



APG777 PHASE 1 DATA

March 2024



Disclaimers and Forward-looking statements



This presentation contains certain "forward-looking statements" within the meaning of applicable securities laws. Other than statements of historical facts, all statements included in this presentation are forward-looking statements, including statements about our plans, objectives, goals, strategies and future events, the efficacy, safety, tolerability, PK and PD profile of APG777, the potential dosing regimen of APG777, the potential superiority of APG777 compared to current therapies, our expectations regarding plans for our current and future product candidates and programs, our plans for our current and future clinical trials, our plans for clinical trial design, the anticipated timing of the initiation of and results from our clinical trials, the potential clinical benefit and half-life of APG777, APG808, APG990, APG222 and any other potential programs, our expected timing for future pipeline updates and estimates of market size. In some cases, you can identify forward-looking statements by terms such as "anticipate," "believe," "can," "could," "design," "estimate," "expect," "intend," "likely," "may," "might," "plan," "potential," "predict," "suggest," "target," "will," "would," or the negative of these terms, and similar expressions intended to identify forward-looking statements. The forward-looking statements are based on our beliefs, assumptions and expectations of future performance, taking into account the information currently available to us. These statements are only predictions based upon our current expectations and projections about future events. Forward-looking statements are subject to known and unknown risks, uncertainties and other factors that may cause our actual results, level of activity, performance or achievements to be materially different from those expressed or implied by such forward-looking statements, including those risks described in "Risk Factors" and "Management's Discussion and Analysis of Financial Condition and Results of Operations" in our Annual Report on Form 10-K for the year ended December 31, 2023, filed with the SEC on March 5, 2024, and subsequent disclosure documents we may file with the U.S. Securities and Exchange Commission. Although we have attempted to identify important factors that could cause actual results to differ materially from those contained in forward-looking statements, there may be other factors that cause results not to be as anticipated, estimated or intended.

This presentation concerns drug candidates that are under clinical investigation, and which have not yet been approved by the U.S. Food and Drug Administration. These are currently limited by federal law to investigational use, and no representation is made as to their safety or effectiveness for the purposes for which they are being investigated.

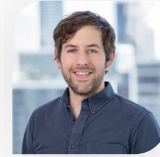
The assumptions used in the preparation of this presentation, although considered reasonable by us at the time of preparation, may prove to be incorrect. You are cautioned that the information is based on assumptions as to many factors and that actual results may vary from the results projected and such variations may be material. Accordingly, you should not place undue reliance on any forward-looking statements contained herein or rely on them as predictions of future events. All forward-looking statements in this presentation apply only as of the date made and are expressly qualified by the cautionary statements included in this presentation. We do not undertake to update any forward-looking statements, except in accordance with applicable securities laws.

The trademarks, trade names and service marks appearing in this presentation are the property of their respective owners. Certain information contained in this presentation relate to or are based on studies, publications and other data obtained from third-party sources as well as our own internal estimates and research. While we believe these third-party sources to be reliable as of the date of this presentation, it has not independently verified, and makes no representation as to the adequacy, fairness, accuracy or completeness of, any information obtained from third-party sources.

Agenda



Introduction & Executive Summary



**Michael Henderson, MD
Chief Executive Officer**

APG777 Phase 1 Interim Results



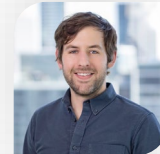
**Carl Dambkowski, MD
Chief Medical Officer**

APG777 Phase 2 Trial in Atopic Dermatitis



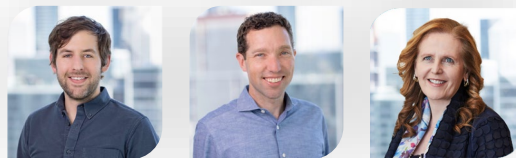
**Kristine Nograles, MD
SVP, Clinical Development**

Building a Leading I&I Company



**Michael Henderson, MD
Chief Executive Officer**

Analyst Q&A



**Michael Henderson, MD, CEO
Carl Dambkowski, MD, CMO
Jane Pritchett Henderson, CFO**

Apogee plans to reshape the current standard of care for inflammatory and immune diseases



Refusing to stop at “good enough”

Focus on developing differentiated biologics with known biologic drivers

Near term priority on treatments for atopic dermatitis (AD), asthma and chronic obstructive pulmonary disease (COPD)

People living with these diseases deserve the best possible treatment

Significant unmet need continues



1Q24 Update reflects significant progress



APG777

- Phase 1 initial data has exceeded all trial objectives

- Phase 2 in AD anticipated to start in 1H 2024 (ahead of schedule)

- Planned, integrated Phase 2 in AD combines Phase 2a and Phase 2b elements with potential for significant timeline acceleration (topline data from Part A remains 2H 2025)

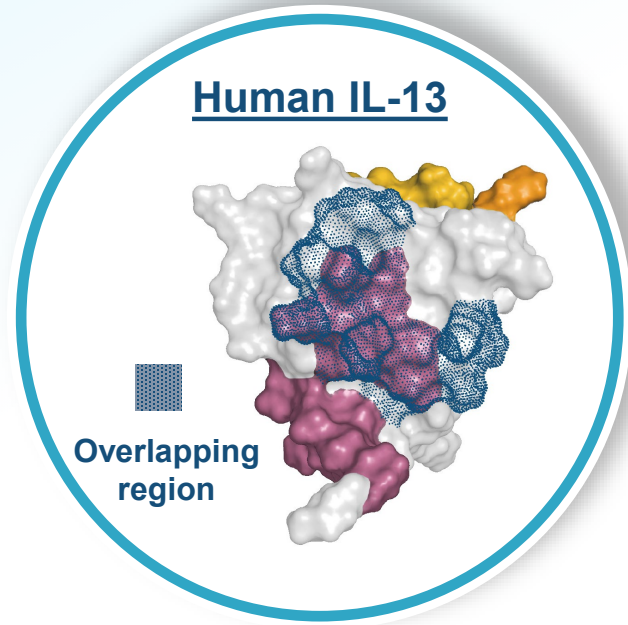
- Enhanced 180 mg/mL formulation enables 44% higher dose vs lebrikizumab in the same volume

APG808

- Expect to initiate Phase 1 in healthy volunteers in 1H'24 (ahead of schedule)

- Phase 1 interim data accelerated to 2H 2024 (from 2025)

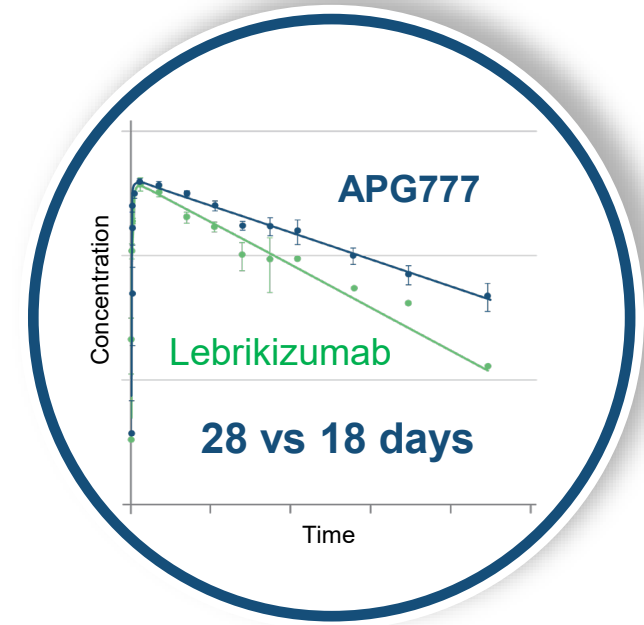
APG777 leverages lebrikizumab's mechanism to deliver a potentially best-in-class, pipeline in a product antibody



APG777's epitope on IL-13 overlaps with lebrikizumab's and leverages proven MoA and biology



APG777 is as potent as lebrikizumab and DUPIXENT in key preclinical assays



APG777 NHP half-life is significantly longer than lebrikizumab

APG777 Phase 1 initial data has exceeded all trial objectives



GOAL

Establish **safety** & **PK** profile

Well-tolerated with at least 33-day half-life

Set **Ph2 induction regimen**

Achieve at least equiv. exposures to lebrikizumab with same or fewer injections

Set **Ph2 maintenance regimens**

Equal lebrikizumab exposure with every 2-month or longer dosing¹

Supplemental

Demonstrate effect on biomarkers pSTAT6 or TARC

RESULT

- **Half life of ~75 days**
- Doses up to 1200mg tested and **well-tolerated**
- Initial **multiple-dose** data consistent with PK & safety profile from SAD cohorts

- Regimen modeled to **exceed lebrikizumab exposure by ~30-40% with potential for improved clinical responses (e.g. EASI-75, EASI-90, IGA 0/1)**
- **~50% fewer injections than lebrikizumab in induction (6 vs 11)**

- **3- or 6- month maintenance dosing enabled** with modeled exposures similar to or greater than lebrikizumab

- **Extended PD effect** on both pSTAT6 and TARC for ~3 months with follow-up ongoing



Exceeded



Exceeded



Exceeded

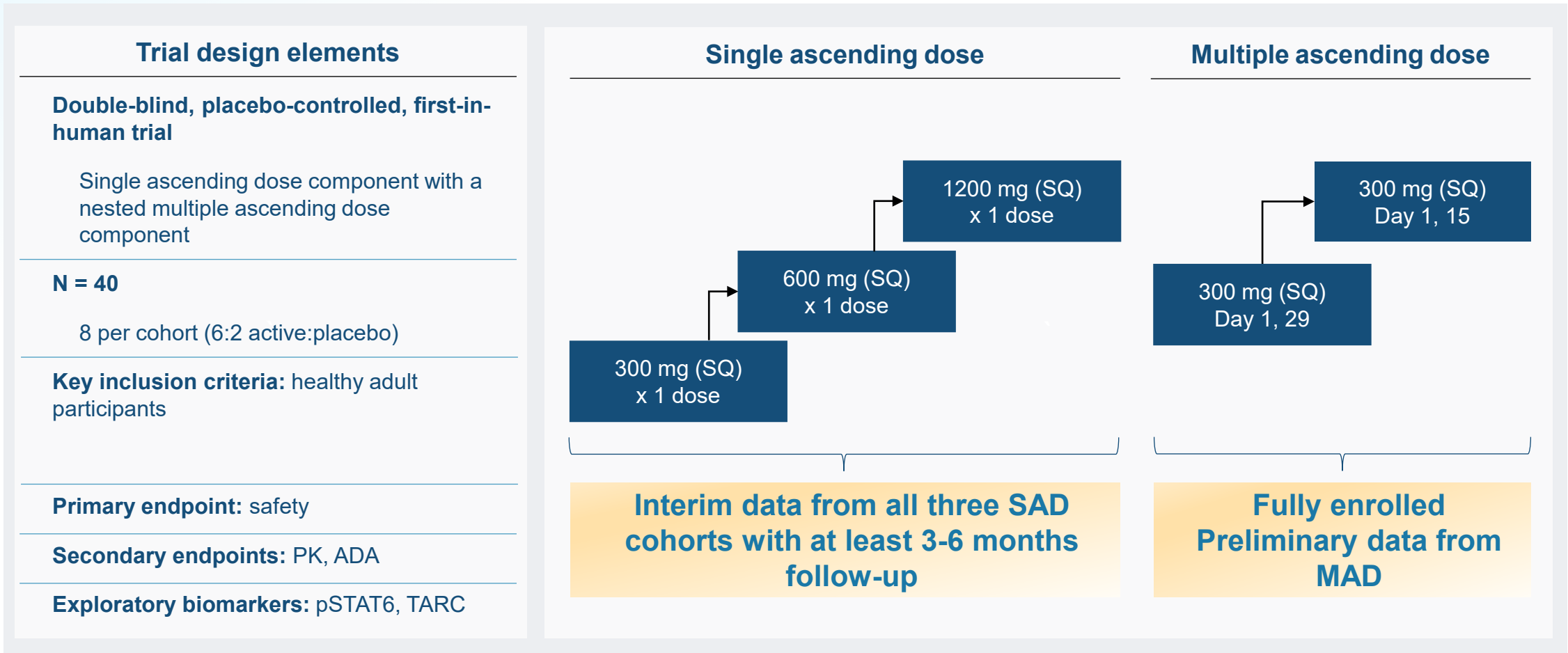


Exceeded



APG777 Phase 1 in
Healthy Volunteers

APG777 interim data from ongoing Phase 1 trial in healthy volunteers



Baseline characteristics are in line with our expectations



	Single dose				Multiple dose		
	Placebo N=6	Cohort 1 300 mg N=6	Cohort 2 600 mg N=6	Cohort 3 1,200 mg N=6	Placebo N=4	Cohort 1 300 mg at Day 1, 300 mg at Day 29 N=6	Cohort 2 300 mg at Day 1, 300 mg at Day 15 N=6
Age (yrs), mean (SD)	41.3 (16.2)	30.2 (12.2)	40.2 (18.4)	29.7 (4.6)	42.0 (12.1)	42.7 (13.9)	40.2 (13.8)
Female	100%	66.7%	83.3%	33.3%	100%	50.0%	50.0%
Caucasian	100%	33.3%	83.3%	100%	75.0%	100%	33.3%
Weight (kg), mean (SD)	72.5 (12.6)	74.3 (14.6)	78.8 (14.0)	77.2 (16.2)	62.3 (9.5)	80.5 (8.9)	66.7 (12.9)

Demographics were well balanced across cohorts

APG777 was well-tolerated with a favorable safety profile



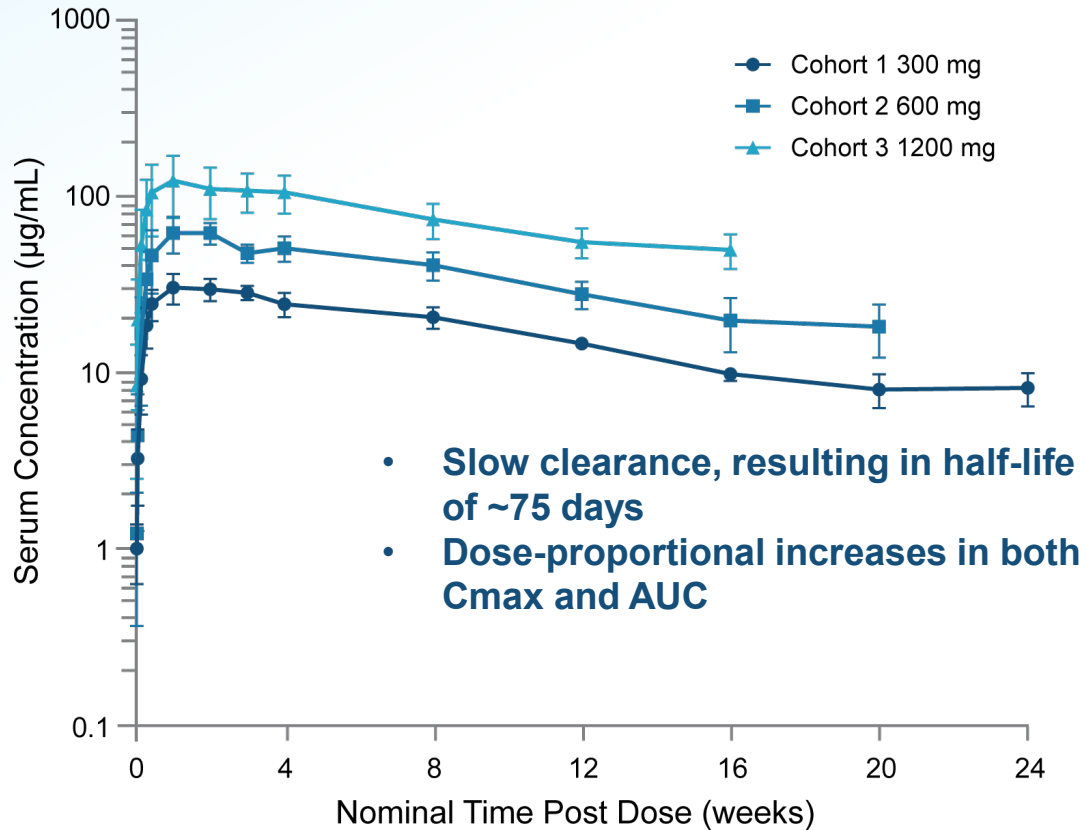
N (%)	Single dose				Multiple dose			Overall trial	
	Placebo N=6	Cohort 1 300 mg N=6	Cohort 2 600 mg N=6	Cohort 3 1,200 mg N=6	Placebo N=4	Cohort 1 300 mg at Day 1, 300 mg at Day 29 N=6	Cohort 2 300 mg at Day 1, 300 mg at Day 15 N=6	APG777 N=30	Placebo N=10
Participants with at least one TEAE	5 (83.3%)	4 (66.7%)	5 (83.3%)	2 (33.3%)	2 (50.0%)	5 (83.3%)	1 (16.7%)	17 (56.7%)	7 (70.0%)
Participants with at least one TE-SAE	0	0	0	0	0	0	0	0	0
Participants with at least one drug-related AE	3 (50.0%)	0	1 (16.7%)	1 (16.7%)	0	1 (16.7%)	0	3 (10.0%)	3 (30.0%)
Participants with at least one ≥Grade 3 TEAE	0	0	0	0	0	0	0	0	0
Participants that discontinued study due to TEAE	0	0	0	0	0	0	0	0	0
Participants that decreased dose due to TEAE	0	0	0	0	0	0	0	0	0

The safety profile is in line with expectations for therapies targeting the IL-13 pathway

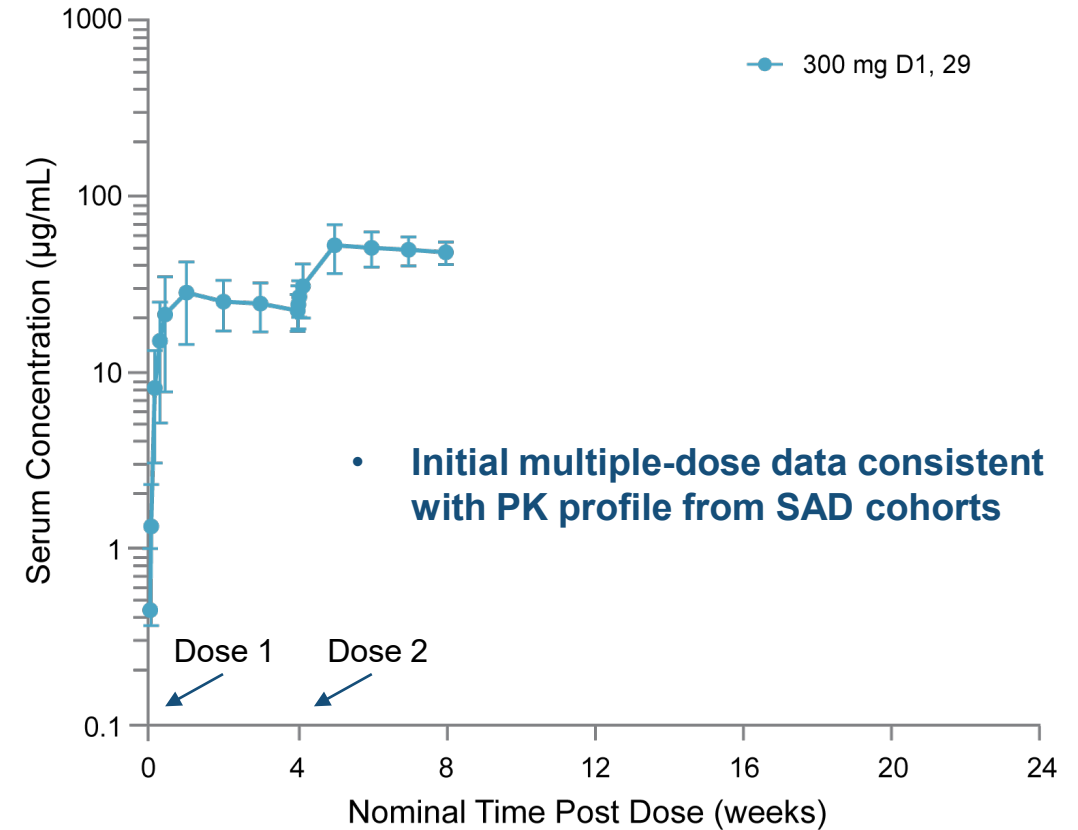


APG777 exhibited a potentially best-in-class PK profile with a half-life of ~75 days

Single-dose concentration-time profile

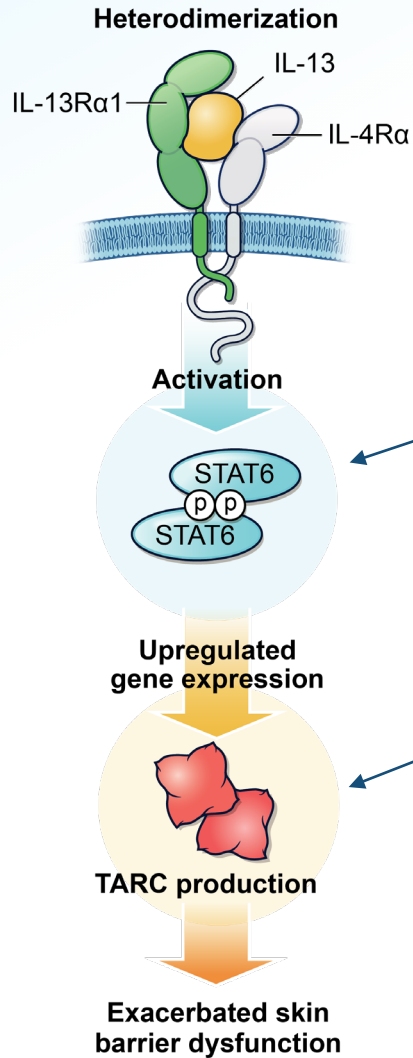


Multi-dose concentration-time profile



PK demonstrated dose-proportionality and half-life of ~75 days (approximately 3x lebrizumab)

pSTAT6 and TARC are biomarkers of IL-13 target engagement and AD severity



APG777 Phase 1 biomarkers

1. pSTAT6 is one of the **earliest markers of IL-13 receptor activation**

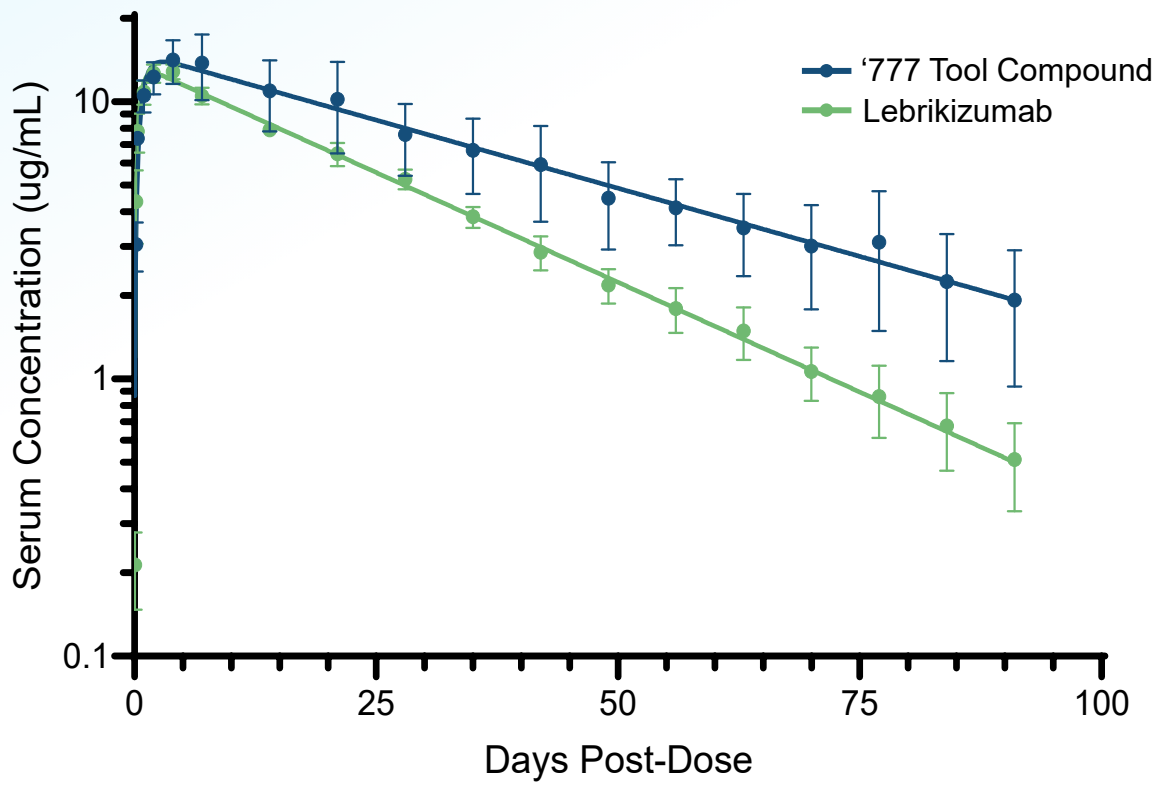
2. TARC levels are the most strongly correlated to AD severity of any biomarker

Taken together, **APG777's reduction of these biomarkers confirms inhibition of IL-13 signaling** and allows comparison to other agents

In a head-to-head NHP study, '777 tool compound inhibited pSTAT6 significantly longer than lebrizumab

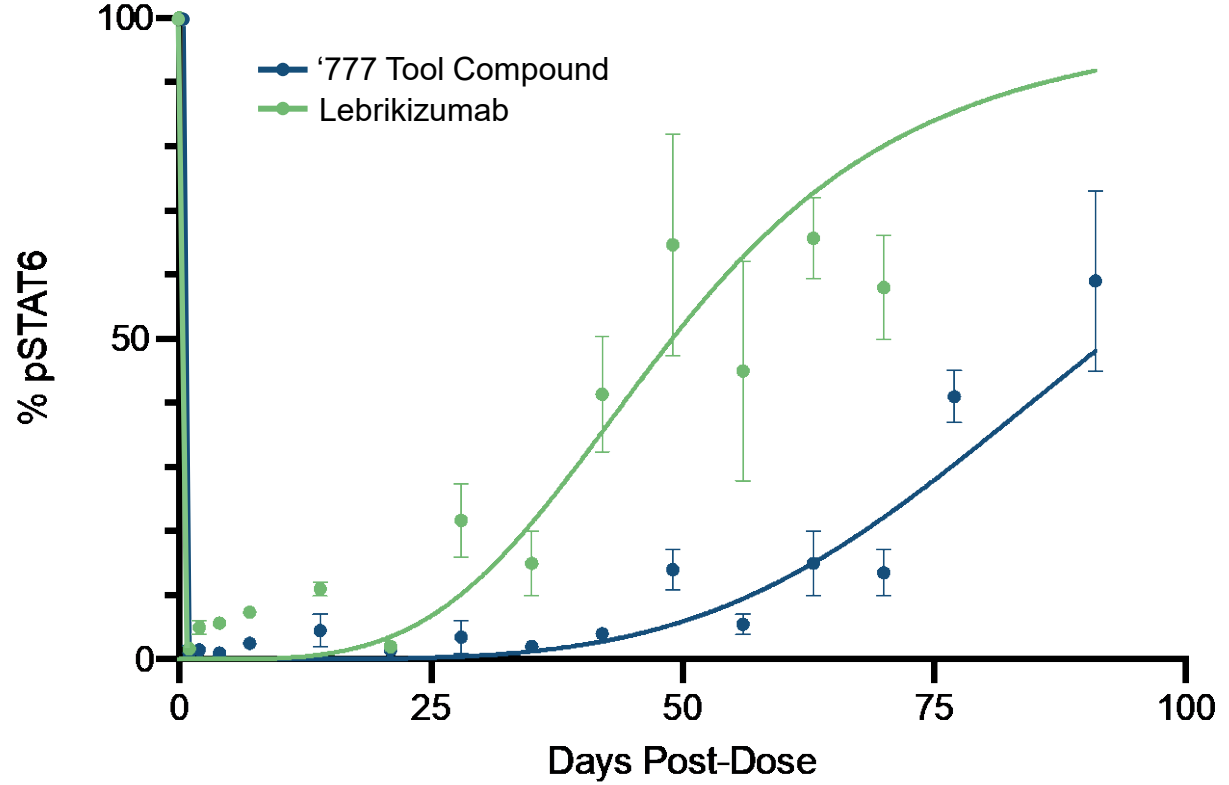


Head-to-head PK data in NHP



777 tool compound had **~60% longer half-life** vs. lebrizumab

Head-to-head PD data in NHP



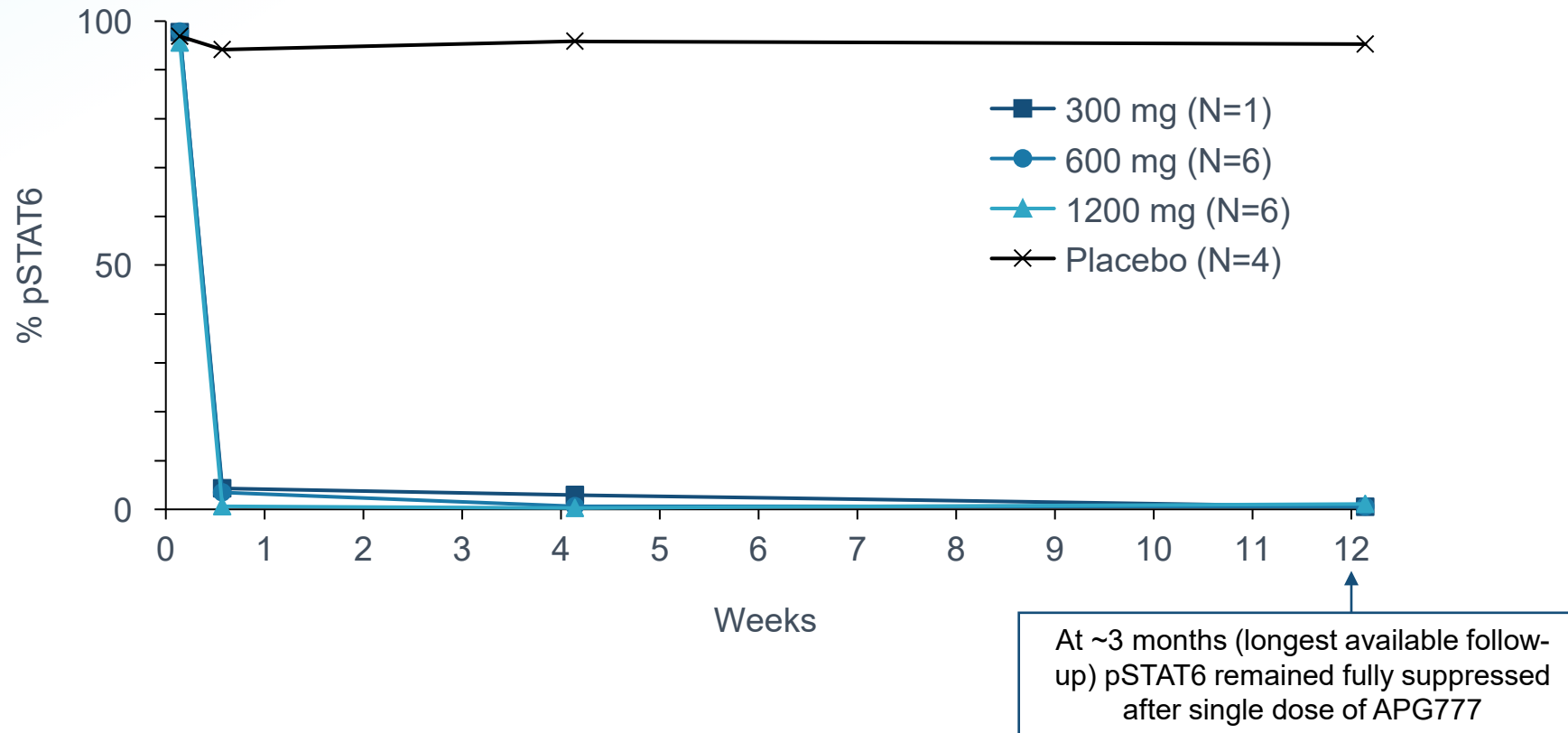
777 tool compound achieved **~2X longer pSTAT6 inhibition** vs. lebrizumab¹

*Note: N = 2 for '777 tool compound arm; N = 3 for lebrizumab arm. Initial pSTAT6 level was normalized to 100% separately for each arm.
1 '777 tool compound sustained at least 50% pSTAT6 inhibition until day 92; lebrizumab sustained 50% pSTAT6 inhibition until day 48

Single dose APG777 showed near complete pSTAT6 inhibition for ~3 months (limit of available follow-up)



Median percent change from baseline in pSTAT6

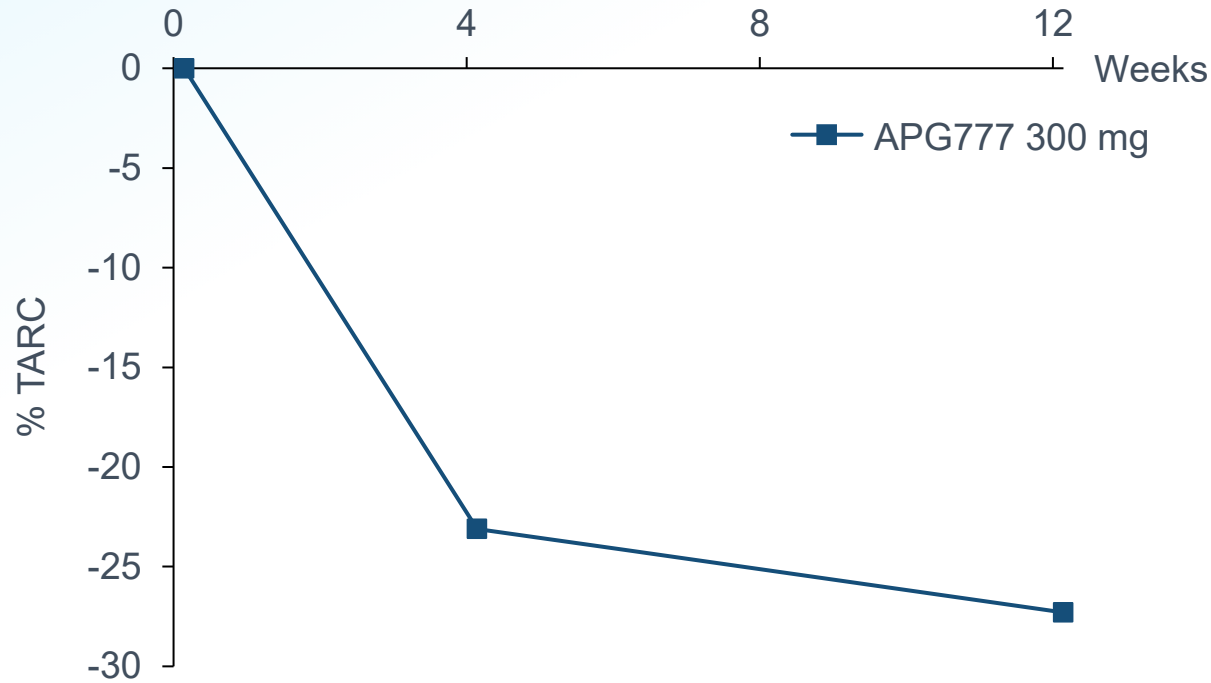


100% pSTAT6 inhibition was demonstrated for approximately 3 months across all doses

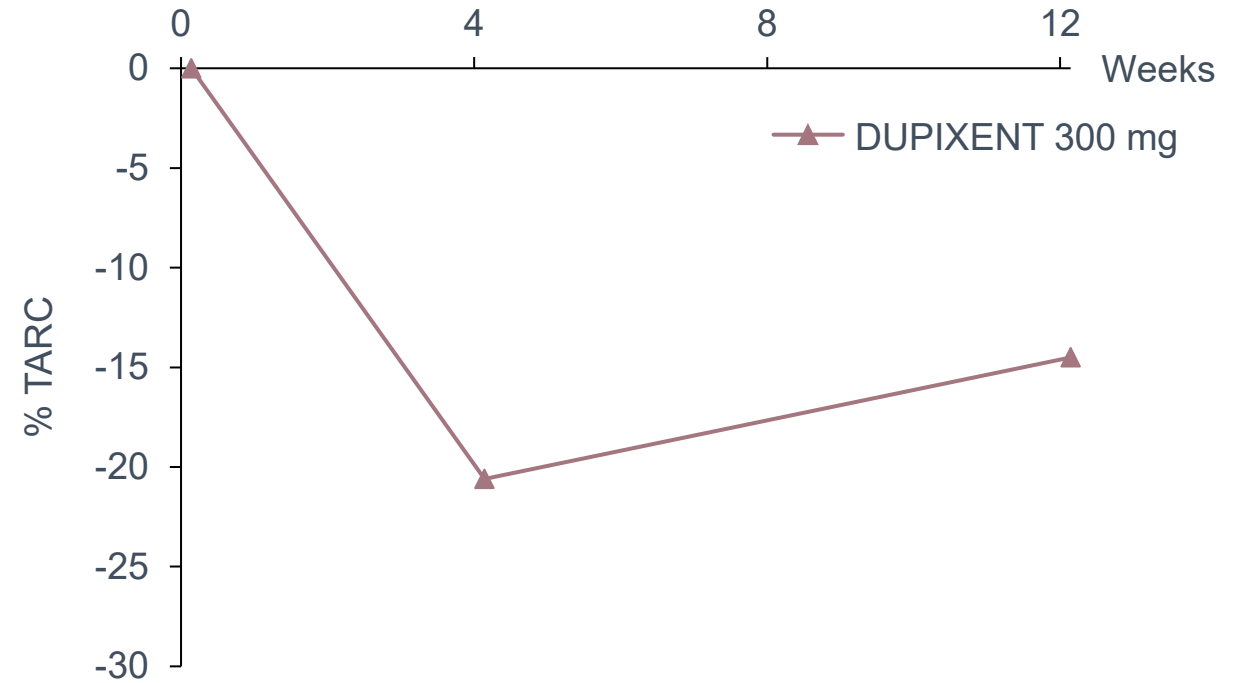


Single dose of APG777 led to deep + sustained TARC inhibition for ~3 months (limit of available follow-up)

Median % changes from baseline in TARC inhibition



Median % changes from baseline in TARC inhibition



- 300 mg APG777 showed similar maximum PD marker changes as DUPIXENT
- APG777 sustained TARC inhibition demonstrates the potential for better durability
- All doses tested of APG777 showed deep TARC inhibition for ~3 months (limit of available follow-up)



APG777 positive PK readout is a key risk-reducing milestone that validates program and pipeline

Antibody attributes

- ✓ Clinically validated IL-13 target
- ✓ Epitope overlaps with lebrikizumab epitope
- ✓ Equivalent or better potency vs. 1st generation mAbs across relevant pre-clinical assays

Clinical profile

- ✓ Well-tolerated with ability to achieve increased exposures in induction for potential improved clinical responses
- ✓ PK data supports every 3- to 6-month maintenance dosing:
 - ~75-day half-life
 - Near maximal pathway suppression for ~3 months (limit of current follow up)

Apogee intends to initiate a Phase 2 in atopic dermatitis in 1H 2024



APG777 Phase 2 in Atopic Dermatitis



APG777 Phase 2 in atopic dermatitis expected to begin 1H 2024 with 16-week efficacy data in 2H 2025



GREATER INDUCTION EXPOSURES

Potential for improved clinical responses (e.g. EASI-75, EASI-90, IGA 0/1) based on ~30-40% greater modeled exposure vs lebrikizumab and ~50% fewer injections



PROLONGED MAINTENANCE DOSING

Every 3- or 6- month maintenance regimens with similar modeled exposure to lebrikizumab Q4W



HIGHER DOSES ENABLED

APG777 180 mg/mL formulation enables 44% greater dose than lebrikizumab in the same volume



INTEGRATED DESIGN

Planned to combine Ph2a and Ph2b elements into a single protocol; significant timeline acceleration over traditional sequenced approach



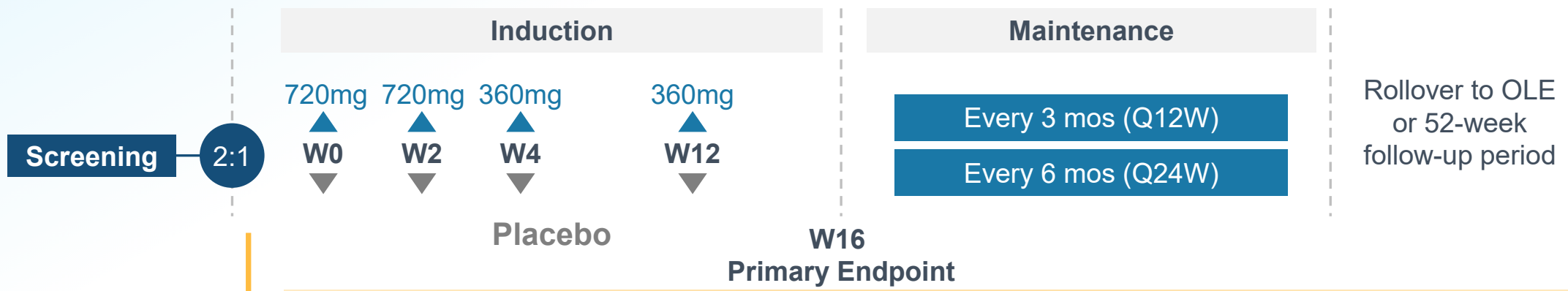
1H 2024 INITIATION

Topline 16 Week data from Part A anticipated in 2H 2025



Planned integrated Phase 2 expected to have 16-week topline data in 2H'25

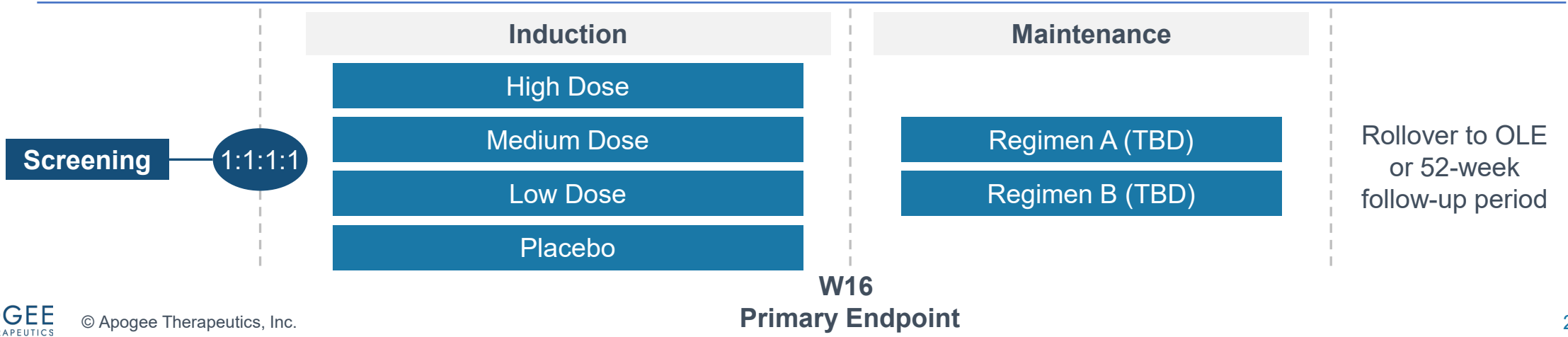
Part A: Proof-of-concept (N ~110)



Integrated Phase 2 design has potential for significant timeline acceleration

- Combines Ph2a and Ph2b elements into a single protocol; All Part A sites are Part B sites (avoiding site startup delays between parts)

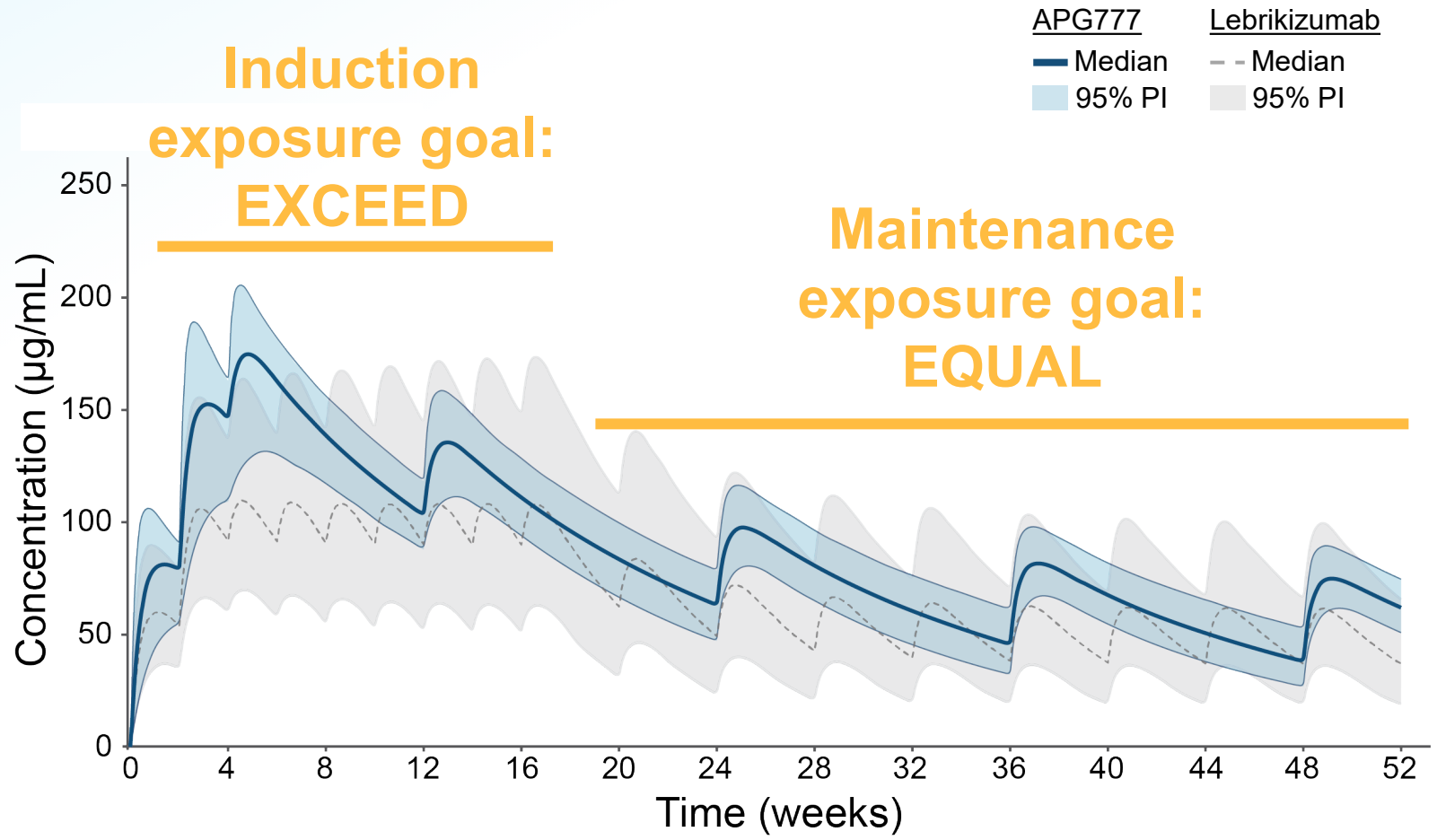
Part B: Dose optimization (N ~360)





APG777 Phase 2 exposures are designed to exceed lebrikizumab in induction and equal in maintenance

Modeled induction and maintenance dosing for APG777¹ and lebrikizumab

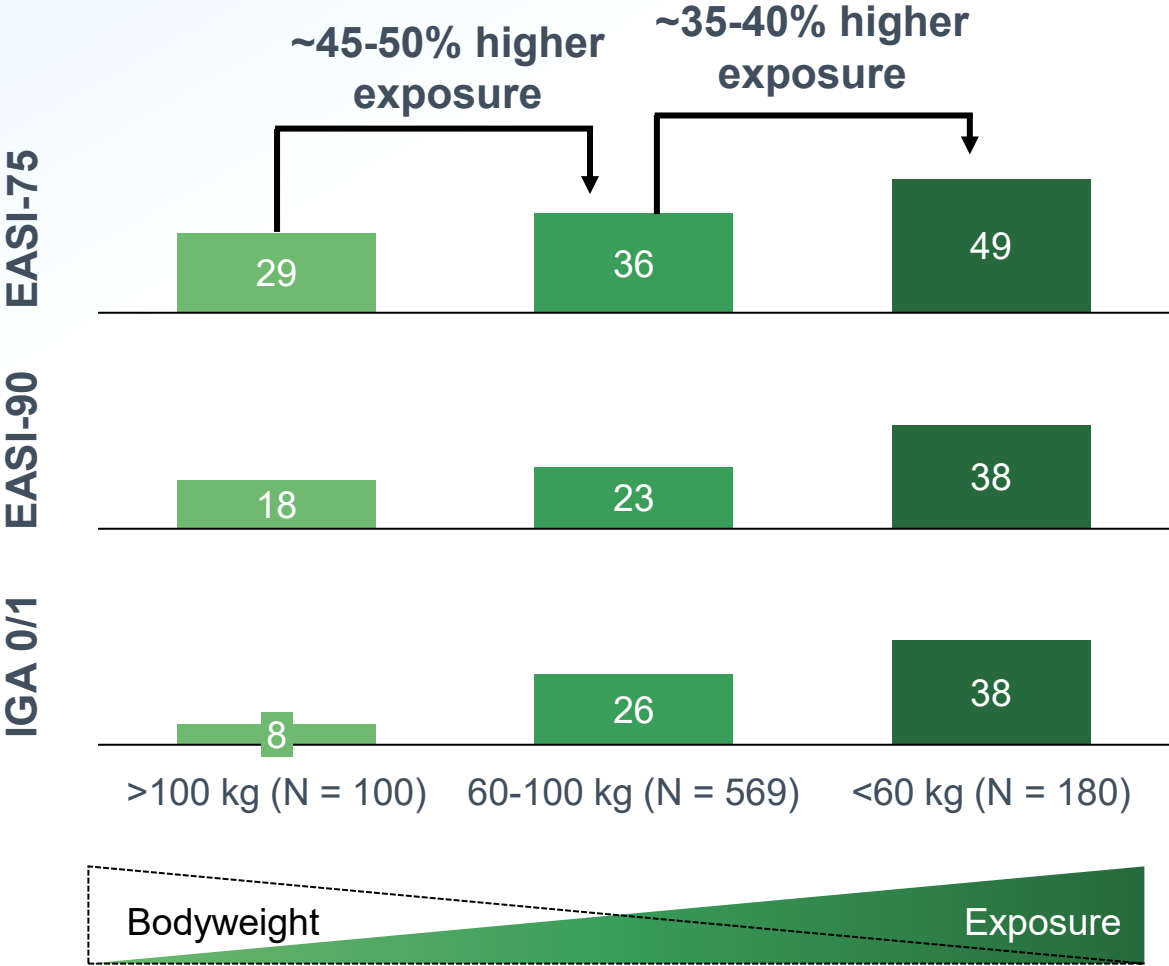


- Lebrikizumab data suggests an **exposure-response (E-R)** for efficacy in induction that underpins our goal to **EXCEED** lebrikizumab induction exposures
- There was **no E-R observed in maintenance for lebrikizumab**; our aim is to **EQUAL** its exposure in maintenance



Lebrikizumab Ph3 appears to show an E-R relationship for efficacy in induction that has not been maximized

Lebrikizumab Ph3 response at Week 16 (Placebo-adjusted), %

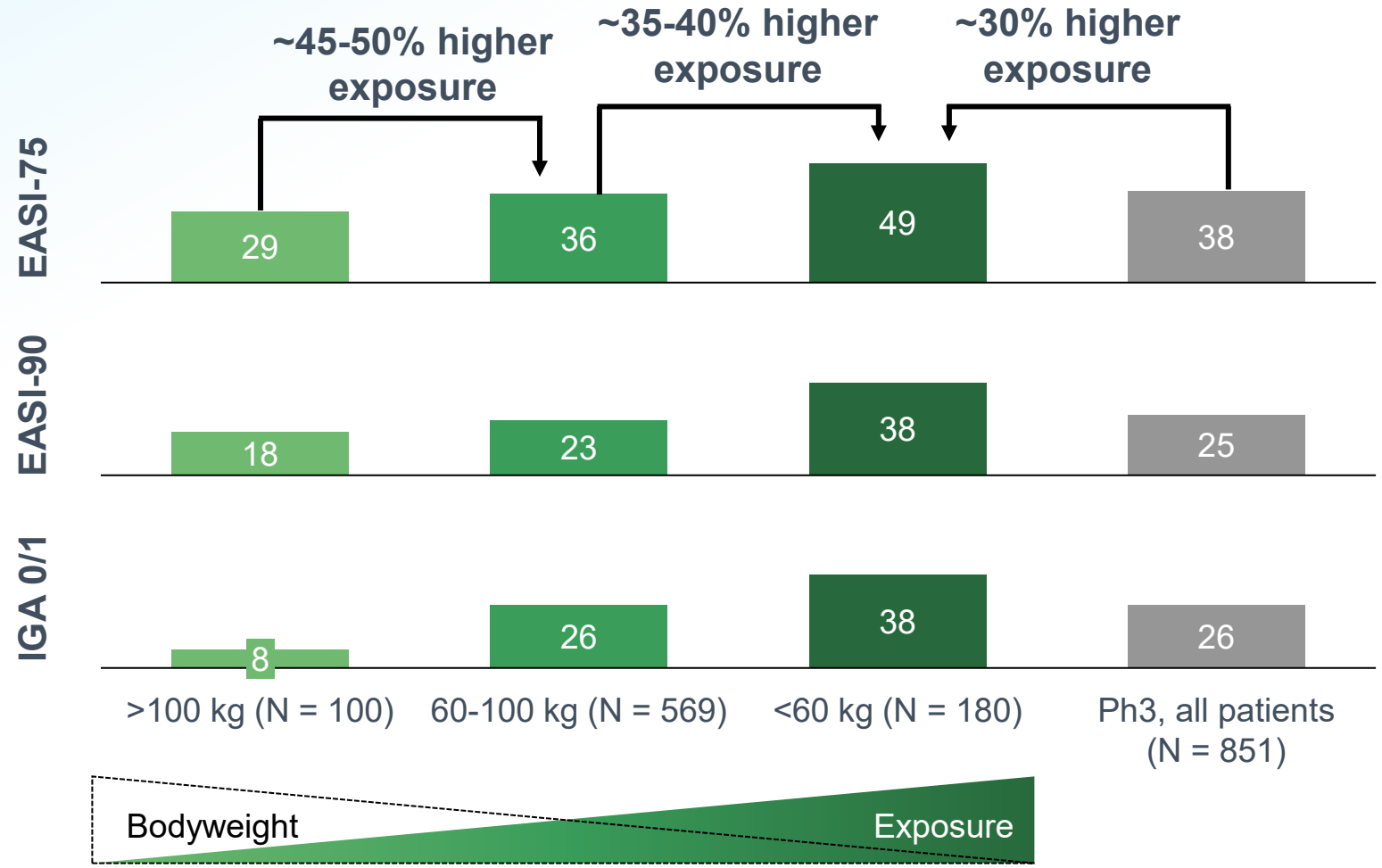


- Lebrikizumab exposure and induction efficacy are both inversely correlated with body weight
- Relationships suggest an **exposure-response for efficacy in induction and support testing higher exposures with APG777**
- In lebrikizumab Ph2b and Ph3 there has been **no dose-AE or exposure-AE relationship**
- **APG777 plans to test ~30-40% higher exposures in induction with ~50% fewer injections**



Lebrikizumab Ph3 appears to show an E-R relationship for efficacy in induction that has not been maximized

Lebrikizumab Ph3 response at Week 16 (Placebo-adjusted), %

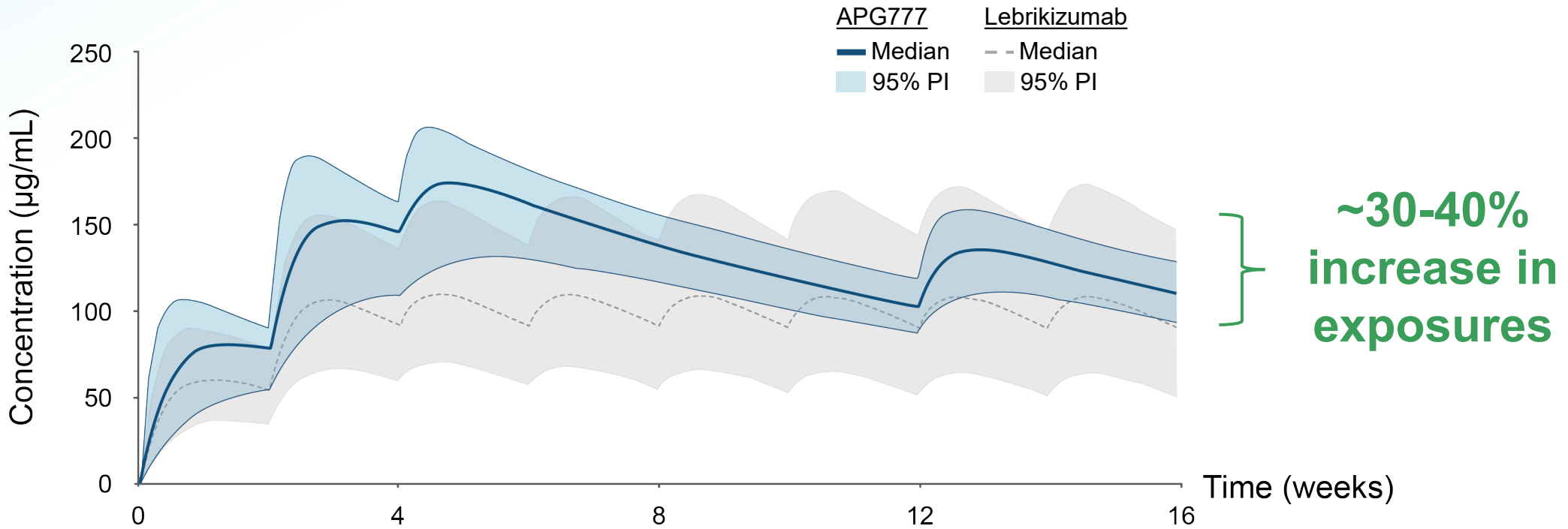


- Lebrikizumab exposure and induction efficacy are both inversely correlated with body weight
- Relationships suggest an **exposure-response for efficacy in induction and support testing higher exposures with APG777**
- In lebrikizumab Ph2b and Ph3 there has been **no dose-AE or exposure-AE relationship**
- **APG777 plans to test ~30-40% higher exposures in induction with ~50% fewer injections**

Modeled Phase 2 induction exposures exceed those of lebrikizumab by ~30-40%



Modeled induction dosing for APG777 and lebrikizumab



Injections

APG777



Lebrikizumab



~30-40% increase in exposures

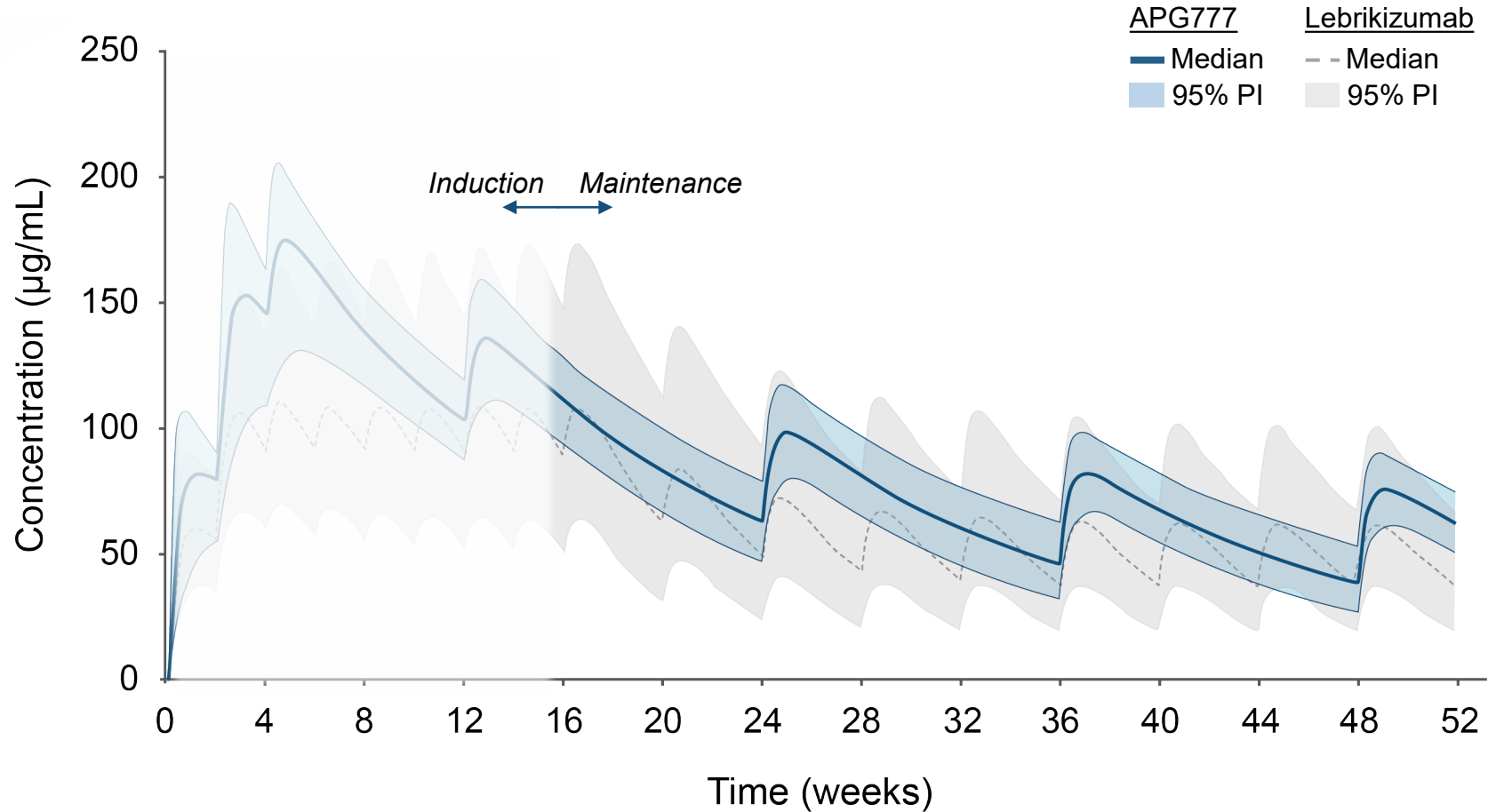
~50% decrease in injections

Modeled Phase 2 Q3M maintenance exposures equal those of lebrikizumab



APG777 Q3M
Aiming for annual maintenance injections:
4 vs 13-26
for
lebrikizumab/
DUPIXENT

Modeled concentration in maintenance



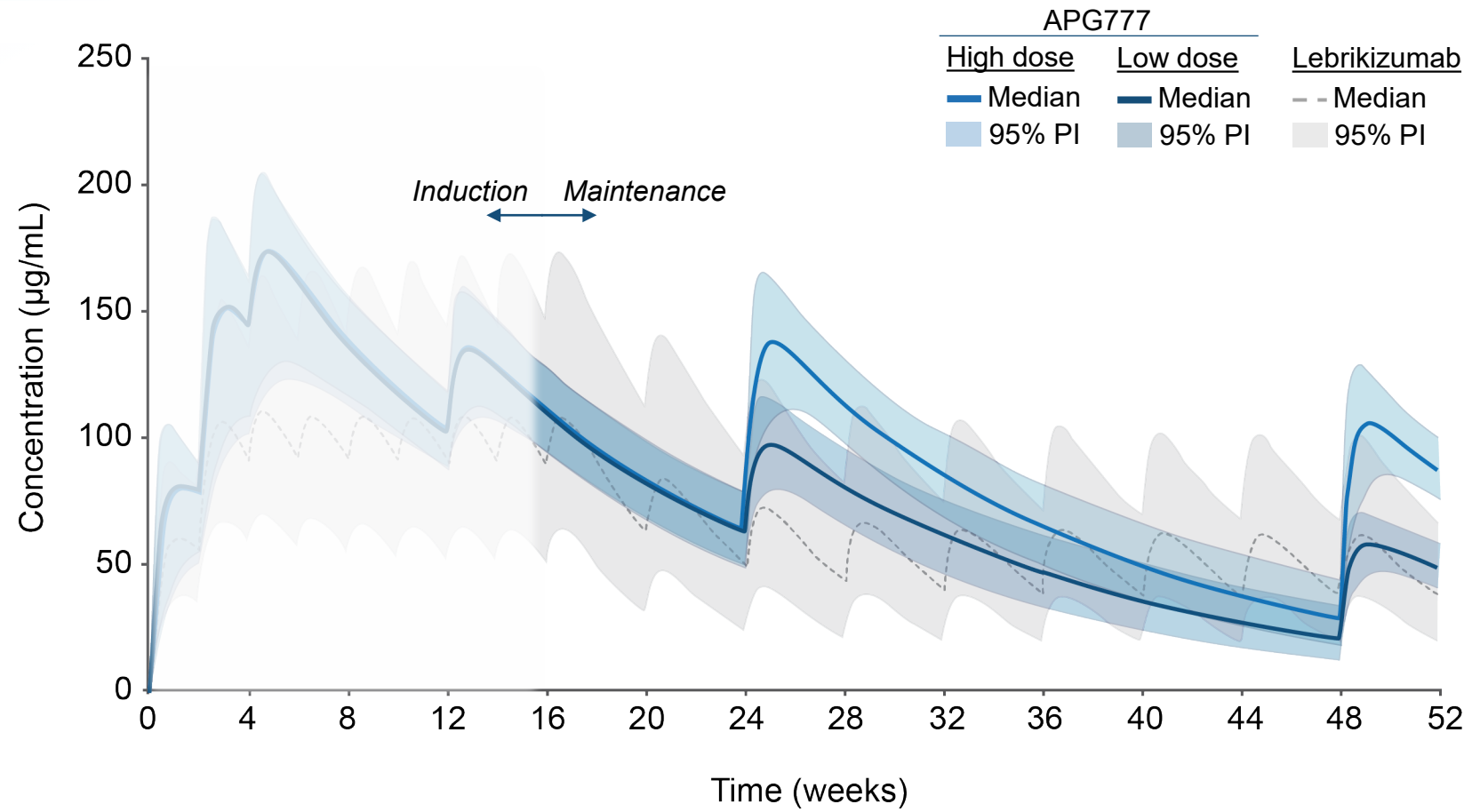
NOTE: The above graph is illustrative only with respect to plans for dosing of APG777 and does not present comparative data.

Modeled Phase 2 Q6M maintenance exposures equal those of lebrikizumab



Modeled concentration in maintenance

APG777 Q6M
 Aiming for annual maintenance injections:
2 vs 13-26
 for
lebrikizumab/
DUPIXENT





Building a Leading I&I Company

APG777's best-in-class Phase 1 PK profile shows potential to be a leading product in the expected \$50B+ AD market¹



Potential for improved clinical responses (e.g. EASI-75, EASI-90, IGA 0/1) based on ~30-40% greater modeled induction exposures than lebrikizumab

- Overlapping epitope and equivalent potency as lebrikizumab ($K_D \leq 100 \text{ pM}$)²
- ~30% higher exposure seen in lebrikizumab low bodyweight group resulted in at least 10 PPT better efficacy than overall study population across all key endpoints

Extended dosing interval addresses clear unmet need

- Potential for every 3- or 6-month dosing to improve patient convenience & compliance

Favorable product characteristics and COGS

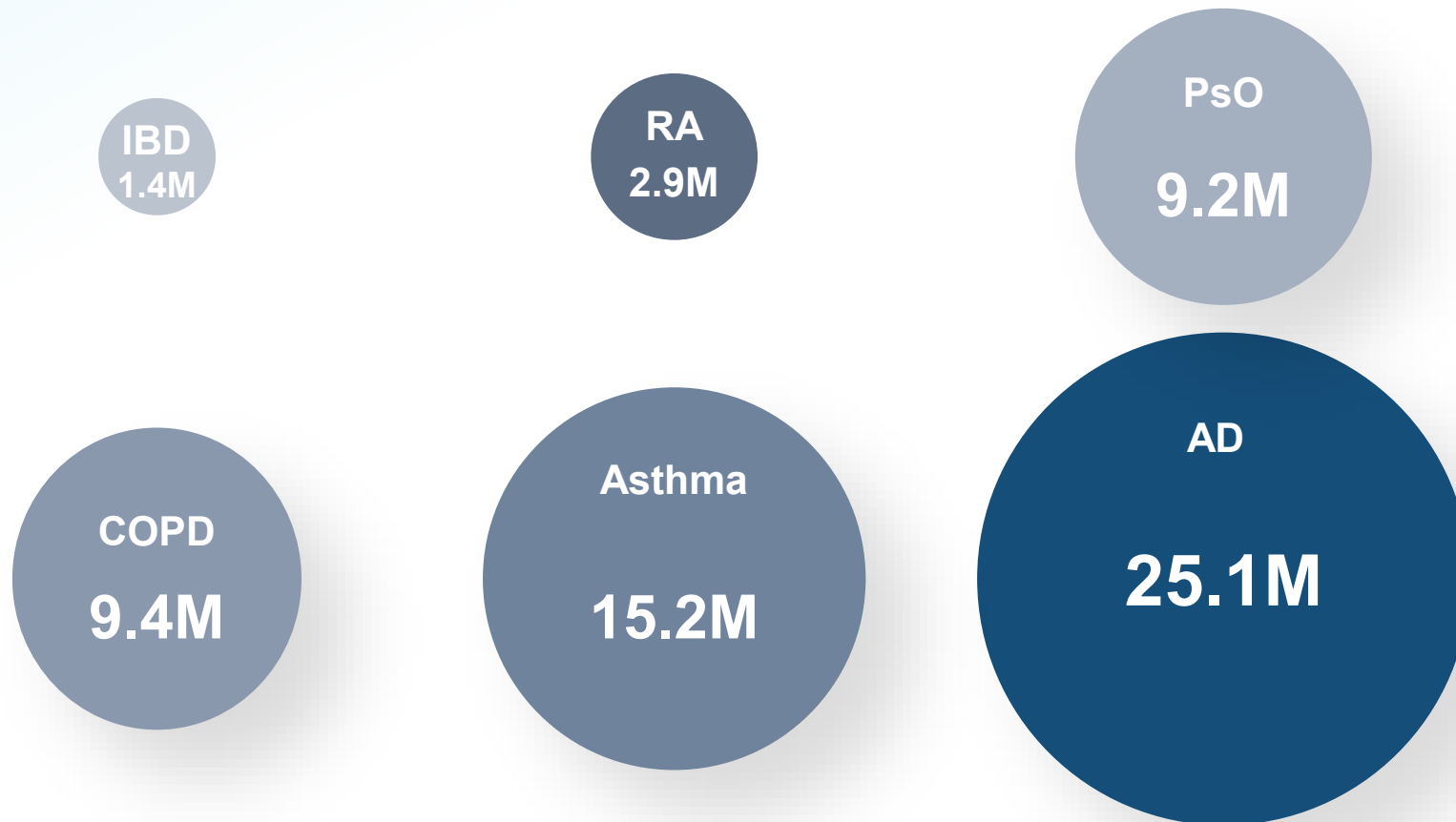
- As few as 2-4 doses per year in maintenance
- Expected improved formulation, manufacturability and viscosity

Novel IP into mid-2040s

AD is the largest of the major I&I markets and projected to grow significantly in the next decade



Estimated population size, MM
Moderate or severe in 7 Major Markets¹

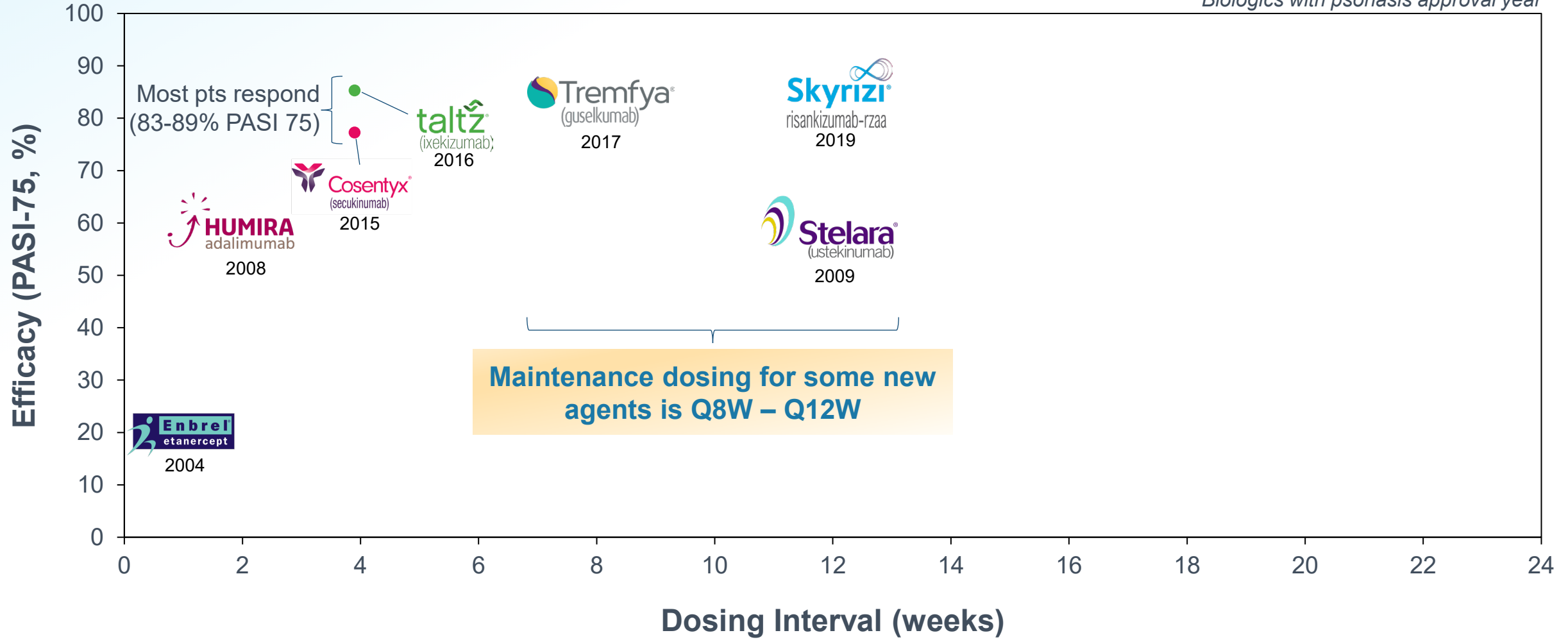


- Psoriasis expected to be a \$30B+ market; **atopic dermatitis (AD) represents a larger opportunity** based on **~3x larger patient population**
- AD biologics penetration is outpacing early years of psoriasis biologics (8% vs 5% at 5 years)
- **AD market is projected to grow more than any other I&I market**



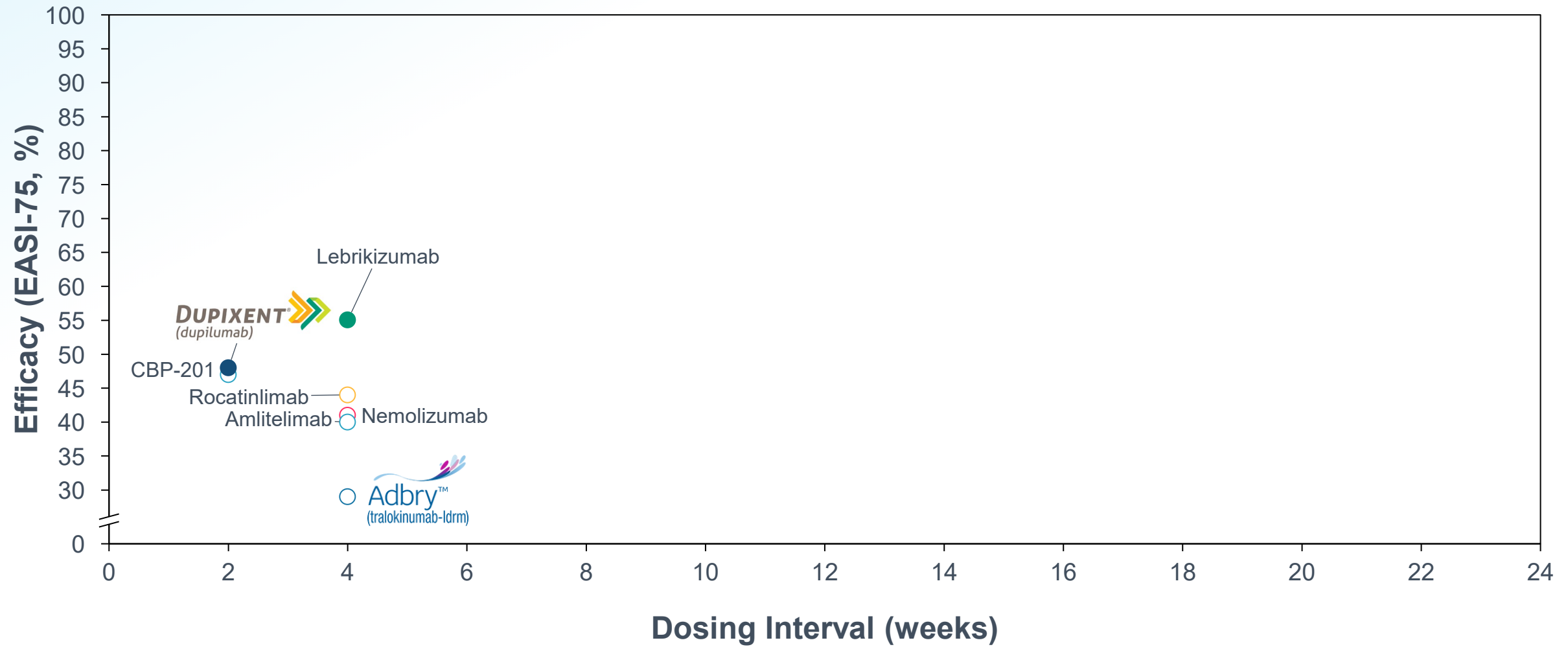
In psoriasis, an analog to AD, Skyrizi has taken the lead with quarterly dosing

Biologics with psoriasis approval year



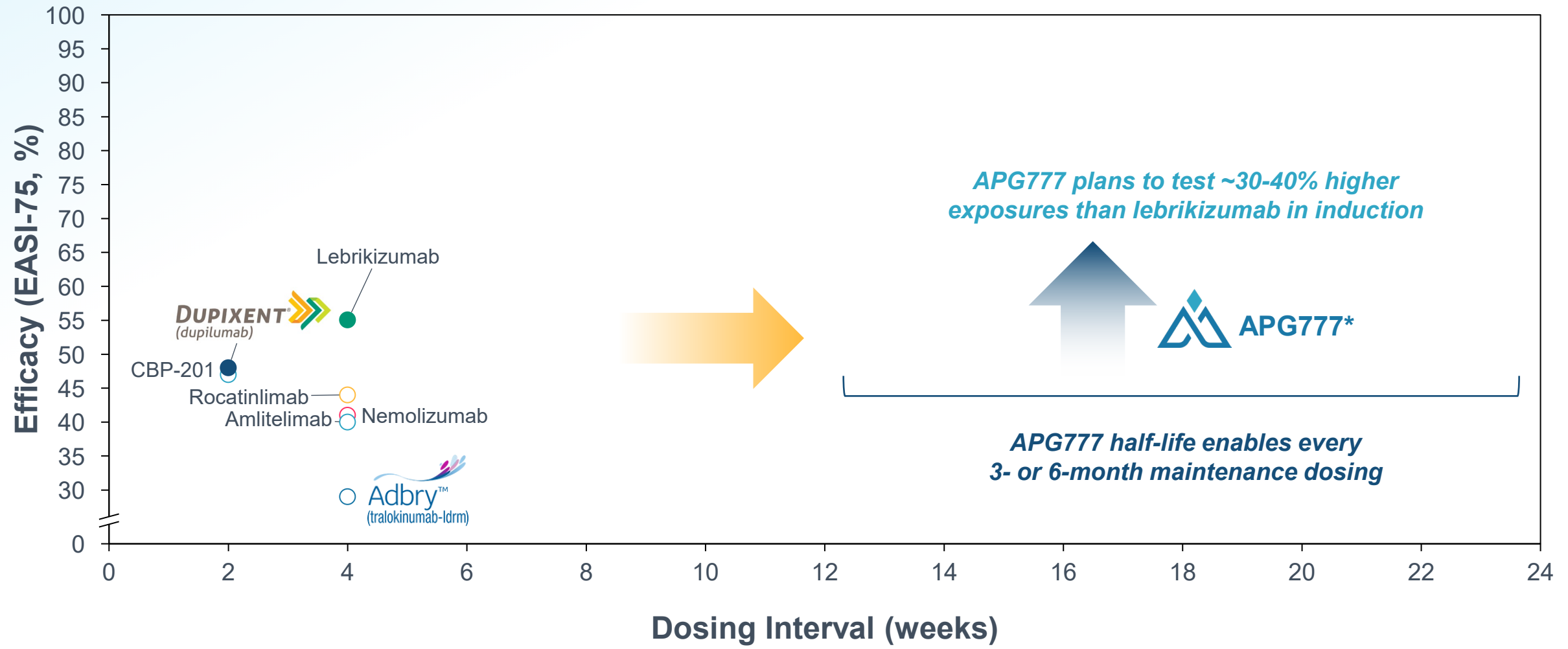


There is significant whitespace in the landscape of approved and in-development biologics for AD





Apogee plans to advance APG777 into a Phase 2 trial with 3- or 6-month maintenance dosing

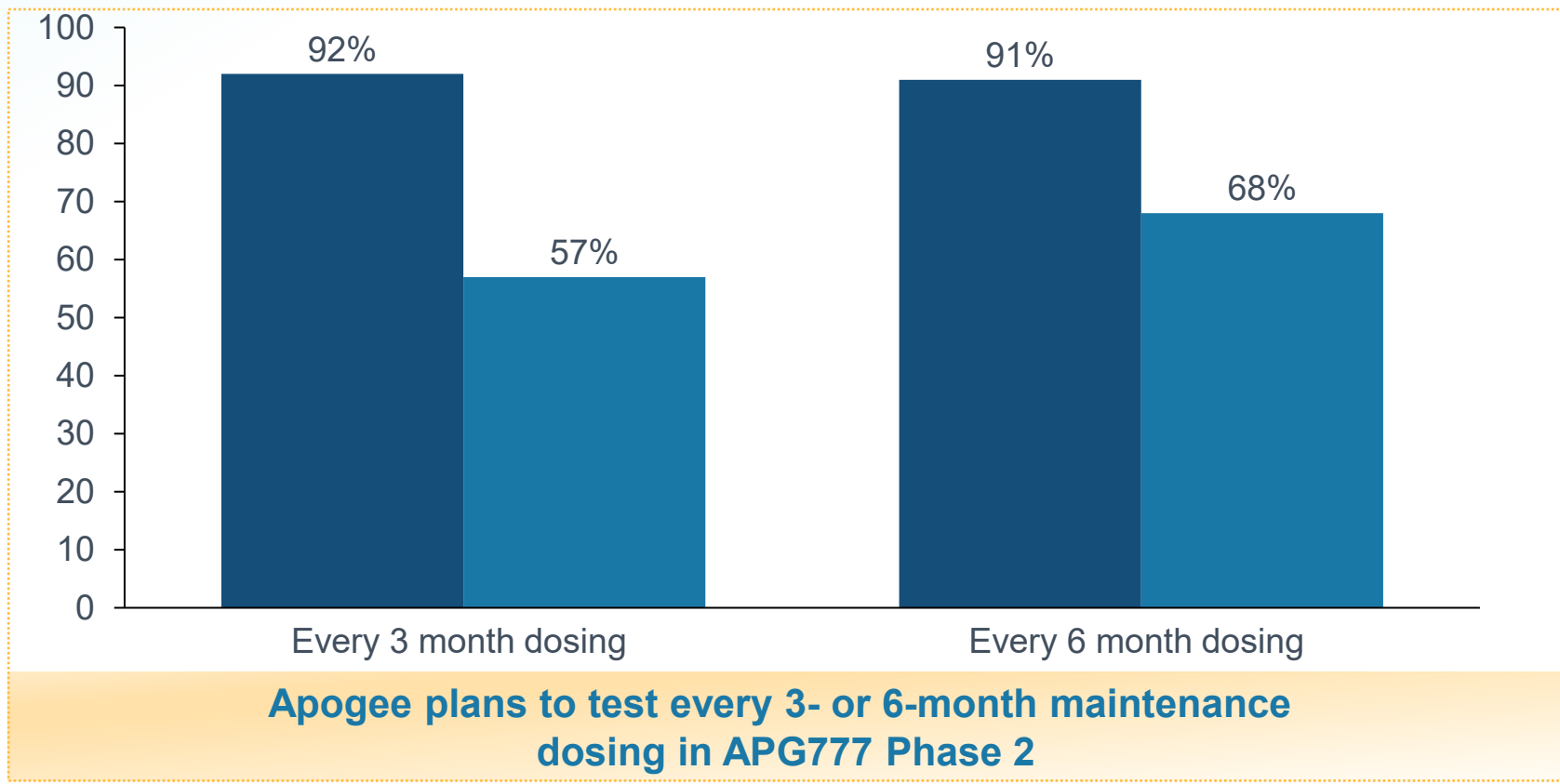




Dermatologists view every 3- or 6-month dosing as highly differentiated

Intent to use a product with APG777 Target Product Profile

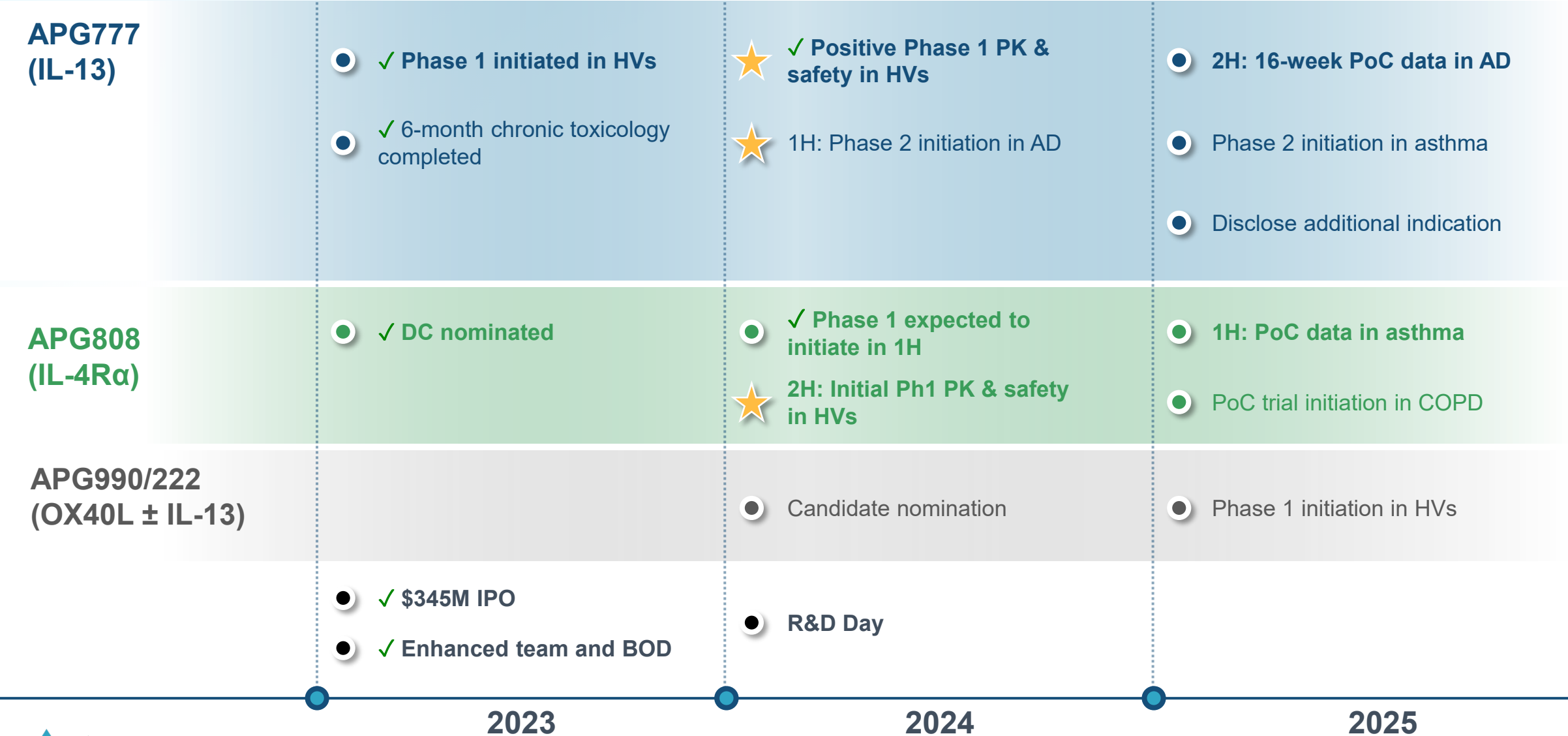
(Assuming every 3-, or 6-month maintenance dosing and equivalent efficacy and safety to DUPIXENT)



- Proportion of new patients (biologic-naïve)
- Proportion of switch patients (currently/formerly on a biologic)



Apogee plans to become a leader in I&I therapeutics





Q&A Backup

APG777 was well-tolerated with a favorable safety profile (TEAEs $\geq 5\%$ across all cohorts, all grades)

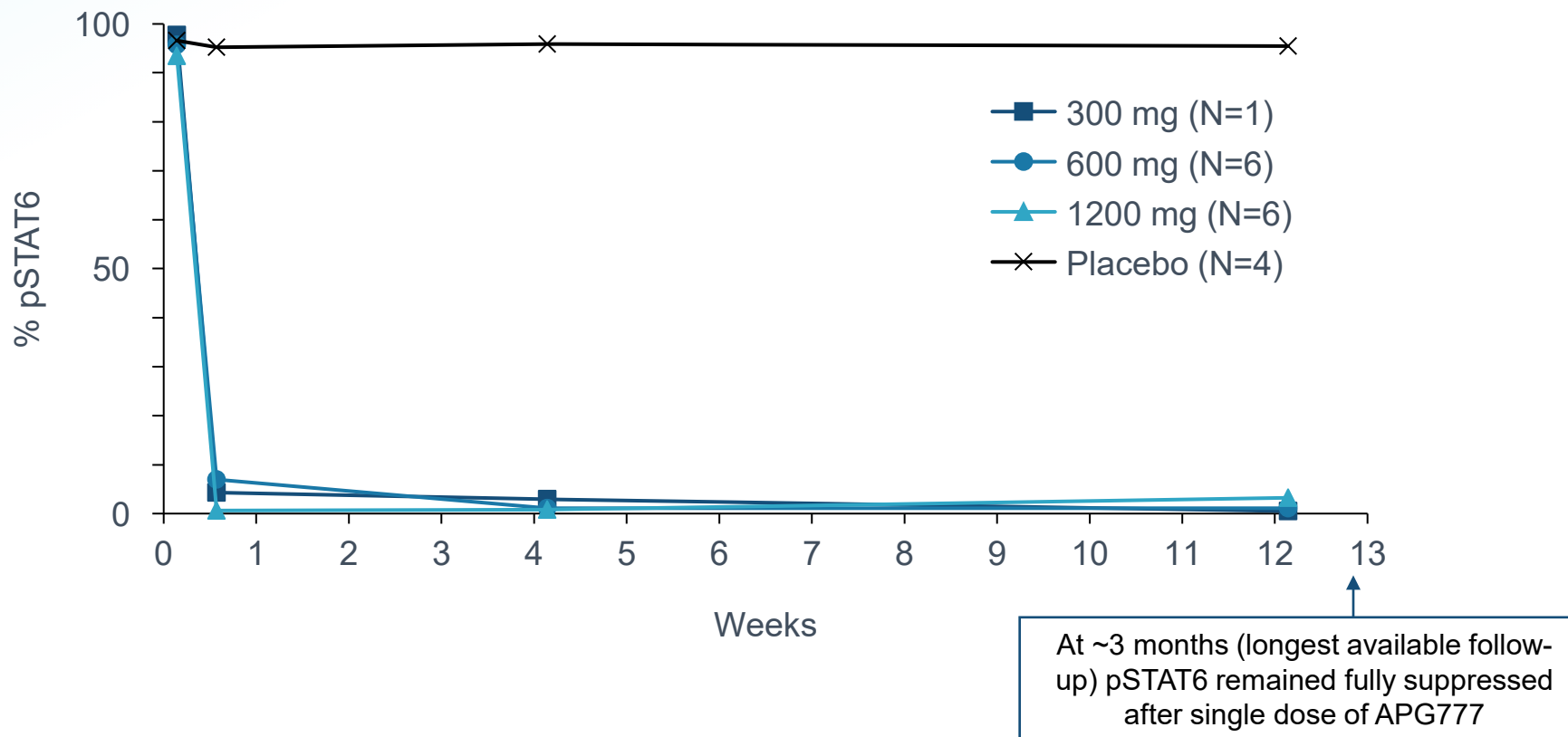


N (%)	Single dose				Multiple dose			Overall trial	
	Placebo N=6	Cohort 1 300 mg N=6	Cohort 2 600 mg N=6	Cohort 3 1,200 mg N=6	Placebo N=4	Cohort 1 300 mg at Day 1, 300 mg at Day 29 N=6	Cohort 2 300 mg at Day 1, 300 mg at Day 15 N=6	APG777 N=30	Placebo N=10
TEAE ($\geq 5\%$ across all cohorts), all grades									
Vascular access site pain*	1 (16.7%)	3 (50.0%)	0	0	0	1 (16.7%)	0	4 (13.3%)	1 (10%)
Vessel puncture site bruise*	2 (33.3%)	0	0	0	1 (25.0%)	2 (33.3%)	0	2 (6.7%)	3 (30%)
Headache	0	0	1 (16.7%)	1 (16.7%)	0	2 (33.3%)	0	4 (13.3%)	0 (0%)
Vascular access site bruising*	1 (16.7%)	1 (16.7%)	1 (16.7%)	0	0	1 (16.7%)	0	3 (10%)	1 (10%)
Back pain	1 (16.7%)	0	1 (16.7%)	1 (16.7%)	0	0	0	2 (6.7%)	1 (10%)
Injection site bruising*	1 (16.7%)	0	0	2 (33.3%)	0	0	0	2 (6.7%)	1 (10%)
Neutrophil count decrease	3 (50.0%)	0	0	0	0	0	0	0	3 (30%)
Contusion	1 (16.7%)	0	0	0	0	1 (16.7%)	0	1 (3.3%)	1 (10%)
Cough	0	1 (16.7%)	0	0	0	1 (16.7%)	0	2 (6.7%)	0 (0%)
Dermatitis contact	1 (16.7%)	0	0	0	1 (25.0%)	0	0	0	2 (20%)
Diarrhea	0	1 (16.7%)	0	1 (16.7%)	0	0	0	2 (6.7%)	0
Nausea	0	0	1 (16.7%)	1 (16.7%)	0	0	0	2 (6.7%)	0
Oropharyngeal pain	0	1 (16.7%)	0	0	0	0	1 (16.7%)	2 (6.7%)	0
Pain in extremity	1 (16.7%)	0	1 (16.7%)	0	0	0	0	1 (3.3%)	1 (10%)
Upper respiratory tract infection	1 (16.7%)	0	0	0	0	1 (16.7%)	0	1 (3.3%)	1 (10%)

Single dose APG777 showed near complete pSTAT6 inhibition for ~3 months (limit of available follow-up)



Mean percent change from baseline in pSTAT6



100% pSTAT6 inhibition was demonstrated for approximately 3 months across all doses



Company Overview

Apogee plans to transform the I&I space



FOCUS

Engineering antibodies with potential **best-in-class** profiles in largest I&I indications with **highly differentiated dosing**

APPROACH

Technology approach **proven** to create antibodies with significantly **extended half-life** and other optimized properties

EXPANSION

Pipeline-in-a-product potential via **indication expansion** and **combination** approaches

PIPELINE

Four programs leveraging **well-established mechanisms** and addressing I&I indications with **multi-billion-dollar potential**

Program / Target	Discovery	Preclinical	Phase 1	Phase 2	Phase 3
APG777 IL-13 <i>Same MOA as lebrikizumab</i>	Atopic Dermatitis			1H 2024: Phase 2 trial initiation ¹ 2H 2025: 16-week proof-of-concept data in AD patients	
	Asthma			2025: Phase 2 trial initiation ¹	
APG808 IL-4Rα <i>Same MOA as DUPIXENT</i>	COPD		1H 2024: Phase 1 initiation in HV 2H 2024: Initial Phase 1 PK and safety in HV 2025: Proof-of-concept trial initiation in COPD		
APG990 OX40L <i>Same MOA as amlitelimab</i>	Atopic Dermatitis	2024: Candidate nomination 2025: Phase 1 initiation in HV			
APG222 Combination IL-13 and OX40L	Atopic Dermatitis				

Apogee mAbs are engineered for best-in-class properties, including half-life extension



Based on clinically-validated epitopes with performance across five properties:

 **Backbone**

 **Potency**

 **PK**

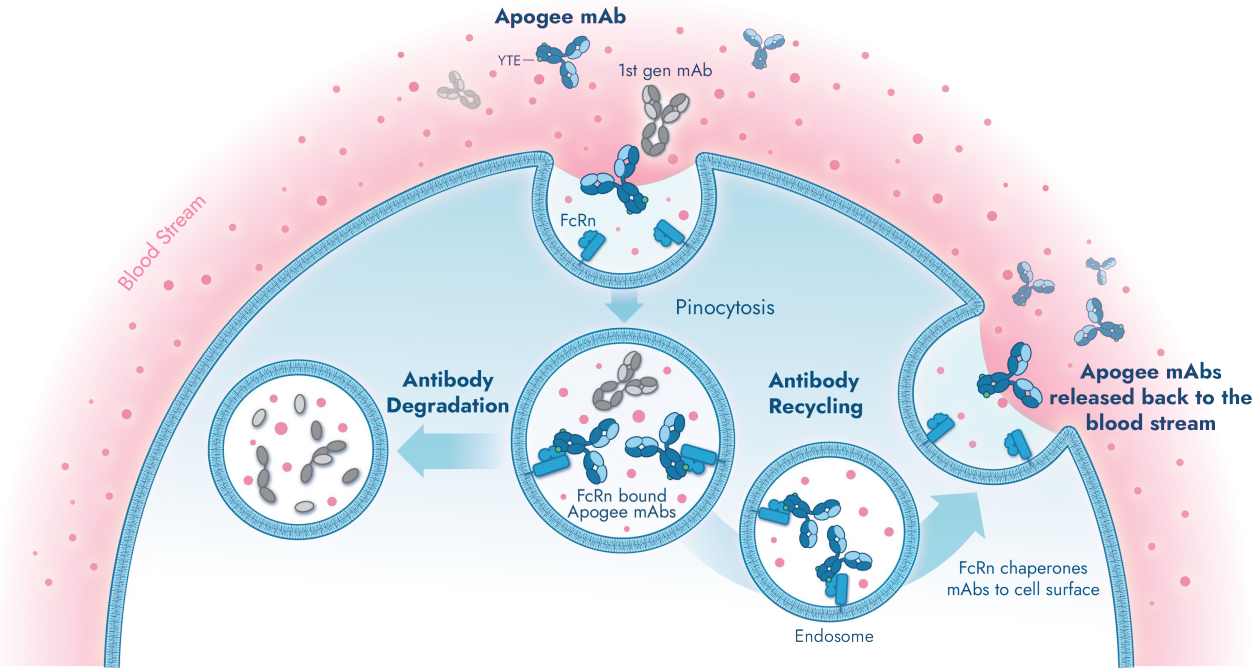
 **Stability**

 **Viscosity**

• Designed to maximize antibody recycling
• Drug exists at higher levels for longer effect

Potential for PK that:

- *Optimizes exposures*
- *Decreases variability*
- *Increases half-life*



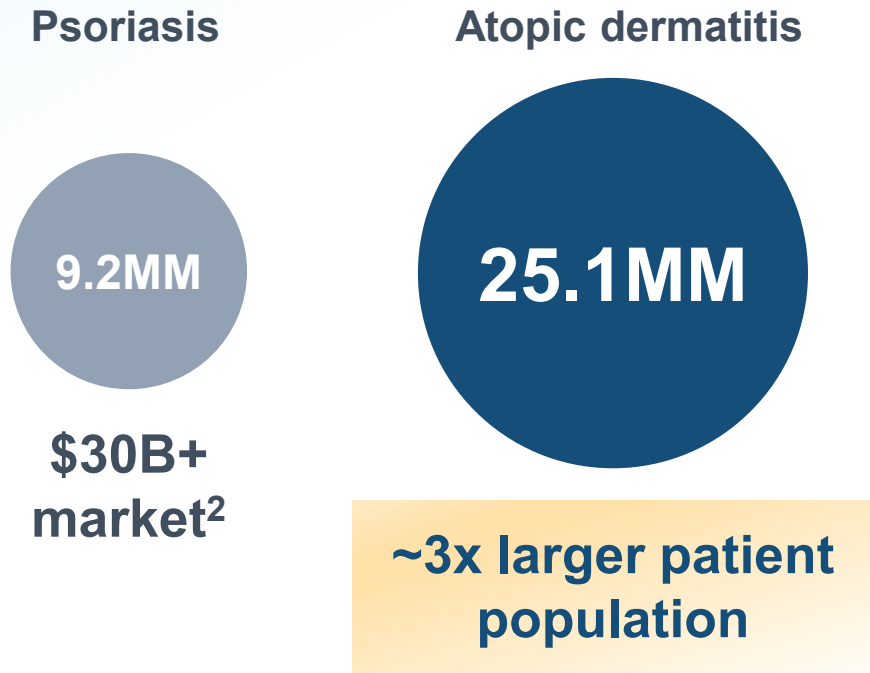


APG777



AD represents a larger opportunity than psoriasis; AD biologics penetration mirrors early years of psoriasis

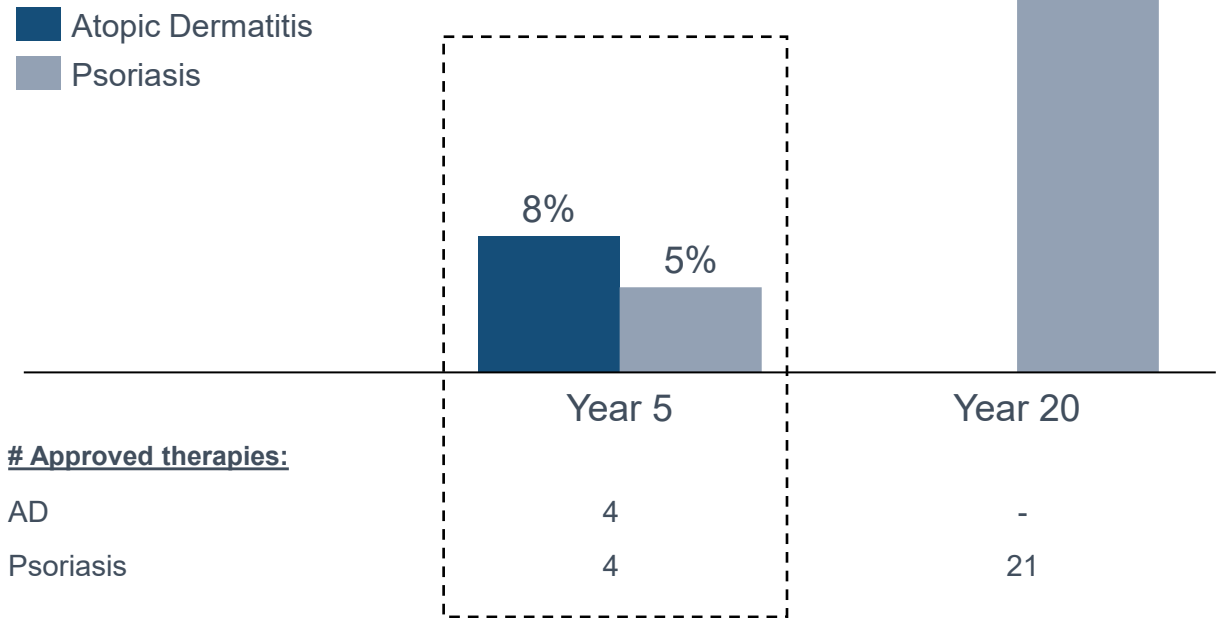
Population size, MM
Moderate or severe in 7 Major Markets¹, 2020



Psoriasis expected to be a \$30B+ market; atopic dermatitis (AD) represents a larger opportunity

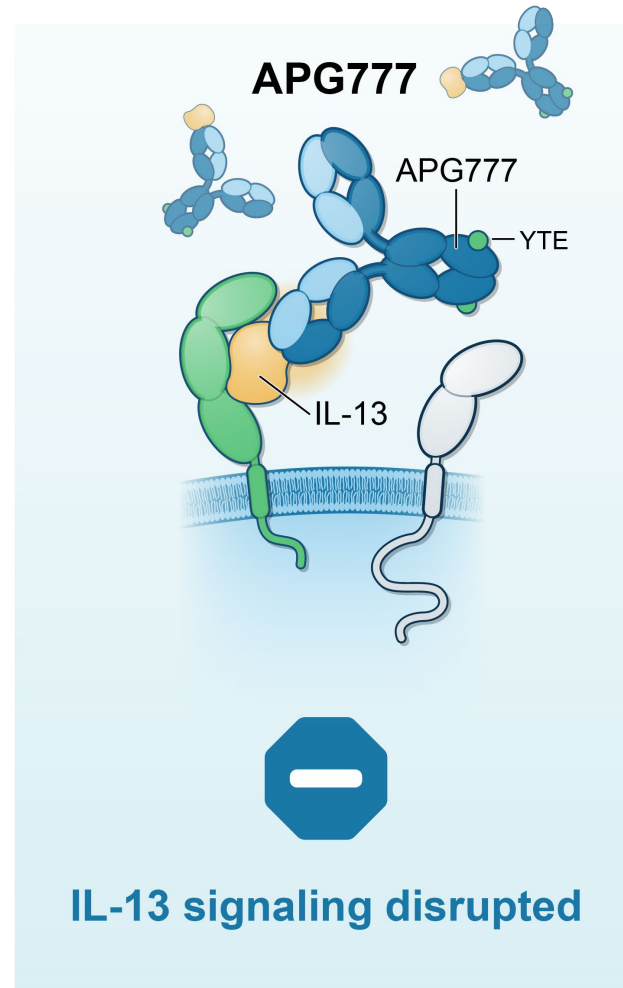
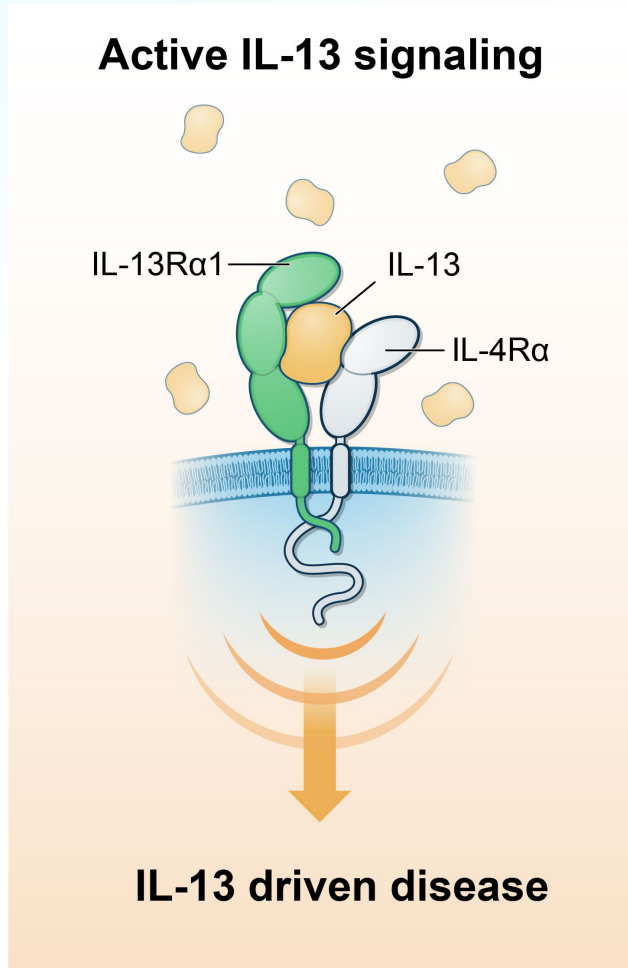
Penetration of approved systemic therapy in AD expected to ramp 8% → 25%+ by 2032

Penetration in years after launch (US)³



More convenient dosing could potentially expand AD biologics' penetration beyond projected 25%+

APG777 is designed to disrupt Th2 signaling by preventing formation of IL-13R α 1 / IL-4R α heterodimer

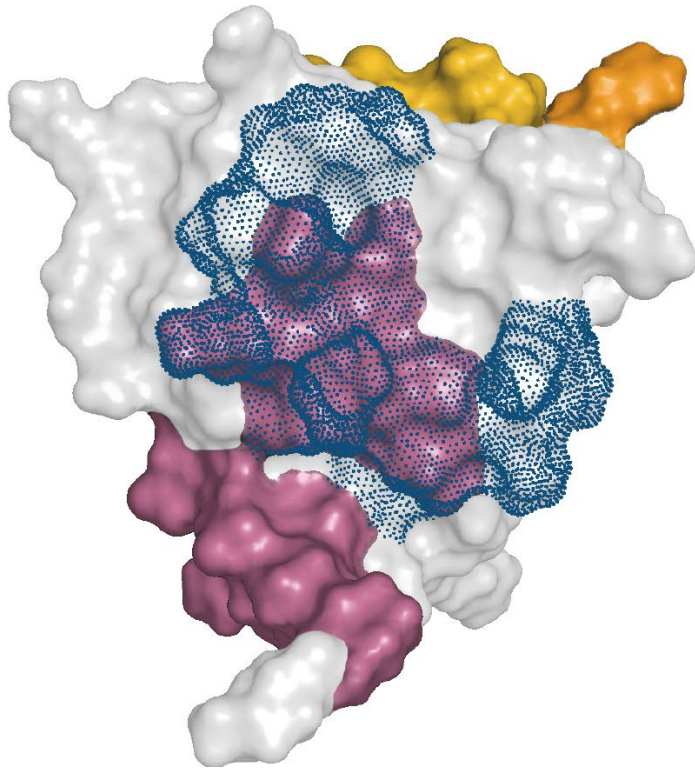


- IL-13 signaling **begins with binding of IL-13 to IL-13R α 1**
- This forms an inactive complex that then **binds to IL-4R α to create a complete, active heterodimer**
- **Active IL-13R α 1 / IL-4R α heterodimer sets off a signaling cascade** that leads to:
 - Skin barrier defects
 - Immune cell recruitment
 - Tissue inflammation
 - Lichenification (skin thickening)
 - Pruritis (skin itching)

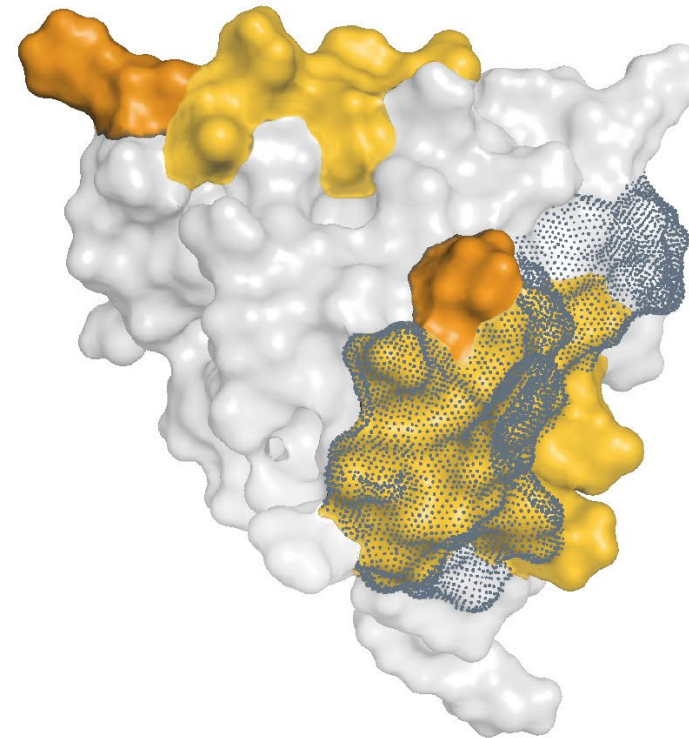
APG777's epitope overlaps with lebrikizumab, differentiating from other approaches to target IL-13



Human IL-13 (side 1)



Human IL-13 (side 2)



180°

Epitope (binding region)

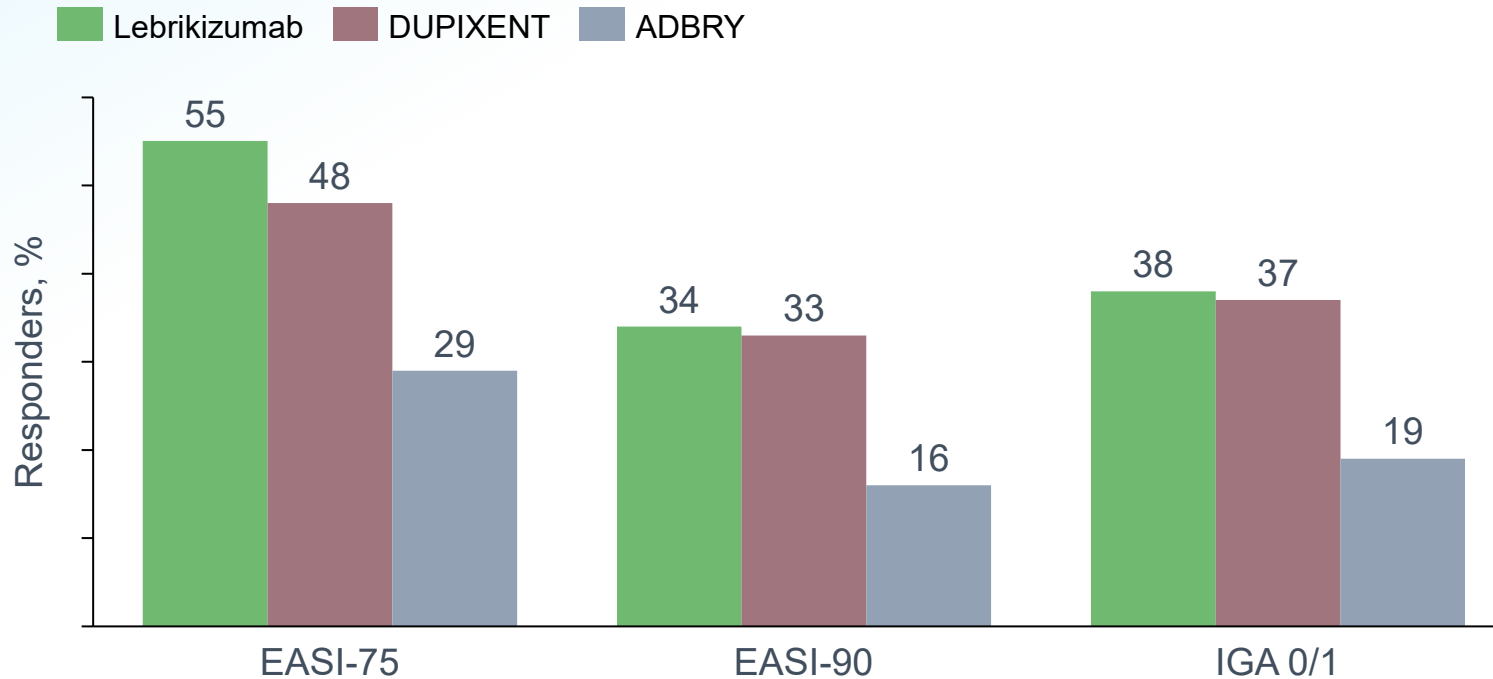
- APG777/lebrikizumab (overlapping region)
- IL-4Rα
- ADBRY
- IL-13Rα1/IL-13Rα2 (overlapping region)
- IL-13Rα2

APG777's mechanism of action disrupts Th2 signaling by blocking IL-4Rα binding and subsequent formation of the IL-13Rα / IL-4Rα heterodimer



Lebrikizumab and DUPIXENT have similar efficacy across key AD endpoints

Efficacy of biologics in AD (week 16)

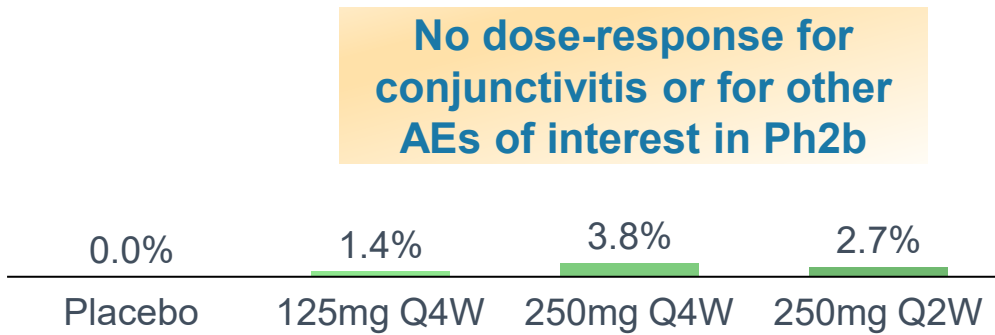


Targeting the key pathogenic step in AD, like lebrikizumab and DUPIXENT, has consistently resulted in high efficacy

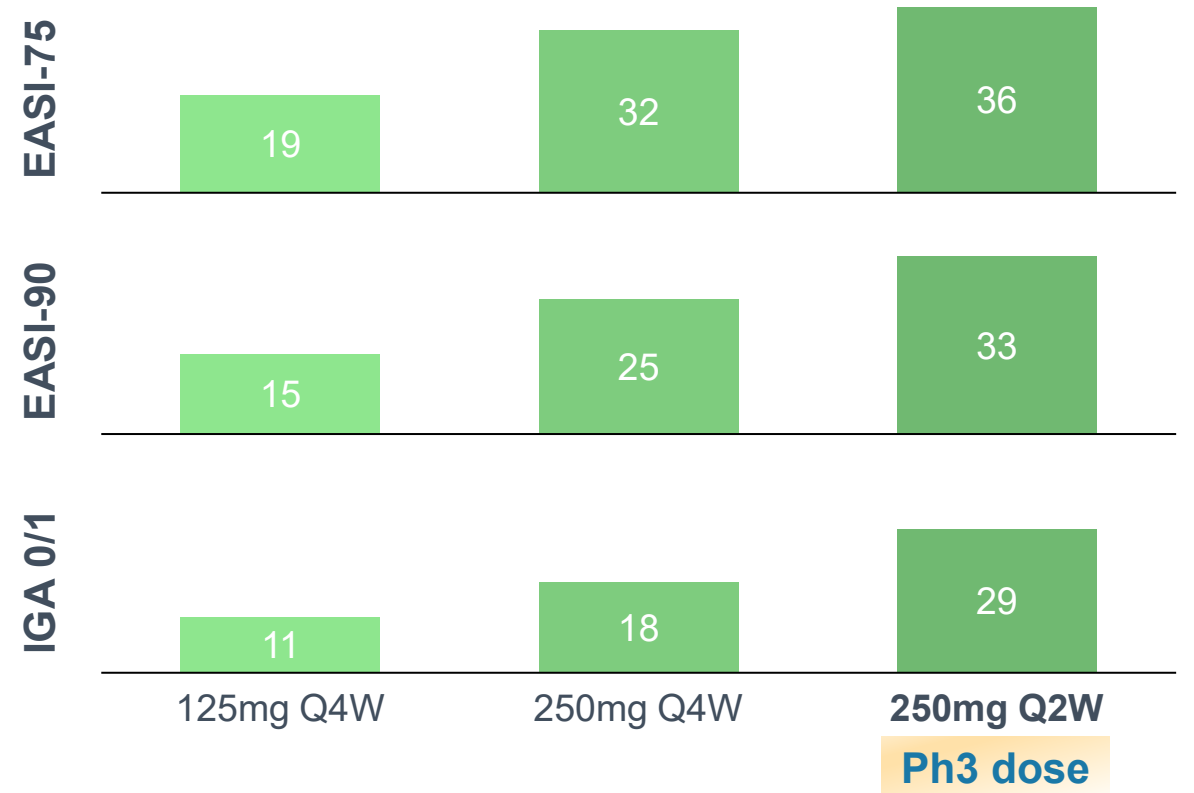
- Lebrikizumab and DUPIXENT show **consistently high results across all important efficacy parameters**
 - Mechanistically, **both target the key pathogenic step in AD**, the heterodimerization of IL-4R α and IL-13R1, which may explain the similar efficacy observed
- However, both are **dosed every other week⁴**, a burden for patients
- Lebrikizumab showed, at minimum, equivalent maintenance efficacy for both Q2W and Q4W dosing, a main differentiator from DUPIXENT

Lebrikizumab showed greater efficacy with higher doses in Ph2b with no dose-dependent increases in AE rates

Conjunctivitis rates by dose level in lebrikizumab Ph2b



Response at 16 weeks (placebo-adjusted), % by dose level in lebrikizumab Ph2b



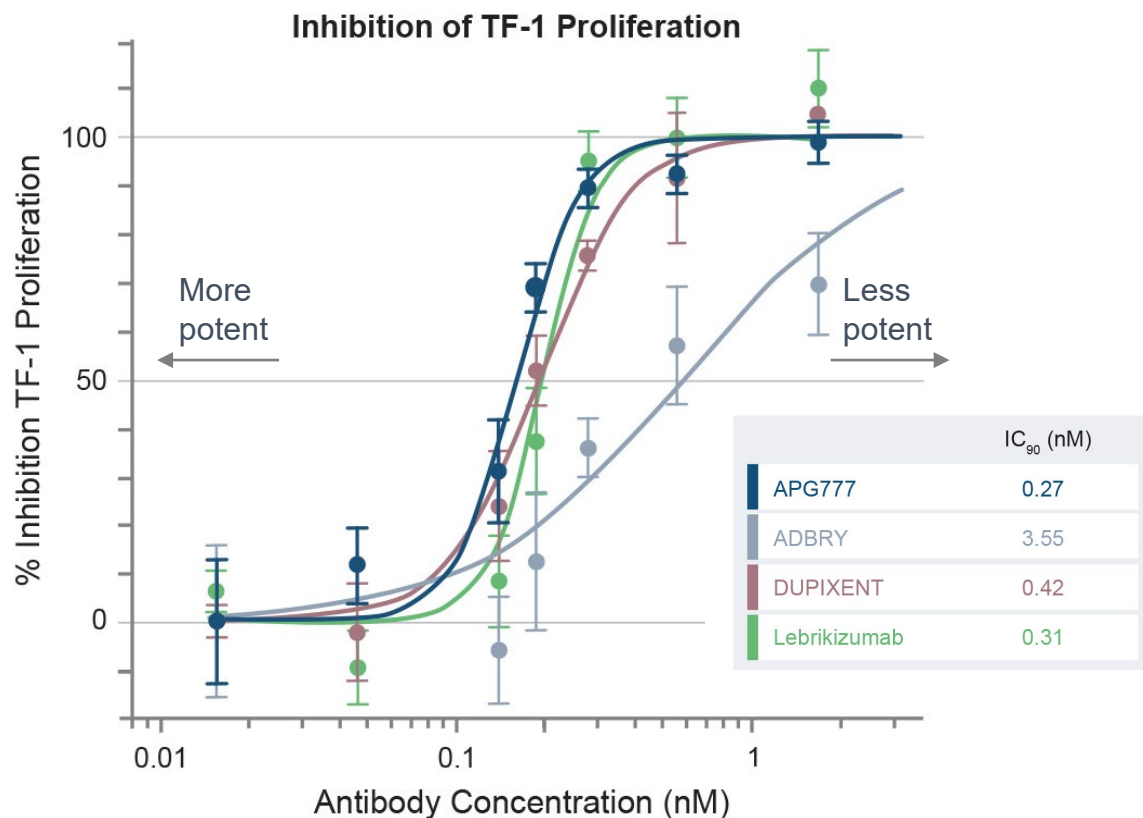
With no plateau in efficacy across doses, a higher dose and/or greater exposures could lead to better efficacy

APG777 is as potent as lebrikizumab and DUPIXENT in key preclinical assays



APG777 vs DUPIXENT, ADBRY, and lebrikizumab on key potency assay

Additional *in vitro* assays support APG777 potency

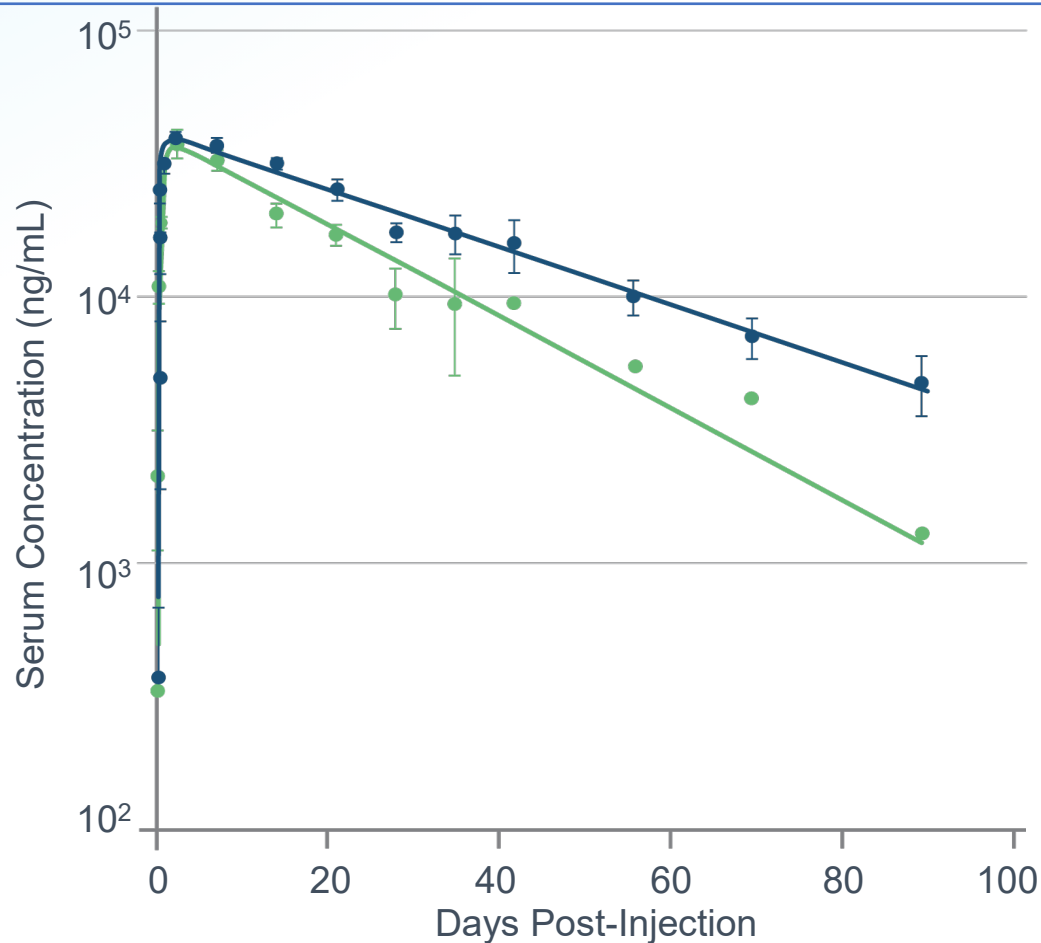


Assay	Affinity to human IL-13 by SPR	Inhibition of STAT-6 phosphorylation	Inhibition of TARC secretion
Measurement	K _D (pM)	IC ₉₀ (nM)	IC ₉₀ (nM)
APG777	78	0.56	1.40
ADBRY	116	1.34	27.96
DUPIXENT		0.58	13.41
Lebrikizumab	131	0.46	1.37

APG777 NHP half-life is significantly longer than lebrikizumab



NHP PK, SQ administration



APG777 has advantages over lebrikizumab in our NHP head-to-head studies

NHP average half-life¹

APG777: 28 days

Lebrikizumab: 18 days

- APG777 shows extended half-life in NHPs
- APG777 had decreased PK variability with potential for greater consistency in response

APG777 can potentially achieve every 2- or 3-month maintenance dosing vs Q4W for lebrikizumab and Q2W for DUPIXENT



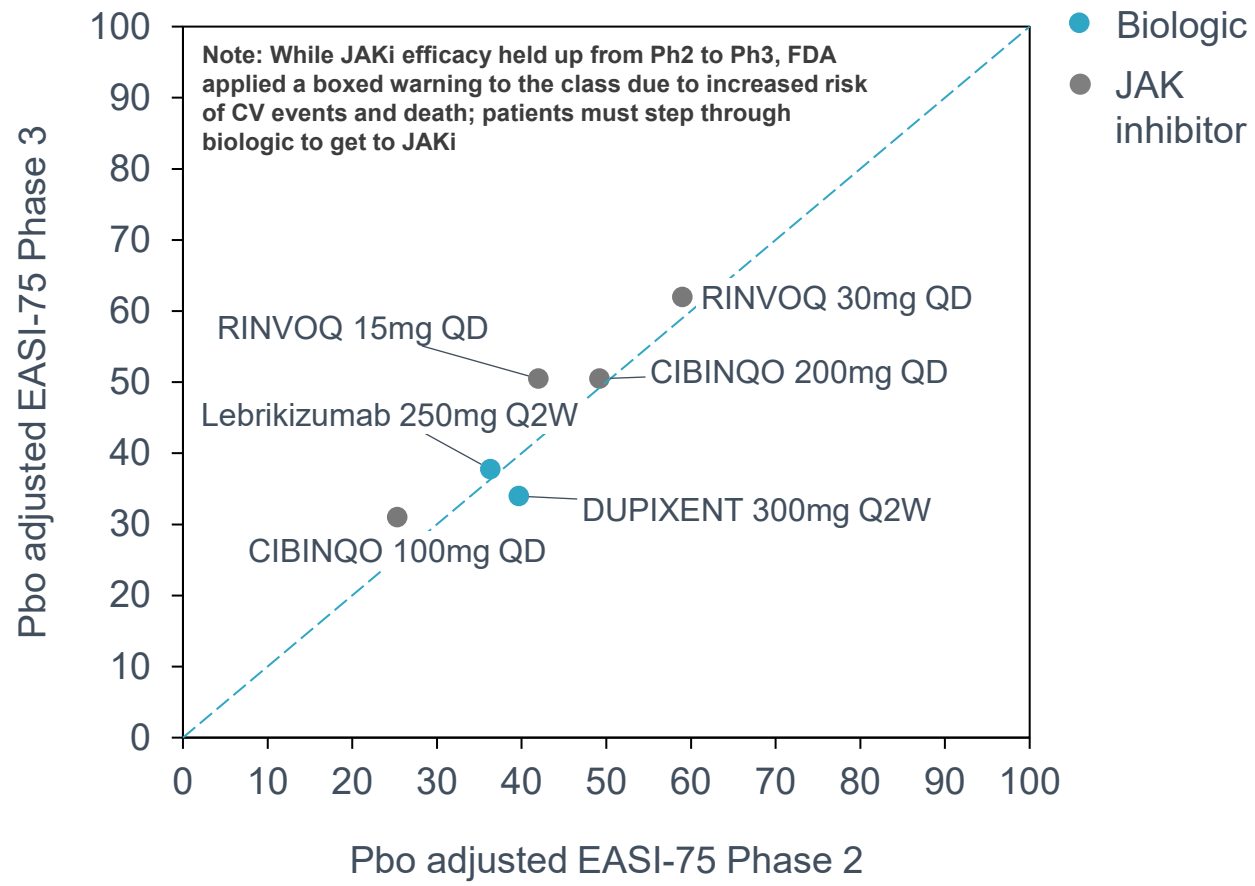
Strong historical correlation between Phase 2 and 3 data makes APG777 16-week AD data a key catalyst

Phase 2 16-week data in atopic dermatitis planned to readout in 2H 2025

Phase 2 objectives

- **2H 2025 POC readout:** % change from baseline in EASI at Week 16 powered >90% to detect effect
 - Induction regimen that exceeds lebrikizumab exposures by ~30%
- **Maintenance POC:** Study every 3- or every 6-month dosing in initial POC study to demonstrate the full potential of APG777 to reduce injection burden of patients
- **Phase 2b dose optimization:** examine range of regimens with exposures at, below, and above lebrikizumab

Strong correlation between Phase 2 and 3 results in AD for validated endpoints EASI-75 and IGA 0/1





APG777 could substantially decrease annual maintenance injections for patients

APG777*

2-4

INJECTIONS

ONE INJECTION EVERY
3- or 6- MONTHS



Lebrikizumab

13

INJECTIONS

ONE INJECTION EVERY
4 WEEKS

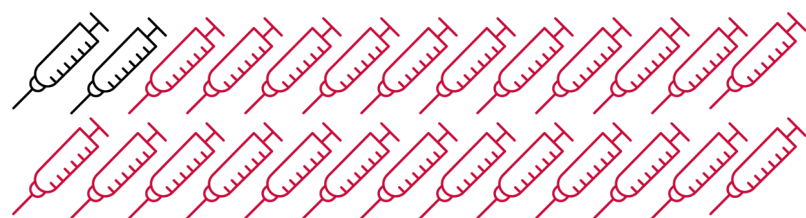



DUPIXENT

26

INJECTIONS

ONE INJECTION EVERY
OTHER WEEK



 Additional injection relative to Q6M APG777



APG808

APG808 targets the same mechanism as DUPIXENT, which has been validated in COPD



COPD represents area of high unmet and a promising opportunity given recent positive DUPIXENT data

10%

of the global population >40 yrs

3rd

Leading cause of death in the US in 2019

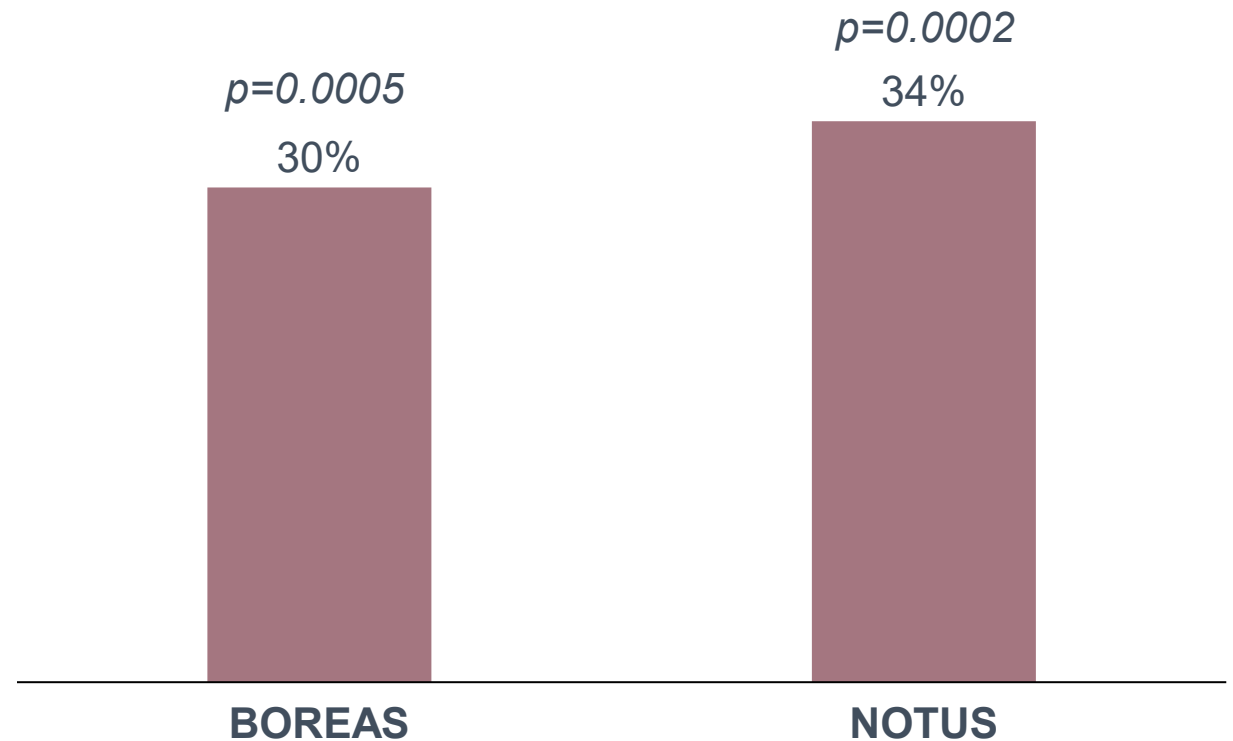
150K+

People die each year in the US

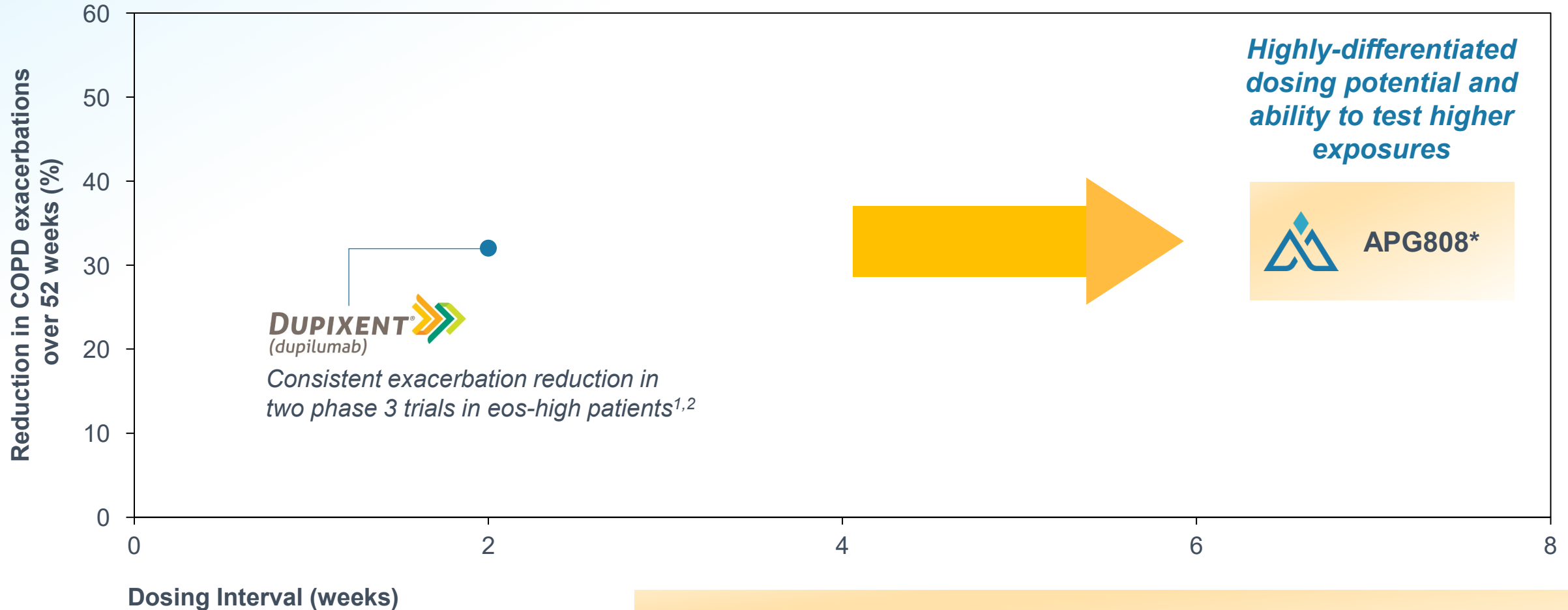
No biologic therapies are approved for COPD, but DUPIXENT demonstrated promise in two Phase 3s:

- Significant, clinically meaningful **reduction in moderate or severe acute COPD exacerbations**
- **Improved lung function from baseline at 12 weeks** compared to placebo with separation from placebo as early as 2 weeks

DUPIXENT produced a significant and clinically meaningful reduction in exacerbations in two Phase 3 studies



Treatments for moderate-severe COPD are limited

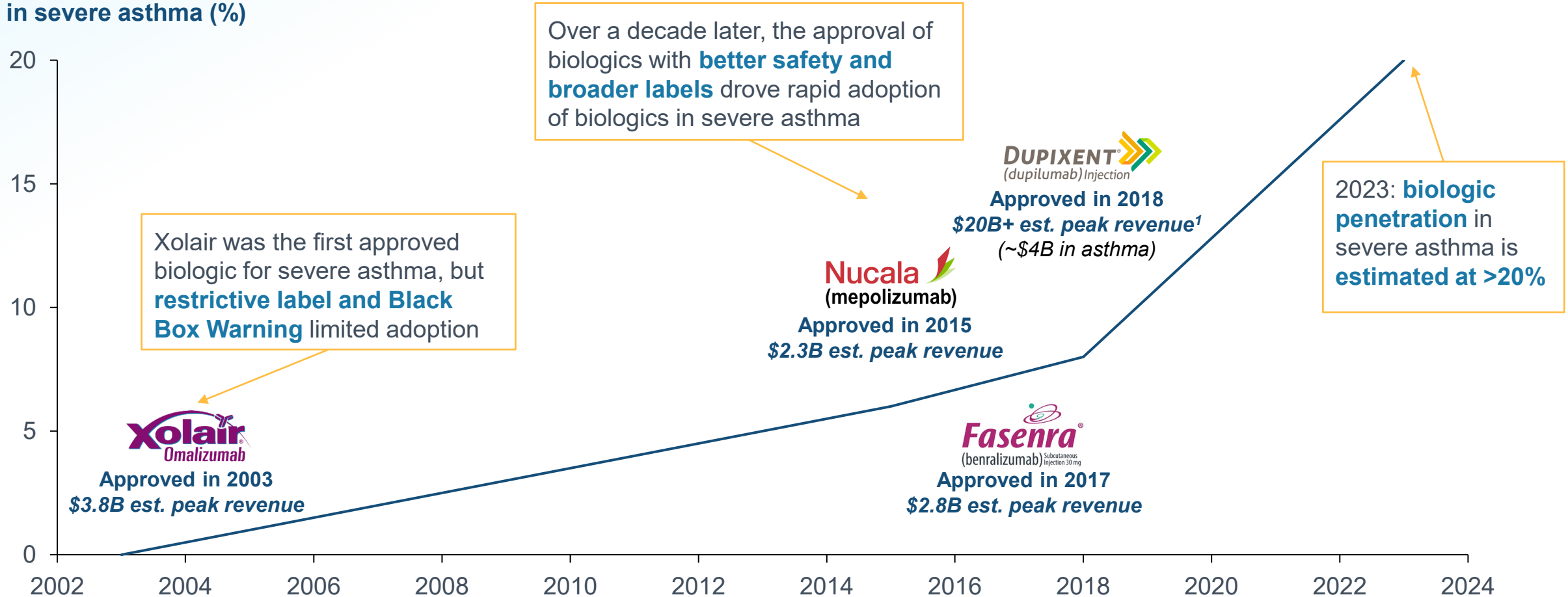


Other than DUPIXENT, no other late-stage biologic for the treatment of COPD has achieved its primary endpoint, leaving a vast unmet need for dosing beyond Q2W

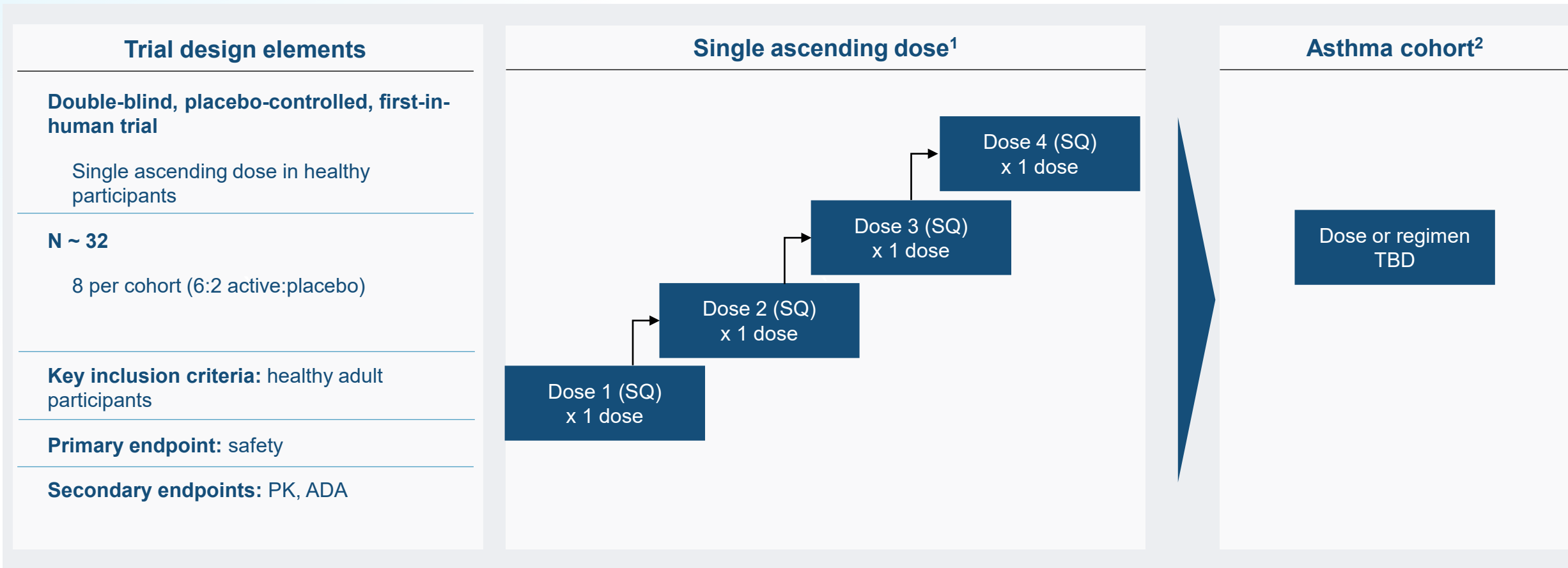
Asthma, an analog for COPD, shows how biologics can be rapidly adopted when they address unmet needs



Biologic penetration in severe asthma (%)



APG808 Phase 1 expected to initiate in 1H 2024 (ahead of schedule) with planned readout in 2H 2024

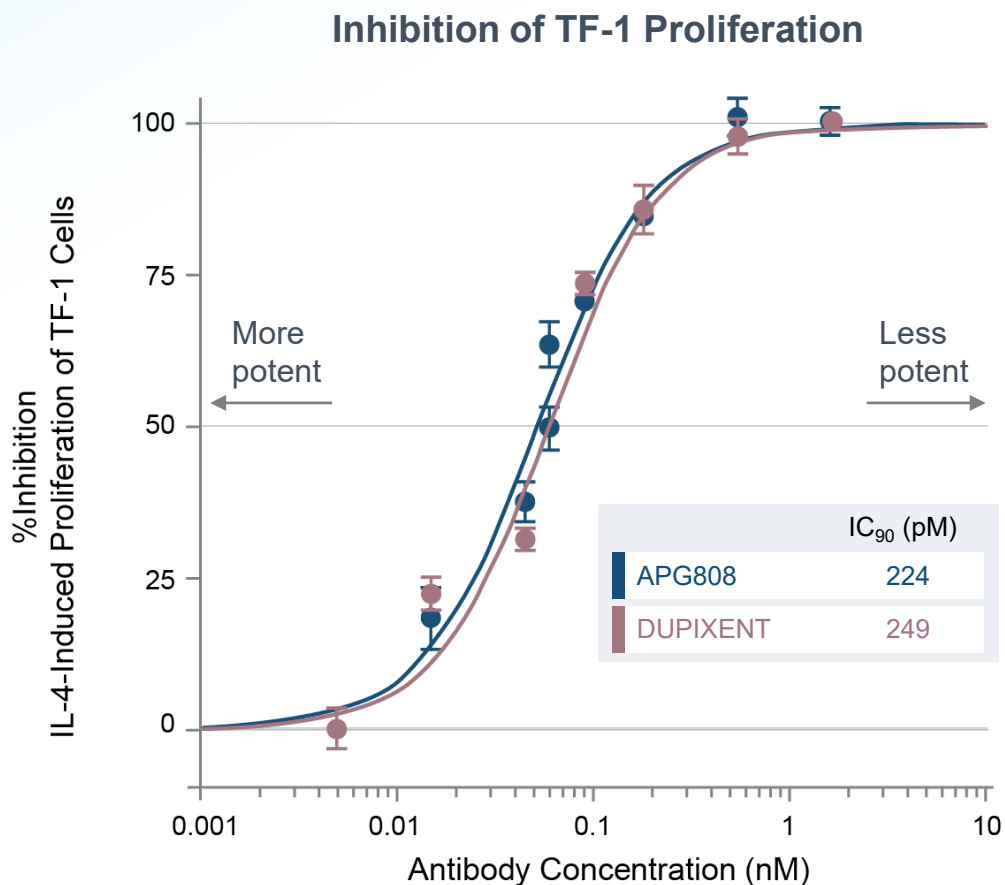


2H 2024: Present APG808 safety and PK, including potentially extended half-life, optimized exposures, and low variability

APG808 is as potent as DUPIXENT in key preclinical assays



APG808 vs DUPIXENT on key potency assay



Additional *in vitro* assays support APG808 potency

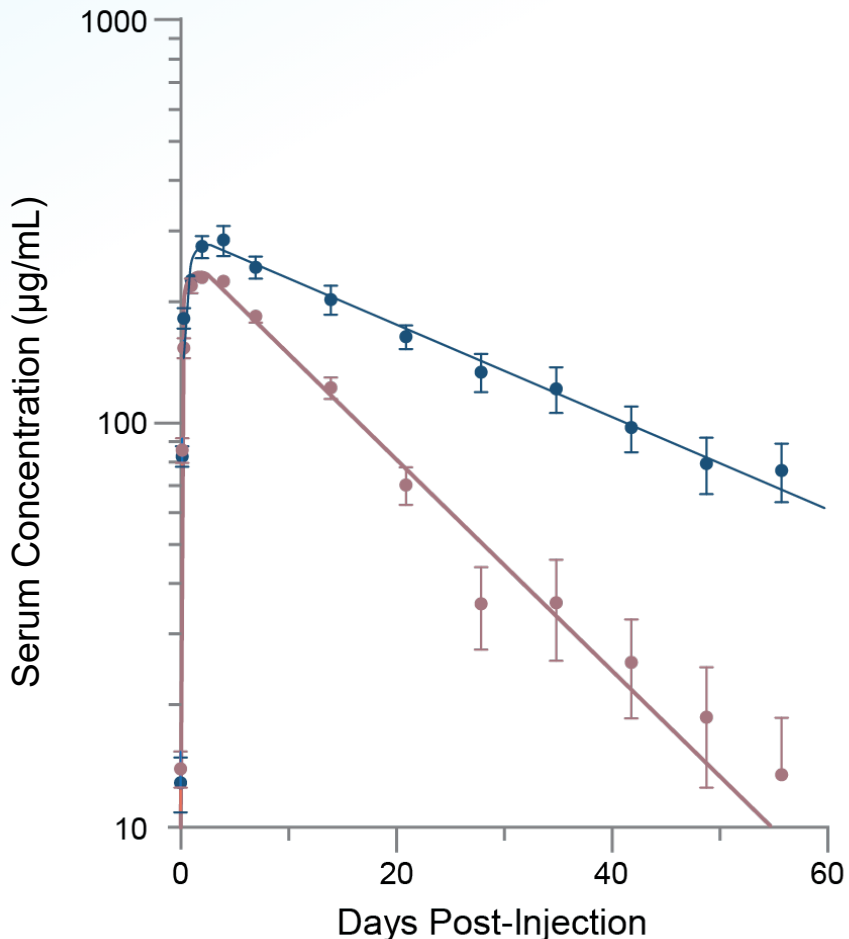
Assay	Affinity to human IL-4R α ^{1,2}	Inhibition of STAT-6 phosphorylation	Inhibition of TARC secretion
Measurement	K _D (pM)	IC ₉₀ (nM)	IC ₉₀ (nM)
APG808	0.4	1.11	1.25
DUPIXENT	12	1.93	1.67

Additional preclinical assays demonstrate APG808 and DUPIXENT have an overlapping binding site on IL-4R α

APG808 NHP half-life is significantly longer than DUPIXENT



NHP PK, SQ administration



APG808 has advantages over DUPIXENT in our NHP head-to-head studies

NHP average half-life¹

APG808: ~26 days

DUPIXENT: ~12 days



APG808 showed extended half-life in NHPs

- APG808 also showed decreased variability on PK and potential for greater consistency in response

APG808 can potentially achieve 6- or 8-week dosing vs Q2W for DUPIXENT



APG808 NHP half-life suggests potential for significant improvement over DUPIXENT in humans

APG808 predicted human half-life vs. observed comparators, days

■ Non-YTE mAb
■/■ YTE mAb

Indication NHP half-life, days

COPD

APG808 (YTE)* 27

DUPIXENT
(dupilumab) 11

Celldex therapeutics CDX-0159 (YTE) (c-KIT/CD117) 22

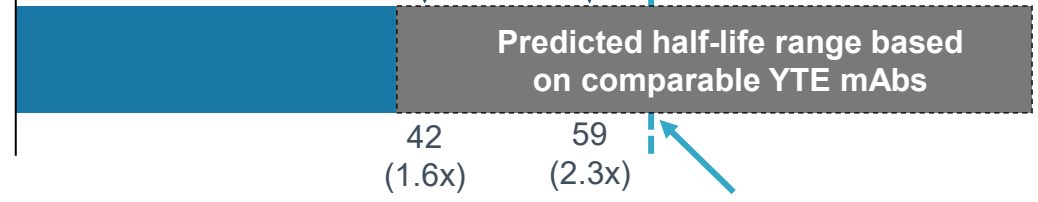
VIRIDIAN VRDN-002 (YTE) (IGF-1R) 14

VRDN-003 (YTE) (IGF-1R) 13

Comparable YTE mAbs for other indications

Every 6 weeks
Every 8 weeks

APG808 dosing interval based on PK modeling¹



Average of YTE mAbs for receptor targets (2.5x)

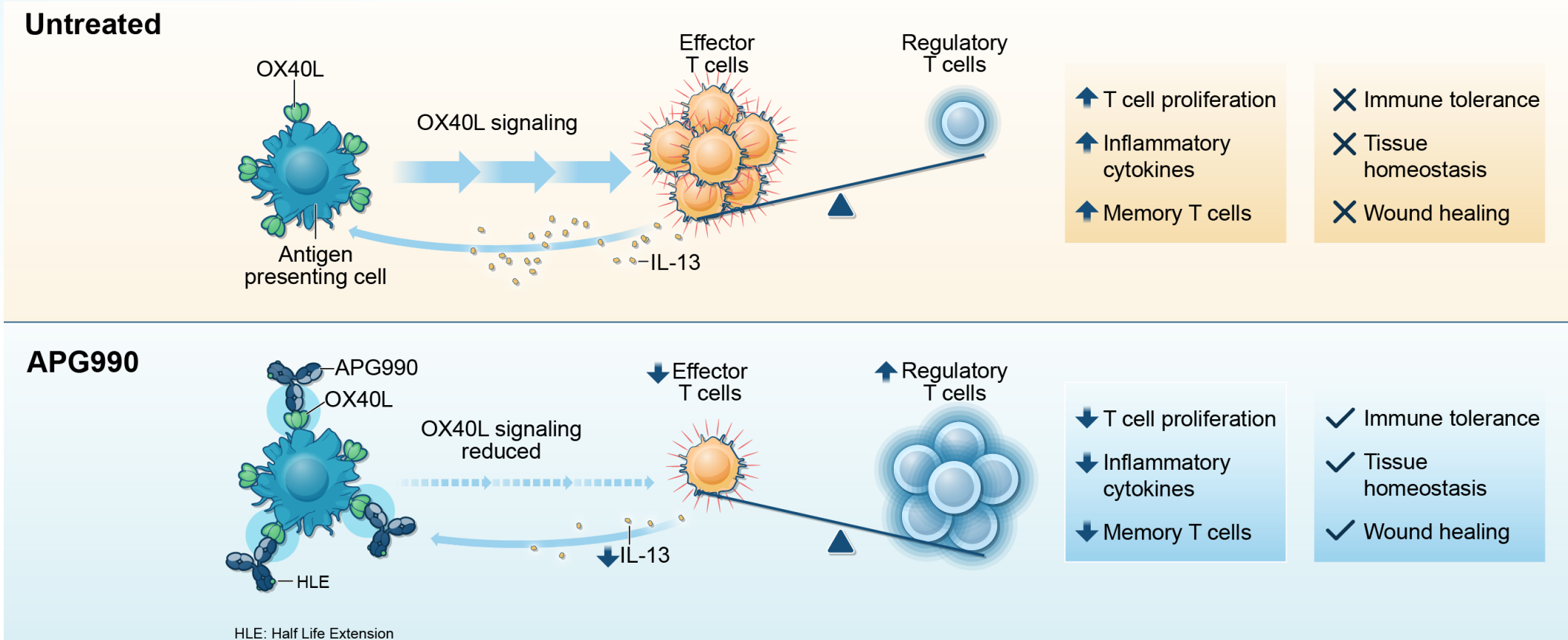
NOTE: Half-lives as reported in studies conducted by the sponsor of each of these product candidates or in the label of approved products. Half-lives are not based on head-to-head studies and are derived from different studies at different points in time, with differences in study design. As a result, cross-study comparisons cannot be made. ¹Based on steady state PK simulations made with parameters for APG808 identical to Dupixent except changes in $k_{elimination}$.
*Positioning of Apogee program is illustrative and not based on clinical trial data and is based only on pre-clinical study results



APG990/APG222



APG990 blocks OX40L and potentially rebalances the immune system



OX40L blockade targets Th2, Th17, and Th22 pathways, which have been implicated in numerous I&I conditions

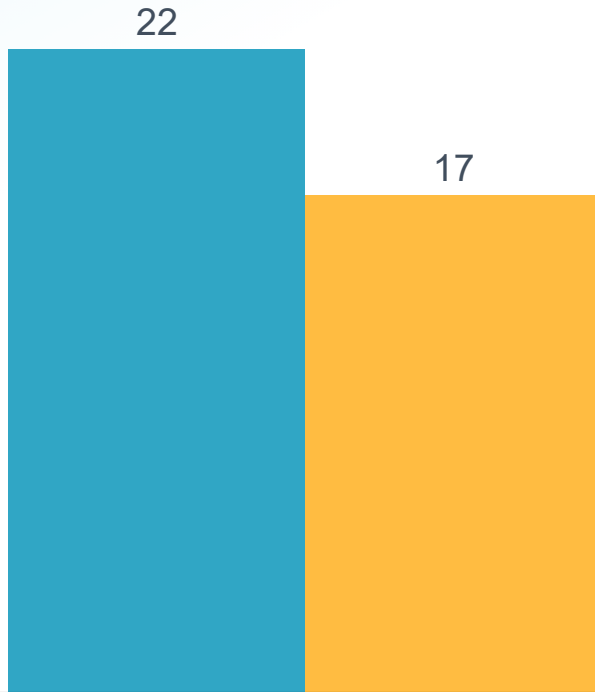
Upcoming clinical trial readouts could provide PoC for OX40L beyond AD including asthma, hidradenitis suppurativa, alopecia areata, celiac disease, and systemic sclerosis



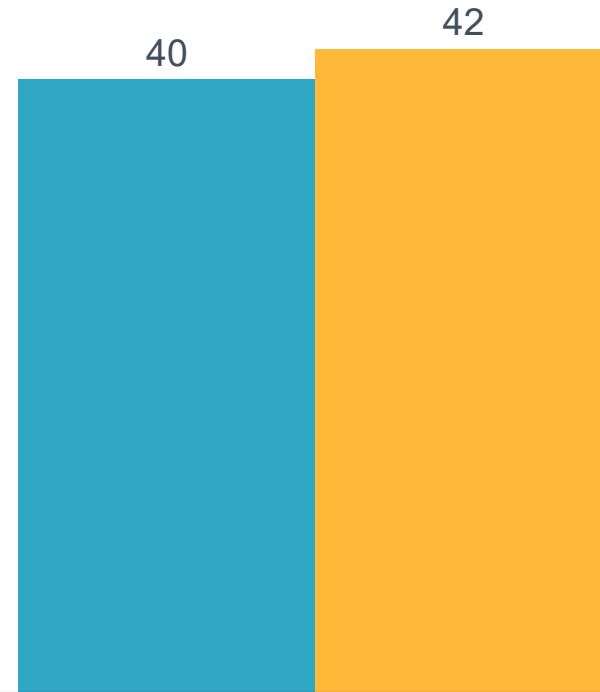
OX40L and OX40 inhibition have shown similar efficacy, but OX40L has a clear advantage on safety

IGA 0/1 response at Week 16

■ Amlitelimab (OX40L)¹
 ■ Rocatinlimab (OX40)²



EASI-75 at Week 16

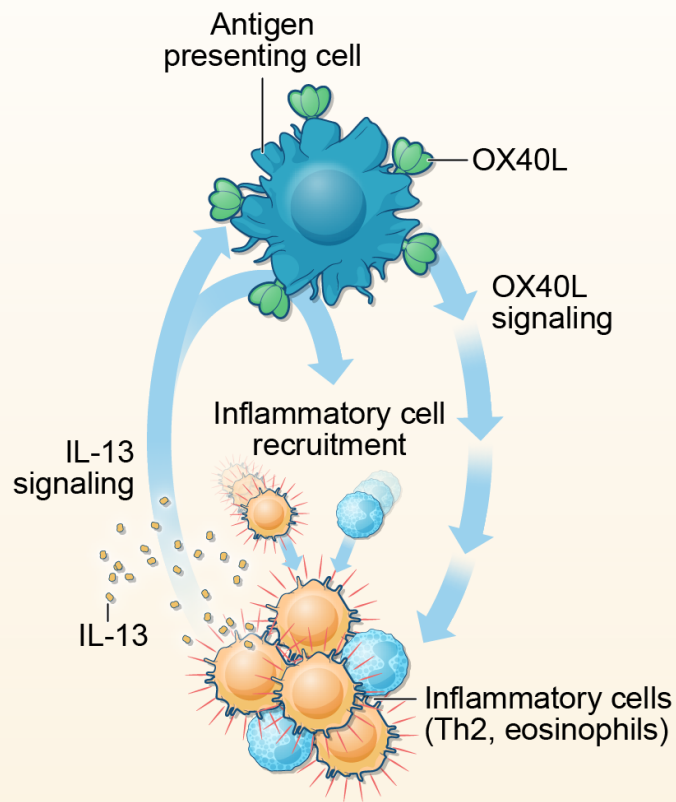


- In Phase 2b, **rocatinlimab (OX40) was associated with pyrexia (17% of patients) and chills (11% of patients)**
- In contrast, **no pyrexia³ or chills for amlitelimab (OX40L) in Phase 2b**

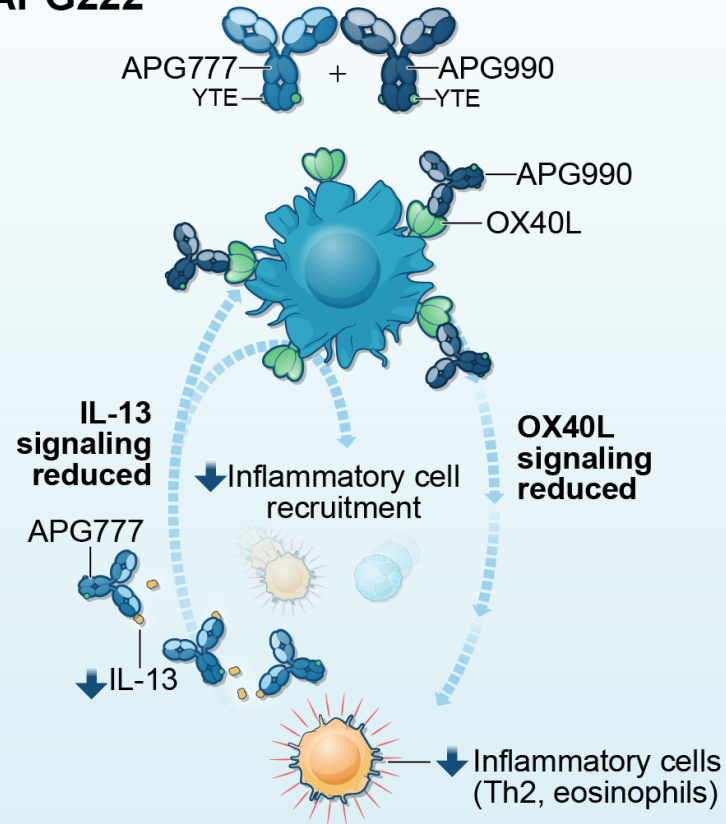
APG222 combines two validated mechanisms and may enhance benefit in AD and other I&I indications



Untreated



APG222



- **OX40L treatment reduces circulating IL-13 levels** supporting the potential for synergy with IL-13 blocker
- Combination potentially enables wider subset of patients to achieve **deeper clinical responses** and **durable remission in AD and other I&I indications**

Given strong mechanistic rationale, APG222 program will explore combination potential



Corporate

Experienced team with proven history of clinical development and commercial execution



Michael Henderson, MD
Chief Executive Officer, Director



Carl Dambkowski, MD
Chief Medical Officer



Jane Pritchett Henderson
Chief Financial Officer



Rebecca Dabora, PhD
Chief Technical Officer



Matt Batters, JD
General Counsel



Wendy Aspden-Curran
SVP of Clinical Operations



Drew Badger, PhD
SVP of Regulatory Affairs & Toxicology



Dan Mulreany
SVP of Business Development & Strategy



Kristine Nograles, MD, MSc
SVP of Clinical Development



Board of Directors with industry-leading development, commercial and management expertise



Mark McKenna
Chairman



Michael Henderson, MD
CEO, Apogee Therapeutics



Jennifer Fox
CFO & CBO, Zenas BioPharma



Andrew Gottesdiener, MD
Venrock



Peter Harwin
Managing Member, Fairmount



BJ Jones
CCO, NewAmsterdam Pharma



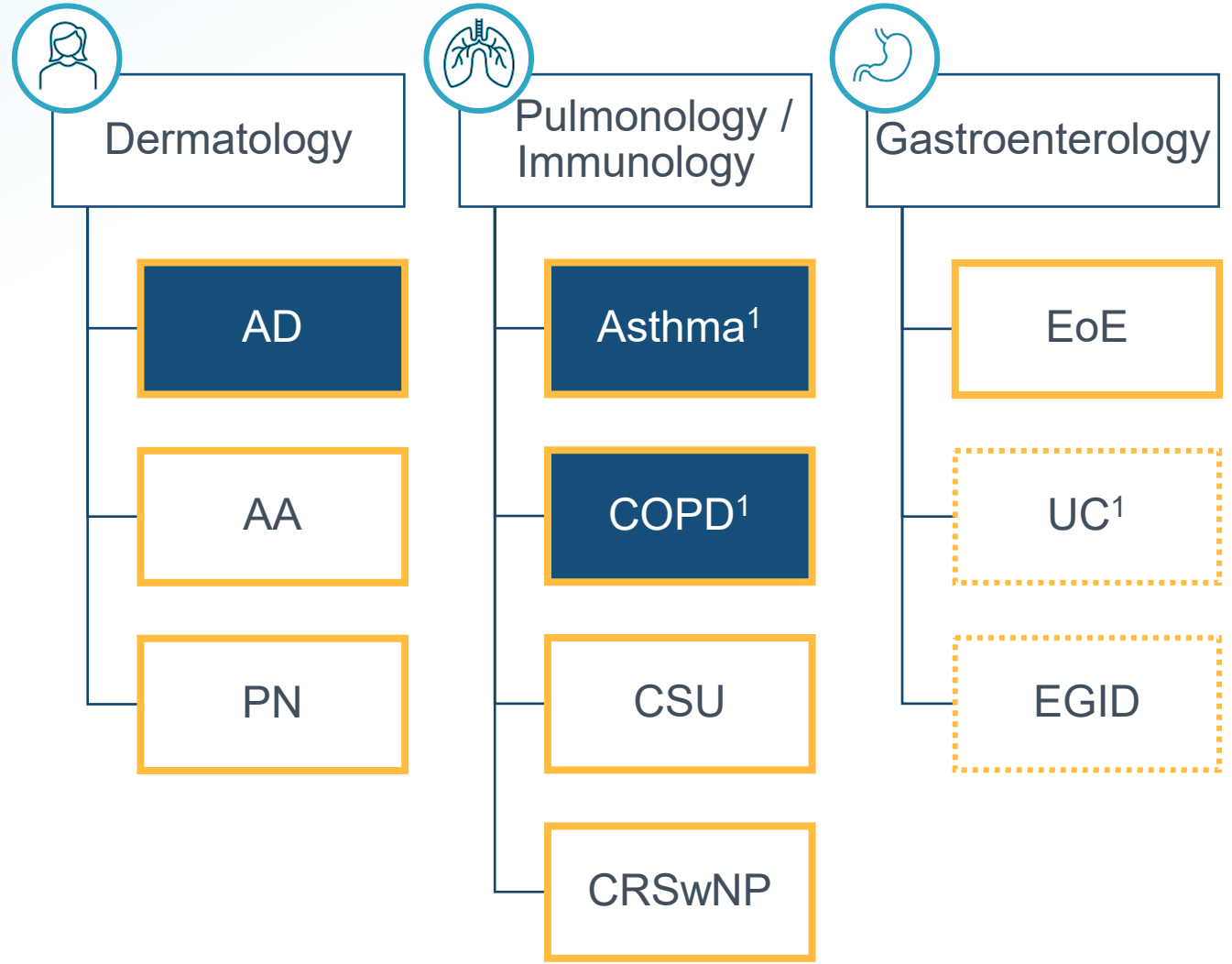
Tomas Kiselak
Managing Member, Fairmount



Nimish Shah
Venrock



Our programs have broad potential to disrupt the I&I space



- Planned Apogee Ph2 based on well-established mechanism
- 3rd party clinical data supporting one or more Apogee targets
- ⋯ Ongoing 3rd party clinical trials which could validate one or more Apogee targets



Apogee /'apəjē/ *noun*

The highest point in the development of something; a climax or culmination