

CORPORATE UPDATE INVESTOR CALL – 20 August 2025

Dr Jeremy Levin – Chairman of the Board

Fred Guerard – Chief Executive Officer

Tom Reilly – Chief Financial Officer

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AGENDA – CORPORATE UPDATE

- Introduction – Dr Jeremy Levin, Chairman of the Board
- Results of Sozinibercept Phase 3 Trials (COAST and ShORe) – Fred Guerard, CEO
- Update on Development Funding Agreement Negotiations – Tom Reilly, CFO
- Current state of the company – Dr Jeremy Levin, Chairman of the Board
- The path forward – Dr Jeremy Levin, Chairman of the Board
- Q&A

Introduction

Dr Jeremy Levin

INTRODUCTION

- The results of our Phase 3 program did not meet their primary endpoints and were deeply disappointing
- Thanks to our CEO's and CFO's efforts, we have now completed negotiations with the Development Funding Agreement investors, gaining clarity on available resources and outcomes for shareholders
- The purpose of this presentation is to share clinical results with shareholders, provide an update on the DFA outcomes and outline potential opportunities for the company moving forward
- While there have been challenges, this marks a new chapter for the company

Results of Sozinibercept Phase 3 Trials in Participants with Neovascular Age-related Macular Degeneration: COAST (Sozinibercept in Combination with Aflibercept) and ShORe (Sozinibercept in Combination with Ranibizumab)

Fred Guerard

Sozinibercept Phase 3 Trials Design

Primary Endpoint

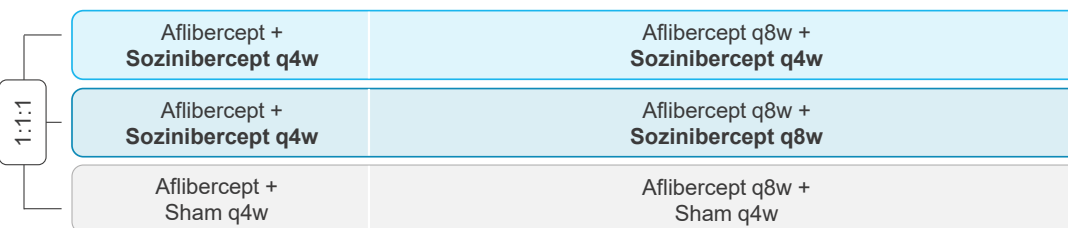
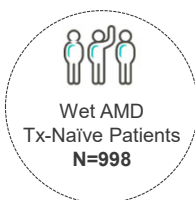
- Mean change in BCVA from baseline to week 52

Key Secondary Endpoints

(Baseline to Week 52)

- Proportion of participants gaining ≥ 15 letters
- Proportion of participants gaining ≥ 10 letters
- Change in CNV area
- Proportion of participants with absence of both SRF and IR cysts

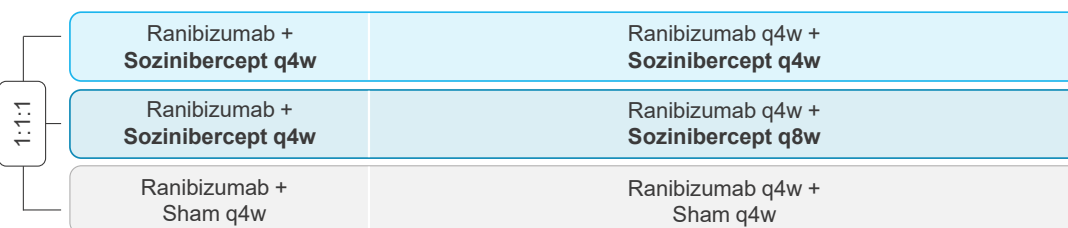
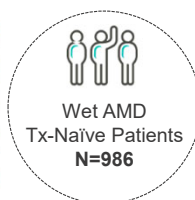
COAST



**Primary Efficacy
Endpoint Week 52**

**Safety Follow-up
Week 100**

ShORE



**Primary Efficacy
Endpoint Week 52**

**Safety Follow-up
Week 100**

Loading Doses

Maintenance Dosing

Standard of care administered according to approved dosing schedule: aflibercept (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.

COAST Baseline Characteristics

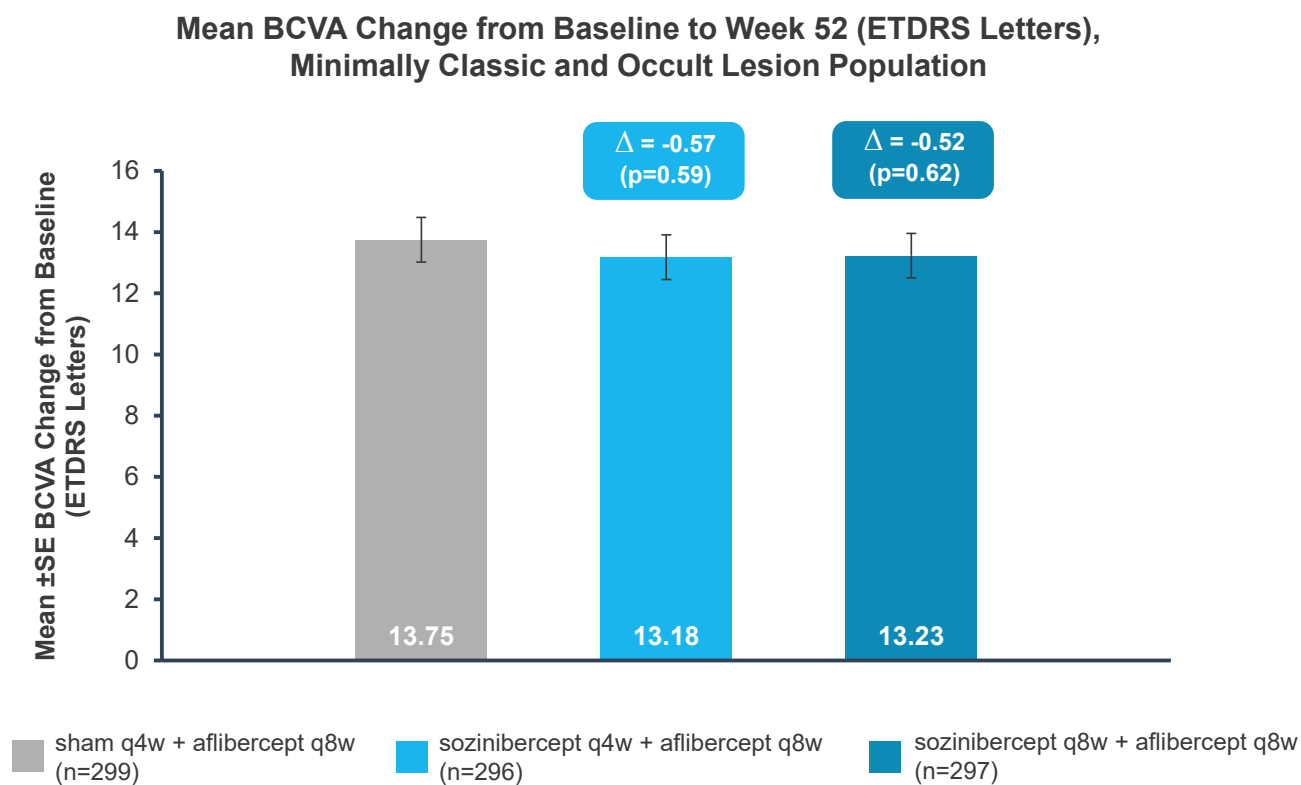
Baseline Characteristics Were Well Balanced Across Arms

Demographic/Baseline Disease Characteristic, Overall Population		sham q4w + aflibercept q8w (n=330)	sozinibercept q4w + aflibercept q8w (n=333)	sozinibercept q8w + aflibercept q8w (n=330)
Mean Age – years ± SD		75.2 (8.28)	74.3 (7.80)	74.9 (7.97)
Sex – n (%)	Male	146 (44.2%)	143 (42.9%)	147 (44.5%)
	Female	184 (55.8%)	190 (57.1%)	183 (55.5%)
Race – n (%)	White	281 (85.2%)	288 (86.5%)	287 (87.0%)
	Asian	29 (8.8%)	29 (8.7%)	25 (7.6%)
	American Indian or Alaska Native	6 (1.8%)	3 (0.9%)	9 (2.7%)
	Black or African American	0	1 (0.3%)	0
	Other	13 (3.9%)	12 (3.6%)	8 (2.4%)
Mean Visual Acuity (BCVA) – letters ± SD		52.4 (9.65)	52.8 (9.04)	52.3 (9.63)
Mean Total CNV Lesion Area - mm ² ± SD		6.54 (3.19)	6.16 (3.28)	6.47 (3.14)
Lesion Type*	Occult - n (%)	186 (56.4%)	186 (55.9%)	182 (55.2%)
	Minimally classic – n (%)	113 (34.2%)	110 (33.0%)	115 (34.8%)
	Predominantly classic – n (%)	31 (9.4%)	37 (11.1%)	33 (10.0%)
Mean central subfield thickness (CST) - mm ±SD		449.46 (136.55)	445.58 (140.09)	444.52 (142.83)
Region	North America	92 (27.9%)	93 (27.9%)	92 (27.9%)
	South America	33 (10.0%)	33 (9.9%)	33 (10.0%)
	Europe/West Asia	176 (53.3%)	179 (53.8%)	178 (53.9%)
	Asia and Pacific	29 (8.8%)	28 (8.4%)	27 (8.2%)

*Lesion type determined by independent reading center

COAST Phase 3 Trial Did Not Achieve Primary Endpoint

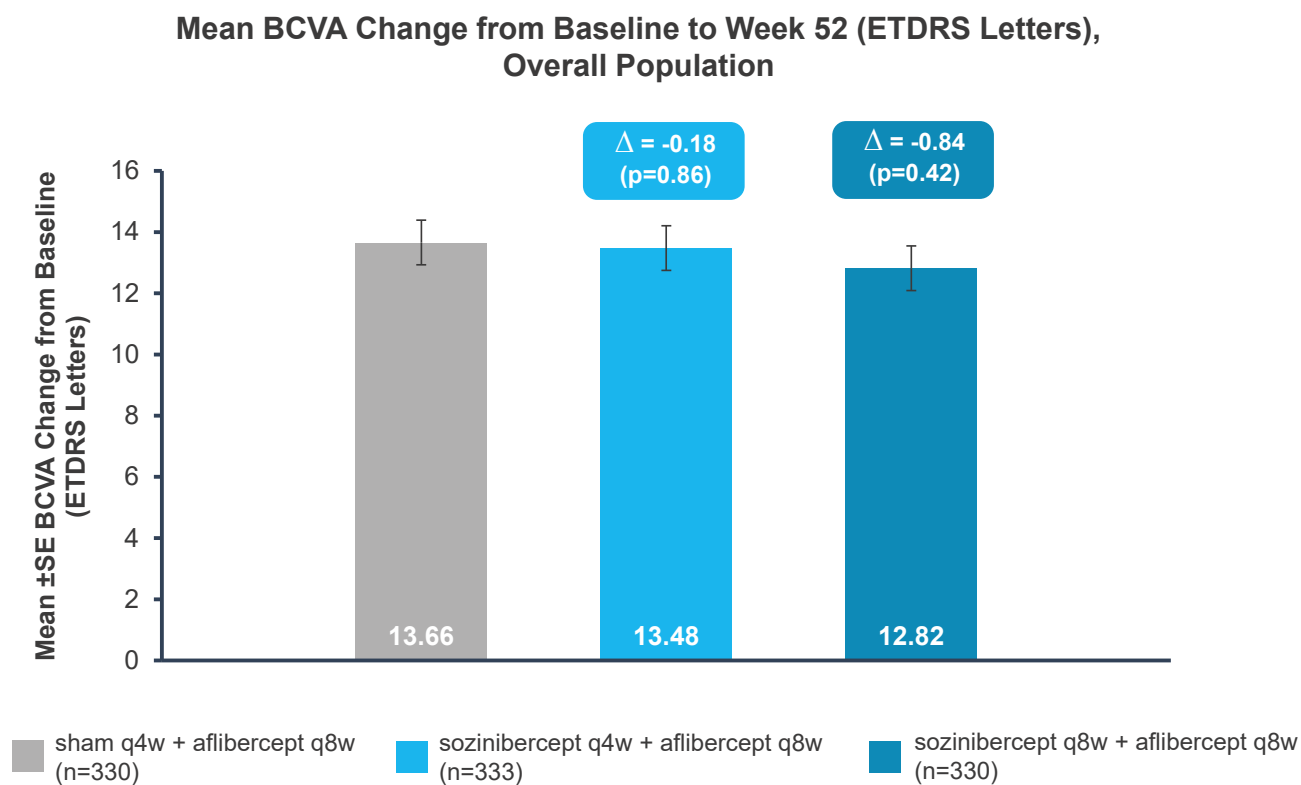
Similar Visual Outcomes Across All Treatment Arms



BCVA: best corrected visual acuity

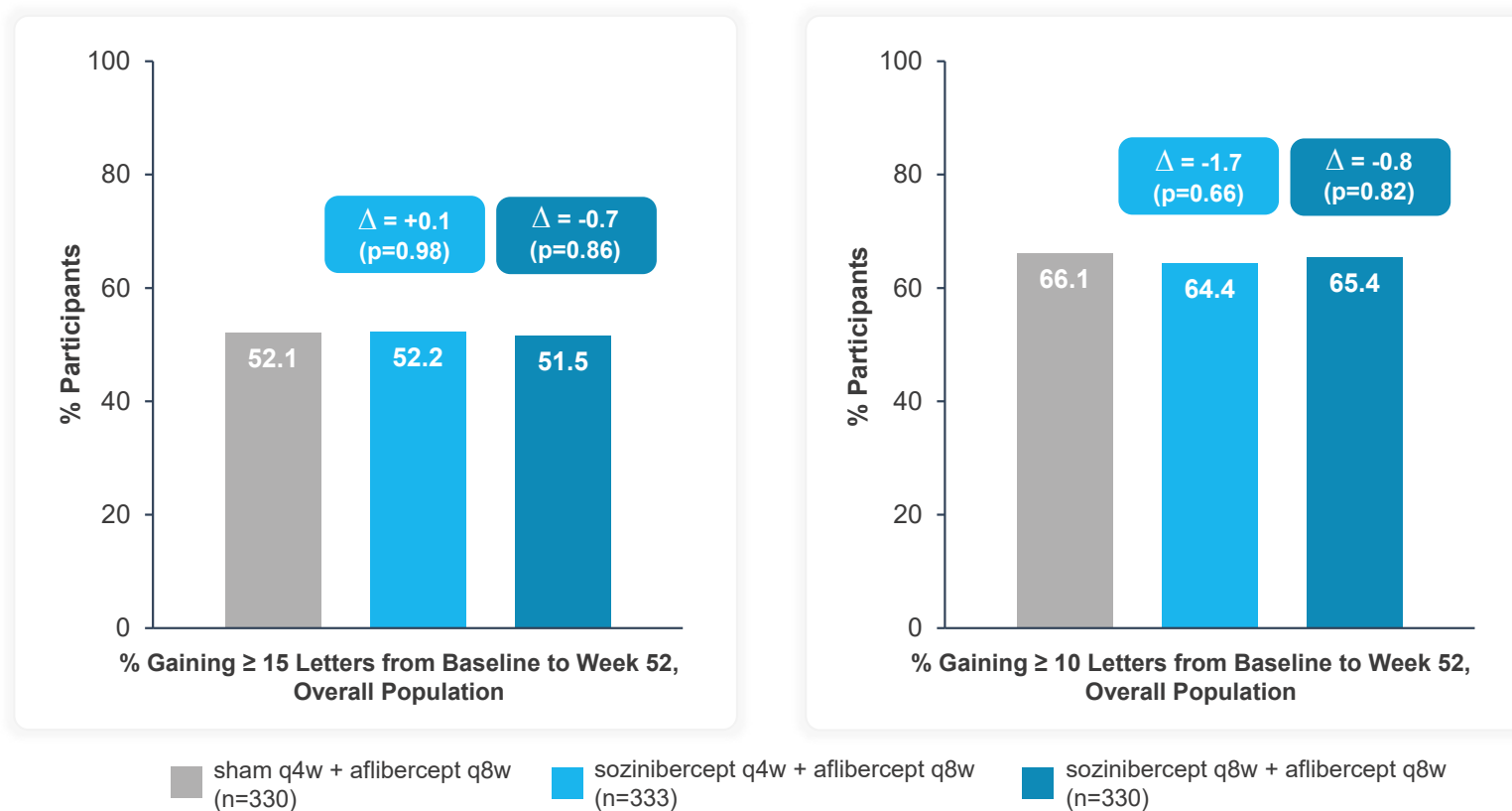
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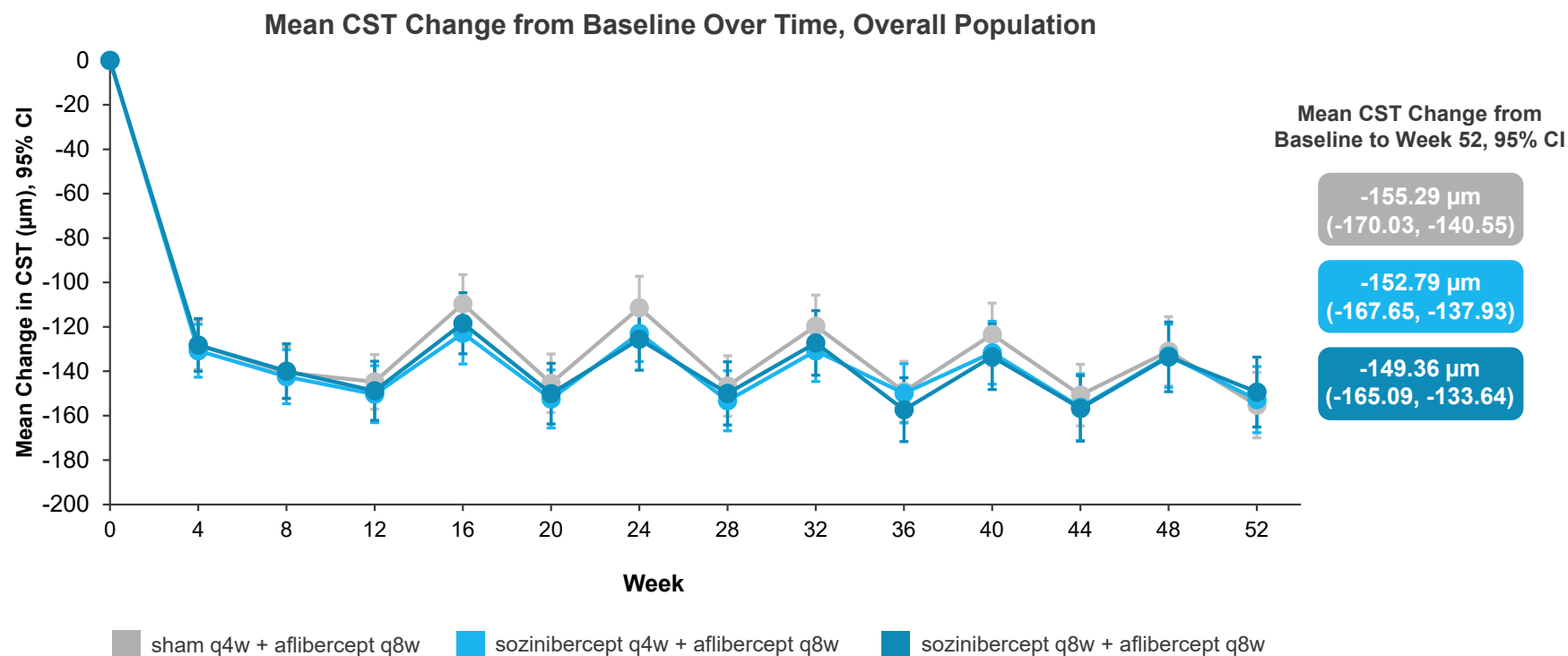
BCVA: best corrected visual acuity
Mixed Model for Repeated Measures (MMRM) Analysis

Similar Results Observed Across Secondary Visual Acuity Endpoints



Multiple Imputation Analysis Assuming Missing at Random

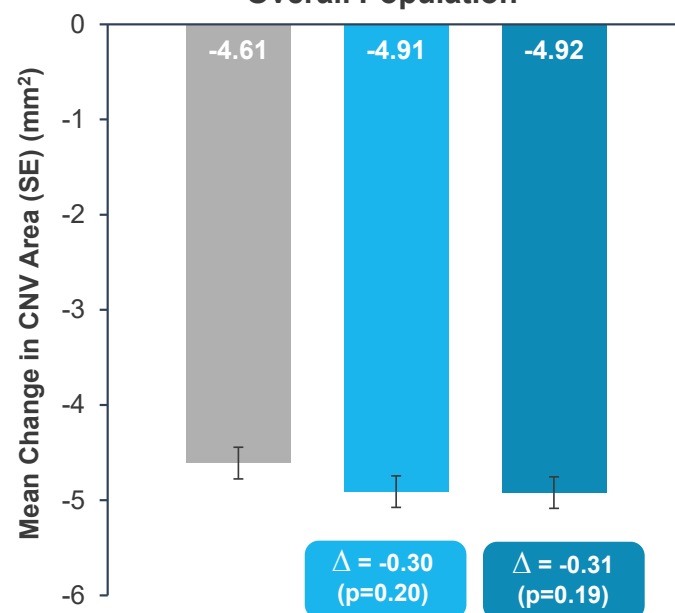
Similar Reductions in CST Observed in All Treatment Arms



CST: central subfield thickness

No Statistically Significant Difference in CNV Regression Across Treatment Arms

LS Mean Change in CNV Area from Baseline to Week 52,
Overall Population



sham q4w + aflibercept q8w
(n=330)

sozinibercept q4w + aflibercept q8w
(n=333)

sozinibercept q8w + aflibercept q8w
(n=330)

CNV: choroidal neovascularization
Mixed Model for Repeated Measures (MMRM) Analysis

Treatment Emergent Adverse Events

Sozinibercept Combination Therapy Demonstrated Similar Safety Profile to Aflibercept

	sham q4w + aflibercept q8w (n=330)	sozinibercept q4w + aflibercept q8w (n=333)	sozinibercept q8w + aflibercept q8w (n=330)
TEAEs, n (%)	256 (77.6%)	260 (78.1%)	255 (77.3%)
Ocular in the study eye	124 (37.6%)	140 (42.0%)	130 (39.4%)
Non-ocular	204 (61.8%)	202 (60.7%)	199 (60.3%)
Serious TEAEs, n (%)	50 (15.2%)	42 (12.6%)	29 (8.8%)
Ocular in the study eye	6 (1.8%)	5 (1.5%)	3 (0.9%)
Non-ocular	44 (13.3%)	37 (11.1%)	25 (7.6%)
TEAEs leading to study discontinuation, n (%)	7 (2.1%)	15 (4.5%)	9 (2.7%)
Ocular in the study eye	2 (0.6%)	10 (3.0%)	1 (0.3%)
Non-ocular	5 (1.5%)	5 (1.5%)	8 (2.4%)

TEAEs: treatment emergent adverse events

Serious Ocular TEAEs

Sozinibercept Combination Therapy Demonstrated Similar Safety Profile to Aflibercept

Serious ocular TEAEs in study eye, n (%)	sham q4w + aflibercept q8w (n=330)	sozinibercept q4w + aflibercept q8w (n=333)	sozinibercept q8w + aflibercept q8w (n=330)
Visual acuity reduced	1 (0.3)	1 (0.3)	0
Retinal detachment	0	1 (0.3)	0
Retinal haemorrhage	2 (0.6)	0	1 (0.3)
Eye inflammation	0	0	1 (0.3)
Uveitis	0	0	1 (0.3)
Ocular hypertension	1 (0.3)	0	0
Rhegmatogenous retinal detachment	1 (0.3)	0	0
Endophthalmitis	1 (0.3)	3 (0.9)	0
Cataract traumatic	0	1 (0.3)	0

TEAEs: treatment emergent adverse events

Cases of Intraocular Inflammation in the Study Eye

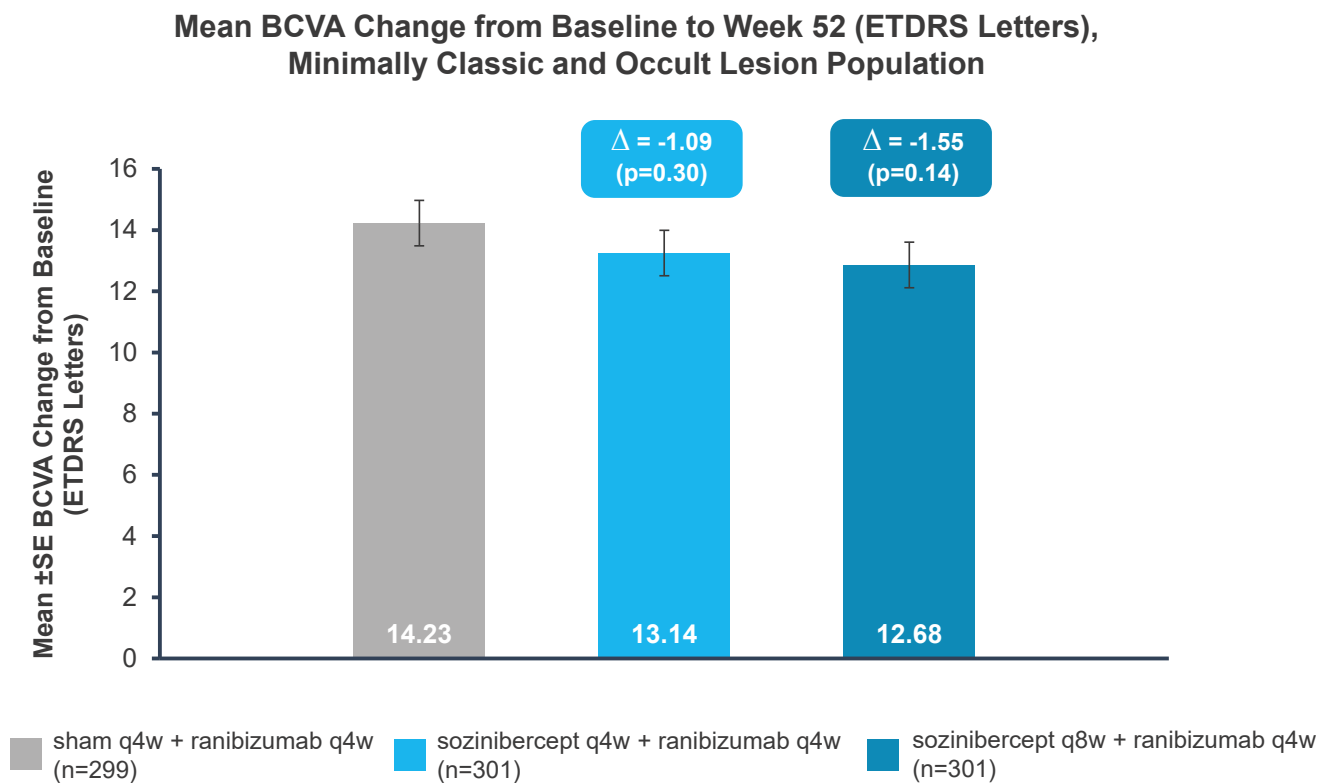
Low Rates of IOI, Majority of Cases Observed in Sozinibercept Combination Arms

	sham q4w + aflibercept q8w (n=330)	sozinibercept q4w + aflibercept q8w (n=333)	sozinibercept q8w + aflibercept q8w (n=330)
Intraocular Inflammation	0	10 (3.0)	6 (1.8)
Uveitis	0	4 (1.2)	2 (0.6)
Iridocyclitis	0	4 (1.2)	1 (0.3)
Eye inflammation	0	2 (0.6)	1 (0.3)
Vitritis	0	2 (0.6)	1 (0.3)
Keratic precipitates	0	1 (0.3)	0
Anterior chamber cell	0	0	2 (0.6)
Anterior chamber inflammation	0	0	1 (0.3)
Endophthalmitis	1 (0.3)	3 (0.9)	0

IOI: intraocular inflammation

ShORe Phase 3 Trial Did Not Achieve Primary Endpoint

ShORe (Sozinibercept in Combination with Ranibizumab) Unmasked Early Following COAST Results

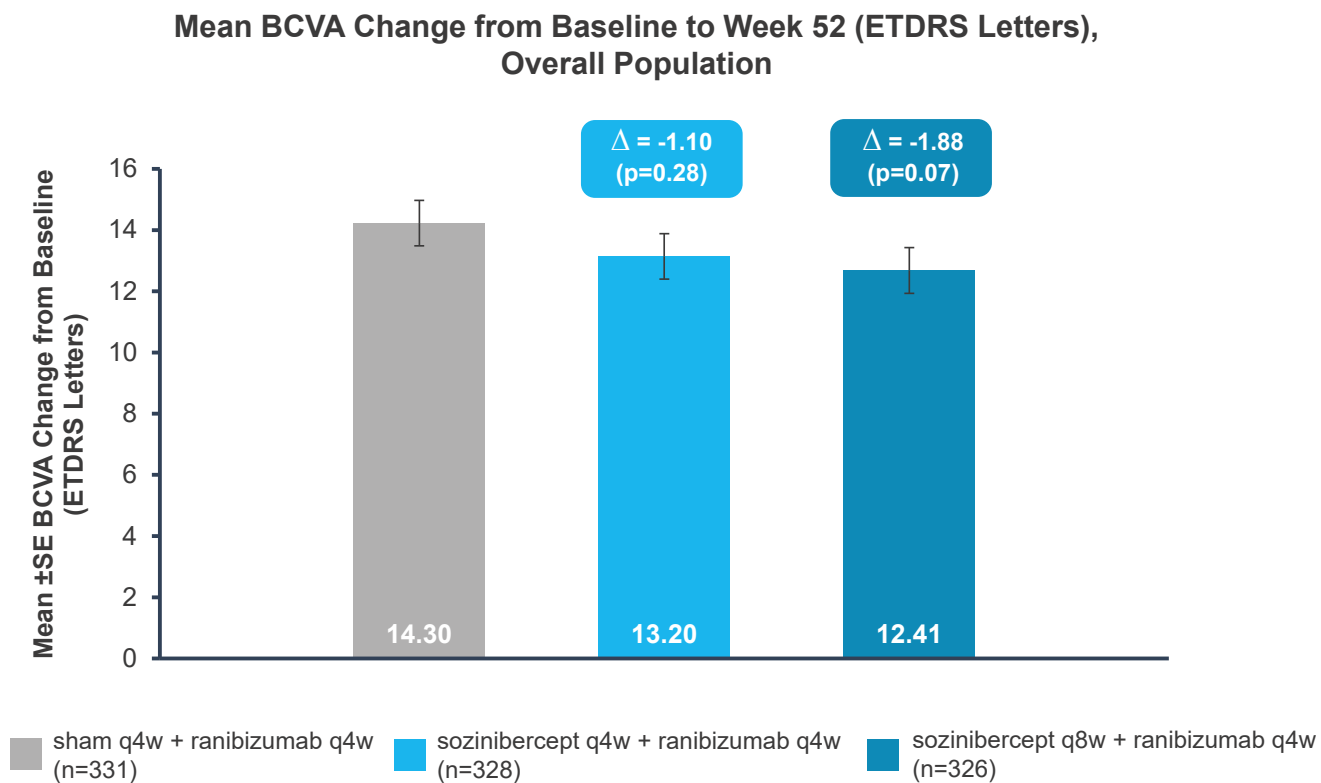


BCVA: Best Corrected Visual Acuity

Because the ShORe trial was unmasked before completion, there are some immaterial differences between the data announced at the time of the trial termination, or presented at subsequent scientific meetings, and the final data presented here

ShORe Phase 3 Trial Did Not Achieve Primary Endpoint

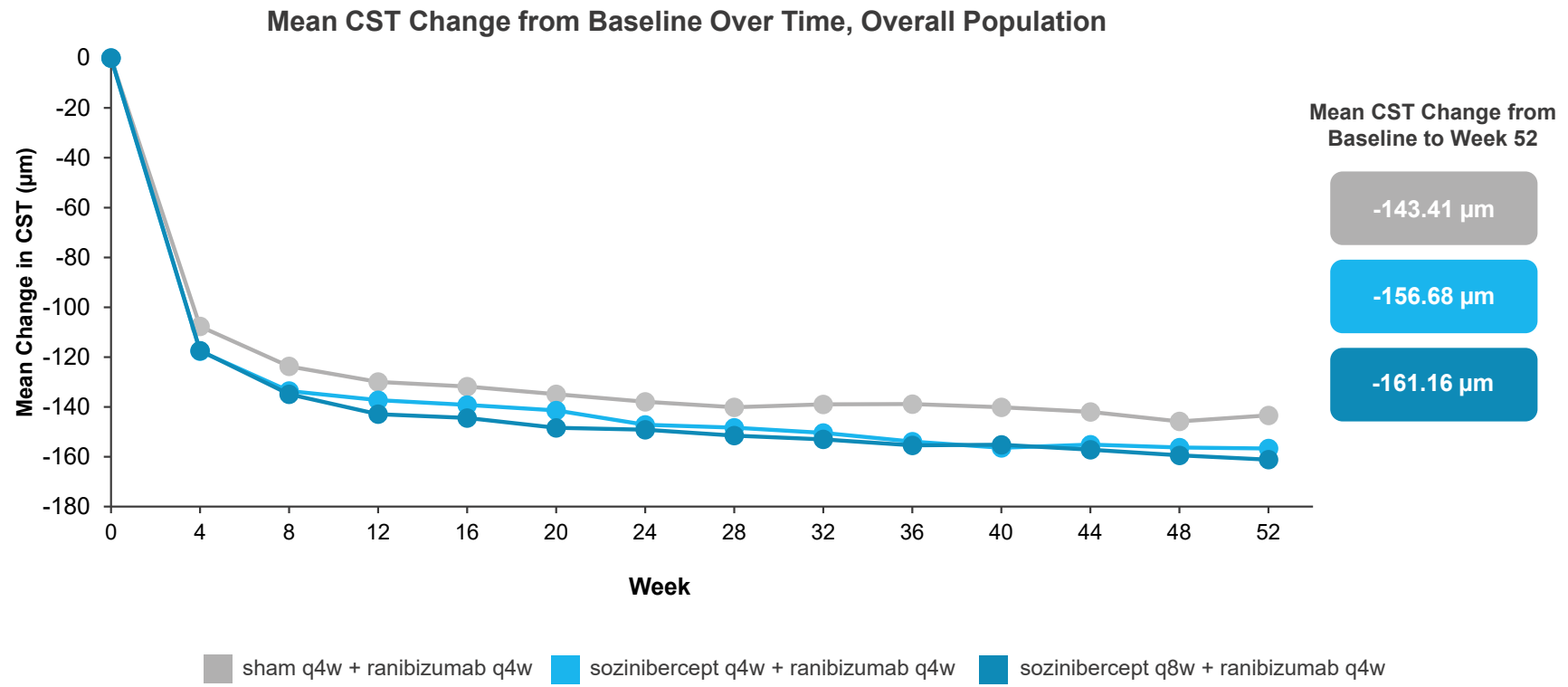
ShORe (Sozinibercept in Combination with Ranibizumab) Unmasked Early Following COAST Results



BCVA: Best Corrected Visual Acuity

Because the ShORe trial was unmasked before completion, there are some immaterial differences between the data announced at the time of the trial termination, or presented at subsequent scientific meetings, and the final data presented here

ShORe Trial: CST Reductions Observed in All Treatment Arms



CST: central subfield thickness
ShORe trial unmasked early following negative COAST results. ShORe CST results are not validated, 95% CI not available.

COAST and ShORe Trials Key Takeaways

- The COAST and ShORe Phase 3 trials did not meet the primary endpoint of superiority in vision improvement from baseline to week 52
- No statistically significant differences in key anatomical outcomes were observed between treatment arms
- Sozinibercept combination therapy was well tolerated
- In this large global Phase 3 program, VEGF-C and VEGF-D inhibition did not provide additional functional benefit beyond standard of care anti-VEGF-A therapy in wet AMD patients

Update on Development Funding Agreement (DFA) Negotiations

Tom Reilly

Original terms of the DFA

- In August 2022, Opthea entered into a Development Funding Agreement (DFA) with Ocelot SPV LP which provided USD\$120M of funding to support our development of sozinibercept for the treatment of wet AMD
- In December 2023, Opthea entered into an Amended and Restated DFA with Ocelot as collateral agent, pursuant to which a new co-investor provided an additional USD\$50 million in funding, bringing the total funding to USD\$170 million
- If sozinibercept was approved, repayment to the DFA investors was 4x investment (USD\$680M)
- Under the DFA, the DFA investors held security over the assets of Opthea in the form of an "all assets" lien
- There are termination clauses in the DFA which involve repayments ranging from USD\$0M up to USD\$680M

Successful DFA Negotiations Allowed The Company to Remain Solvent Which Is a Better Outcome For Shareholders

- Following the negative topline data of the COAST trial, Opthea's management and Board of Directors began discussions with the DFA investors related to the next steps of the Sozinibercept Wet AMD program
- In consultation with the DFA investors, Opthea determined the most appropriate course of action for wet AMD patients, Opthea shareholders, and other stakeholders was to accelerate the timing of the ShORE trial topline data
- Following the negative topline data of the COAST and ShORE trial, Opthea in consultation with the DFA investors determined to terminate the Sozinibercept wet AMD program. Opthea determined that this was in the best interests of Opthea shareholders, including to preserve cash
- Since these negative Phase 3 results, Opthea has reduced the work force by over 80%, reduced the Board of Directors by 50%, renegotiated all contracts related to the clinical trials & had active discussions with the DFA investors to settle the DFA arrangements
- As announced on August 18th/August 19th (Australia), Opthea has agreed to settle with the DFA investors with a cash payment of USD20M & 9.99% equity stake in the Company (equivalent to 136.7M common shares). These settlement arrangements include termination of the DFA and all liens⁽¹⁾
- As of August 19th, 2025 (after payment of USD20M to the DFA investors) the company has cash of approximately USD20M and after issue of the new shares to the DFA investors ~1.4B common shares outstanding

⁽¹⁾ For further detail regarding the material terms of the Settlement Agreement and Subscription Deed, see the Company's ASX Announcement dated August 19th, 2025

Current State of the Company

Dr Jeremy Levin

Successful DFA Negotiations Resulted in Positive Outcome for Shareholders

- Opthea settled arrangements with the DFA investors removing financial uncertainty
- Opthea retains meaningful assets:
 - Significant cash position
 - No debt
 - No lien on any asset ⁽¹⁾
 - Dual-listing on ASX and NASDAQ. Trading in Opthea's listed securities remains suspended by ASX under ASX Listing Rule 17.3. Opthea is currently engaging with the ASX regarding these matters.
 - Clinical, preclinical, scientific knowledge and assets and IP related to VEGF-C and VEGF-D
 - Existing API and materials to allow potential new investigations
- Streamlining of operations has been achieved. Only 3 employees to remain as of September 15th
- Board of Directors was reduced by half to four directors. Sujal Shah to step down as of September 15th.
- As announced to the market on August 19th, given settlement of the DFA arrangements, Opthea is no longer relying on the 'safe harbour' provisions in section 588GA of the *Corporations Act 2001* (Cth).

⁽¹⁾ For further detail regarding the material terms of the Settlement Agreement and Subscription Deed, see the Company's ASX Announcement dated August 19th 2025

The Path Forward

Dr Jeremy Levin

Leadership and Governance

Board of Directors:

- Dr. Jeremy Levin
- Kathy Connell
- Lawrence Gozlan
- Sujal Shah, who will step down as of September 15th, 2025

Planned Executive Departures:

- Fred Guerard, CEO – stepping down effective September 1st, 2025
- Tom Reilly, CFO – stepping down effective September 15th, 2025
- Karen Adams, Corporate Secretary – stepping down effective November 1, 2025

Management: We have implemented a streamlined, cost-efficient structure that aligns with the Company's current scale and strategic priorities. This includes active engagement of our Board, whose deep expertise across areas including science, business development, finance, commercial operations, and investment is being fully leveraged to ensure effective governance, operational oversight



Strategy To Maximize Shareholder Return

The Board will continue to focus on maximizing shareholder value and will assess the following:

- Full strategic review over the next six months
- Targeted internal development
- Strategic partnerships or potential BD/licensing, where appropriate
- Return of capital to shareholders, where appropriate



Executing with Focus and Accountability

Strategic Transition: Clear Priorities, Steady Progress

- A comprehensive business and asset review is actively underway
- The Board
 - Is focused on delivering long-term shareholder value
 - Will provide additional support to the company during these transitional stages
 - Expects to provide shareholders with a further update in CY Q4

Thank you for your continued trust and support

Q&A



Thank you

