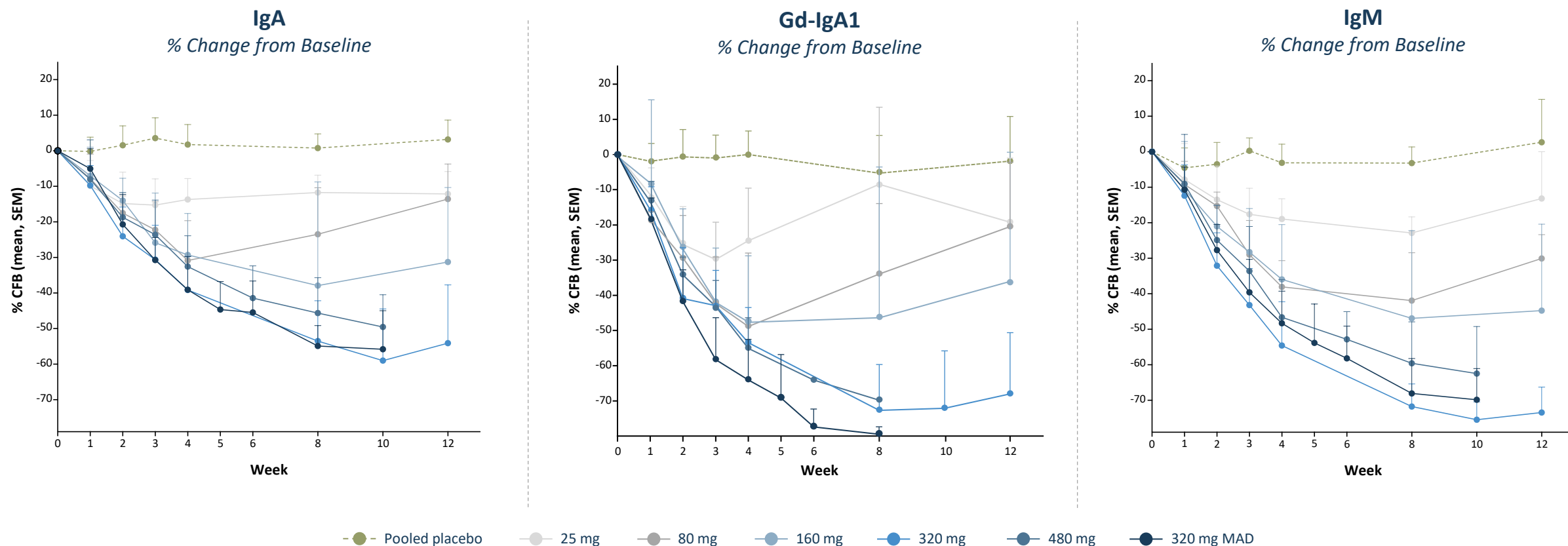
Longer half-life and exposure and greater *in vivo* potency relative to sibeprenlimab translated to robust and prolonged suppression of APRIL through 12 weeks



Above reflects cross-trial comparisons of observed single-dose healthy-volunteer data and not data from head-to-head studies; cross-trial comparisons are limited by differences in study design, populations, assays, regimens, and follow-up.

CLYM116 Achieved Deep, Durable IgA, Gd-IgA1, and IgM Reductions

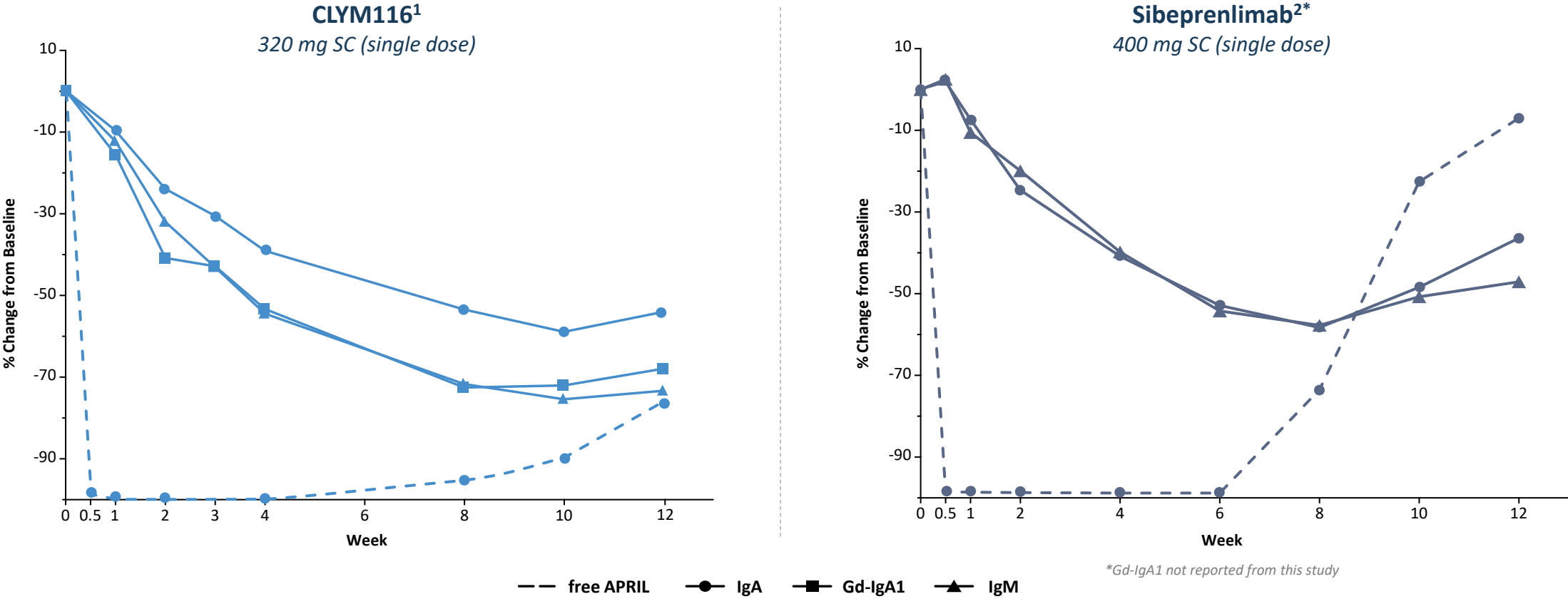
~60-75% suppression of IgA, Gd-IgA1, and IgM through 12 weeks with a single 320 mg dose



Follow-up ongoing; nadir not yet reached in the 480 mg SAD or 320 mg MAD cohorts

CLYM116 Demonstrated a Robust Q12W PD Profile

Prolonged APRIL suppression translates into sustained IgA, Gd-IgA1, and IgM reductions through 12 weeks, differentiating CLYM116 from once-monthly first-gen approaches

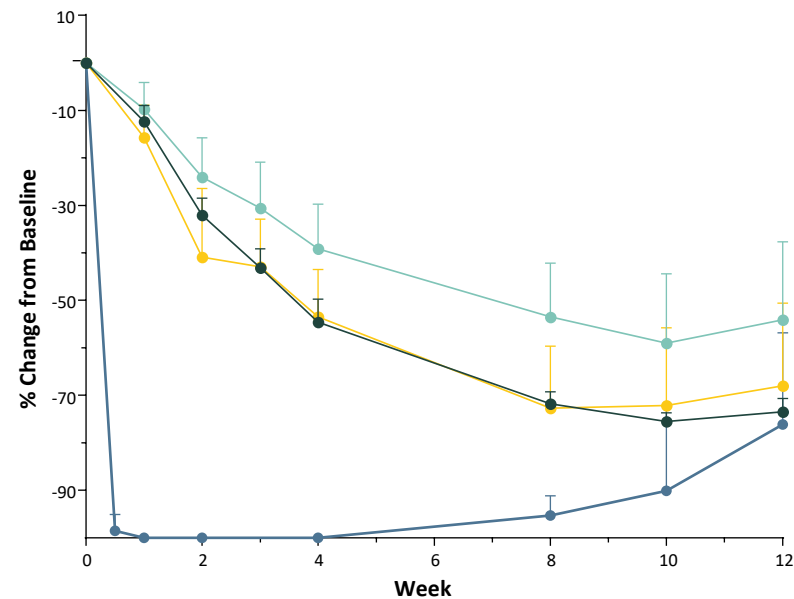


Above reflects cross-trial comparisons of observed single-dose healthy-volunteer data and not data from head-to-head studies; cross-trial comparisons are limited by differences in study design, populations, assays, regimens, and follow-up.

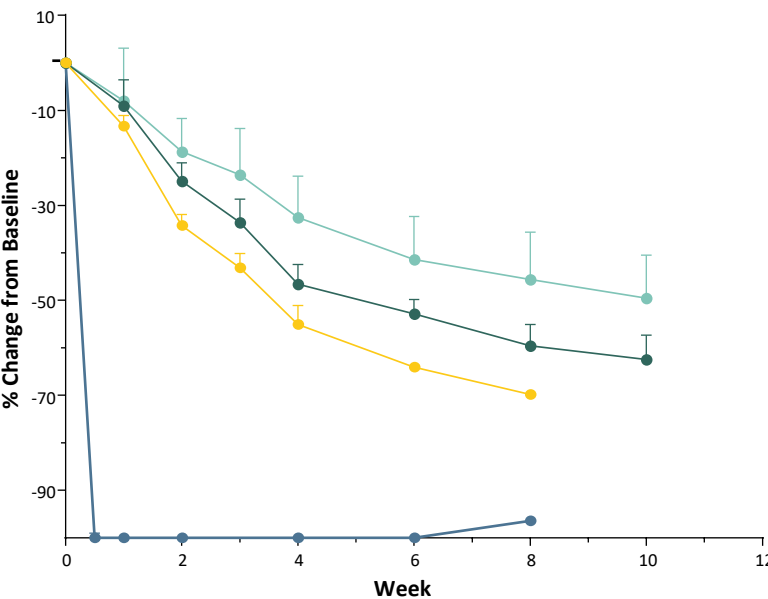
CLYM116 PD Effects Followed Consistent Trends Across Cohorts

Robust and sustained effects observed across all relevant biomarkers, supporting advancement of the 400 mg Q12W dosing regimen

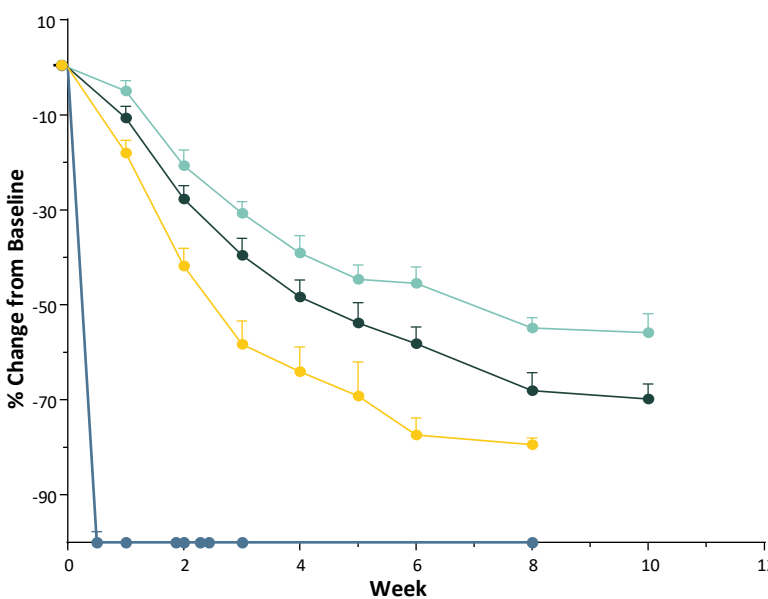
320 mg SAD



480 mg SAD



320 mg MAD



● free APRIL ● IgA ● Gd-IgA1 ● IgM

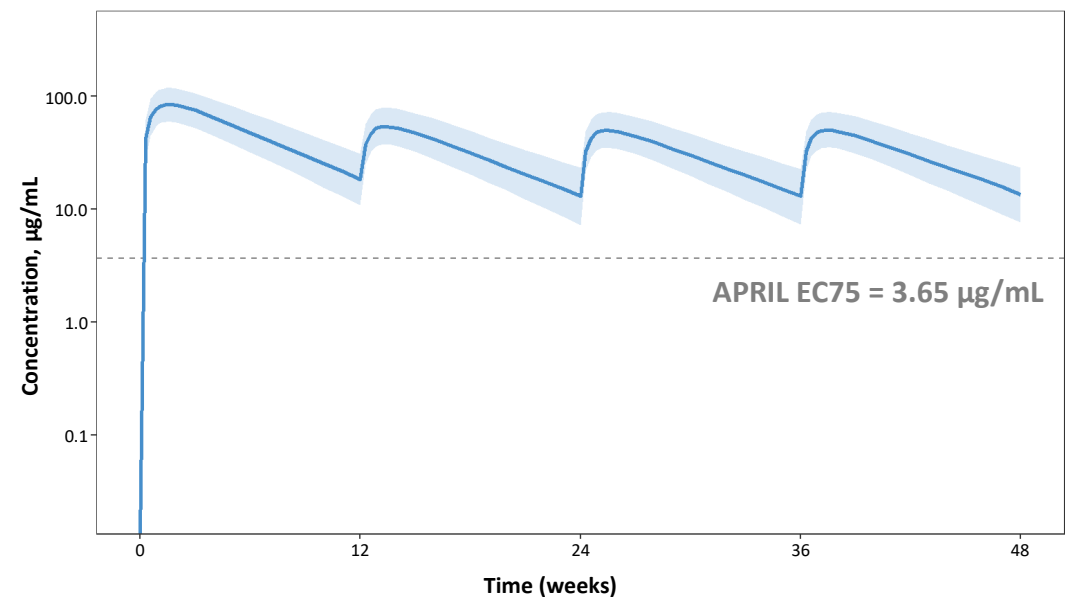
Follow-up ongoing; nadir not yet reached in the 480 mg SAD or 320 mg MAD cohorts

CLYM116 PK/PD Modeling Supports Q12W Dosing

400 mg Q12W regimen predicted to maintain exposure above APRIL EC75 throughout the dosing interval and produce near-complete APRIL suppression out to 48 weeks

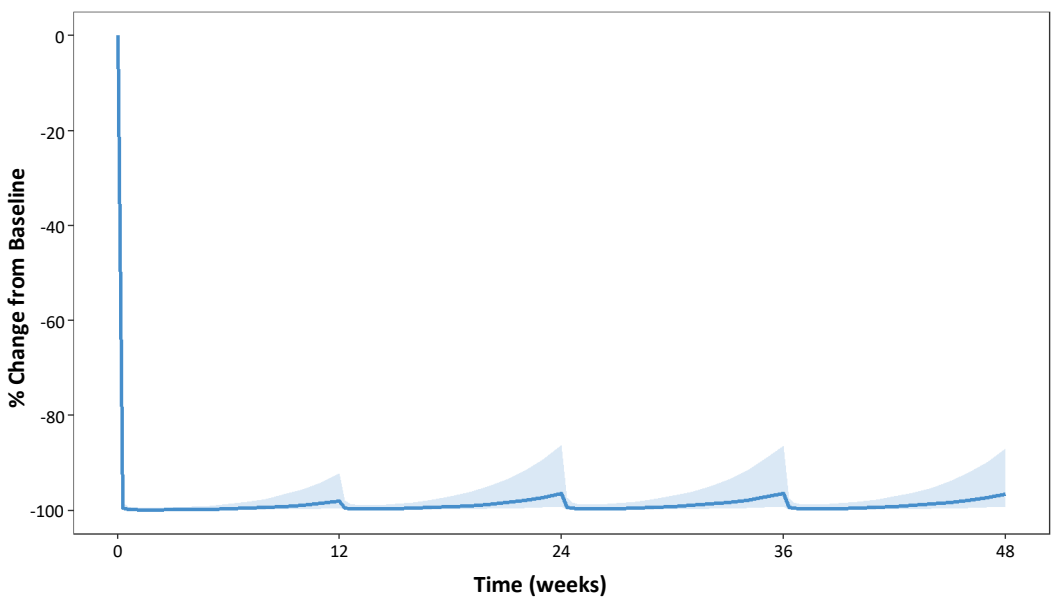
Simulated Serum Concentration Over Time

800 mg loading dose + 400 mg Q12W



Simulated Free APRIL Over Time

800 mg loading dose + 400 mg Q12W



Median 25-75th Percentile

Data as of 26 August 2026. Simulated serum concentrations are derived from a population pharmacokinetic model, and free APRIL levels are derived from a population pharmacokinetic / pharmacodynamic model. Both models were developed with interim data from two Phase 1 studies in healthy subjects. The simulations show the median and 25-75th percentiles of 1000 virtual subjects with the indicated dosing regimen.
APRIL = a proliferation-inducing ligand, PD = pharmacodynamic, PK = pharmacokinetic

CLYM116 NAVIGATE-2 Phase 2 Study in IgAN Ongoing



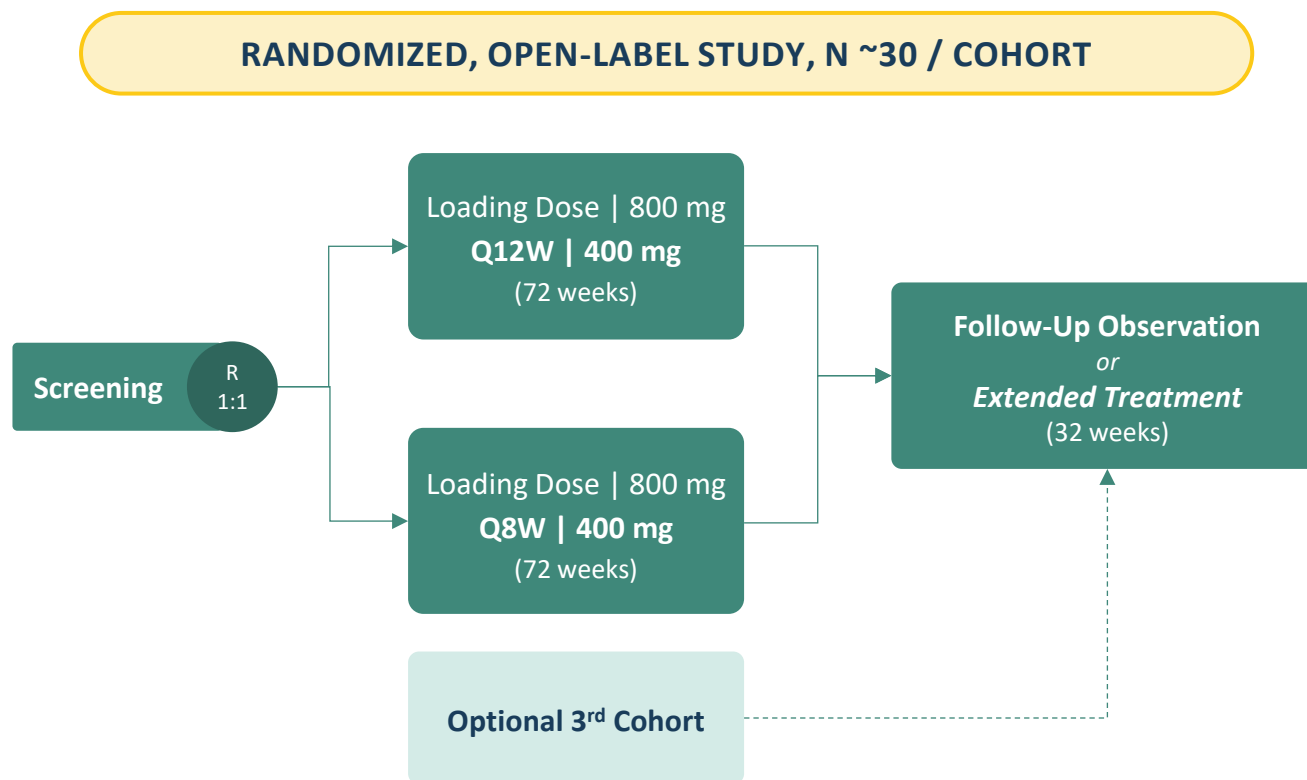
Initial data anticipated in H1 2027; designed to confirm integrated Phase 1 PK/PD modeling and support advancing Q12W regimen into Phase 3

Population:

- Adults with biopsy-proven IgAN
- 24-hour UPCR ≥ 0.5 g/g or 24-hour urine protein ≥ 0.75 g
- eGFR ≥ 30 mL/min/1.73 m²
- Stable, maximum tolerated ACEi/ARB for ≥ 12 weeks

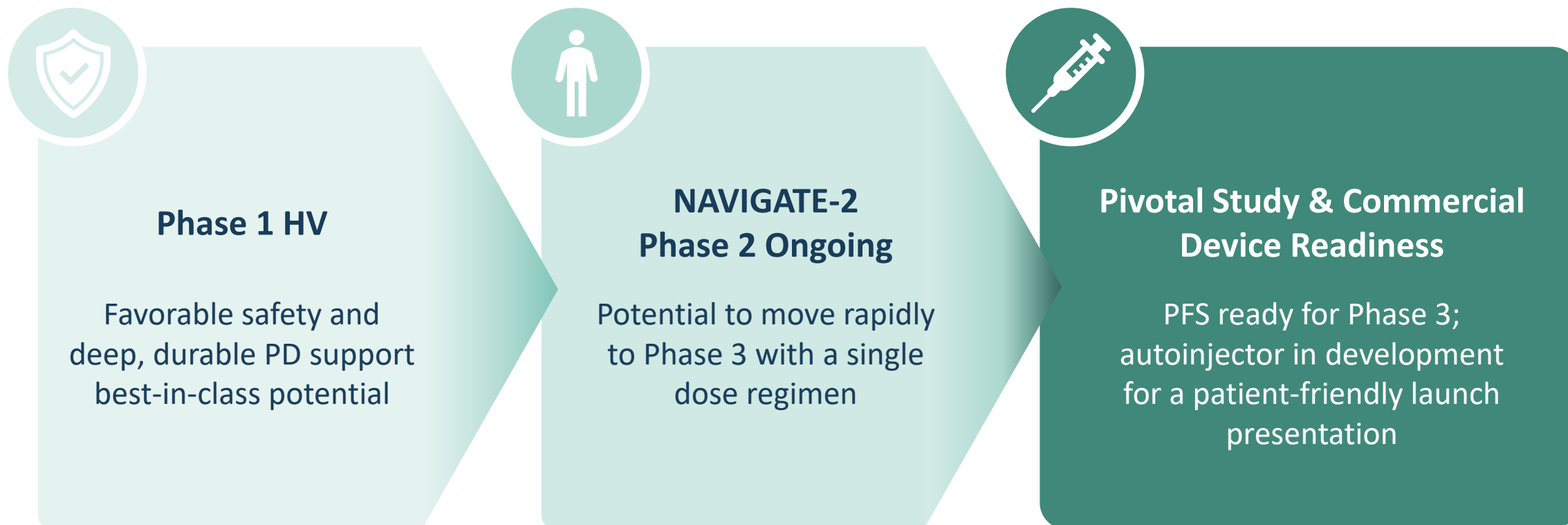
Designed to evaluate:

- Safety and tolerability
- Gd-IgA1
- 24-hour UPCR
- eGFR slope
- Serum immunoglobulin profile
- Free APRIL



CLYM116: Compelling Phase 1 Data, Clear Path Ahead

CLYM116 offers the potential for durable disease control with convenient Q12W dosing;
NAVIGATE-2 Phase 2 ongoing and device development advancing





Closing Remarks



CLYM116: Positioned to Accelerate

Depth, durability, and convenience in a single molecule, with multiple upcoming readouts

Compelling Phase 1 Results

- ✓ Deep & durable APRIL, IgA and Gd-IgA1 suppression
- ✓ Potential for every-12-week dosing
- ✓ Favorable safety & tolerability profile

2026

2027

CLYM116

Additional Australia Phase 1 HV data
Mabworks China Phase 1 HV data
Medical meeting, Q4 2026

CLYM116

Initial Phase 2 IgAN data
1H 2027

CLYM116

Phase 3 initiation, pending
regulatory feedback
2027

Q&A Session



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Perrin Wilson, Ph.D.

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