



# Transforming Patient Outcomes with Superior Vision Gains

**Corporate Overview | September 2024**

NASDAQ (OPT); ASX (OPT.AX)

# Disclaimer and Forward-looking Statements

This presentation includes general background information about the activities of Opthea Limited (ABN 32 006 340 567) (“Opthea” or “Company”) and its affiliates and subsidiaries (together, the “Opthea Group”). The information contained in this presentation is in summary form and does not purport to be complete or to contain all material information about the Opthea Group which a prospective investor or purchaser may require in evaluating a possible investment in Opthea or acquisition of securities in Opthea. The information in this presentation remains subject to change without notice. No member of the Opthea Group nor any director, officer, employee, adviser, agent or representative of any member of the Opthea Group (each an Opthea Party and together, the Opthea Parties) has any obligation to update or correct this presentation.

This presentation contains forward-looking statements within the meaning of the U.S. federal securities laws that involve substantial risks and uncertainties. All statements, other than statements of historical facts, contained in this presentation, including statements regarding the therapeutic and commercial potential and size of the estimated market opportunity of the Company’s product in development, the viability of future opportunities, future market supply and demand, the expected timing of top-line data, our expectations about topline data based on masked pooled data, the future cash runway, the financial condition, results of operations and business of Opthea, certain plans, objectives and strategies of management of Opthea, including with respect to the current and planned clinical trials of its product candidate and the future performance of Opthea, are forward-looking statements. The words “anticipate,” “believe,” “estimate,” “expect,” “intend,” “may,” “might,” “plan,” “predict,” “project,” “target,” “potential,” “will,” “would,” “could,” “should,” “continue,” and similar expressions are intended to identify forward-looking statements, although not all forward-looking statements contain these identifying words. Opthea may not actually achieve the plans, intentions or expectations disclosed in the forward-looking statements, and you should not place undue reliance on the forward-looking statements as predictions of future events. Actual results or events could differ materially from the plans, intentions and expectations disclosed in the forward-looking statements we make. The forward-looking statements contained in this presentation reflect Opthea’s current views with respect to future events, and we assume no obligation to update any forward-looking statements except as required by applicable law. Please refer to information, including risk factors, set forth in Opthea’s Annual Report on Form 20-F filed with the U.S. Securities and Exchange Commission on August 30, 2024, and other future filings with the U.S. Securities and Exchange Commission for key factors that could cause actual results to differ materially from those projected in the forward-looking statements contained herein including risks associated with: future capital requirements and ability to continue as a going concern, the development, testing, production, marketing and sale of drug treatments, regulatory risk and potential loss of regulatory approvals, ongoing clinical studies to demonstrate sozinibercept safety, tolerability and therapeutic efficacy, additional analysis of data from Opthea’s Phase 3 clinical trials, including once masked pooled data is unmasked, clinical research organization and labor costs, intellectual property protections, future capital requirements, the development, testing, production, marketing and sale of drug treatments, regulatory risk and potential loss of regulatory approvals, ongoing clinical studies to demonstrate sozinibercept safety, tolerability and therapeutic efficacy, and other factors that are of a general nature which may affect the future operating and financial performance of the Company.

This presentation includes statistical and other industry and market data that we obtained from industry publications and research, surveys and studies conducted by third parties as well as our own estimates of potential market opportunities. All of the market data used in this presentation involves a number of assumptions and limitations, and you are cautioned not to give undue weight to such data. Industry publications and third-party research, surveys and studies generally indicate that their information has been obtained from sources believed to be reliable, although they do not guarantee the accuracy or completeness of such information. Our estimates of the potential market opportunities for our product candidates include several key assumptions based on our industry knowledge, industry publications, third-party research and other surveys, which may be based on a small sample size and may fail to accurately reflect market opportunities. While we believe that our internal assumptions are reasonable, no independent source has verified such assumptions.

The information contained in this presentation does not constitute investment or financial product advice (nor taxation or legal advice) and is not intended to be used as the basis for making an investment decision. The presentation is for informational purposes only and is not a prospectus or other disclosure document under Australian law or the law of any other jurisdiction and does not contain all the information which would be required to be disclosed in a prospectus or other disclosure document. The information presented in this presentation may differ materially from that presented in any disclosure document prepared in connection with any offer of securities. It does not take into account the investment objectives, financial situation, taxation position or needs of any particular investor, which should be considered when deciding if an investment is appropriate. You must consider your own investment objectives, financial situation and needs and conduct your own independent investigations and enquiries, including obtaining taxation, legal, financial or other professional advice in relation to the information contained in this presentation as appropriate to your jurisdiction. This presentation should not be relied upon by the Recipient in considering the merits of any particular transaction.

This presentation does not constitute an offer to sell, or the solicitation of an offer to buy, any securities in the United States or any other jurisdictions in which such an offer would be unlawful prior to registration or qualification under the U.S. Securities Act of 1933, as amended, or the securities laws of any state or other jurisdiction of the United States.

This presentation may contain trademarks and trade names of third parties, which are the property of their respective owners. Third party trademarks and trade names used in this presentation belong to the relevant owners and use is not intended to represent sponsorship, approval or association by or with any of the Opthea Group.

# Sozinibercept Has the Potential to Be the First Product in 20 Years to Improve Visual Outcomes

## Addressing High Unmet Need

- Despite wide use of anti-VEGF-A therapy, wet AMD patients still experience loss in vision long term<sup>1</sup>
- Every letter of vision counts to improve quality of life and reduce mortality

## Proprietary Technology

- First-in-class VEGF-C/D ‘trap’ inhibitor intended for combination with standard of care anti-VEGF-A therapies
- Composition of Matter and Methods of Use Patents through 2034; opportunities to extend beyond 2034\*

## Superior Lead Asset

- Phase 2b demonstrated superiority in combination with SOC therapy, with well tolerated safety profile
- Sozinibercept has the potential to improve vision for millions of patients with wet AMD

## Topline Data from Pivotal Trials in 2025

- Topline data anticipated for COAST (n=998) in early 2Q CY2025 and ShORe (n=986) in mid-CY2025
- Current cash expected to fund operations into 3Q CY2025<sup>2</sup>

## Substantial Market Opportunity

- Multibillion dollar commercial opportunity in a growing market with an established clinical practice
- Sozinibercept developed for use in combination with any anti-VEGF-A; will not compete directly with SOC therapies

AMD – age-related macular degeneration; MOA – Mechanism of Action; SOC – Standard of care

<sup>1</sup>CATT Research Group; Maguire MG et al. Ophthalmology. 2016 Aug.

<sup>2</sup>Additional funding will be required to reach commercialization of sozinibercept and to meet obligations under the Development Funding Agreement ("DFA"). As a result of obligations under the DFA and applicable law regarding liquidity, the Company expects to raise or obtain additional capital in one or more transactions, earlier than 3Q CY 2025 or anticipated topline data readout dates.

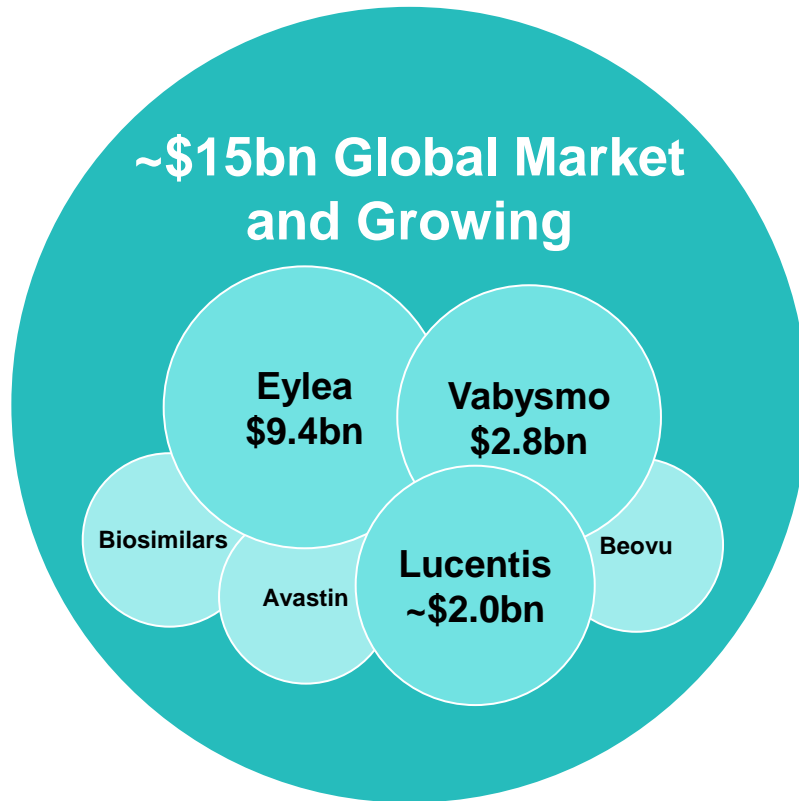
\*Potential for Patent Term Extensions & Data and Market Exclusivity (12 Years for Biologic)

# Sozinibercept Designed to Improve Visual Outcomes in Combo with VEGF-A Inhibitors; Potential to Create New Multi-Billion Dollar Class

Global Marketed VEGF-A Inhibitors



Sozinibercept is a VEGF-C/D “Trap” Inhibitor



Potential value proposition:

**Targeting Improved Visual Function**

*Critical for Patients, Physicians and Payors*

**Fits Seamlessly into Physician Practice**

**Potential Use with Any VEGF-A Inhibitor**

**Multi-Billion Dollar Commercial Opportunity**

# Experienced Leadership Team

Expertise and Track Record to Make a Positive Impact on the Retinal Community

## Management Team



**Fred Guerard, PharmD, MS**  
Chief Executive Officer



**Daniel Geffken**  
Interim Chief Financial Officer



**Megan Baldwin, PhD, MAICD**  
Founder, Chief Innovation  
Officer & Executive Director



**Mike Campbell**  
Chief Commercial Officer



## Chief Medical Advisor



**Arshad M. Khanani, MD, MA, FASRS**  
Managing Partner, Director of Clinical Research  
and Director of Fellowship at Sierra Eye  
Associates, and Clinical Professor at the University  
of Nevada, Reno School of Medicine

## Clinical Advisory Board



**Charles C. Wykoff, MD, PhD**  
Director of Research, Retina Consultants of Texas,  
Chairman of Research and Clinical Trials  
Committee, Retina Consultants of America



**Tim Jackson, PhD, MB, ChB, FRCophth**  
National Health Service, Consultant at Kings  
Hospital College Hospital, London



**Jason Slakter, MD**  
Clinical Professor at New York University School  
of Medicine and partner at Vitreous Retina Macula  
Consultants of New York

# Despite Treatment with Standard of Care Anti-VEGF-A Therapies, the Majority of Patients Achieve Suboptimal Vision Outcomes

## Despite treatment with anti-VEGF-A therapy\*

**>45%** do not achieve significant vision gains

**>60%** will have **persisting macular fluid**

**25%** will have **further vision loss at 12+ months**



The majority of patients fail to achieve 20/40 vision<sup>1</sup>



Suboptimal vision is associated with decrease in Instrumental Activities of Daily Living (IADL) skills<sup>2</sup>

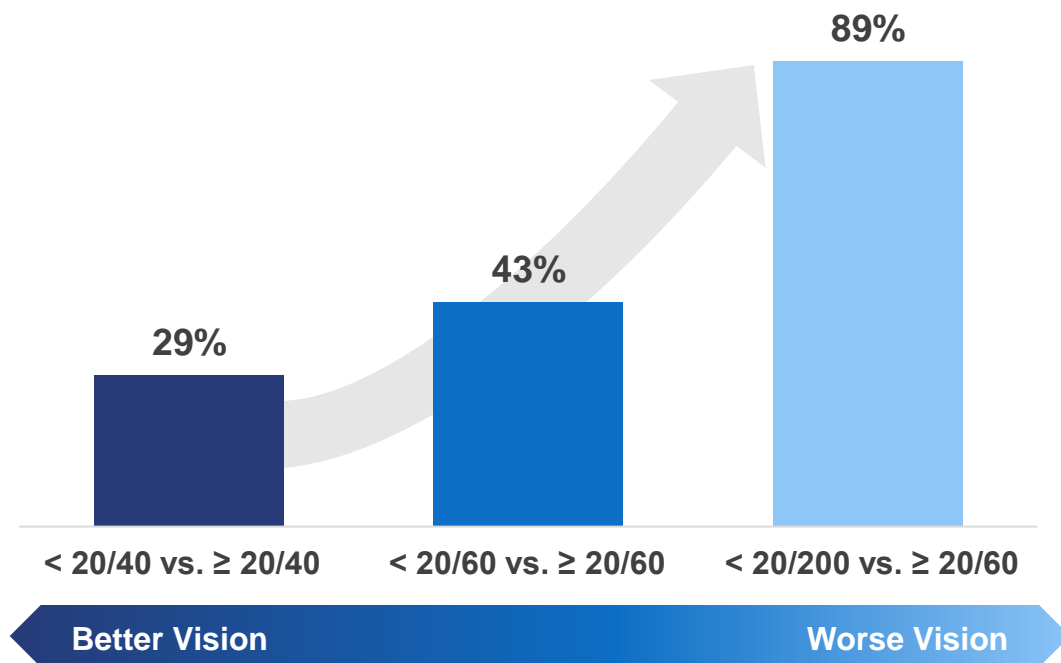
\*Based on randomised, controlled clinical trial data; >45% fail to achieve  $\geq 2$  lines improvement in Best Corrected Visual Acuity (BCVA); Persisting fluid: SD-OCT CST  $\geq 300$   $\mu\text{M}$  or Time-Domain OCT CST  $\geq 250$   $\mu\text{M}$   
IADL: Instrumental activities of daily living (complex activities related to the ability to live independently)

<sup>1</sup>Mettu PS, et al. Prog Retin Eye Res. 2021

<sup>2</sup>Hochberg C, et al. Invest Ophthalmol Vis Sci. 2012 May 31.

# Every Letter Counts When Loss of Vision Potentially Leads to Increased Mortality Risk

## Hazard for All-cause Mortality<sup>1</sup> Higher in People with Vision Impairment



**Decrease of 1 ETDRS letter per year  
increases mortality risk by 16%<sup>2</sup>  
associated exclusively with IADL levels**

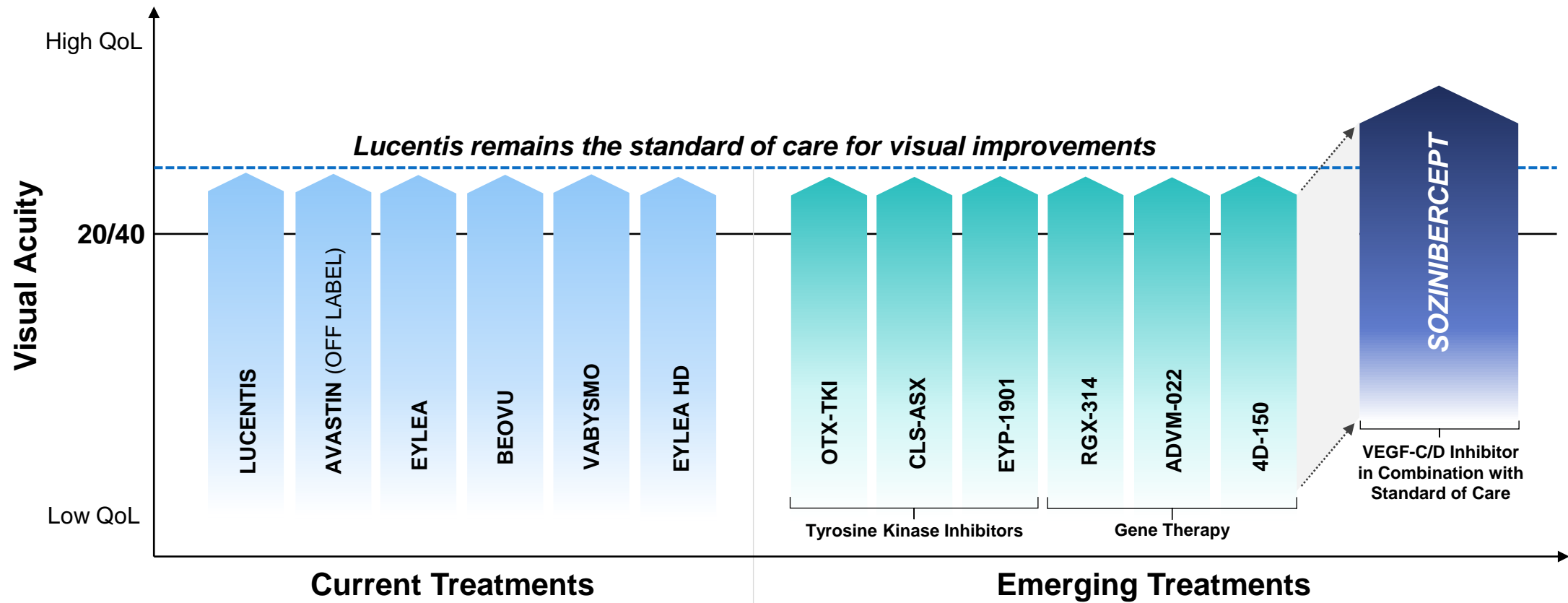
IADL – Instrumental activities of daily living; ETDRS – Early Treatment Diabetic Retinopathy Study chart

<sup>1</sup>Ehrlich JR et al. "Association between vision impairment and mortality: a systematic review and meta-analysis." Lancet Glob Health. 2021.

<sup>2</sup>Christ SL, et al. "Longitudinal relationships among visual acuity, daily functional status, and mortality: the Salisbury Eye Evaluation Study." JAMA Ophthalmol. 2014.

# Sozinibercept Has Demonstrated Improvement in Vision Gains and Reduction in Vision Loss

Opportunity in Wet AMD Market for an *Overall Shift* Towards Superior Visual Outcomes



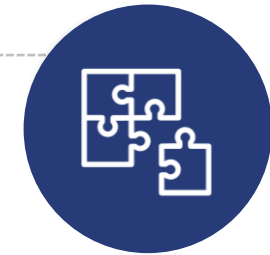
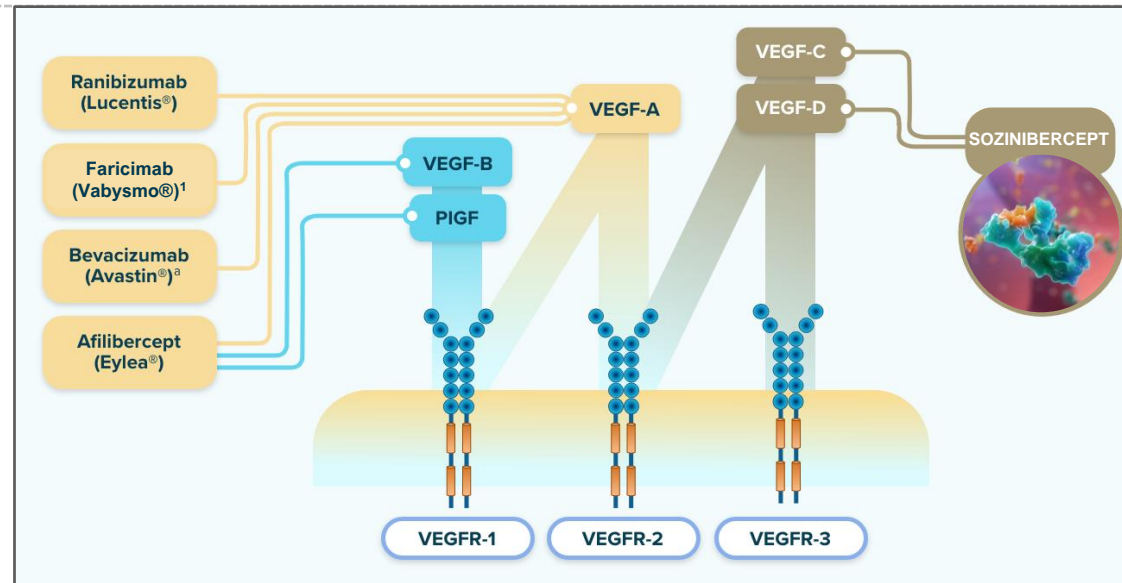


# Sozinibercept, a Proprietary VEGF-C/D “Trap” Inhibitor, Has the Potential to Address the Limitations of Anti-VEGF-A Therapies



## The Problem

Wet AMD is a **multi-factorial disease**. Treatment with VEGF-A inhibitors **upregulates VEGF-C/D**, driving angiogenesis and vascular permeability.



## The Solution

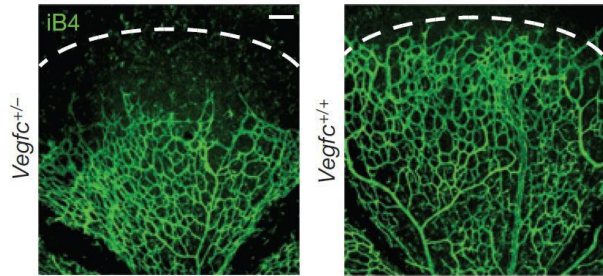
When used in combination with any VEGF-A inhibitor, **sozinibercept completely blocks VEGFR-2 and VEGFR-3 signaling**.

<sup>1</sup> Faricimab also has inhibitory effect on Ang-2.

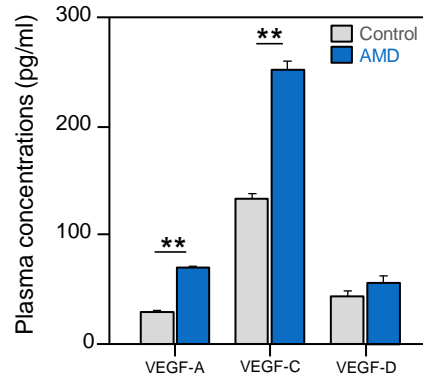
<sup>a</sup> Bevacizumab is used 'off-label' for the treatment of neovascular (wet) AMD

# Published Evidence Supports Broader VEGF Pathway Inhibition with Sozinibercept

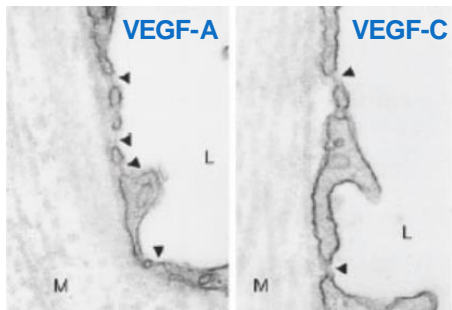
VEGF-C Stimulates Retinal Angiogenesis<sup>^</sup>



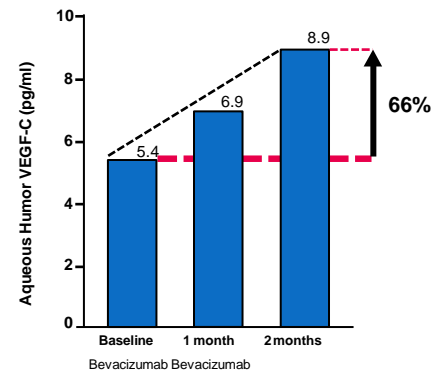
Circulating VEGF-C Levels Significantly Elevated in AMD Patients<sup>†</sup>



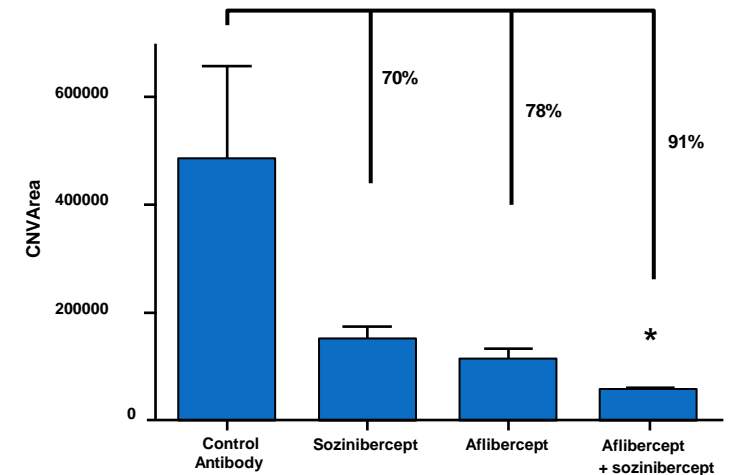
VEGF-A and VEGF-C Induce Vascular Leakage/permeability<sup>#</sup>



Elevated VEGF-C in Aqueous Humor Following Anti-VEGF-Atherapy in Wet AMD Patients<sup>\*</sup>



Additive Benefit of VEGF-A and VEGF-C/D Inhibition in Mouse Wet AMD Model



# Sozinibercept Is Designed to Integrate into Current Anti-VEGF-A Clinical Practice



## Patients

- Superior visual outcomes meaningfully improves patients' lives
- Intended to be administered at same anti-VEGF-A visit



## Retina Specialists

- Better vision outcomes is a high unmet medical need
- Designed to be agnostic to anti-VEGF-A treatment type, including biosimilars

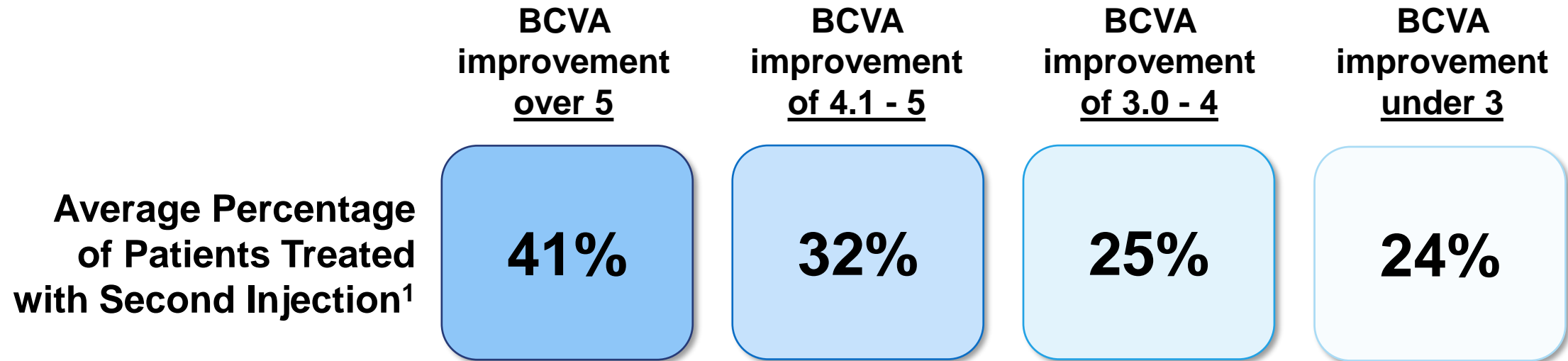


## Payors / Insurance Companies

- Better clinical outcomes represent better health economics
- Visual benefits a key driver in reimbursement

**Concentrated prescriptions in U.S. enables potential self-commercialization opportunity with lean and targeted organization**

# Physicians Willing to Administer Second Injection to up to 41% of Their Patients for Additional BCVA Improvement



*What percentage of your Wet AMD patients would you use a second injection (anti-VEGF C/D) immediately after an anti-VEGF-A injection at various levels of BCVA improvement of the combination over SoC? (Among Total Respondents, Avg. % of Patients\*, n=125)*

**Estimate 1% Share of Wet AMD TAM Equals ~\$100M+ in Sales Per Annum**

TAM – Total Addressable Market

<sup>1</sup>Source: InCrowd Awareness Trial and Usage (ATU) Report, June 2024

\*Averages calculated using the midpoints of each % prescribing allocation group. Callouts indicate statistical significance between groups at 90% CI.

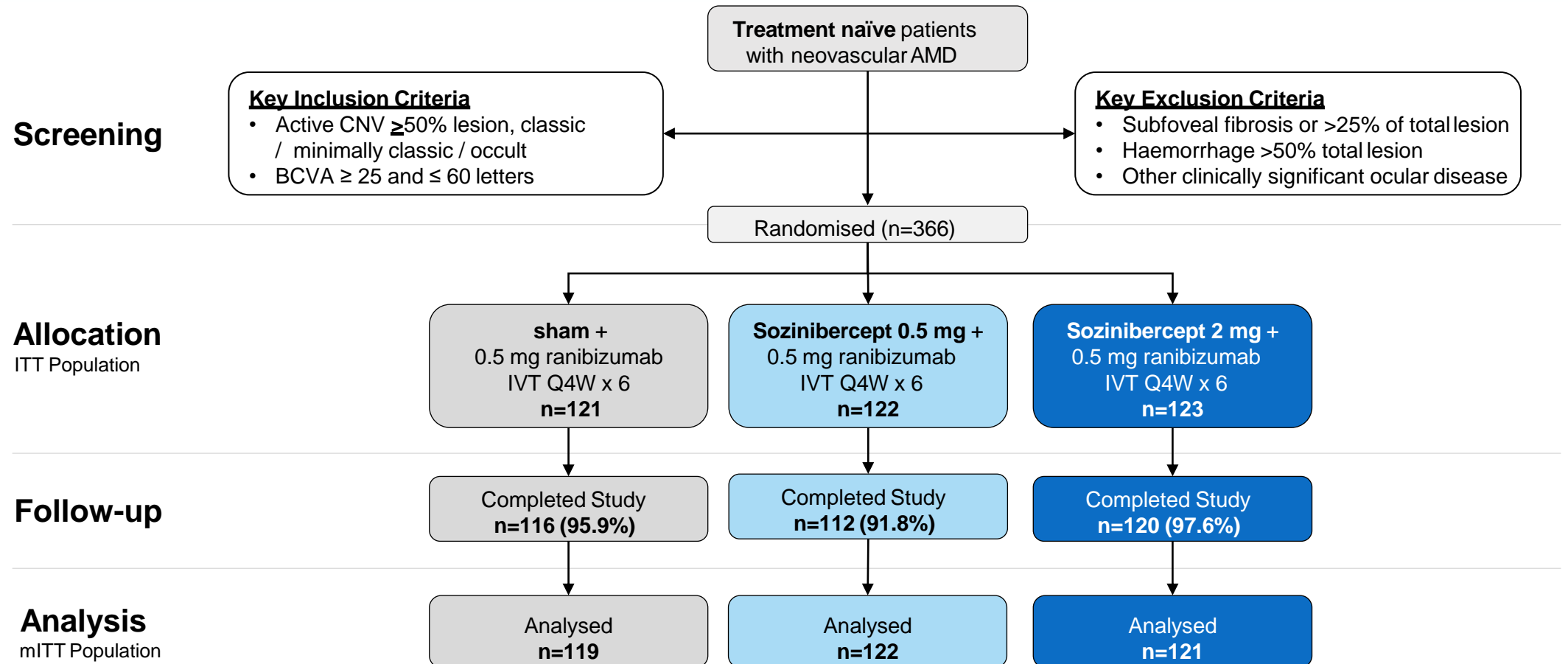
# Long-term Value Opportunities for Sozinibercept

Main Patent Family Extends through 2034, with Expansion Opportunities Beyond 2034\*

PROGRAM	DEVELOPMENT PHASE				ANTICIPATED MILESTONES
	RESEARCH / PRECLINICAL	PHASE 1	PHASE 2	PHASE 3	
<b>Wet Age-Related Macular Degeneration (Wet AMD)</b>					
<b>Sozinibercept</b> For use in combination with anti-VEGF-A therapies					<b>Topline data:</b> COAST (in early 2Q CY2025) ShORe (in mid-CY2025)
<b>Diabetic Macular Edema (DME)</b>					
<b>Sozinibercept</b> For use in combination with anti-VEGF-A therapies					<b>Phase 3 ready</b>
<b>Co-formulation (Sozinibercept + VEGF-A Inhibitor)</b>					
<b>Sozinibercept</b> Co-formulation with VEGF-A Inhibitor					<b>Feasibility underway</b>

\*Potential for Patent Term Extensions & Data and Market Exclusivity (12 Years for Biologic)

# Robust Phase 2b Trial in Wet AMD Demonstrated Superiority in Visual Outcome



CNV – choroidal neovascularisation; IVT – intravitreal; Q4W – once every 4 weeks; ITT – Intent to Treat Population, all participants who were randomised into the study irrespective of whether study medication was administered or not; Safety Population - all participants in the ITT but excluding those who did not receive at least one dose of study medication; mITT – Modified ITT Population, all participants in the Safety Population but excludes any participant without a Baseline VA score and/or any participant who did not return for at least one post-baseline visit

# Phase 2b Primary and Secondary Endpoints

## Primary Endpoint

**Mean change from baseline in BCVA at week 24**

## Key Secondary Endpoints

Proportion of patients gaining  $\geq 15$  letters from baseline at week 24

Change in central subfield thickness (CST) from baseline at week 24

Change in intra-retinal and sub-retinal fluid from baseline to week 24

Safety and tolerability

## Select Pre-specified Subgroups

**Predominantly classic, minimally classic, & occult lesions**  
(Stratification Factor)

**Retinal Angiomatous Proliferation (RAP)**  
detected/not detected at baseline

**Polypoidal Choroidal Vasculopathy (PCV)**  
detected/not detected at baseline

# Phase 2b Trial Demographics and Baseline Characteristics

Demographic/Baseline Disease Characteristic	Sham + ranibizumab n=121	0.5 mg sozinibercept + ranibizumab n=122	2 mg sozinibercept + ranibizumab n=123
Mean Age – years ± SD	76.1 ± 9.48	78.8 ± 8.16	77.8 ± 8.82
Sex – n (%)	Male	48 (39.7%)	45 (36.6%)
	Female	73 (60.3%)	78 (63.4%)
Caucasian Race – n (%)	117 (99.2%)	119 (99.2%)	117 (97.5%)
Mean Visual Acuity (BCVA) – letters ± SD	50.7 ± 10.21	51.1 ± 8.96	49.5 ± 10.26
Mean Total Lesion Area - mm <sup>2</sup> ± SD	6.08 ± 3.21	6.48 ± 3.30	6.62 ± 3.39
Lesion Type	Predominantly classic – n (%)	15 (12.4%)	15 (12.3%)
	Minimally classic – n (%)	53 (43.8%)	51 (41.8%)
	Occult - n (%)	53 (43.8%)	56 (45.9%)
	PCV detected <sup>1</sup> – n (%)	20 (16.5%)	24 (19.7%)
	RAP detected <sup>2</sup> – n (%)	15 (12.7%)	22 (18.5%)
Mean central subfield thickness (CST) - mm ±SD	412.10 ± 110.62	425.18 ± 120.45	414.12 ± 123.25
Sub-retinal fluid (SRF) present – % participants	89.3%	84.4%	87.8%
Intra-retinal cysts present – % participants	57.9%	63.9%	56.1%

Intent-to-Treat (ITT) population; SD – standard deviation; BCVA – Best Corrected Visual Acuity

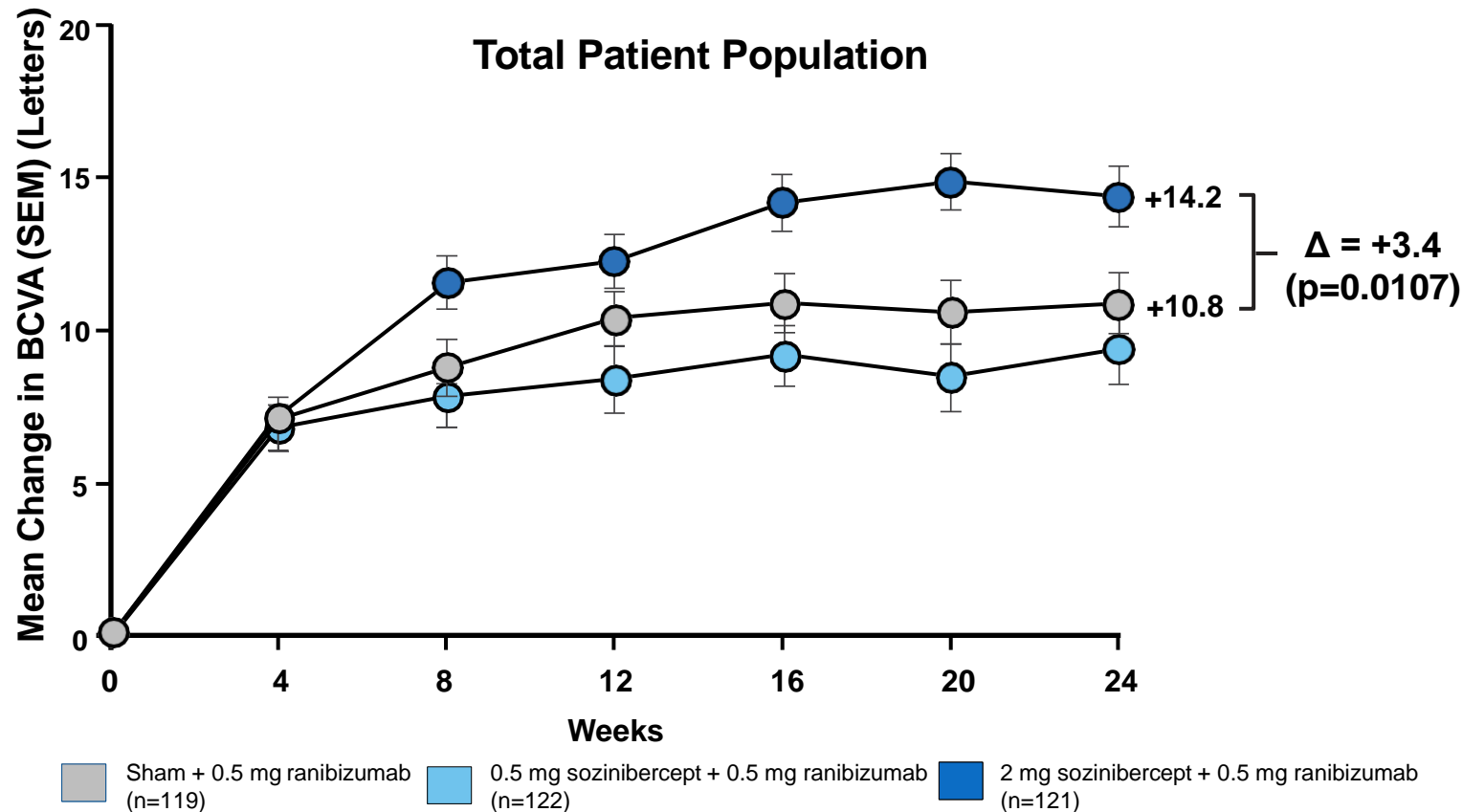
<sup>1</sup>PCV - polypoidal choroidal vasculopathy, detected by SD-OCT, FA and fundus photography.

<sup>2</sup>RAP - retinal angiomatous proliferation, detected by SD-OCT, FA and fundus photography.

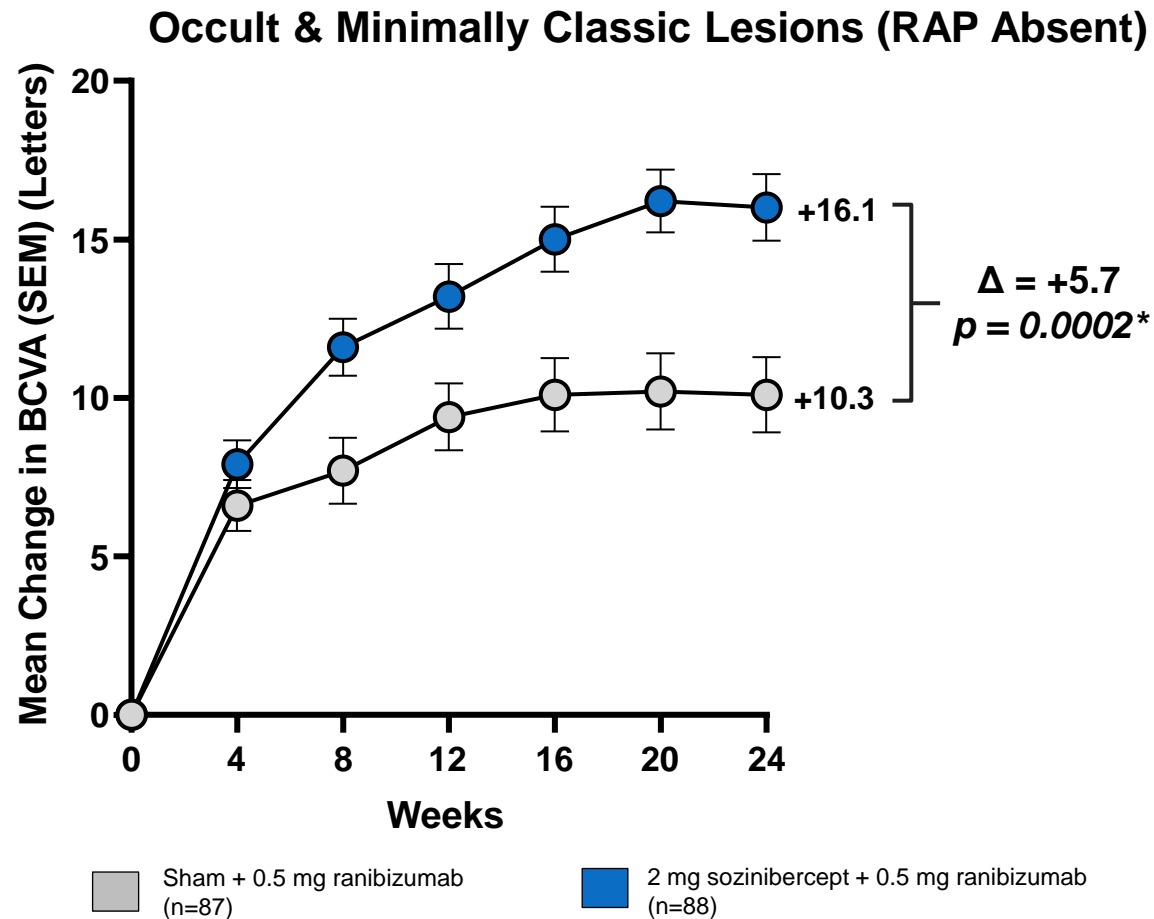


# Sozinibercept 2 mg Combination Therapy Demonstrated Superiority in Visual Acuity over Ranibizumab Monotherapy

## Phase 2b Primary Endpoint Achieved



# To Maximize Probability of Success, Best Responding Phase 2b Patients Represents Primary Analysis Population in the Pivotal Phase 3 Trials



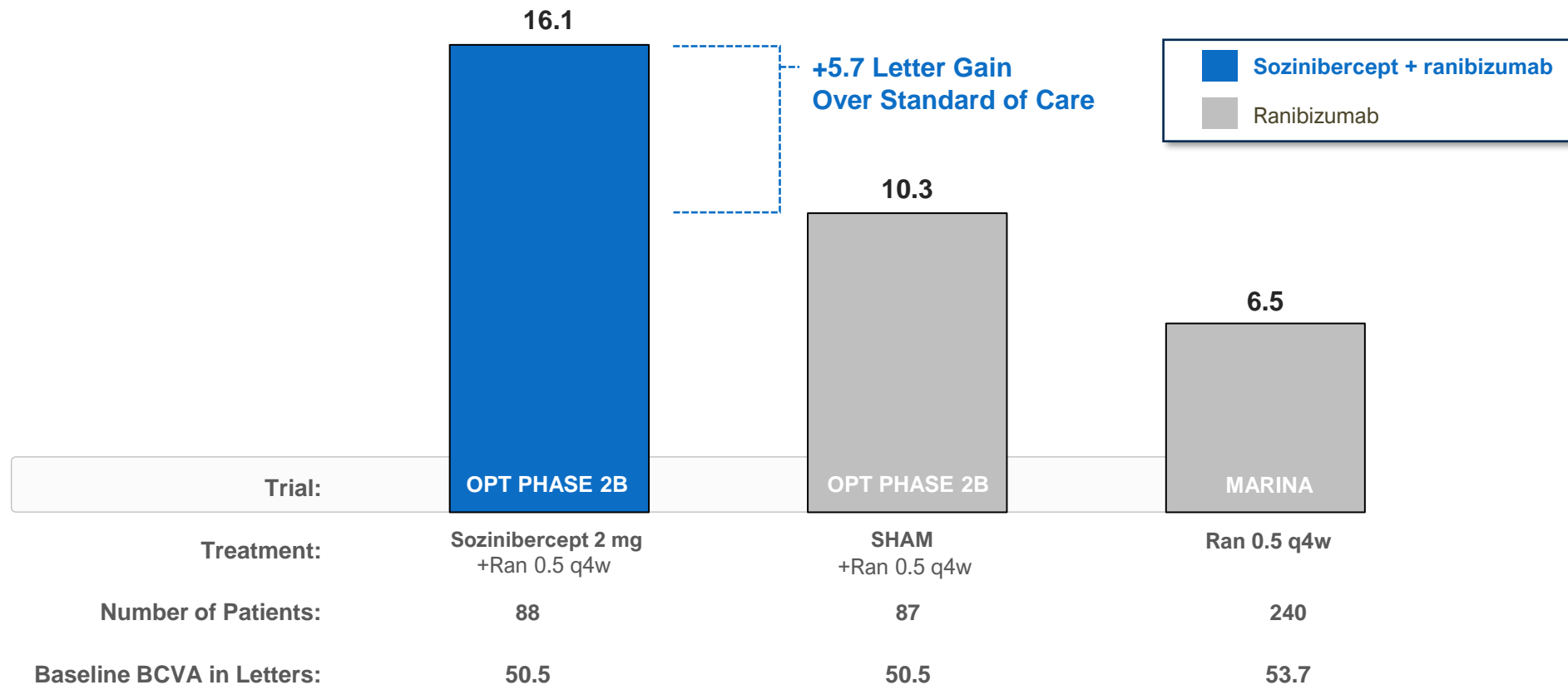
Phase 2b demonstrated **superior efficacy** of **+5.7 letter gain over standard of care**, based on a **pre-specified analysis**

This patient population (minimally classic & occult) represents **~75% of wet AMD patients**

\*Unadjusted p-value

# Control Arm in Phase 2b Overperformed MARINA Trial at Week 24 in Similar Lesion Type Patient Population

**Mean Change in BCVA from Baseline at Week 24 – Sozinibercept Phase 2b vs. MARINA Trial**  
Occult and Minimally Classic Lesions

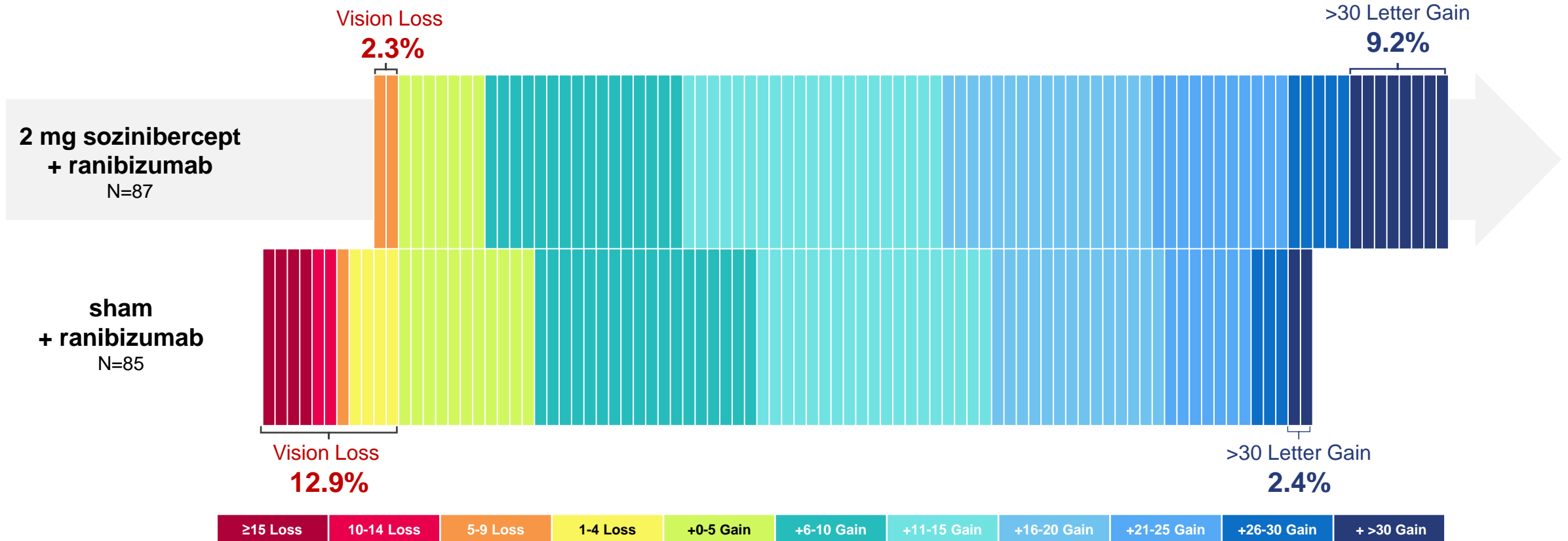


MARINA was a Phase 3 registrational trial. Baseline BCVA values across trials vary. Number of patients randomised to treatment group (n, bottom table). Mean change in Best Corrected Visual Acuity (BCVA) from baseline shown in ETDRS letters (top of bars).

# In a Disease Where Every Letter Counts, a Greater Proportion of Sozinibercept Patients Gained Substantial Vision and Fewer Experienced Vision Loss

## Change from Baseline to Week 24 (ETDRS Letters, Individual Participants)

Occult and Minimally Classic Lesions (RAP Absent)

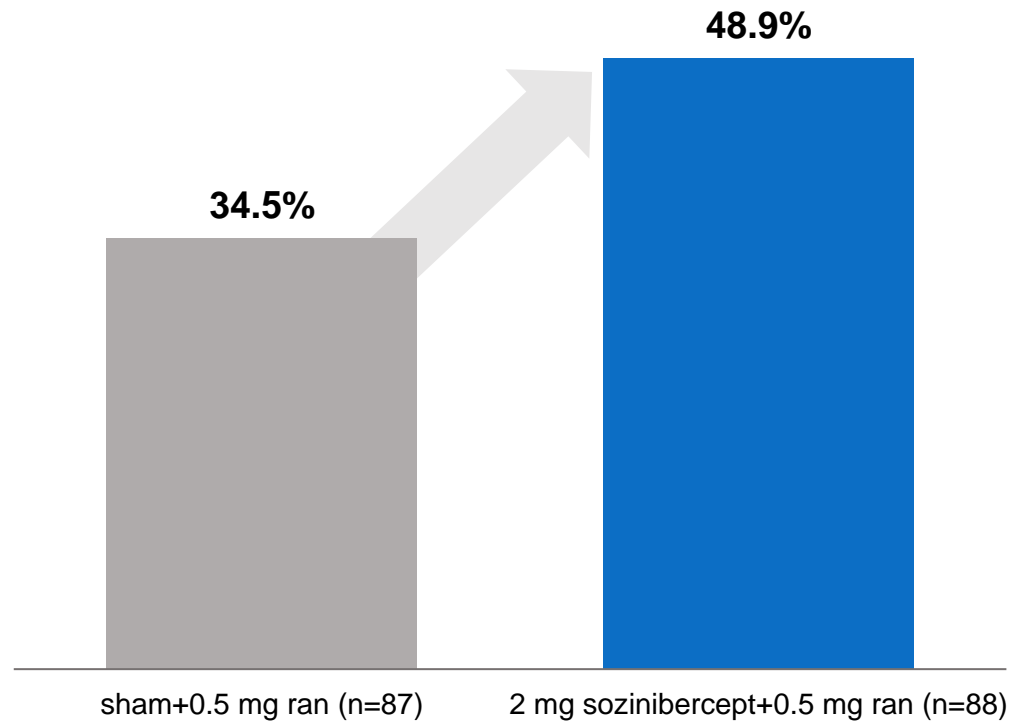


Legend (ETDRS Letter Change from Baseline to Week 24)

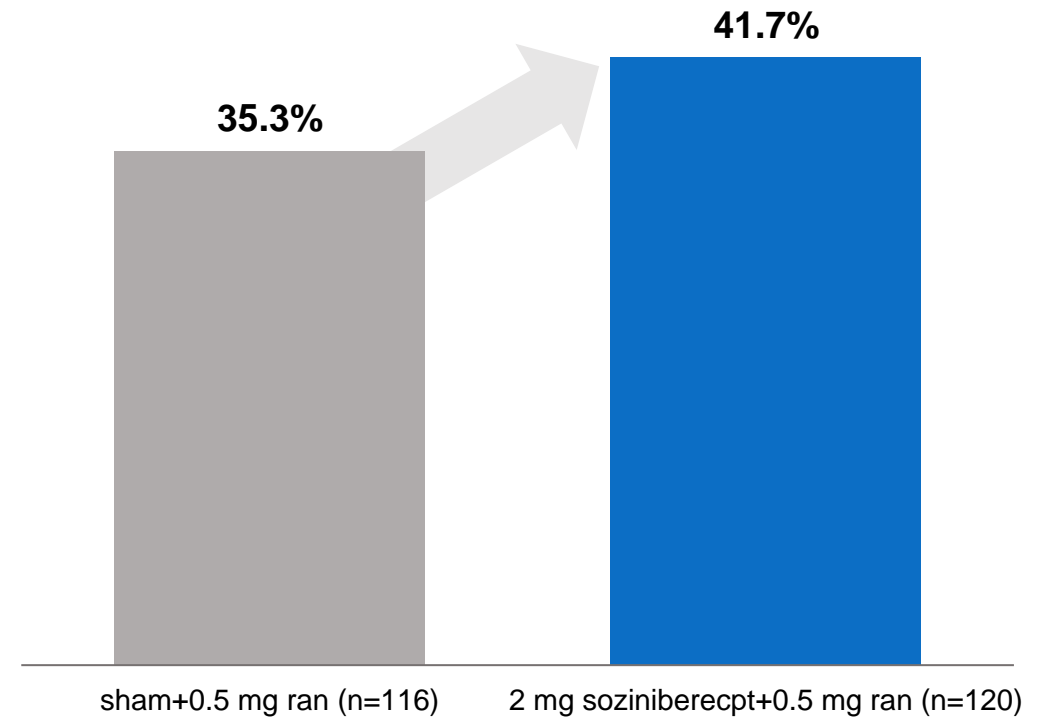
# Greater Proportion of Sozinibercept Patients Achieved Minimum Driving-level of Vision ( $\geq 20/40$ )

## Percentage of Participants with 20/40 Vision or Greater at Week 24

**Occult and Minimally Classic, RAP Absent**  
42% relative increase compared to ranibizumab control



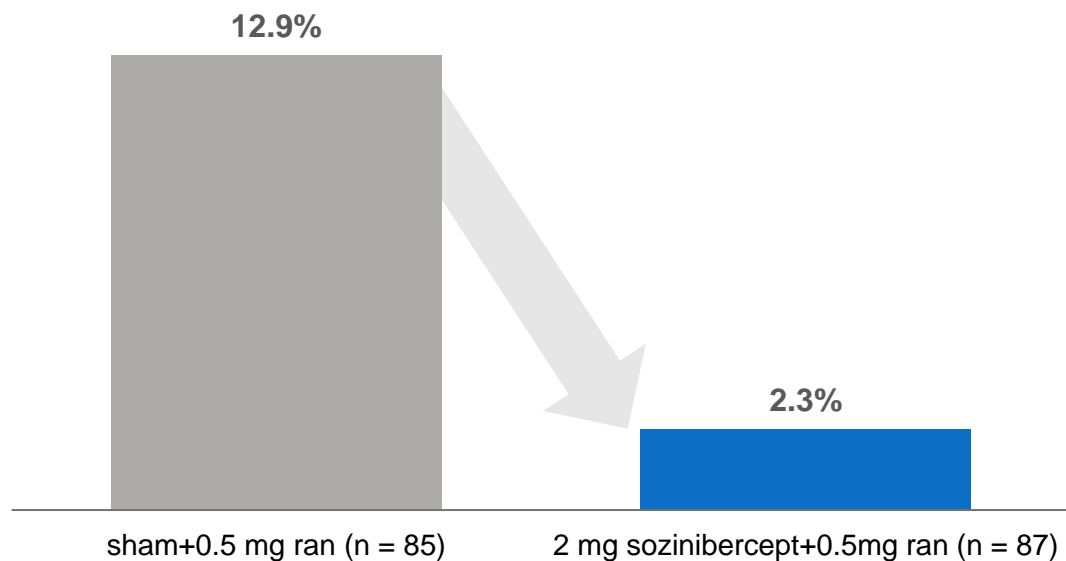
**Total Patient Population**  
18% relative increase compared to ranibizumab control



# Sozinibercept Reduced the Proportion of Patients Experiencing Vision Loss by 82%

## Percentage of Participants with any Vision Loss $\geq 1$ ETDRS Letter at Week 24

**Occult and Minimally Classic Lesions (RAP Absent)**  
82% relative decrease compared to ranibizumab control



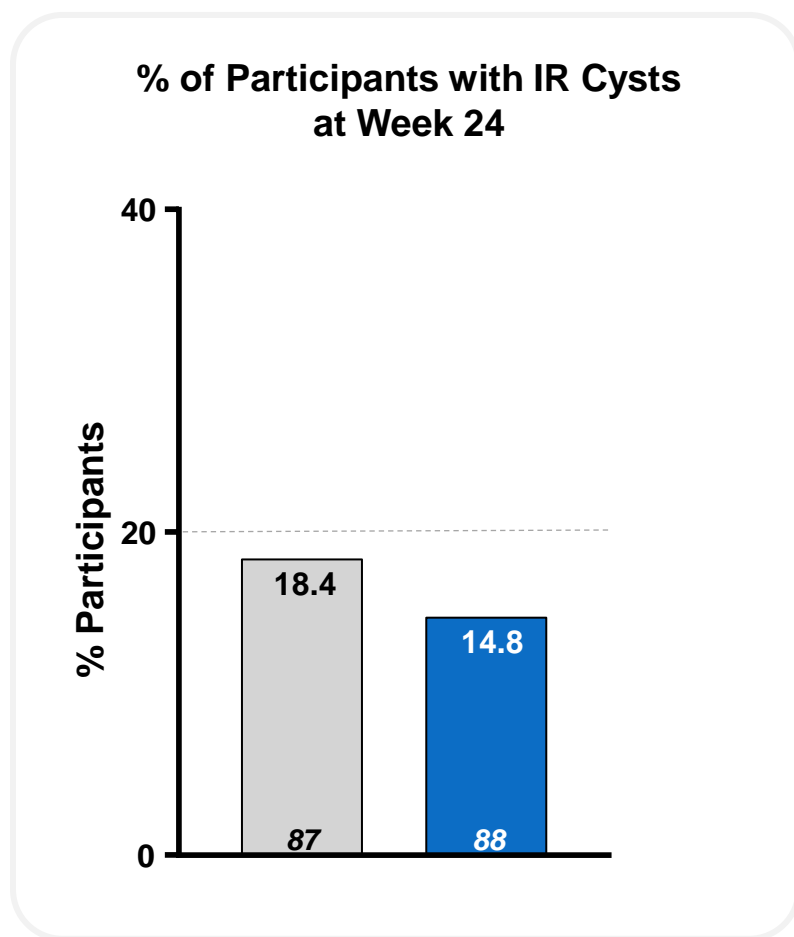
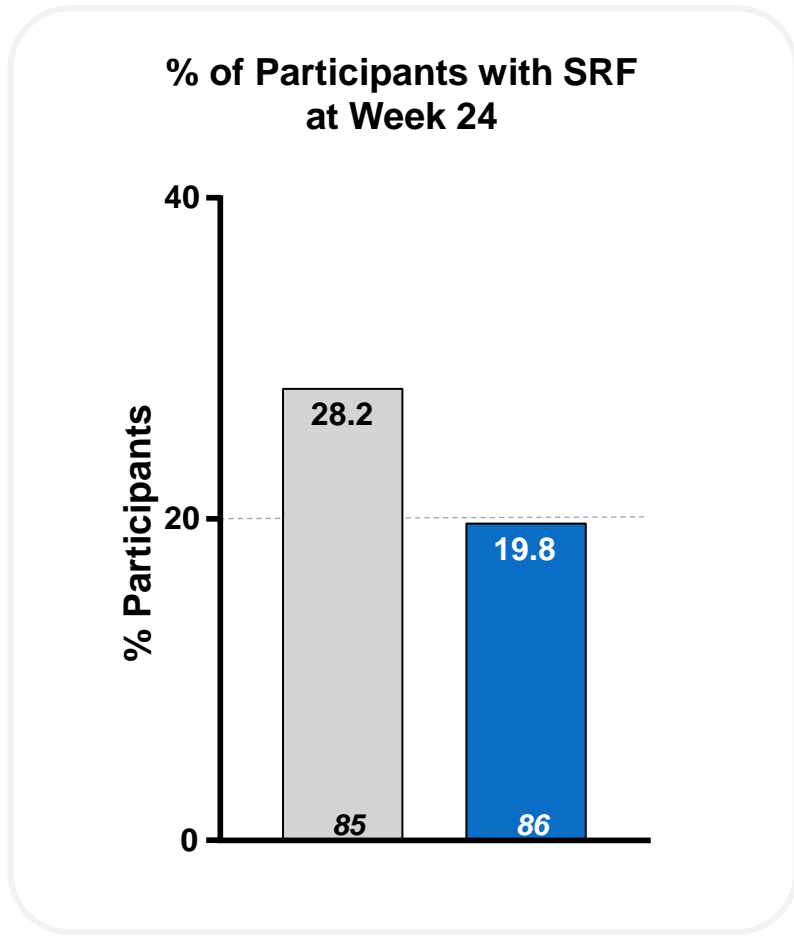
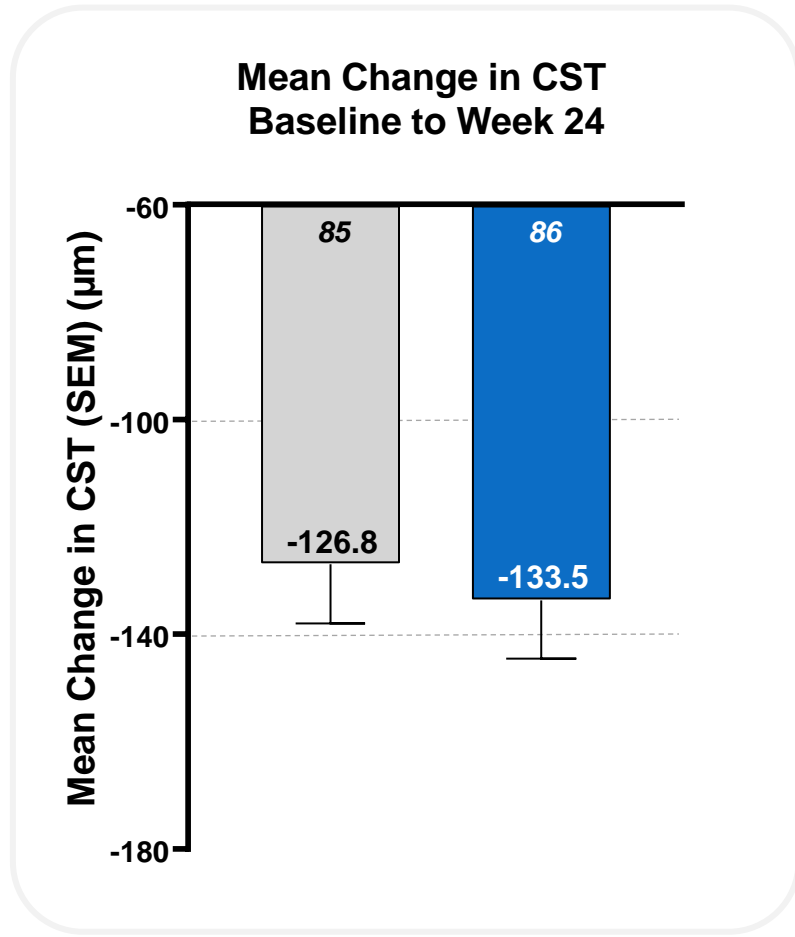
**Decrease of 1 ETDRS letter per year increases mortality risk by 16%<sup>2</sup> associated exclusively with IADL levels**

Modified Intent-to-Treat (mITT) population; as observed.

IADL – Instrumental activities of daily living; ETDRS – Early Treatment Diabetic Retinopathy Study chart

<sup>1</sup>Christ SL, et al. "Longitudinal relationships among visual acuity, daily functional status, and mortality: the Salisbury Eye Evaluation Study." JAMA Ophthalmol. 2014.

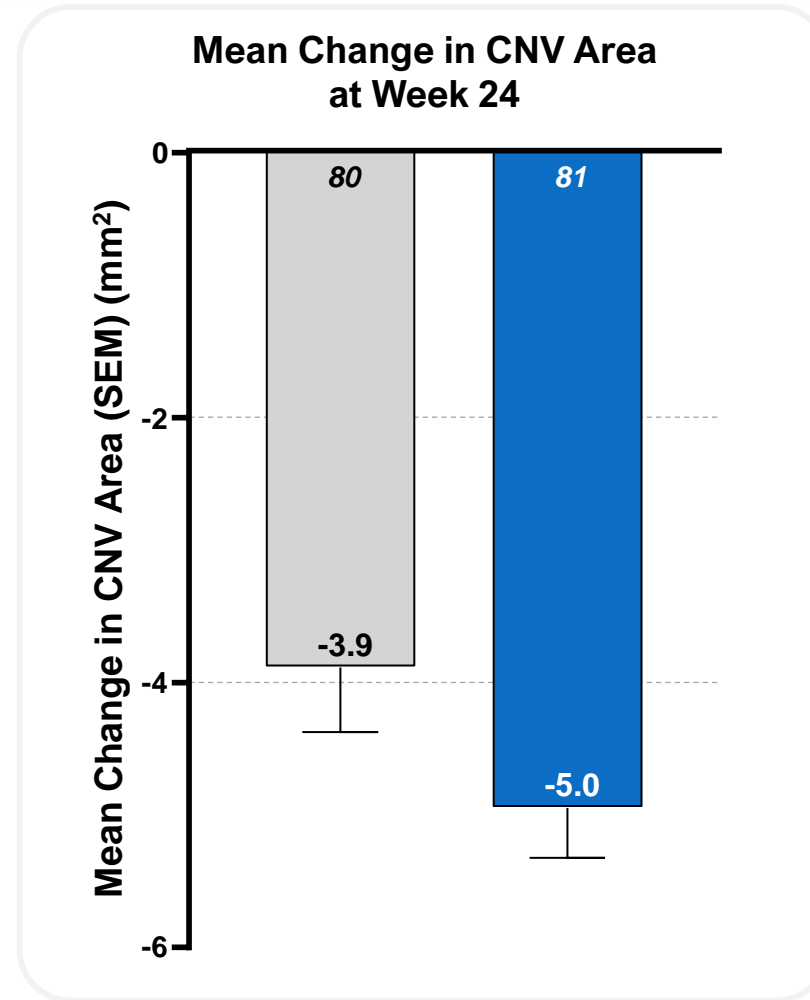
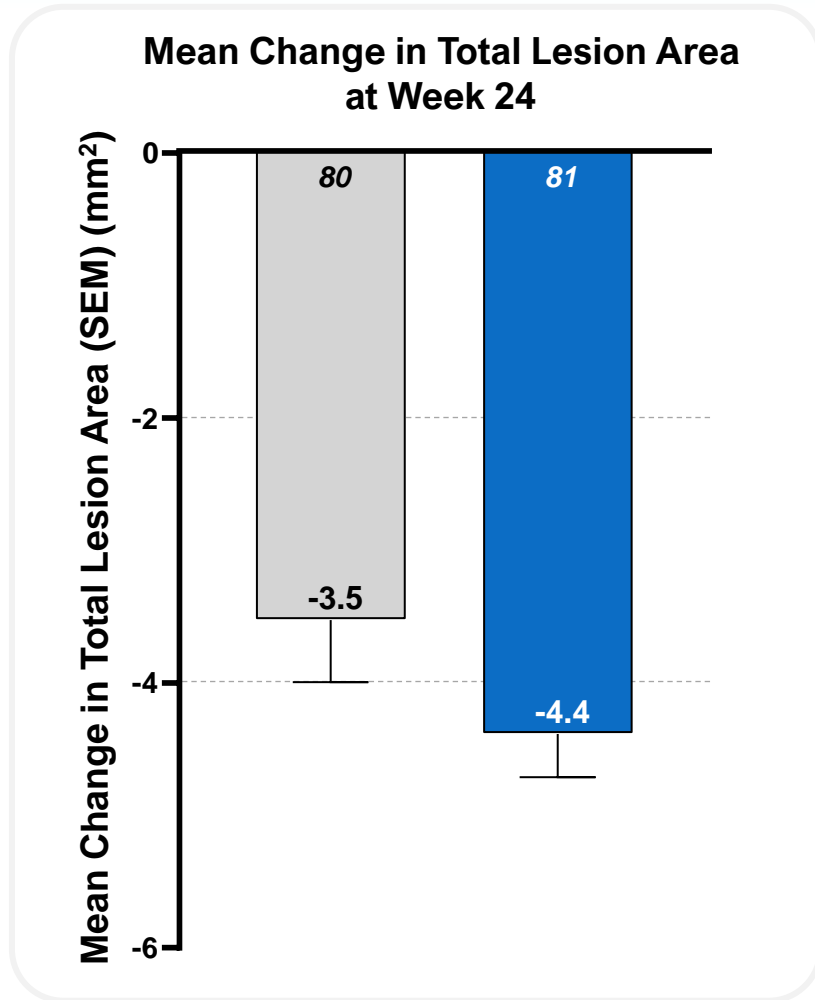
# Sozinibercept Reduced Retinal Thickness and Dried the Retina Better With Combination Therapy in Occult & Minimally Classic (RAP Absent) Patients



Sham + 0.5 mg ranibizumab      2 mg sozinibercept + 0.5 mg ranibizumab

mITT; as observed; top of bar – statistic, bottom of bar – n.  
CST: Central Subfield Thickness; SRF: Subretinal fluid; IR: Intra-retinal.

# Sozinibercept Demonstrated Greater CNV and Lesion Regression With Combination Therapy in Occult & Minimally Classic (RAP Absent) Patients

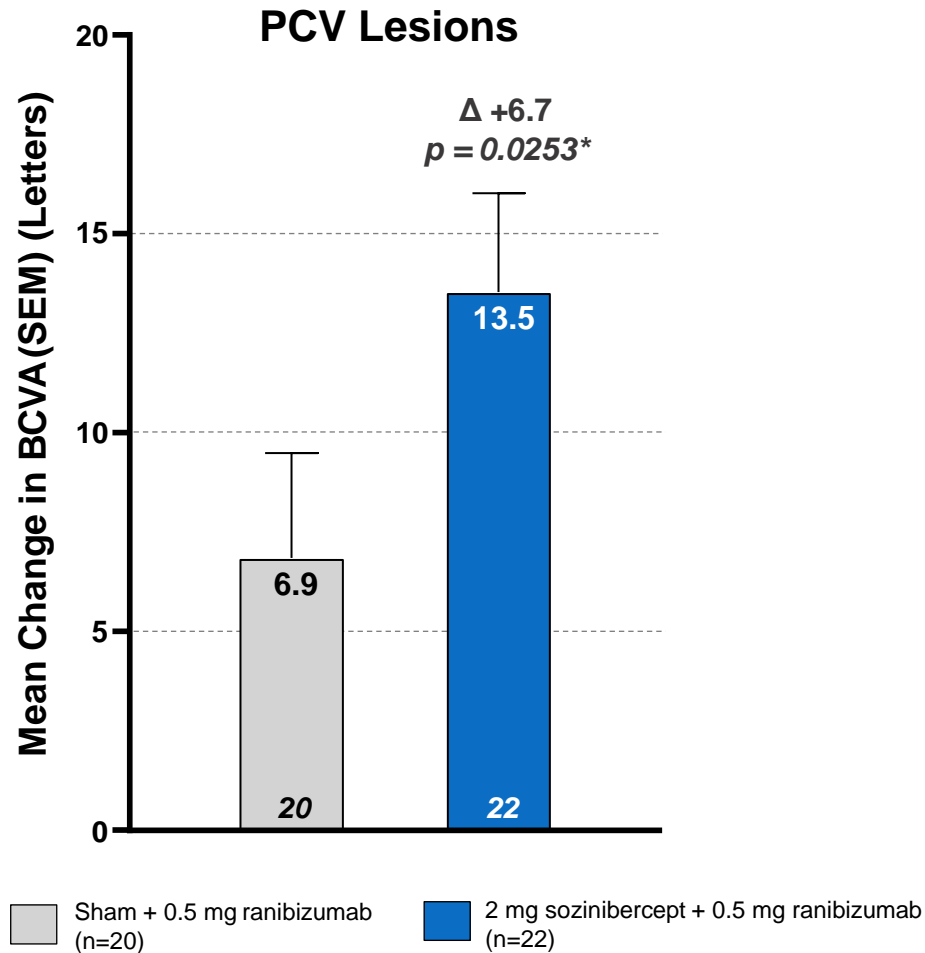


Sham + 0.5 mg ranibizumab (n=80)

2 mg sozinibercept + 0.5 mg ranibizumab (n=81)



# Sozinibercept Demonstrated Superior Vision Gains in a Pre-Specified Subgroup of Hard-To-Treat PCV Lesion Patients



Polypoidal Choroidal Vasculopathy (**PCV**) is a difficult-to-treat wet AMD subtype; it is often described as the **most prevalent form of wet AMD worldwide**

PCV is **highly prevalent in Asian populations** (up to ~60%), while ~8-13% prevalence in Caucasians

**Phase 3 ShORe and COAST trials enrolled patients with PCV<sup>1</sup>**

\*Unadjusted p-value

<sup>1</sup> Evaluated by color fundus photography (FP), fluorescein angiography (FA), and spectral domain optical coherence tomography (SD-OCT)

# Pooled Safety for Completed Sozinibercept Trials

Combination Therapy Well Tolerated and Comparable to Standard of Care Monotherapy

N Participants (%)	Sozinibercept Any dose* N=399 (N=1,842 injections)	Sozinibercept 2 mg N=263 (N=1,121 injections)	Sham + anti-VEGF-A control N=170 (N=854 injections)
Ocular TEAEs - Study Eye – related to study product(s)	41 (10.2%)	22 (8.4%)	20 (11.8%)
Ocular TEAEs - Study Eye – Severe	4 (1.0%)	2 (0.8%)	2 (1.2%)
Intraocular inflammation – Study Eye	7 <sup>1,2,3</sup> (1.8%)	3 <sup>1</sup> (1.1%)	3 <sup>1</sup> (1.8%)
Participants with AEs leading to treatment discontinuation	4 <sup>2,4-6</sup> (1.0%)	1 <sup>4</sup> (0.4%)	2 <sup>7,8</sup> (1.2%)
Any APTC event	4 <sup>4,5,9,10</sup> (1.0%)	3 <sup>5,9,10</sup> (1.1%)	2 <sup>11,12</sup> (1.2%)
Deaths	2 <sup>10,13</sup> (0.5%)	2 <sup>10,13</sup> (0.8%)	2 <sup>14,15</sup> (1.2%)

- Pooled safety analysis of 399 patients for completed sozinibercept trials
- Data Monitoring Committee (“DMC”) regularly reviews data from ongoing Phase 3 COAST and ShORe studies
- Safety data from our completed sozinibercept trials show sozinibercept combination therapy has a safety and tolerability profile comparable to standard of care anti-VEGF-A monotherapy.
- Masked data from patients that have completed the week 52 visit in the ongoing Phase 3 clinical trials show greater mean BCVA increases from baseline than results with standard of care anti-VEGF-A monotherapy from Opthea’s Phase 2b study\*\*

<sup>1</sup>Transient anterior chamber cell (trace 1-4 cells); <sup>2</sup>SAE of endophthalmitis, with AE’s of hypopyon and anterior chamber cell (n=1; 0.5 mg); <sup>3</sup>SAE of vitritis (n=1; 0.5 mg); <sup>4</sup>Non-fatal myocardial infarction; <sup>5</sup>Cerebrovascular accident; <sup>6</sup>Enteritis; <sup>7</sup>Abdominal pain;

<sup>8</sup>Increased IOP; <sup>9</sup>Non-fatal angina pectoris; <sup>10</sup>Fatal congestive heart failure/myocardial infarction; <sup>11</sup>Non-fatal arterial embolism; <sup>12</sup>Embolic stroke; <sup>13</sup>Metastatic ovarian cancer; <sup>14</sup>Pneumonia; <sup>15</sup>infective endocarditis.

\*Any dose (sozinibercept 0.3 mg, 0.5 mg, 1 mg or 2 mg)

\*\*Masked data represent pooled data from both sozinibercept combination and standard of care monotherapy treatment arms. The Phase 3 clinical trial masked data are incomplete and subject to additional analysis once unmasked. There is no assurance that standard of care monotherapy in our Phase 3 clinical trials will yield similar results to our prior clinical trials or previously published clinical trials with anti-VEGF-A monotherapies. As a result, there can be no assurance that topline results for sozinibercept from the Phase 3 clinical trial, if completed, will be consistent with results from masked data available to date.

# Intraocular Inflammation Observed in Combination Therapy Across Completed Sozinibercept Trials Similar to Standard of Care

<b>N Participants (%)</b>	<b>Sozinibercept Any dose* N=399 (N=1,842 injections)</b>	<b>Sozinibercept 2 mg N=263 (N=1,121 injections)</b>	<b>Sham + anti-VEGF-A control N=170 (N=854 injections)</b>
<b>Intraocular Inflammation<sup>1</sup></b>	7 (1.8%)	3 (1.1%)	3 (1.8%)
<b>OPT-302-1001 (Phase 1/2a wet AMD)</b>	2	0	0
Uveitis with anterior chamber cell 1+	1	0	0
Uveitis with anterior chamber cell 2+	1	0	0
<b>OPT-302-1002 (Phase 2b wet AMD)</b>	3	1	2 <sup>a</sup>
Endophthalmitis with anterior chamber 1+ and hypopyon	1	0	0
Vitritis	1	0	0
Anterior chamber cell, trace	1	1	2 <sup>a</sup>
<b>OPT-302-1003 (Phase 1b/2a DME)</b>	2 <sup>b</sup>	2 <sup>b</sup>	1
Iritis with keratic precipitates and anterior chamber cell 2+	1	1	0
Iritis with anterior chamber cell 2+	0	0	1
Anterior chamber cell 4+, associated with cataract extraction/ intraocular lens implant and hyphema	1 <sup>b</sup>	1 <sup>b</sup>	0

Safety population

<sup>1</sup>AEs observations considered to be indicative of intraocular inflammation, defined prior to database lock

<sup>a</sup>Observed during ophthalmic examination, but not reported as TEAEs

<sup>b</sup>Considered associated with lens extraction and not reported as TEAEs

# Phase 3 Clinical Program Is Informed by Phase 2b Results and Optimized for Success



**Hierarchical primary analysis first conducted in the high-responding occult and minimally classic population (RAP absent), followed by total patient population**



**Two robust pivotal trials studying sozinibercept in combination with Eylea<sup>®</sup> and Lucentis<sup>®</sup> in treatment naïve patients with wet AMD**



**Phase 3 designed to support broad label for use in combination with any VEGF-A inhibitor for all wet AMD patients (treatment naïve and prior treated)**

# Phase 3 Wet AMD Trials COAST and ShORe Are Well Advanced

Topline Data Anticipated for COAST in Early 2Q CY 2025 and ShORe in Mid-CY2025

## Design

- Multi-center, double-masked, randomized (1:1:1), sham control
- Treatment naïve wet AMD patients

## Sample Size

- COAST n=998; ShORe n=986

## Comparators

- 2 mg Eylea<sup>®</sup> q8w (COAST) & 0.5 mg Lucentis<sup>®</sup> q4w (ShORe)

## Regulatory Quality

- ~90% power, 5% type I error rate

# Phase 3 Primary and Secondary Endpoints

Primary Efficacy Endpoint at Week 52 to Support BLA Submission

## Primary Endpoint

**Mean change from baseline in BCVA at week 52**

## Key Secondary Endpoints (Baseline to Week 52)

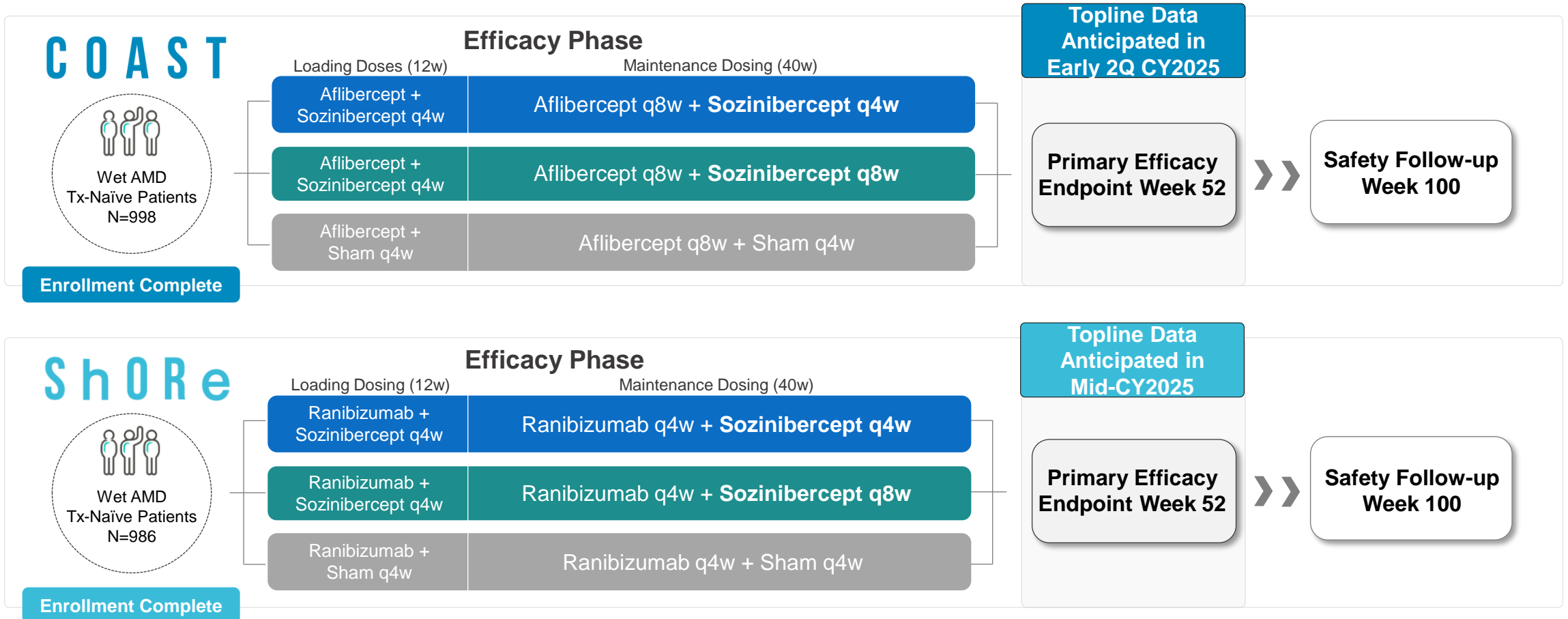
Proportion of participants gaining  $\geq 15$  letters

Proportion of participants gaining  $\geq 10$  letters

Change in choroidal neovascularization area

Proportion of participants with absence of both sub-retinal fluid and intra-retinal cysts

# Phase 3 Trial Design Supports Potential Broad Label for Use With Any Anti-VEGF-A Therapy



Standard of care administered according to approved dosing schedule: **afibercept** (2 mg IVT q8w after 3 loading doses) and **ranibizumab** (0.5 mg IVT q4w after 3 loading doses). Sozinibercept dosed at 2 mg. Note that Sham administered at visits when sozinibercept is not administered. Maintenance dosing continued through end of the safety follow-up.

# Advancing Therapeutic Innovations to Transform Patient Outcomes with Superior Vision Gains

We are dedicated to advancing sozinibercept to **improve patients' visual outcomes**

## Next Steps

### Clinical Milestones

- Phase 3 program enrolled 1,984 patients across COAST and ShORe
- Topline data anticipated for COAST in early 2Q CY2025 and ShORe in mid-CY2025

### Manufacturing Scale-up

- Production of validation batches supportive of BLA filing and launch

### Regulatory Preparations

- FDA Fast Track designation allows rolling submission of completed BLA modules

### Commercial Readiness

- Strengthen medical expert engagement and develop market access strategy
- Complete development of product launch plan



# Recent Financings Anticipated to Provide Cash Runway Through Both Pivotal Topline Data Readouts

## Financial Overview

<b>Ticker</b>	OPT (ASX/NASDAQ)
<b>Shares Outstanding<sup>1</sup></b>	Ordinary Shares: 1,231.1M ADS equivalents: 153.9M
<b>Cash/Cash Equivalents<sup>2</sup></b>	US\$207.3M
<b>Offices</b>	Melbourne, Australia Princeton, NJ

## Development Funding Agreement (DFA)

- Total funding drawn under DFA: US\$170M
- Provides non-dilutive funding for development of sozinibercept
- If sozinibercept is approved, repayment is capped at 4x investment and split between fixed payments and variable payments at 7% of revenues
- No amounts owed if the clinical trials do not meet the primary endpoint or if regulatory approval is not received<sup>3</sup>

<sup>1</sup>As of June 30, 2024, pro-forma for the 2024 Retail Entitlement Offer which closed in July 2024.

<sup>2</sup>Includes \$172.5M as of June 30, 2024 and \$34.8M net proceeds from the 2024 Retail Entitlement Offer which closed in July 2024.

<sup>3</sup>In certain circumstances, upon or following the termination of the DFA, the Company may owe the DFA investors a multiple of amounts paid to the Company under the DFA. Please refer to the description of the DFA included in the Company's Form 6-K filed with the SEC on August 15, 2022 and the DFA filed as Exhibit 4.14 to the Company's Annual Report on Form 20-F filed with the SEC on September 29, 2022 for more information.

Note: Additional funding will be required to reach commercialization of sozinibercept and to meet obligations under the Development Funding Agreement ("DFA"). As a result of obligations under the DFA and applicable law regarding liquidity, the Company expects to raise or obtain additional capital in one or more transactions, earlier than 3Q CY 2025 or anticipated topline data readout dates.

# Sozinibercept Will Not Compete Head-to-Head with Anti-VEGF-A

Differentiated Combination Approach Targeting Better Visual Outcomes Drives Commercial Value

1

**Addressing unmet medical need of improved efficacy** in large wet AMD patient population in a potential ~\$15B market

2

**First and only therapy to have demonstrated superior visual outcomes** over anti-VEGF-A therapy with a novel and highly differentiated MOA

3

**Only asset in near or long-term pipeline with potential to disrupt treatment paradigm** on basis of efficacy in wet AMD

4

**Concentrated prescriptions in U.S. enables potential self-commercialization opportunity** with lean and targeted organization