

Redefining the Management of nAMD

The Impact of Durability and Sustained Disease Control

August 29, 2026

Asia-Pacific Vitreo-retina Society Congress | Gold Coast, Australia



Forward-Looking Statements

Any statements in this presentation about future expectations, plans, and prospects for the Company, including statements regarding the development and regulatory status of the Company's or its competitors' product candidates, including statements regarding the SOL-1 Phase 3 superiority trial being conducted under a Special Protocol Assessment (SPA) agreement with the U.S. FDA and aligned with FDA draft guidance for neovascular AMD drug development; the timing, design, enrollment, randomization, conduct and retention of subjects in the Company's clinical trials, including the Company's SOL-1 trial and its SOL-R Phase 3 non-inferiority trial and SOL-X open-label extension trial of OTX-TKI for the treatment of wet age-related macular degeneration (wet AMD), also known as neovascular age-related macular degeneration (nAMD); statements regarding the potential utility or adoption of OTX-TKI or its competitors, if approved; statements regarding the Company's intention to submit a new drug application for OTX-TKI, based on data from the Company's SOL-1 trial in wet AMD, subject to planned formal discussions with the FDA; and other statements containing the words "anticipate", "become", "believe", "estimate", "expect", "intend", "designed", "goal", "may", "might", "plan", "position", "predict", "project", "target", "potential", "will", "would", "could", "should", "continue", and similar expressions, constitute forward-looking statements within the meaning of The Private Securities Litigation Reform Act of 1995. Actual results may differ materially from those indicated by such forward-looking statements as a result of various important factors. Such forward-looking statements involve substantial risks and uncertainties that could cause the Company's development programs, future results, performance, or achievements to differ significantly from those expressed or implied by the forward-looking statements. Such risks and uncertainties include, among others, the design, timing, conduct and outcomes of ongoing and planned clinical trials, including the SOL-R trial, the second year of the SOL-1 trial and the SOL-X trial; the risk that the FDA will not agree with the Company's interpretation of the written agreements under the Special Protocol Assessments for OTX-TKI, including for the SOL-1 trial; uncertainty as to whether the FDA will accept a new drug application for OTX-TKI on the basis of a single pivotal clinical trial, namely SOL-1; uncertainty as to the minimum clinical data required to demonstrate the safety of a proposed product candidate such as OTX-TKI, even if the FDA recognizes that only one pivotal clinical trial may be required to demonstrate efficacy; the risk that even though the FDA has agreed with the overall design of the SOL-1 trial, the FDA may not find that the data generated by the trial and submitted by the Company are sufficient to demonstrate the safety and efficacy of OTX-TKI to the degree necessary to support marketing approval for wet AMD; uncertainty as to whether the Company will be able to timely satisfy the FDA's other requirements for regulatory approval of OTX-TKI, including the FDA's Chemistry, Manufacturing and Control's requirements, even if the Company can satisfy the FDA's clinical requirements to demonstrate safety and efficacy; uncertainty as to what restrictions, if any, may be imposed on the label for OTX-TKI, if approved, pending the receipt of additional clinical data or otherwise; the risk that the FDA might not agree to the Company's design, protocol, and statistical analysis plan of the SOL-R trial; the risk that the Company and the FDA may not agree on the registrational pathway for any of its product candidates, including OTX-TKI; uncertainty as to whether the data from earlier clinical trials will be predictive of the data of later clinical trials, particularly later clinical trials that have a different design or utilize a different formulation than the earlier trials; uncertainty as to whether preliminary or interim data from a clinical trial will be predictive of final data from such trial; uncertainty as to whether data from the Company's SOL-X trial will demonstrate additional clinically meaningful, long-term benefits; uncertainties regarding the potential commercial advantages and/or position of the Company's product candidates; availability of data from clinical trials and expectations for regulatory submissions and approvals; the Company's scientific approach and general development progress; uncertainties inherent in translating, or in the applicability of, laboratory results to clinical applications; uncertainties inherent in estimating the Company's cash runway, future expenses and other financial results, including its ability to fund future operations, including clinical trials; and other factors discussed in the "Risk Factors" section contained in the Company's quarterly and annual reports on file with the Securities and Exchange Commission. In addition, the forward-looking statements included in this presentation represent the Company's views as of the date of this presentation. The Company anticipates that subsequent events and developments may cause the Company's views to change. However, while the Company may elect to update these forward-looking statements at some point in the future, the Company specifically disclaims any obligation to do so, whether as a result of new information, future events or otherwise, except as required by law. These forward-looking statements should not be relied upon as representing the Company's views as of any date subsequent to the date of this presentation.

Disclaimer

The following presentation discusses OTX-TKI - an investigational product candidate that has not been approved by the FDA or any other regulatory body; safety and effectiveness have not been established

Faculty



Prof. Adrian Fung
MMBS (Hons1), MMed
(Ophthal Sci), MMed
(Clin Epi), FRANZCO

Clinical Professor at the
Macquarie University
Hospital and Clinical
Associate Professor at the
University of Sydney,
Australia



Prof. Adnan Tufail
MBBS, MD, FRCOphth

Consultant Ophthalmic
Surgeon at Moorfields Eye
Hospital, London, UK



Prof. Darius Moshfeghi
MD, FASRS

Chief of the Retina Division and
Professor of Ophthalmology at the
Stanford University School of
Medicine
Palo Alto, CA, USA



Dr. Jeffrey Heier
MD, FASRS

Chief Scientific Officer of
Ocular Therapeutix, Inc.,
Bedford, MA, USA



Prof. Peter K. Kaiser
MD, FASRS

Chief Development Officer of
Ocular Therapeutix, Inc.,
Bedford, MA, USA

Chaney Family Endowed
Chair in Ophthalmology
Research and Professor of
Ophthalmology, Cleveland
Clinic Lerner College of
Medicine

Disclosures

Adrian Fung

- **Consultant/Advisor:** Alcon, Apellis, Astellas, Bayer, Roche, Samsara, Zeiss
- **Lecture honoraria:** 4DMT, AbbVie, Alcon, Bayer, Novartis, Ocular Therapeutix, Roche
- **Research support:** AbbVie, Bayer, Ionis Pharmaceuticals, Roche
- **Travel support:** AbbVie, Bayer, Novartis
- **Equity:** Macuject, Opthea

Adnan Tufail

- **Consultant:** 4DMT, Adverum, Annexon, Apellis, Aviceda, Boehringer Ingelheim, Eyebio, Astellas, Janssen, Kanghong, Nanoscope, Novartis, OcuTerra, Ocular Therapeutix, Regenxbio, Roche/Genentech
- **Stockholder:** Oculogics, Cellio
- **Research support:** Novartis, Bayer

Darius Moshfeghi

- **Consultant:** Boehringer Ingelheim, Feliqs, Plenoptika, Vision Labs AI
- **Stockholder:** Ainsly Limited, Plenoptika, Vision Labs AI
- **Research Support:** Ingel Therapeutics, Ocular Therapeutix, Osanni Bio

Jeffrey Heier

- **Chief Scientific Officer for Ocular Therapeutix**
- **Consultant:** Aavantgarde, Abbvie/Regenxbio, Alzheon, Annexon, Astellas, Aviceda, Bayer, Beacon, Breye Therapeutics, Caeregen, Cogent, Cognition, Complement Therapeutics, Endogena, Focus Biosciences, Frontera, Galimedix, Inflammx, Kaigene, Kanghong, Lilly, Manistee, Nanoscope, Notal Vision, Novartis, Ocugen, Ocuphire, Ora, Inc., Osanni Bio, Perceive Biotherapeutics, Ray Therapeutics, Samsung Bioepis, Sanofi, Stealth Biotherapeutics, Laboratoires Thea, Vanotech, Visgenx
- **Research support:** 4DMT, Abbvie/Regenxbio, Annexon, Apellis, Astellas, Bayer, Beacon, Boehringer Ingelheim, Cognition Therapeutics, Curacle, Janssen R&D, Kodiak, Notal Vision, Novartis, Oculis, Perceive Bio, Sanofi, Skyline, Stealth Biotherapeutics, Vanotech
- **Stock/Stock options:** Adverum, Aldeyra, Alzheon, Aviceda, Caeregen, Inflammx, jCyte, Ocular Therapeutix, Osanni Bio, Ray Therapeutics, RevOpsis, Vinci, Visgenx, Vitranu

Peter K. Kaiser

- **Chief Development Officer for Ocular Therapeutix**
- **Consultant:** Abbvie, Alexion, Alkeus, Allgenesis, Alzheon, Amaros, Annexon Biosciences, Astellas, Augen Therapeutics, Aviceda, Bayer, Biogen Idec, Carl Zeiss Meditec, Celltrion Healthcare Co., Cognition Therapeutics, Complement Therapeutics, Eli Lilly, Endogena Therapeutics, Eyestem, Frontera Therapeutics, Galimedix, Innovent, Invirsa, Isarna, Janssen, jCyte, Kanaph Therapeutics, Kanghong, Kera Therapeutics, Kriya Therapeutics, Movu/Santec, Nanoscope Therapeutics, Ocugenix, Oculis, Omeros, Ophthalmx, Osanni Bio, Panther Pharmaceuticals, Ray Therapeutics, RegenxBio, Resonance Medicine Inc., Restore Vision, Retinal Sciences, ReVana, RevOpsis, Roivant, Samsung Bioepis, Sandoz, SGN Nanopharma Inc., SmileBiotech Zhuhai Ltd, Stealth Biotherapeutics, Stuart, Sudo Biosciences, Sustained Nano Systems, Théa, Tilak, Vanotech, VisgenX
- **Stock/Stock Options:** AAVAntgarde Bio, Alzheon, jCyte, Ocular Therapeutix, Oculis, Osanni Bio, Ray Therapeutics, RevOpsis, Vitranu
- **Board of Directors:** AAVAntgarde Bio

Agenda

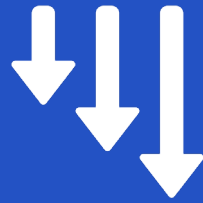
- **The Unmet Need**
Adrian Fung
- **Science Behind the Therapy**
Peter K. Kaiser
- **SOL-1 Results**
Darius M. Moshfeghi

- **SOL-1 Clinical Evidence**
Adrian Fung
- **Key Insights**
Adnan Tufail
- **Clinical Program Update**
Jeffrey S. Heier

The Unmet Need

Adrian Fung, MD

The Unmet Needs of Intravitreal Therapy for nAMD



**Reduced
Treatment
Burden¹**



**Better
Visual
Outcomes²**

Neovascular AMD Prevalence: Asia & Australia



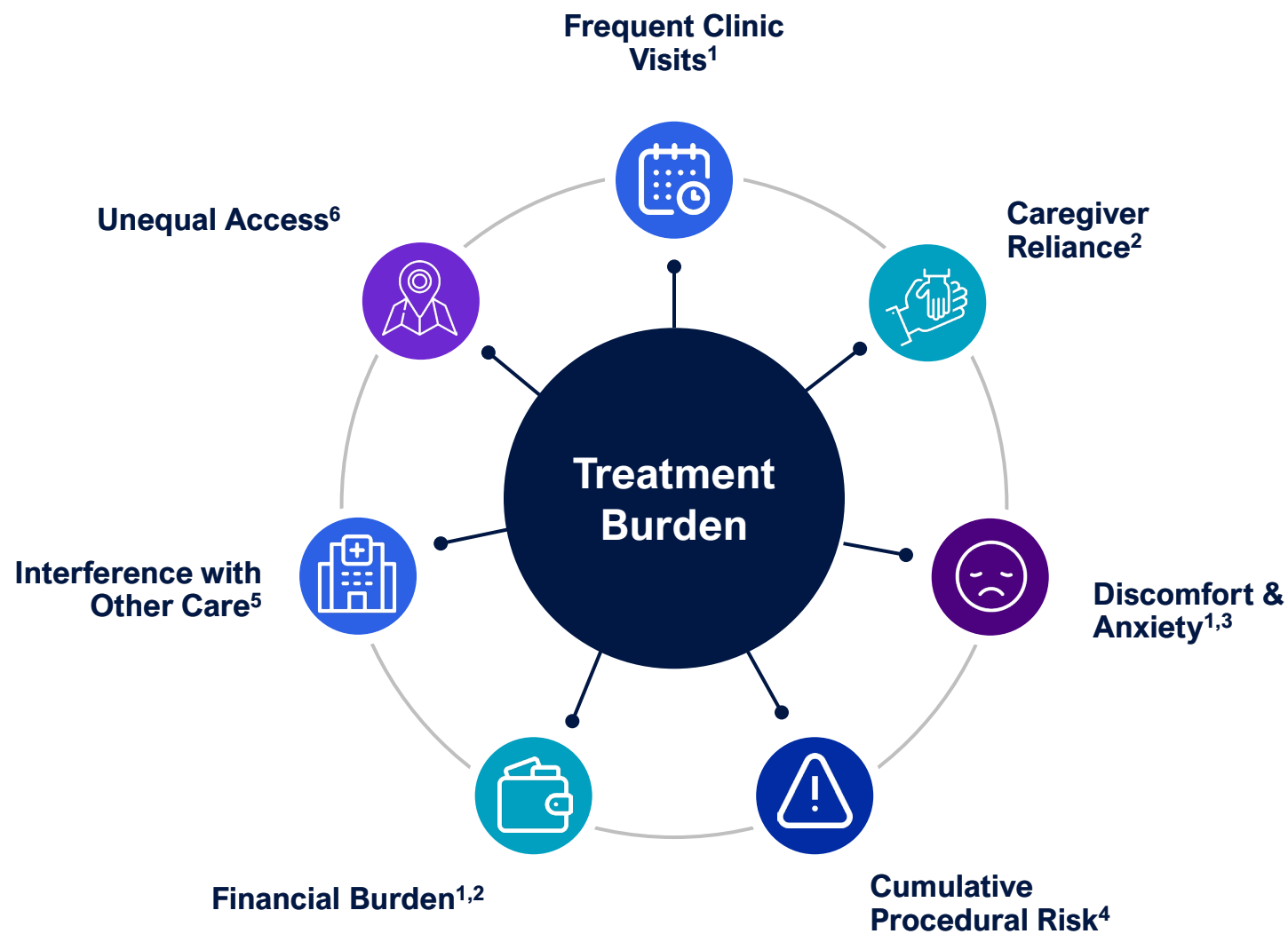
Asia

~8 million^{1,2}

Australia

~160,000³

Treatment Burden of Anti-VEGF Therapy



Australian Approved Anti-VEGF Dosing Intervals in Neovascular AMD

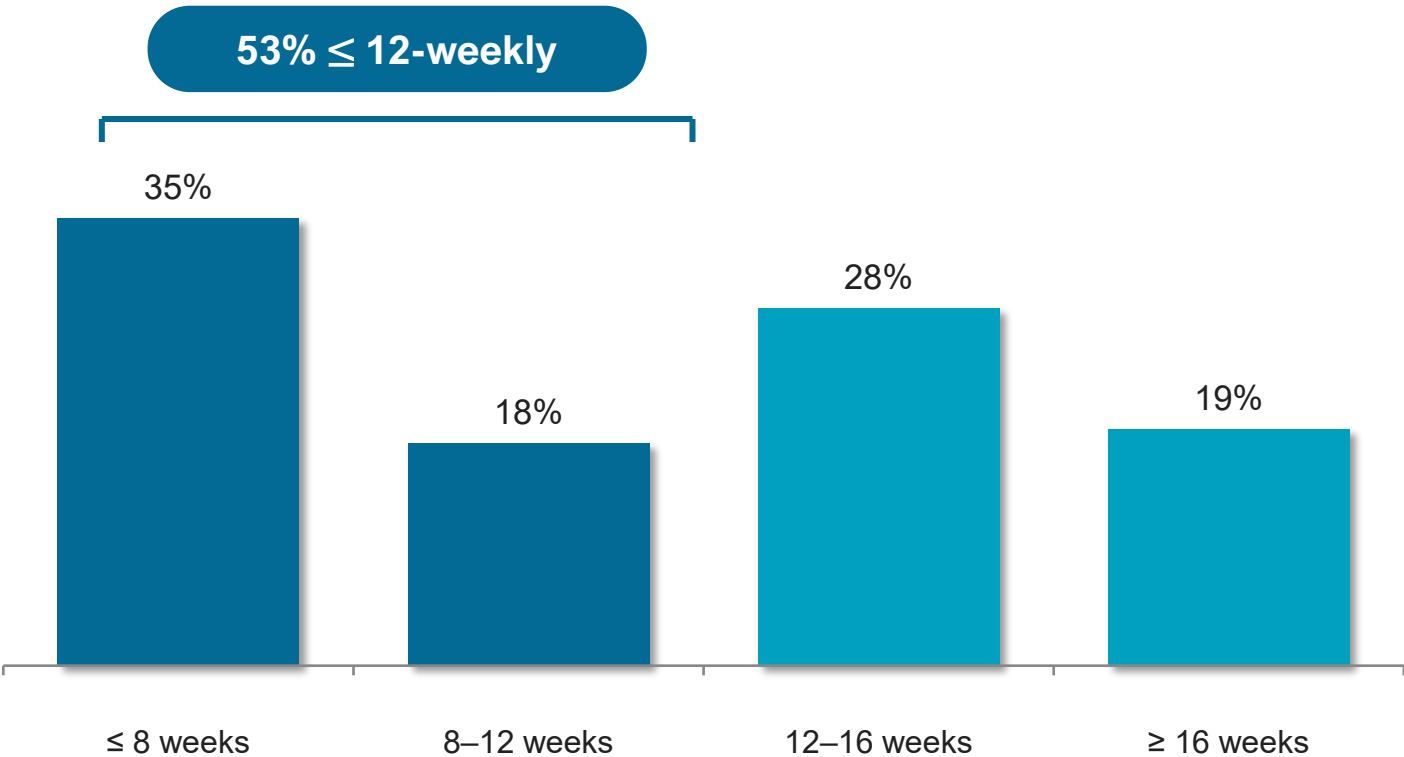
Approved maintenance interval following initial monthly loading doses



But in the Real-World We Inject More Frequently...

Fight Retinal Blindness! Registry Injection Intervals in Neovascular AMD

Percentage of Treatment-naïve Eyes by Injection Interval at 24 Months^{1*}
(First-generation Anti-VEGF: Ranibizumab / Aflibercept 2 mg)



Faricimab Treatment Naïve (12 mo)^{2†}

52% on
≤12 week dosing

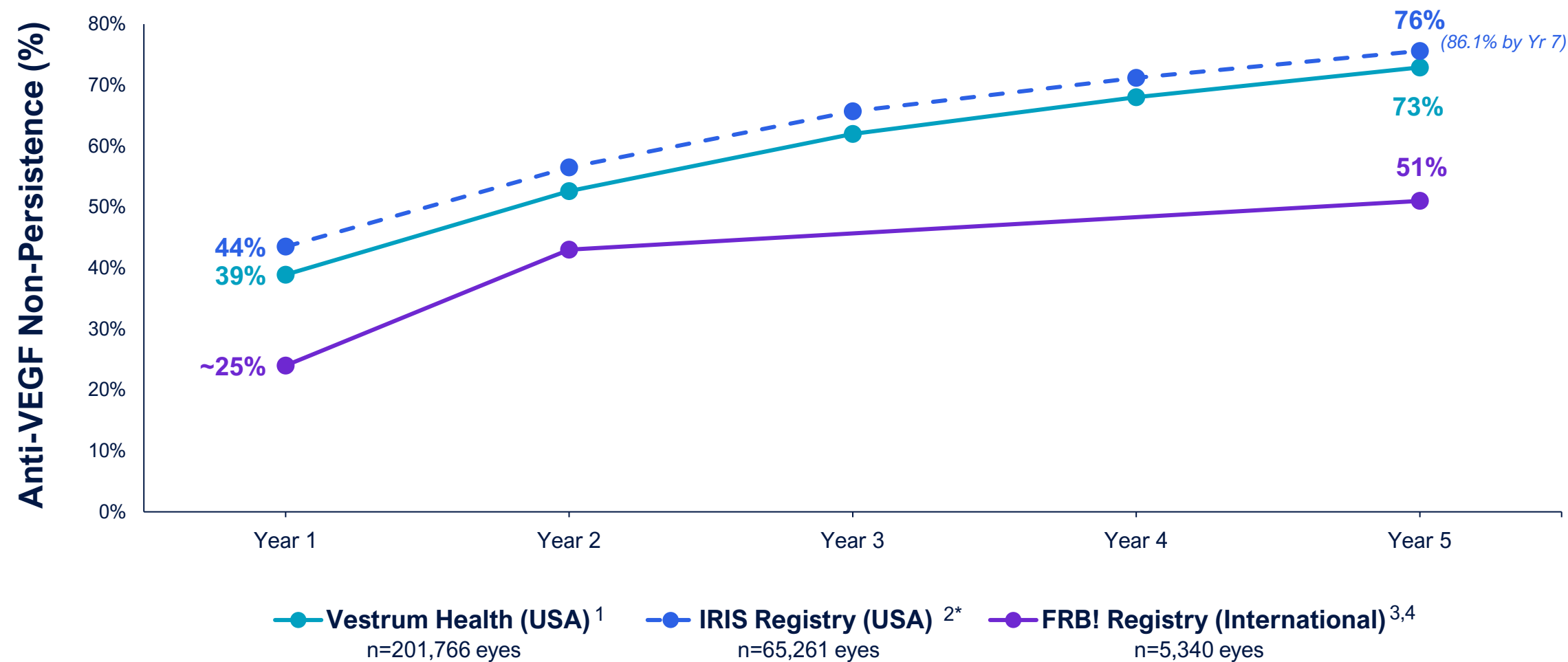
Faricimab Switch Cohort (12 mo)^{3‡}

7.2 → 10.5 weeks

^{*}National retrospective FRB Spain study of treatment-naïve nAMD eyes treated with ranibizumab or aflibercept 2 mg. The 24-month distribution reflects 557 eyes meeting additional treat-and-extend-range criteria and their final observed injection interval.
[†]Retrospective Australian FRB registry study of 160 treatment-naïve nAMD eyes initiating faricimab across 23 practitioners and followed for 12 months. Outcomes were compared with historical aflibercept 2 mg controls using overlap weighting; 48% of faricimab-treated eyes attained a dosing interval of ≥12 weeks.
[‡] Retrospective Australian FRB registry study of previously treated nAMD eyes switched—primarily from aflibercept 2 mg—to faricimab and followed for 12 months. The 7.2-to-10.5-week result represents the cohort-level change in mean injection interval before versus after switching.
1. Zarranz-Ventura J, et al. *Eye (Lond)*. 2025;39(18):3306-3313. 2. Gillies M, et al. *Clin Exp Ophthalmol*. 2026;54(4):500-506. 3. Hunt A, et al. *Clin Exp Ophthalmol*. 2025;53(9):1156-1167.
AMD, age-related macular degeneration; FRB, Fight Retinal Blindness; nAMD, Neovascular AMD; VEGF, vascular endothelial growth factor;

Real-World Anti-VEGF Non-Persistence Increases Over Time

Cumulative non-persistence over time – three independent real-world registries/databases



*Includes eyes that experienced a treatment gap (≥ 180 days gap with restart of treatment) or treatment discontinuation (≥ 180 days without injections, not lost to follow up)

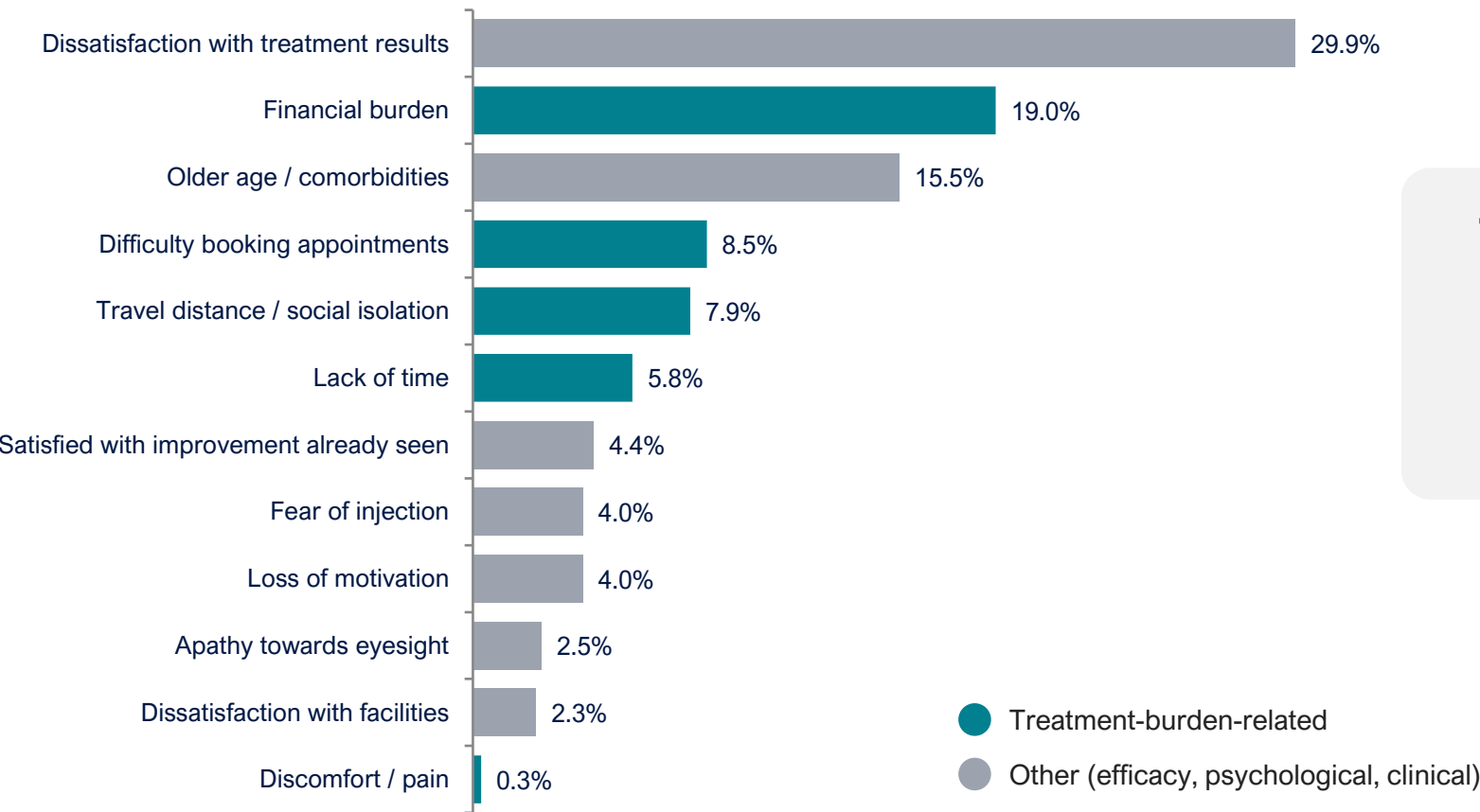
1. Moshfeghi AA, et al. Real-world anti-VEGF treatment utilization and vision outcomes in neovascular age-related macular degeneration: analysis of the Vestrum database. Poster presented at: Association for Research in Vision and Ophthalmology Annual Meeting; May 7, 2026; Denver, CO. 2. Weng CY, et al. Anti-VEGF treatment patterns and long-term vision outcomes in patients with nAMD in the Academy IRIS Registry. Poster presented at: Association for Research in Vision and Ophthalmology Annual Meeting; May 7, 2026; Denver, CO.. 3. Teo KYC, et al. *Eye (Lond)*. 2023;37:467-473. 4. Bhandari S, et al. *Eye (Lond)*. 2023;37:1145-1154.

FRB, Fighting Retinal Blindness; VEGF, vascular endothelial growth factor

Many Patients Stop Due to Treatment Burden

Patient-Reported Reasons for Non-Adherence/Non-Persistence

Meta Analysis of 22 Studies (N=409,215 patients)

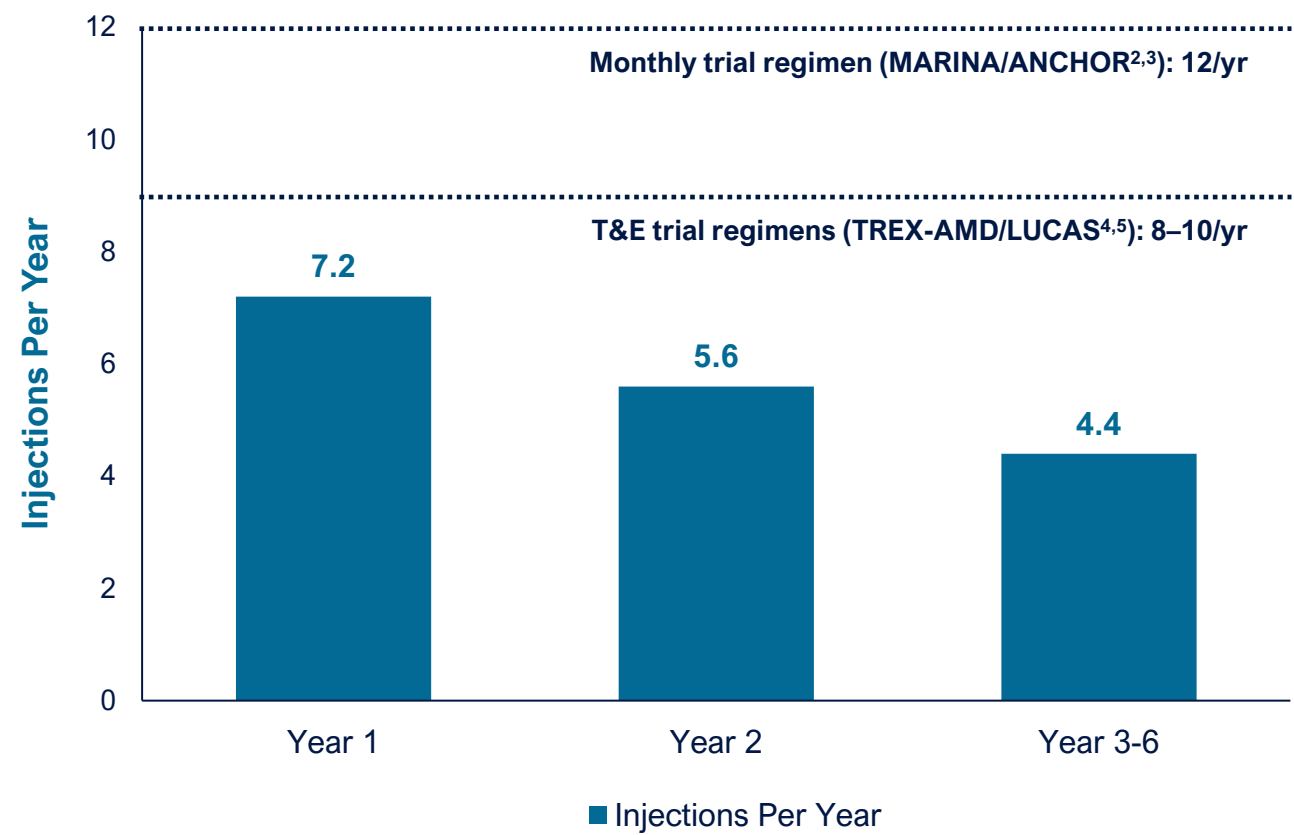


Treatment burden-related reasons combined

~41%

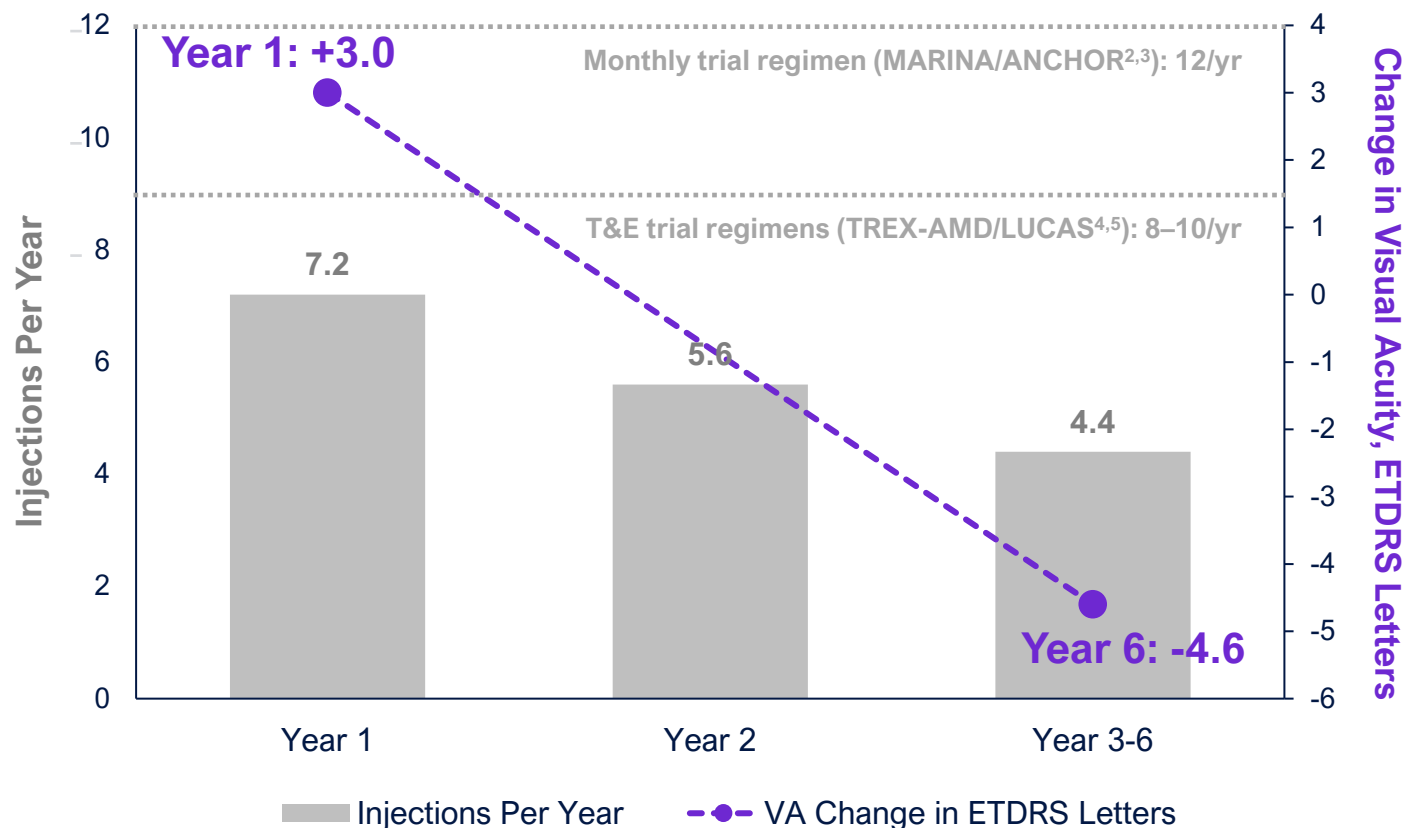
...Or Are Under-Treated, Which Leads to Worse Visual Outcomes

Mean Anti-VEGF Injections per Year:
Real-World (IRIS Registry¹) vs. Trial Benchmarks



...Or Are Under-Treated, Which Leads to Worse Visual Outcomes

Mean Anti-VEGF Injections per Year: Real-World (IRIS Registry¹) vs. Trial Benchmarks



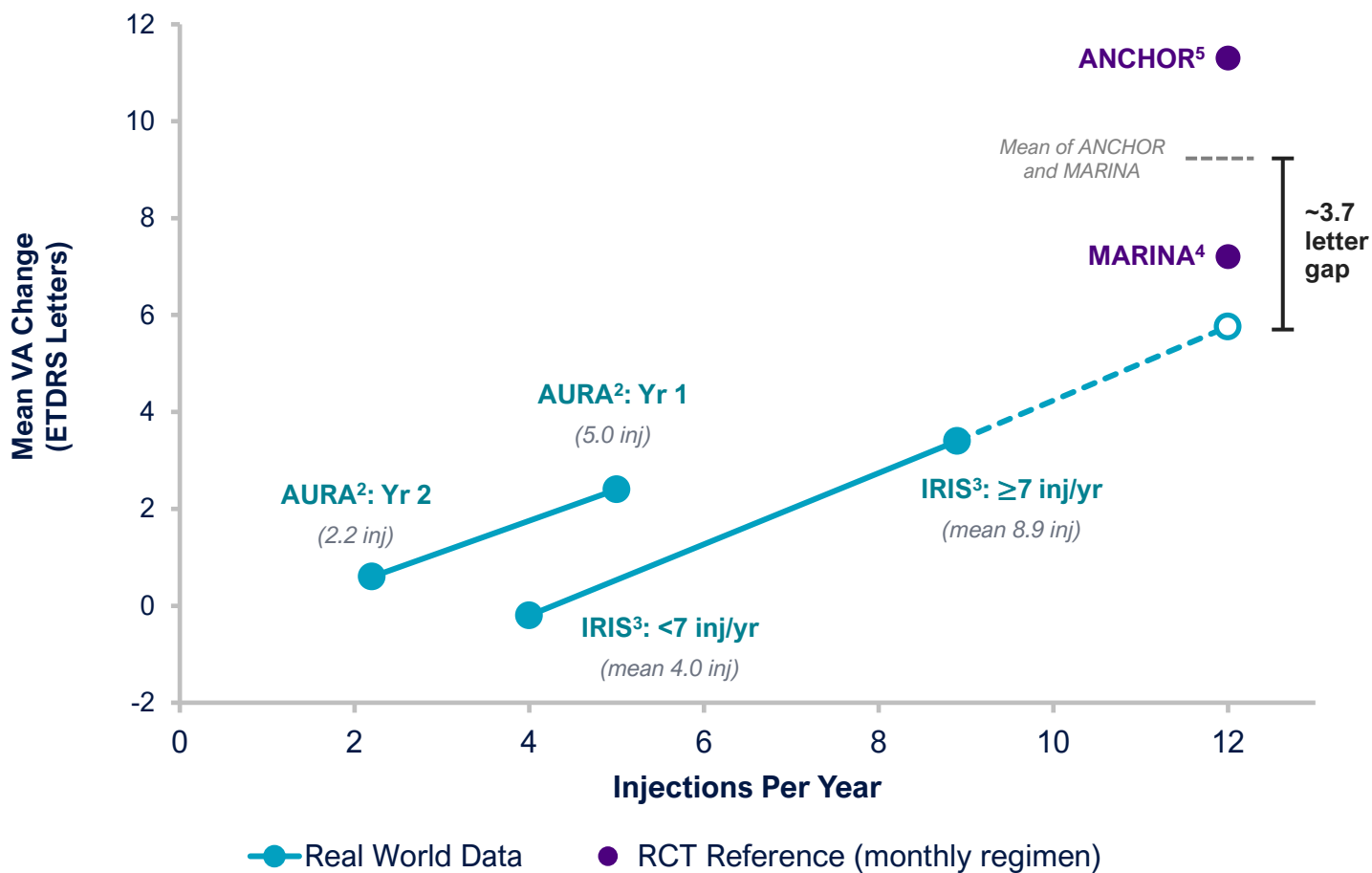
Same IRIS cohort: VA trajectory over time¹

+3.0 → -4.6 letters

Mean VA change from baseline: a +3.0 letter gain at Year 1 erodes to a net loss of 4.6 letters by Year 6

Each Additional Year-1 Injection Was Associated With a 0.68-Letter VA Gain

More injections per year is associated with better mean visual acuity outcomes



Adjusted Regression Estimate¹

+0.68 letters

Mean VA gain per additional anti-VEGF injection in Year 1, adjusted for baseline differences

1. Wyckoff CC, et al. *Ophthalmol Sci.* 2024;4(2):100421. 2. Holz FG, et al. *Br J Ophthalmol.* 2015;99(2):220-226. 3. Barikian A, et al. *Ophthalmol Retina.* 2026;10(1):71-80. 4. Rosenfeld PJ, et al. *N Engl J Med.* 2006;355(14):1419-1431. 5. Brown DM, et al. *N Engl J Med.* 2006;355(14):1432-1444.
ETDRS, Early Treatment Diabetic Retinopathy Study; VA, visual acuity; RCT, randomized controlled trial; VEGF, vascular endothelial growth factor

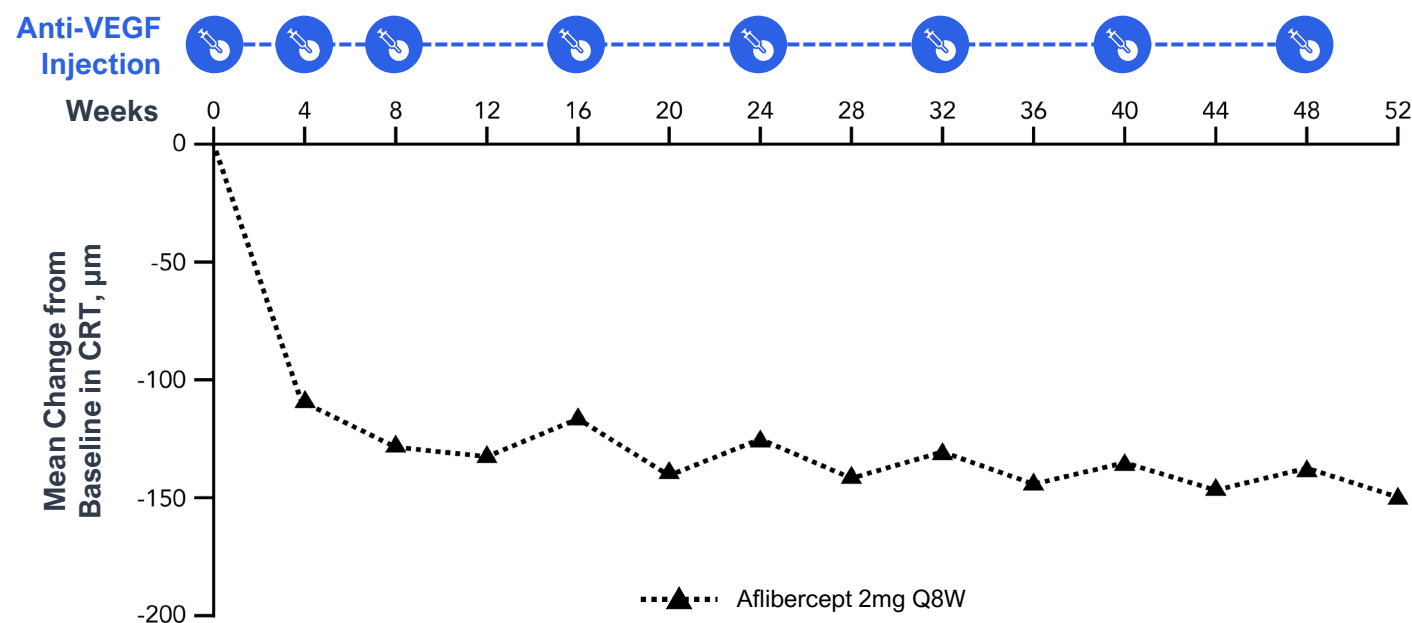
Greater Retinal Thickness Variability During Anti-VEGF Therapy is Associated with Suboptimal Anatomic and Visual Outcomes

Retinal Thickness
Fluctuations

Development of
Fibrosis and Atrophy

Poorer Long-Term
Visual Outcomes

Repeated cycles of fluid resolution and recurrence are reflected in fluctuations in retinal thickness over time



Greater Retinal Thickness Variability During Anti-VEGF Therapy is Associated with Suboptimal Anatomic and Visual Outcomes

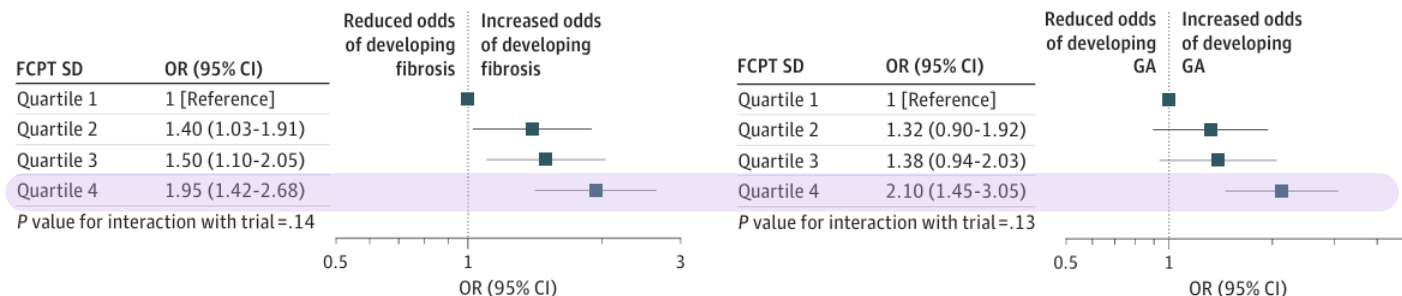
Retinal Thickness
Fluctuations

Development of
Fibrosis and Atrophy

Poorer Long-Term
Visual Outcomes

JAMA Ophthalmology | Original Investigation

Associations of Variation in Retinal Thickness With Visual Acuity and Anatomic Outcomes in Eyes With Neovascular Age-Related Macular Degeneration Lesions Treated With Anti-Vascular Endothelial Growth Factor Agents



In a post hoc analysis, highest retinal thickness variability had approximately **double the odds** of developing fibrosis/macular atrophy

Greater Retinal Thickness Variability During Anti-VEGF Therapy is Associated with Suboptimal Anatomic and Visual Outcomes

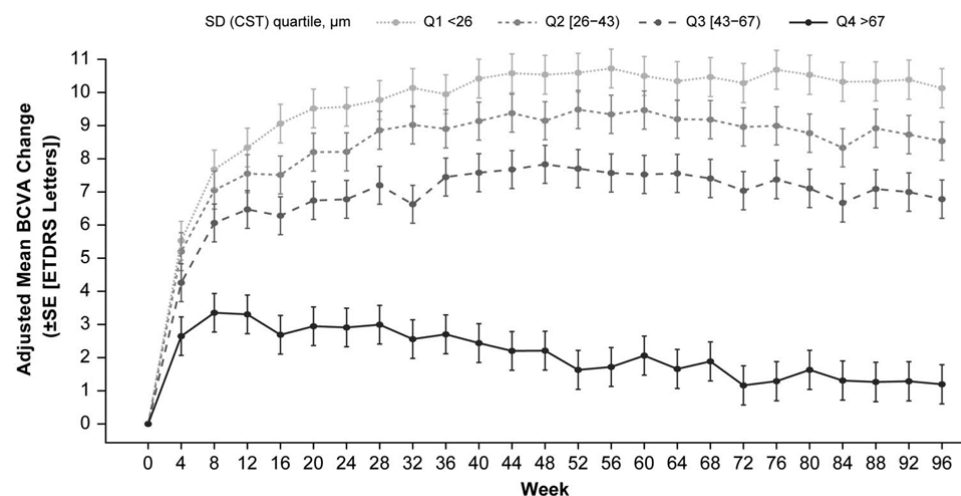
Retinal Thickness
Fluctuations

Development of
Fibrosis and Atrophy

Poorer Long-Term
Visual Outcomes

EFFECT OF RETINAL THICKNESS VARIABILITY ON VISUAL OUTCOMES AND FLUID PERSISTENCE IN NEOVASCULAR AGE-RELATED MACULAR DEGENERATION

A Post Hoc Analysis of the HAWK and HARRIER Studies



Vision deteriorates
by ~9 letters
between patients
with most and least
stable retinal
thicknesses

Two Decades of Innovation Have Improved Durability, But Treatment Burden Persists





2019 | Phase 1 Trial in Australia⁵

OTX-TKI

29 participants, 6 sites

A new approach:

Sustained intraocular delivery⁶



Can durability be extended without sacrificing disease control?

1. Lucentis (ranibizumab) [Australian product information]. Novartis Pharmaceuticals Australia Pty Limited; revised December 4, 2025. Accessed August 14, 2026. https://medsinfo.com.au/api/searchapi/Lucentis_PI?format=pdf 2. EYLEA (aflibercept) [Australian product information]. Bayer Australia Ltd; revised August 7, 2025. Accessed August 14, 2026. <https://www.safetyandquality.gov.au/medicine-finder/eylea> 3. Beovu (brolucizumab) [Australian product information]. Novartis Pharmaceuticals Australia Pty Limited; revised June 19, 2024. Accessed August 14, 2026. https://medsinfo.com.au/api/searchapi/Beovu_PI?format=pdf 4. Vabysmo (faricimab) [Australian product information]. Roche Products Pty Limited; revised September 30, 2024. Accessed August 14, 2026. 5. CLN-0046: Treatment of AMD Subjects With OTX-TKI. ClinicalTrials.gov. NCT03630315. Updated July 10, 2024. Accessed August 19, 2026. <https://clinicaltrials.gov/study/NCT03630315> https://medsinfo.com.au/api/searchapi/Vabysmo_PI?format=pdf 6. Data on File 033. Ocular Therapeutix. AMD, age-related macular degeneration; VEGF, vascular endothelial growth factor

Key Takeaways

Treatment burden is an issue and contributes to discontinuation and under-treatment.¹

Fluid fluctuations are associated with worse vision.^{2,3}

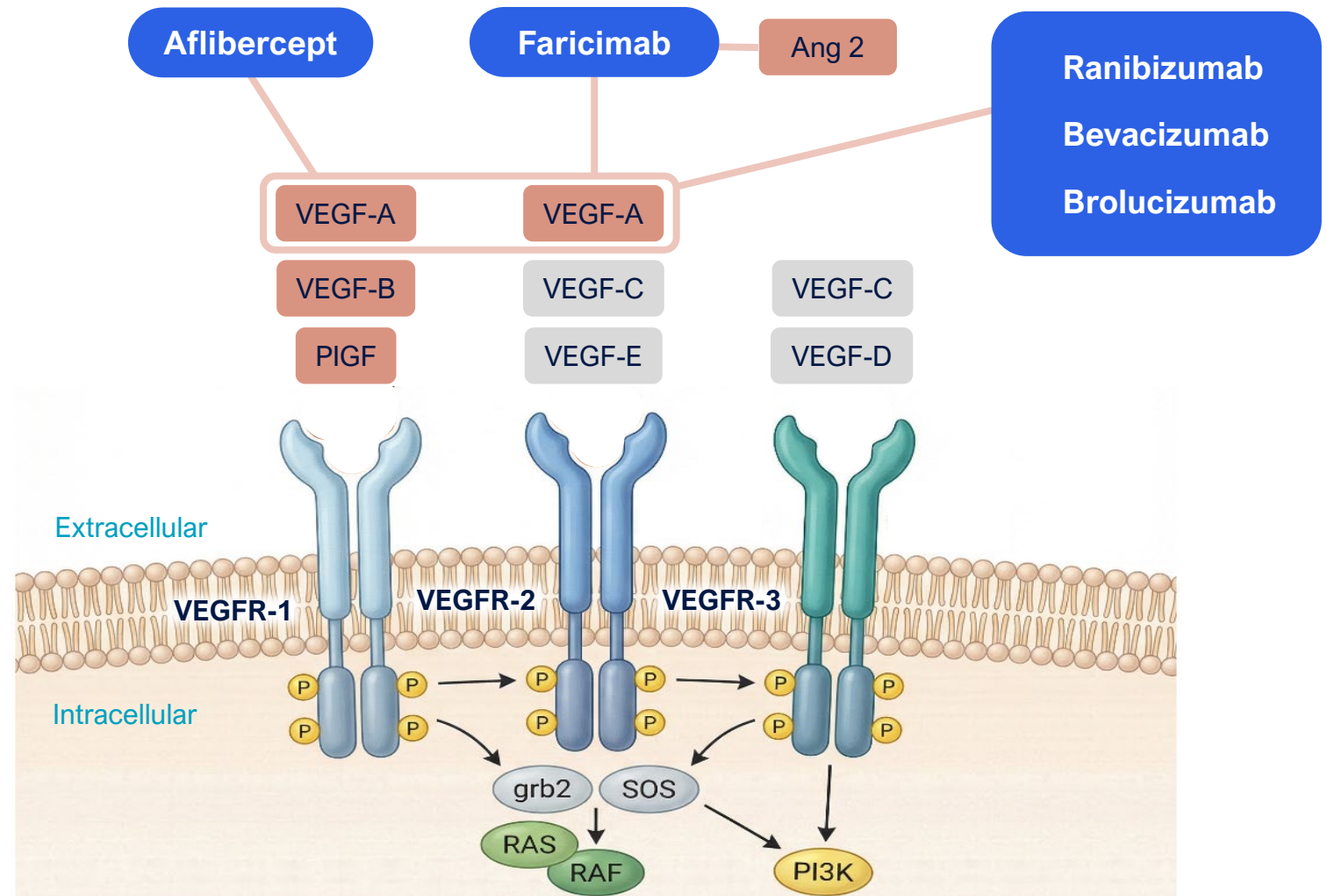
More durable agents don't just reduce visits – they may support better visual outcomes too.³

Science Behind the Therapy

Peter K. Kaiser, MD

Current Anti-VEGF Therapies Selectively Target Only Extracellular VEGF

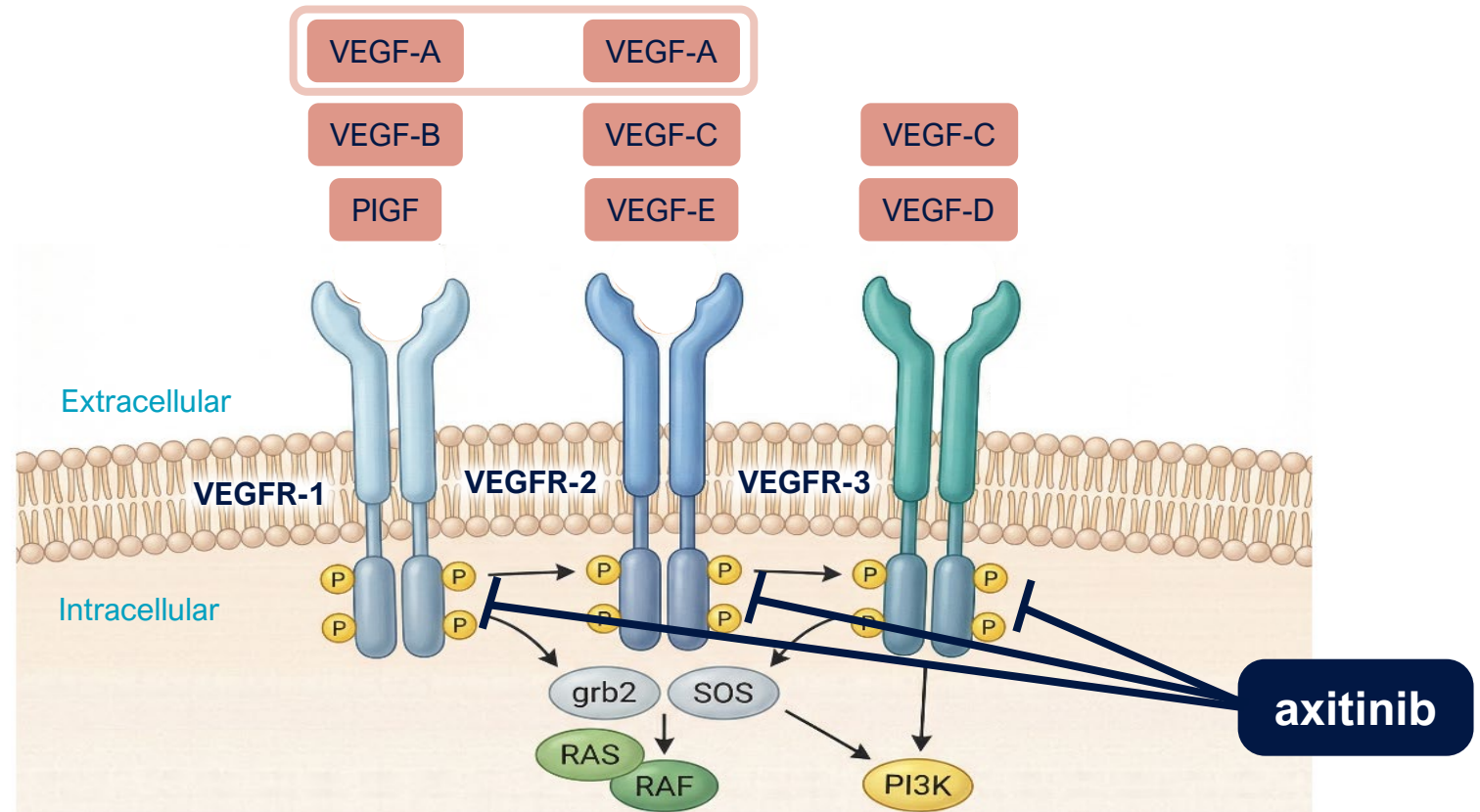
Current **Anti-VEGF** agents act **extracellularly by binding selective ligands**, like VEGF-A, preventing receptor binding and activation^{1,2}



Select extracellular target binding of VEGF, PlGF and Ang-II to inhibit angiogenesis. Adapted from Zhao et al. 2015.^{2,4}

Tyrosine Kinase Inhibitors Act Intracellularly to Inhibit Receptor Signaling

Tyrosine Kinase Inhibitors (TKI) bind at the **intracellular tyrosine kinase domains** of VEGF receptors, inhibiting ATP binding and preventing activation of pro-angiogenic signaling^{1,2}

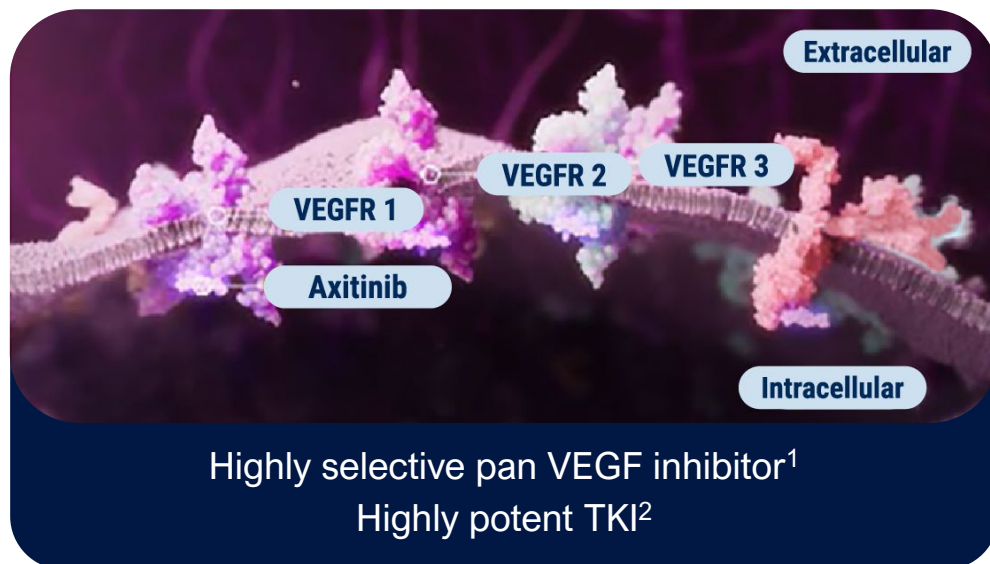


Select angiogenesis intracellular targets of inhibition for tyrosine kinase inhibitors. Adapted from Zhao et al. 2015.²⁻⁴

OTX-TKI: Axitinib via Hydrogel Allows for Tunable Sustained Release

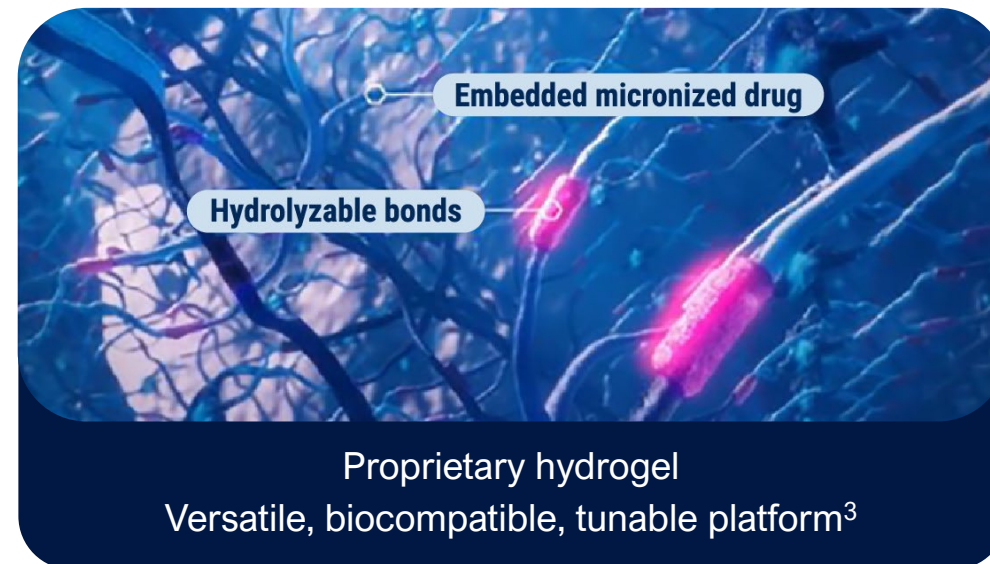
AXITINIB

Multi-target Tyrosine Kinase Inhibitor (TKI)



HYDROGEL TECHNOLOGY

Bioresorbable, Sustained Drug Delivery



+

OTX-TKI

is being developed with the aim to provide:

A single hydrogel insert^{3,4}

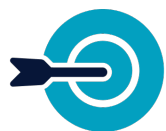
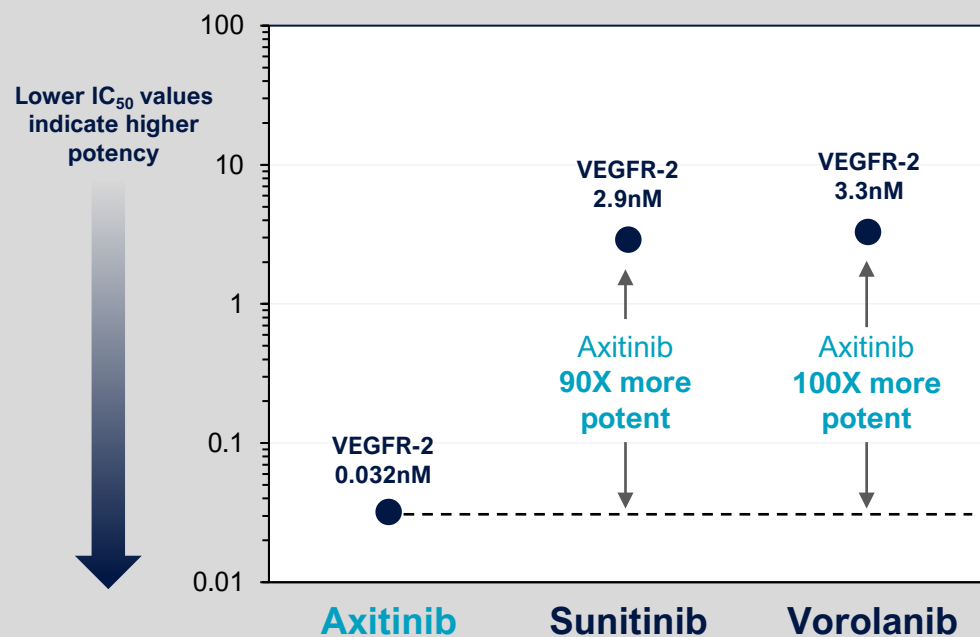
Continuous and consistent
delivery up to 12 months^{3,5}

Complete and predictable
bioresorption^{3,5}

OTX-TKI is an investigational product candidate which has not been approved by the FDA or any regulatory authority.

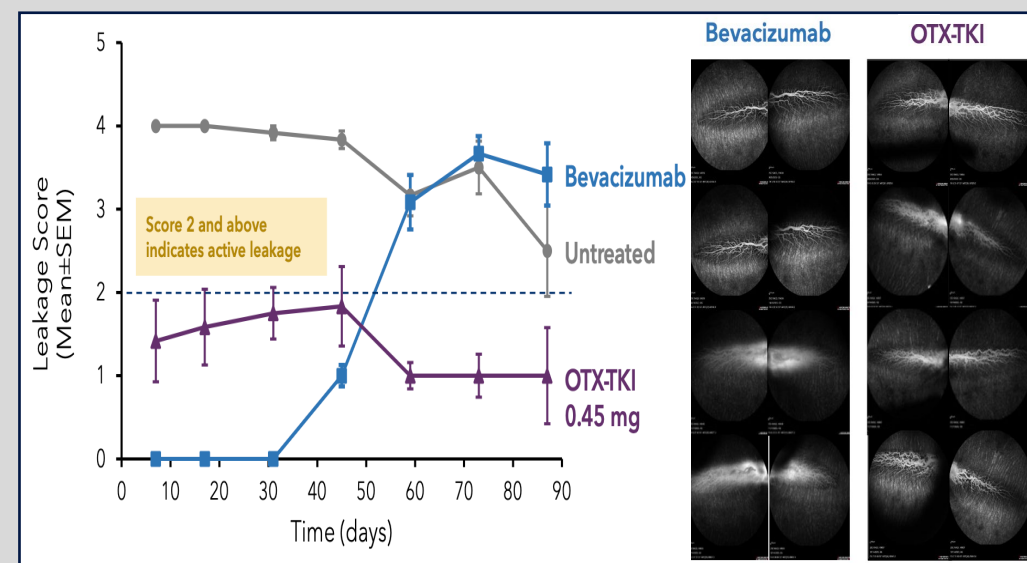
Axitinib: Highly Selective, Very Potent VEGFR Inhibitor

IC₅₀ for VEGF Receptor 2 (nM)¹⁻³



Highly selective for all VEGF receptors, ~100x more potent for VEGFR-2¹⁻³

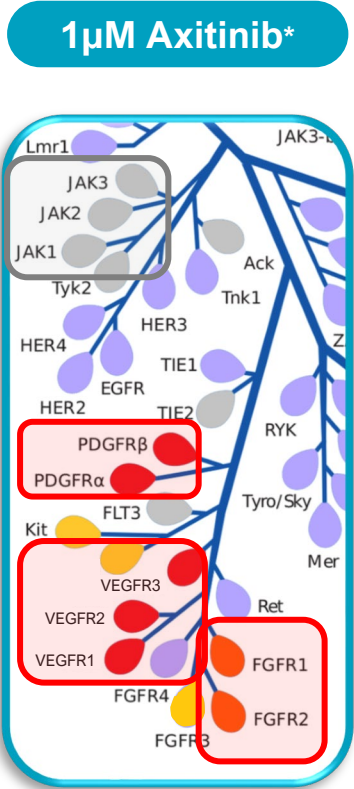
VEGF Challenge Rabbit Model²



Sustained leakage control in repeat VEGF challenge animal model²

Axitinib demonstrates strongest inhibition of all VEGF/PDGF/FGF Receptors

AssayQuant (Fluorescence Assay)*



1 mM ATP

Physiologic Testing Conditions

1 μ M*

Axitinib intraocular conc = 1.03 to 1.13 μ M
Vorolanib intraocular conc = 0.23 to 0.95 μ M^{3,4,6}

Kinase ^{1,2}	Axitinib	Vorolanib ³⁻⁵	Sunitinib
VEGF RECEPTORS			
VEGFR1	99%	39%	83%
VEGFR2	97%	28%	71%
VEGFR3	95%	59%	72%
OTHER RECEPTORS			
PDGFR α	79%	42%	70%
PDGFR β	88%	41%	87%
FGFR1	76%	0%	45%
FGFR2	76%	16%	54%
FGFR3	67%	15%	63%
JAK1	18%	21%	52%

Reaction Biology (Radiometric Assay)**

1 mM ATP

Physiologic Testing Conditions

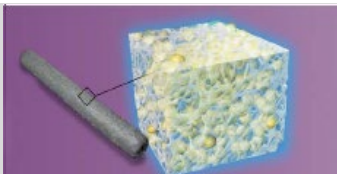
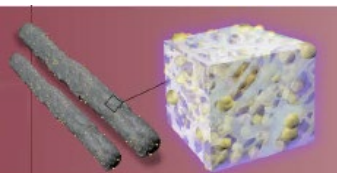
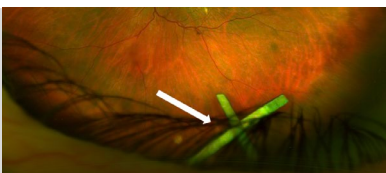
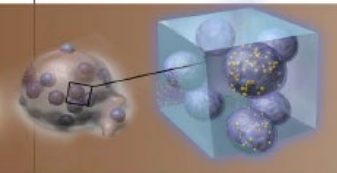
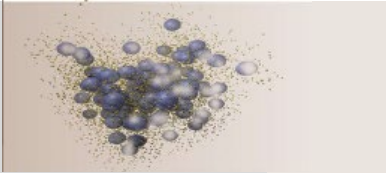
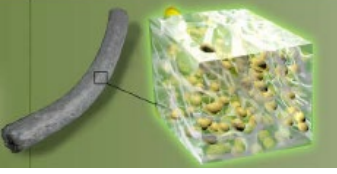
1 μ M*

Axitinib intraocular conc = 1.03 to 1.13 μ M
Vorolanib intraocular conc = 0.23 to 0.95 μ M^{3,4,6}

Kinase ^{1,2}	Axitinib	Vorolanib ³⁻⁵	Sunitinib
VEGF RECEPTORS			
VEGFR1	72%	6%	12%
VEGFR2	76%	6%	14%
VEGFR3	63%	27%	18%
OTHER RECEPTORS			
PDGFR α	54%	37%	37%
PDGFR β	79%	75%	76%
FGFR1	69%	12%	28%
FGFR2	62%	17%	28%
FGFR3	87%	40%	58%
JAK1	5%	14%	0%

Inhibition Key: <50% 50-70% 70-85% 85-100% Not tested

Comparison of Hydrogel vs Other Retinal Polymer Implants

	Implant (Full View)	Implant Dimensions (Length x Diameter)	Needle Size	Matrix Degradation & Release ^{^^}	Polymer Byproduct	Implant Remnants at 12 Months
OZURDEX abbvie		6mm x 0.46mm* 	22-gauge*		Lactic acid and glycolic acid [†]	Remnant lasts ~12 months [‡]
EYP-1901 EYEPOINT [™] PHARMACEUTICALS		Two implants with length of 8mm each [¶] 	22-gauge [#]		>75 KDa polyvinyl alcohol	Two remnants last 18-24 months ^{**}
GB-102 graybug vision		N/A	27-gauge ^{††}		Lactic acid and glycolic acid ^{††}	N/A
OTX-TKI OCULAR THERAPEUTIX		7mm x 0.40mm [^] 	25-gauge [^]		Inert hydrogel PEG chains [^]	No remnants^{^^}

OTX-TKI is an investigational product candidate which has not been approved by the FDA or any regulatory authority.

Schematic drawings, not drawn to scale or color, for illustrative purposes only; yellow or gray dots/spheres depict therapeutic agents released by implants. Images of EYP-1901 implants from Abbey, AM. Topline Data from DAVIO 2: a Phase 2 Study of a Single Injection of DURAVYU (Vorolanib Intravitreal Insert). Presented at Retina World Congress. Fort Lauderdale, FL; May 9-12, 2024.

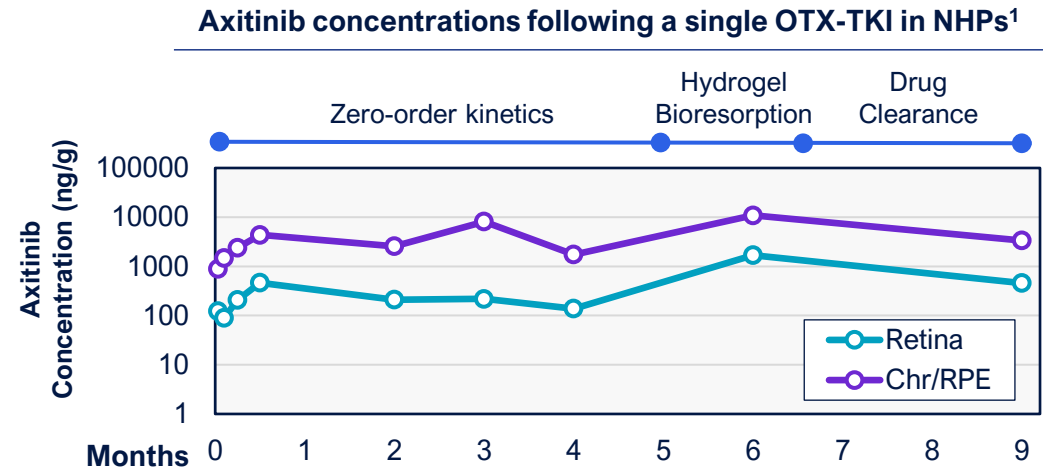
*. Li M, et al. *Heliyon*. 2022;8(12):e12219. †. Pellegrini GABP, et al. *Int J Retina Vitreous*. 2025;11(1):7. ‡. Kim JT, et al. *Retina*. 2020; 40(11):p 2226-2231. §. Schmit-Eilenberger VK. *Clin Ophthalmol*. 2015;9:801-811. ¶. Diameter of EYP-1901 unconfirmed; EyePoint Pharmaceuticals. Clinical Study Protocol. Protocol No. EYP-1901-201. Published January 25, 2024. ClinicalTrials.gov identifier: NCT05381948.. #. EyePoint Pharmaceuticals DAVIO2 Topline Results Conference Call, December 4, 2023. ||. Wykoff CC, et al. *J Vitreoretin Dis*. 2024;8(5):577-586. **EyePoint Pharmaceuticals. TD Cowen 44th Annual Health Care Conference. March 5, 2024. <https://investors.eyepointpharma.com/events-and-presentations>. ††. Graybug Vision Inc. Clinical Study Protocol. Protocol No. GBV-102-002. Published September 8, 2020. ClinicalTrials.gov identifier: NCT03953079. ^. Blizzard CD, et al. Ocular Implant Containing a Tyrosine Kinase Inhibitor. US Patent 11,439,592 B2. September 13, 2022. ^^ Data on File 044. Ocular Therapeutix. 2026. PEG, polyethylene glycol

OTX-TKI is Designed to Release Axitinib and Maintain Intraocular Drug Levels Over Time

OTX-TKI In Vivo Delivery Profile¹

Rapid, continuous and consistent axitinib delivery

Targeted drug delivery to retina and choroid



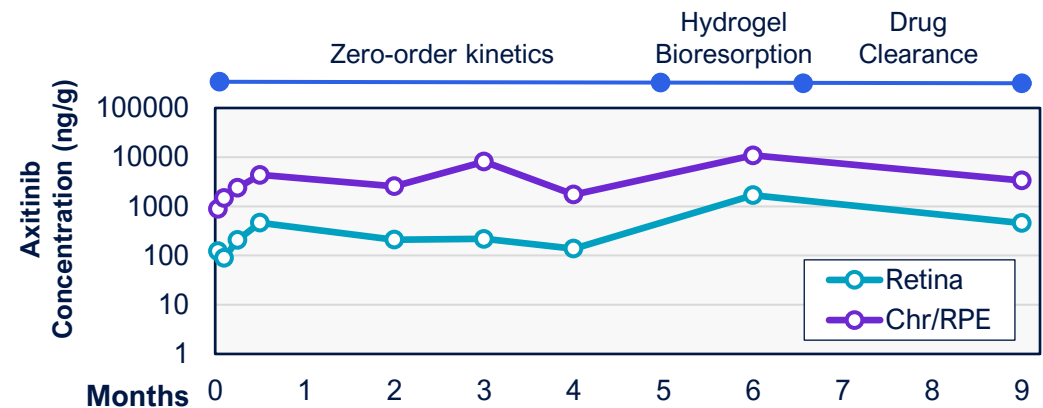
OTX-TKI is Designed to Release Axitinib and Maintain Intraocular Drug Levels Over Time with Complete Bioresorption

OTX-TKI In Vivo Delivery Profile¹

Rapid, continuous and consistent axitinib delivery

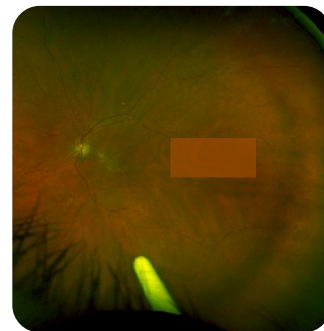
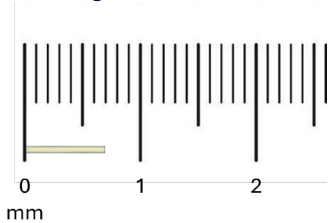
Targeted drug delivery to retina and choroid

Axitinib concentrations following a single OTX-TKI in NHPs¹

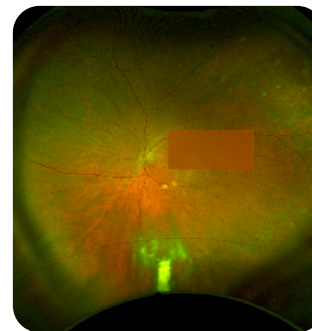


Hydrogel bioresorbs completely and predictably via hydrolysis for potential redosing¹⁻³

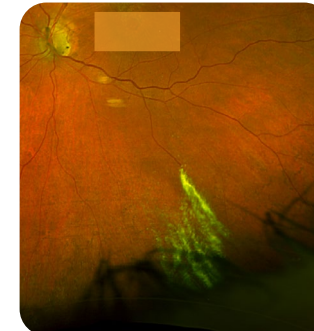
Dry OTX-TKI Dimensions
7mm length, 0.4mm diameter



Day 1



Week 20



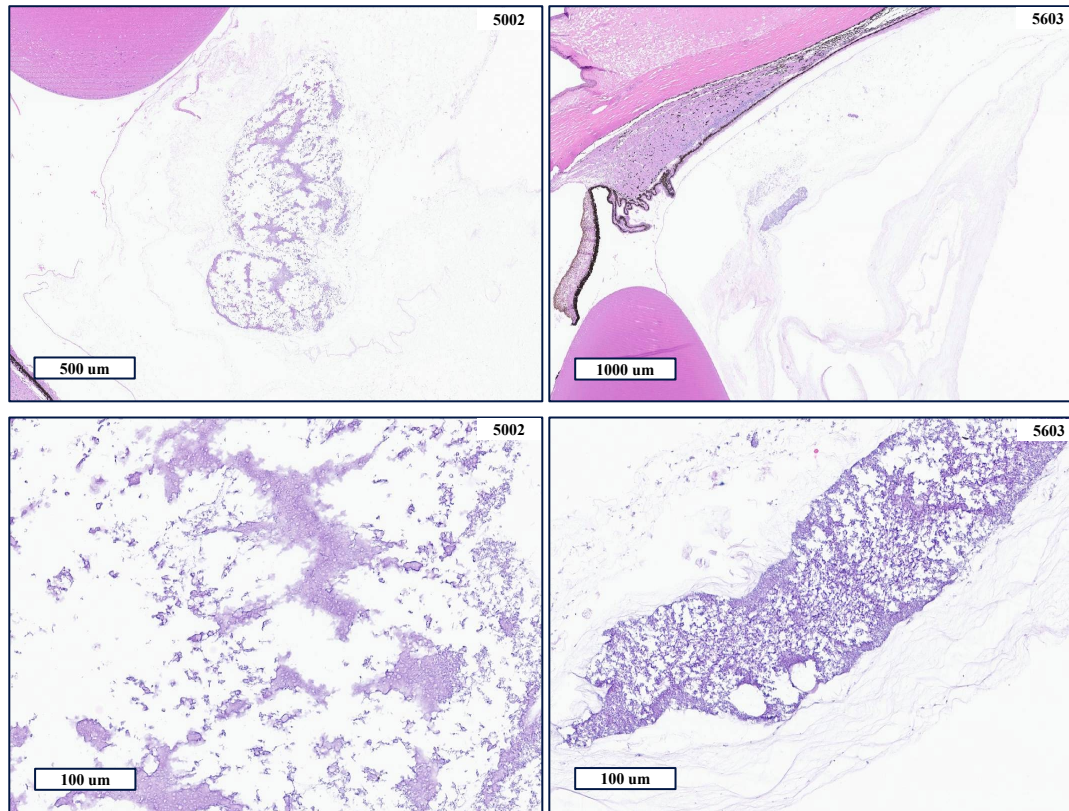
Week 32



Week 44

Histology of Ocular Tissues Demonstrates Axitinib Elution with No Inflammatory Cells

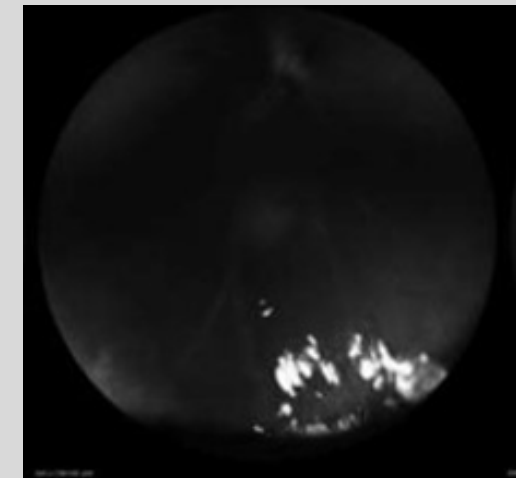
Histology of Ocular Tissues from 0.7 mg OTX-TKI in NHPs



Microscopic Findings

White, refractile, basophilic, granular axitinib in the vitreous

No inflammatory cells in vitreous, retina, or AC



From NHP Toxicity Study, not human picture

OTX-TKI is Designed for Seamless, Immediate Adoption

Ideal Target Product Profile

Pan-VEGF and PDGF
inhibitory activity¹

Axitinib has best-in-class
(TKI) potency¹

Extended durability
up to 12 months²

Single Bioresorbable Hydrogel

No remnants³

Tunable hydrogel⁴

Hydrogel used
in over 5 million patients⁵

Optimizing for Retina Practices

Familiar IVT
injection with **25g needle**⁶

Predictable
schedule for patients²

Designed for **improved**
treatment adherence²

OTX-TKI is an investigational product candidate which has not been approved by the FDA or any regulatory authority.

SOL-1: Results

Darius Moshfeghi, MD

SOL: OTX-TKI Phase 3 Clinical Program in nAMD

Evaluating the efficacy, durability and safety of OTX-TKI in nAMD

Registrational Trials

SOL-1

Phase 3 Superiority Trial

Durability of a single
OTX-TKI injection

SOL-R

Phase 3 Non-Inferiority Trial

Repeat dosing
every 6 months

Complementary studies designed to
provide a comprehensive characterization of
OTX-TKI across patient populations

Extension Study

SOL-X

Open-Label Extension

Long-term safety and disease
modifying potential of continuous
VEGF suppression

Eligible patients who
complete end of Year 2 visit
in SOL-1 or SOL-R

SOL: OTX-TKI Phase 3 Clinical Program in nAMD

Evaluating the efficacy, durability and safety of OTX-TKI in nAMD

Registration Trials

SOL-1

Phase 3 Superiority Trial

Durability of a single
OTX-TKI injection

SOL-R

Phase 3 Non-Inferiority Trial

Repeat dosing
every 6 months

Extension Study

SOL-X

Open-Label Extension

Long-term safety and disease
modifying potential of continuous
VEGF suppression

Eligible patients who
complete end of Year 2 visit
in SOL-1 or SOL-R

Complementary studies designed to
provide a comprehensive characterization of
OTX-TKI across patient populations

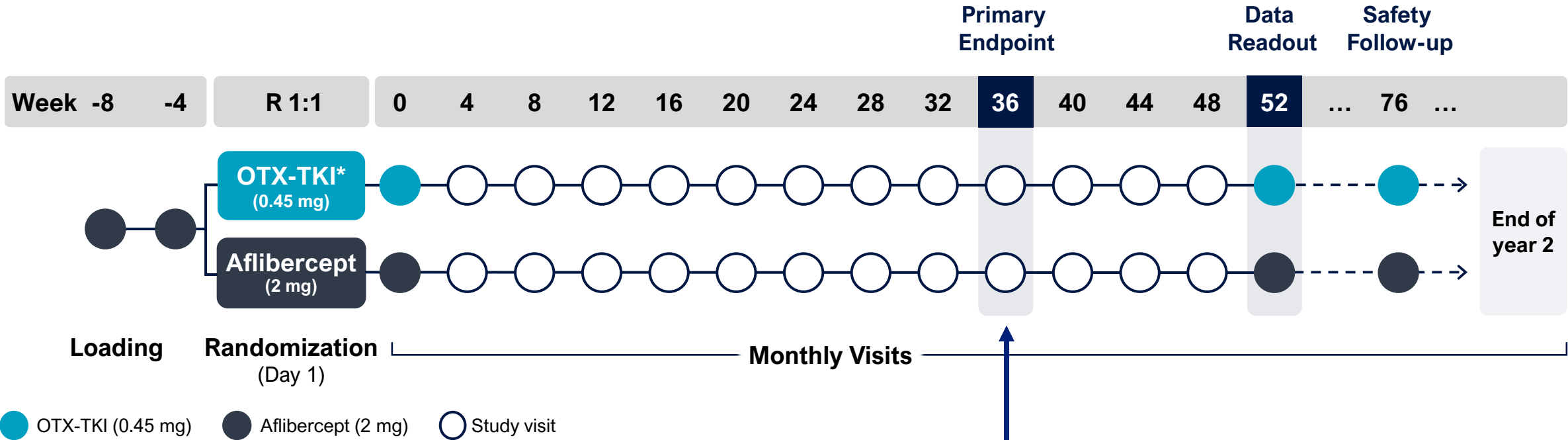
OTX-TKI Evaluated in a Superiority nAMD Trial Designed to Align with FDA Guidance

	FDA Guidance for nAMD Trials		SOL-1 Trial Design
Dosing Schedule	One comparator arm should have the same dosing schedule as the investigational drug ¹	✓	Both study arms have same dosing schedule
Sham Injections	Sham injections are not recommended; considered inadequate masking ^{2,3}	✓	No sham injections
Superiority Endpoint	Decrease  Increase  Difference ≥15-Letter ¹	✓	≥15-Letter Decrease Proportion with <15 ETDRS letters of BCVA loss from baseline at Week 36



- Compares a single OTX-TKI (0.45 mg) dose to a single aflibercept (2 mg) dose
- Designed to show superiority on efficacy and durability
- SOL-1 is being conducted under a **Special Protocol Assessment**

Study Design: OTX-TKI and Aflibercept Direct Comparison

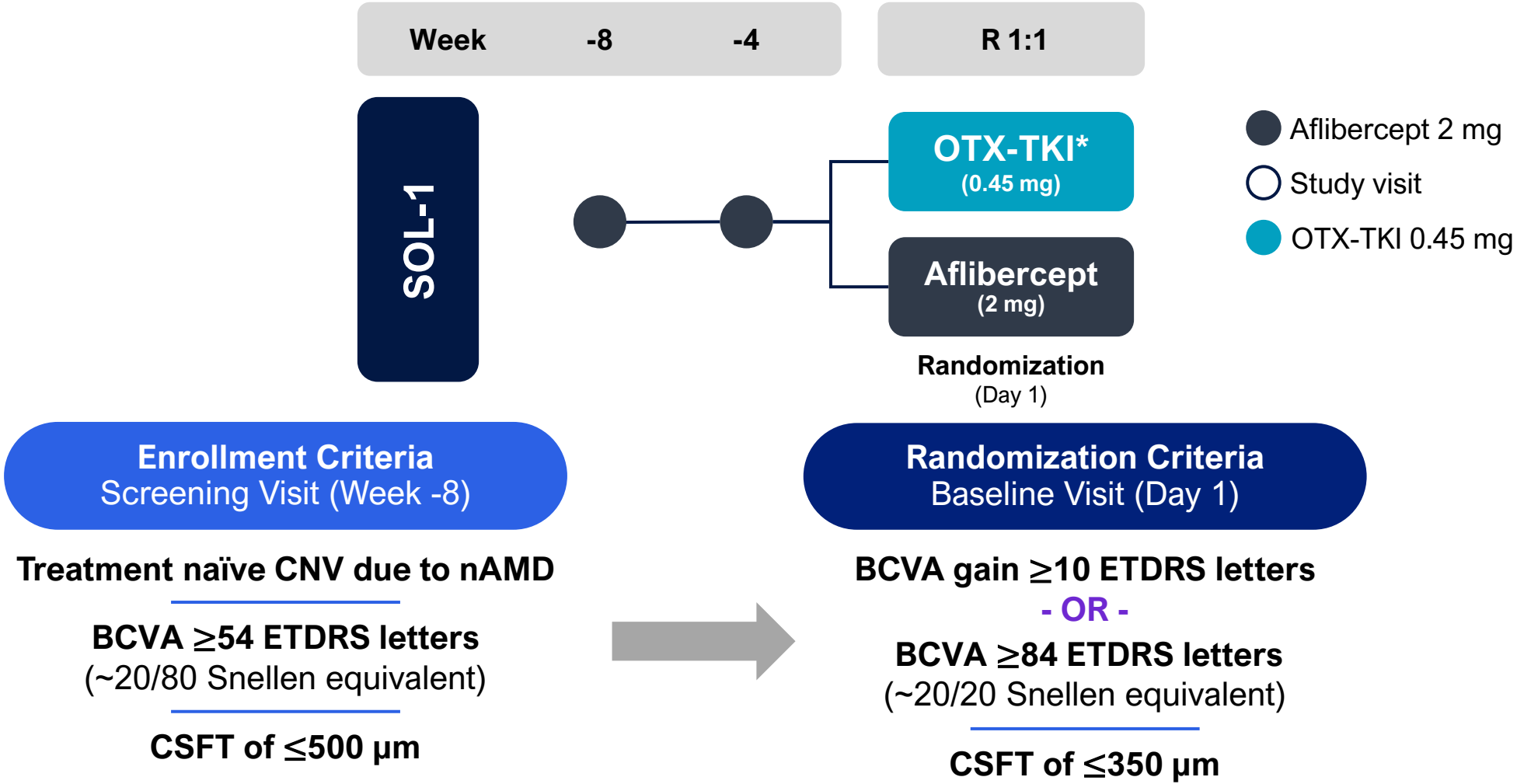


Rescue Treatment Criteria
BCVA loss ≥ 15 ETDRS letters from baseline due to nAMD
— OR —
New macular hemorrhage that would likely lead to irreversible vision loss if left untreated in the opinion of the investigator after discussion with Medical Monitor

Primary Endpoint (Week 36)
Proportion of subjects who maintained visual acuity, defined as <15 ETDRS letters of BCVA loss from baseline at Week 36

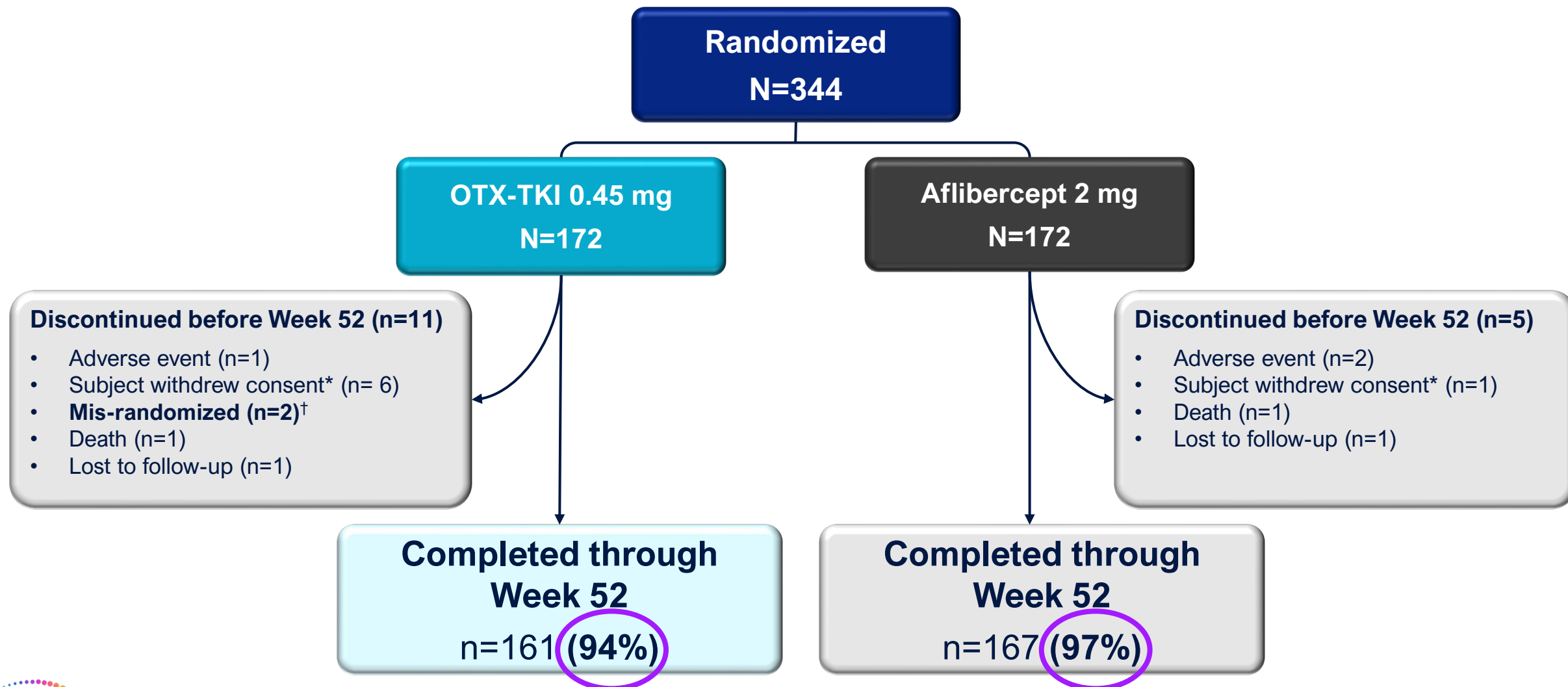
Eligibility and Enrollment Criteria

Key Inclusion Criteria



Strong Subject Retention Maintained through Week 52

Subject Disposition



Excellent Vision at Baseline; Majority Demonstrated ≥ 10 -letter Gain from Screening

SOL-1

Best-Corrected Visual Acuity

Screening Visit (Week -8)

Study Eye Characteristics	OTX-TKI 0.45 mg N = 172	Aflibercept 2 mg N = 172
BCVA, ETDRS letter score, mean (SD) Snellen Equivalent	70.9 (11.3) ~20/40	69.5 (10.8) ~20/40

Baseline (Randomization)

Study Eye Characteristics	OTX-TKI 0.45 mg N = 172	Aflibercept 2 mg N = 172
BCVA, ETDRS letter score, mean (SD) Snellen Equivalent	80.8 (7.6) ~20/25	79.2 (7.9) ~20/25
Change from Screening to Baseline, n (%)		
≥ 10 ETDRS letter gain	107 (62.9)	114 (67.1)
≥ 84 ETDRS letters (~20/20)	63 (37.1)	56 (32.9)

Anatomic Improvement Observed After Two Aflibercept Injections

Central Subfield Thickness

Screening Visit (Week -8)

Study Eye Characteristics	OTX-TKI 0.45 mg N = 172	Aflibercept 2 mg N = 172
BCVA, ETDRS letter score, mean (SD) Snellen Equivalent	70.9 (11.3) ~20/40	69.5 (10.8) ~20/40
CSFT, μm , mean (SD)	303.6 (72.5)	302.7 (78.3)

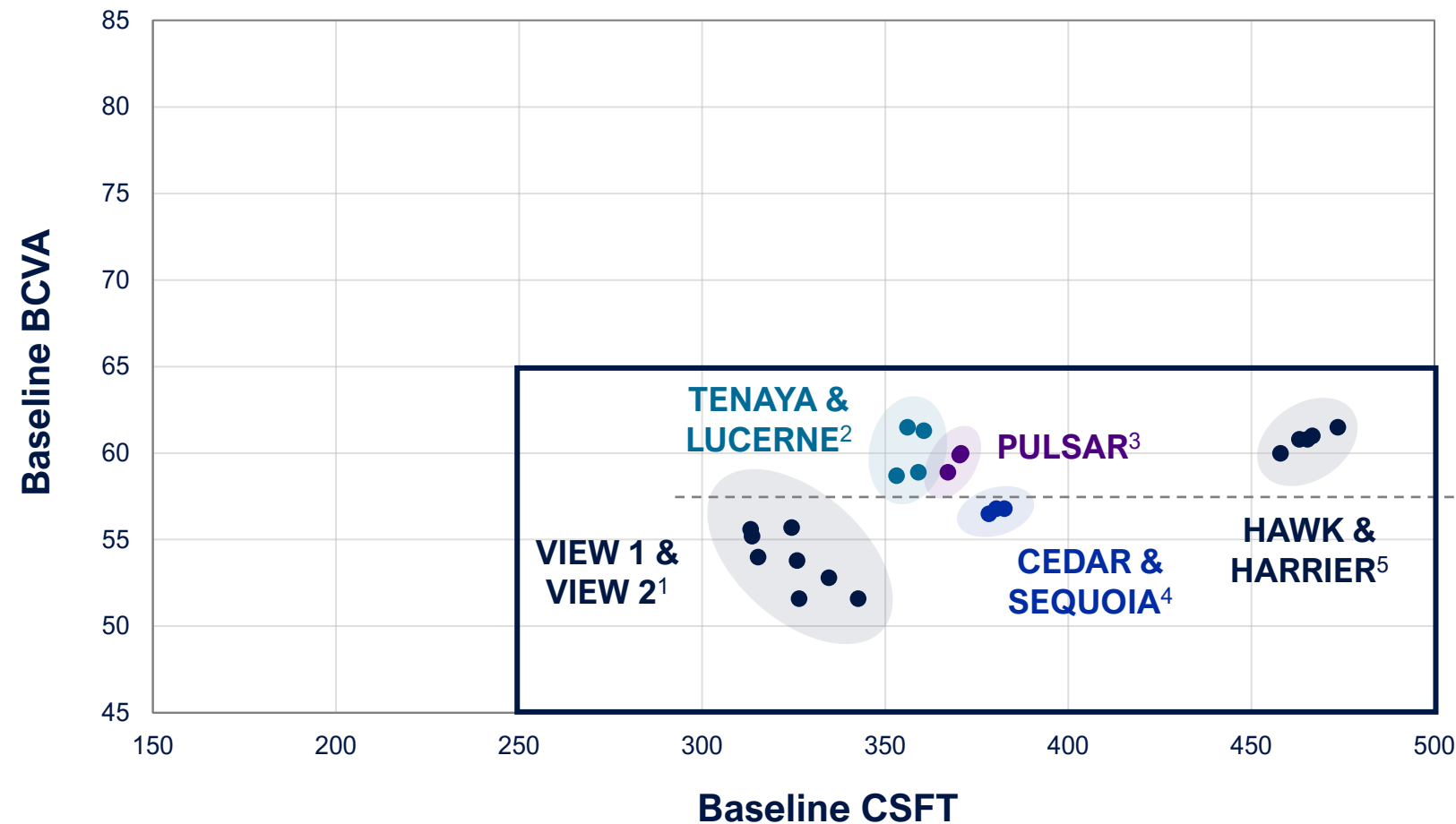
Baseline (Randomization)

Study Eye Characteristics	OTX-TKI 0.45 mg N = 172	Aflibercept 2 mg N = 172
BCVA, ETDRS letter score, mean (SD) Snellen Equivalent	80.8 (7.6) ~20/25	79.2 (7.9) ~20/25
CSFT, μm , mean (SD)	219.3 (37.1)	226.8 (42.1)

SOL-1 Evaluates a Distinct nAMD Cohort with Excellent Baseline Vision



Baseline BCVA Across nAMD Trials



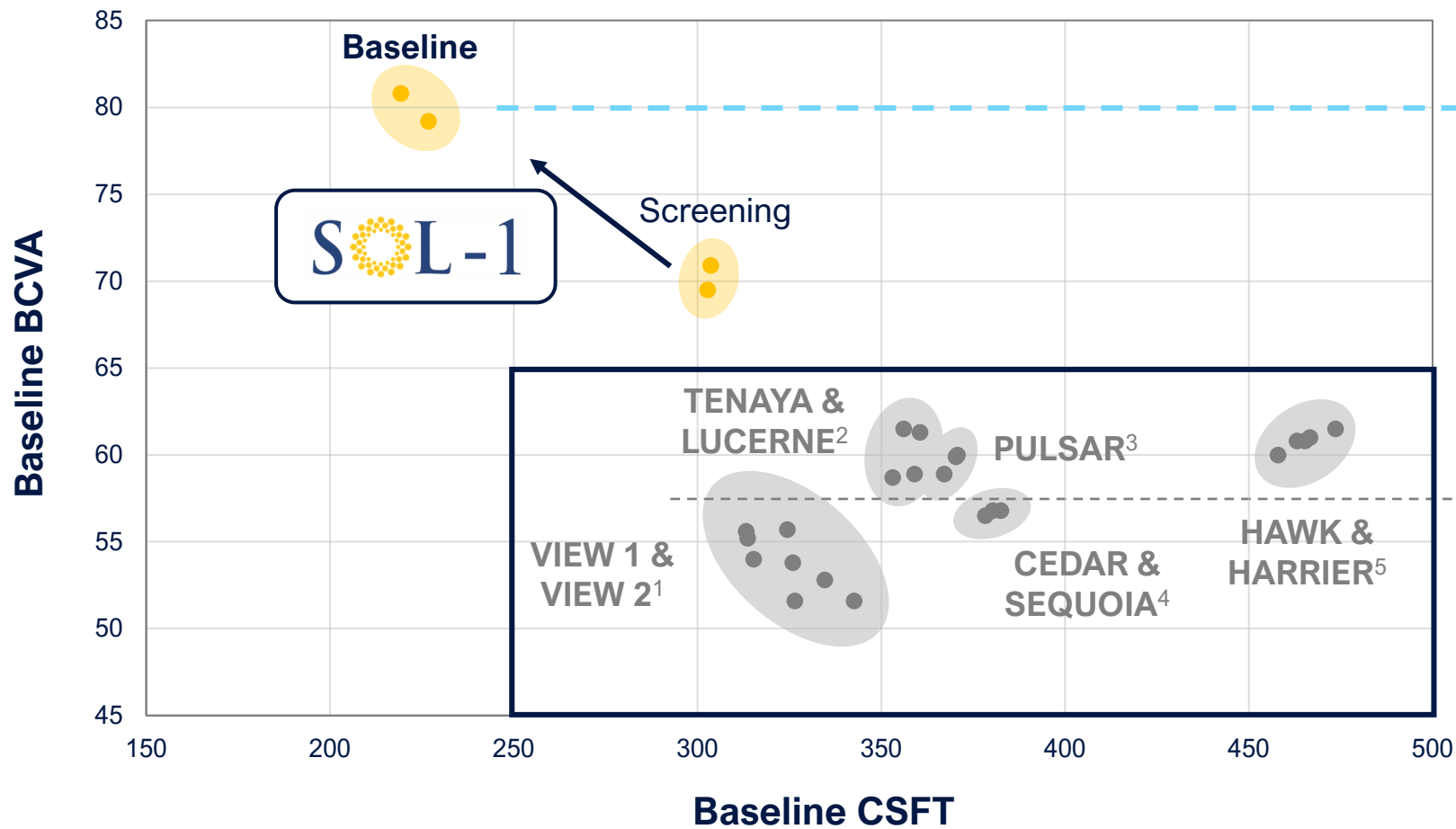
nAMD Trials
mean BCVA: 52 to 62 letters
(~20/100 to 20/60)

1. Heier JS, et al. *Ophthalmology*. 2012;119(12):2537-2548. 2. Heier JS, et al. *Lancet*. 2022;399(10326):729-740. 3. Lanzetta P, et al. *Lancet*. 2024;403(10432):1141-1152. 4. Kunimoto D, et al. *Ophthalmology*. 2020;127(10):1331-1344. 5. Dugel PU, et al. *Ophthalmology*. 2020;127(1):72-84.
BCVA, best-corrected visual acuity; CSFT, central subfield thickness; nAMD, neovascular age-related macular degeneration

SOL-1 Evaluates a Distinct nAMD Cohort with Excellent Baseline Vision



Baseline BCVA Across nAMD Trials

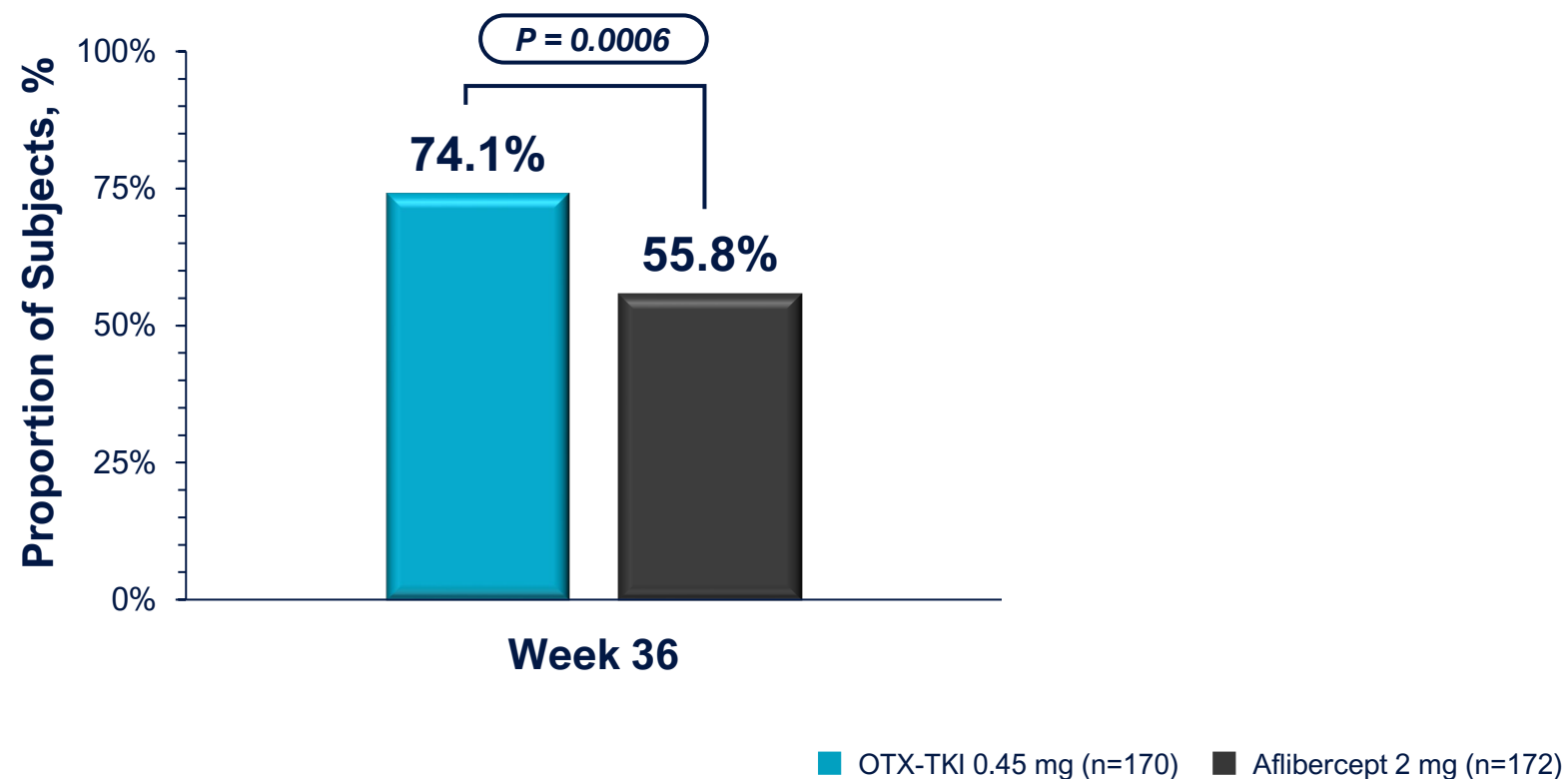


SOL-1
mean BCVA: **81 letters (~20/25)**
mean CSFT: **219 μM**

nAMD Trials
mean BCVA: **52 to 62 letters (~20/100 to 20/60)**

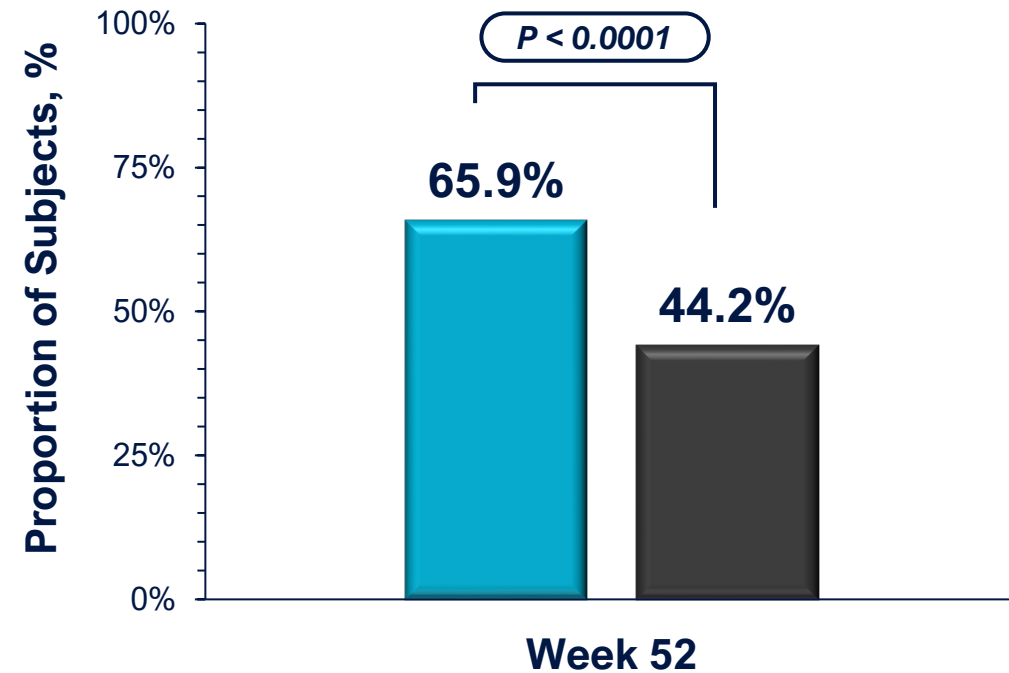
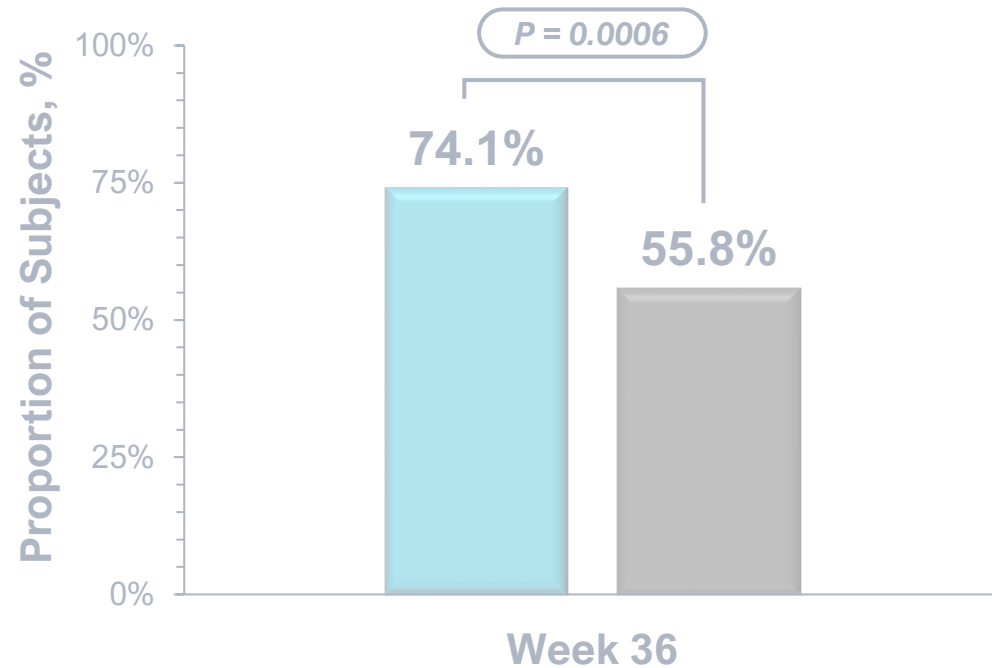
Primary Endpoint Successfully Met

Statistically Significant Higher Proportion of Subjects Maintained Visual Acuity* with OTX-TKI vs. Aflibercept 2 mg at Week 36



Difference in Proportion of Subjects Maintaining Visual Acuity Sustained Through Week 52

Key Secondary Endpoint: Maintenance of Visual Acuity* at Week 52

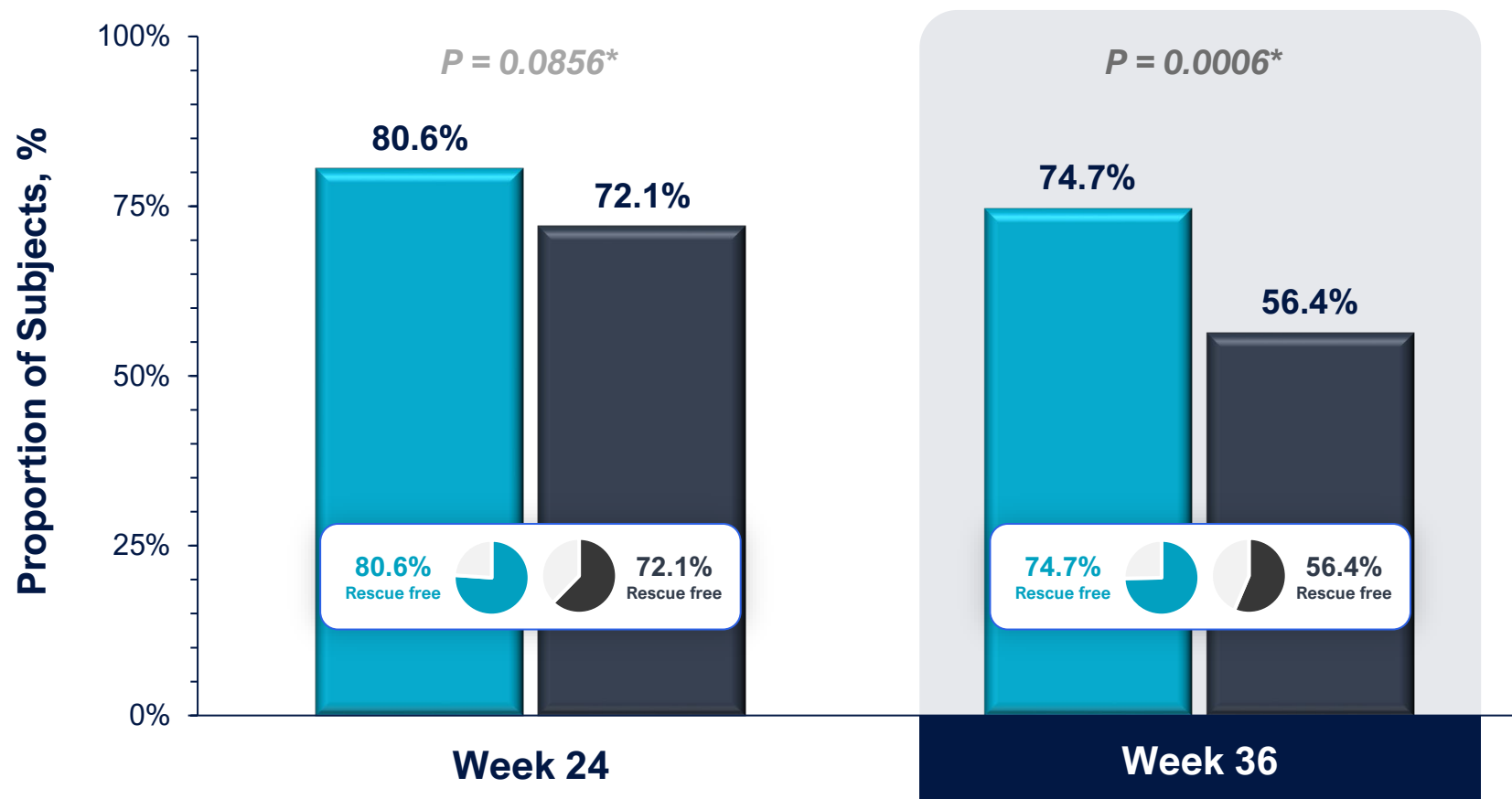


■ OTX-TKI 0.45 mg (n=170) ■ Aflibercept 2 mg (n=172)

Most Subjects (~75%) in the OTX-TKI Arm Did Not Receive Rescue Injections at Week 36

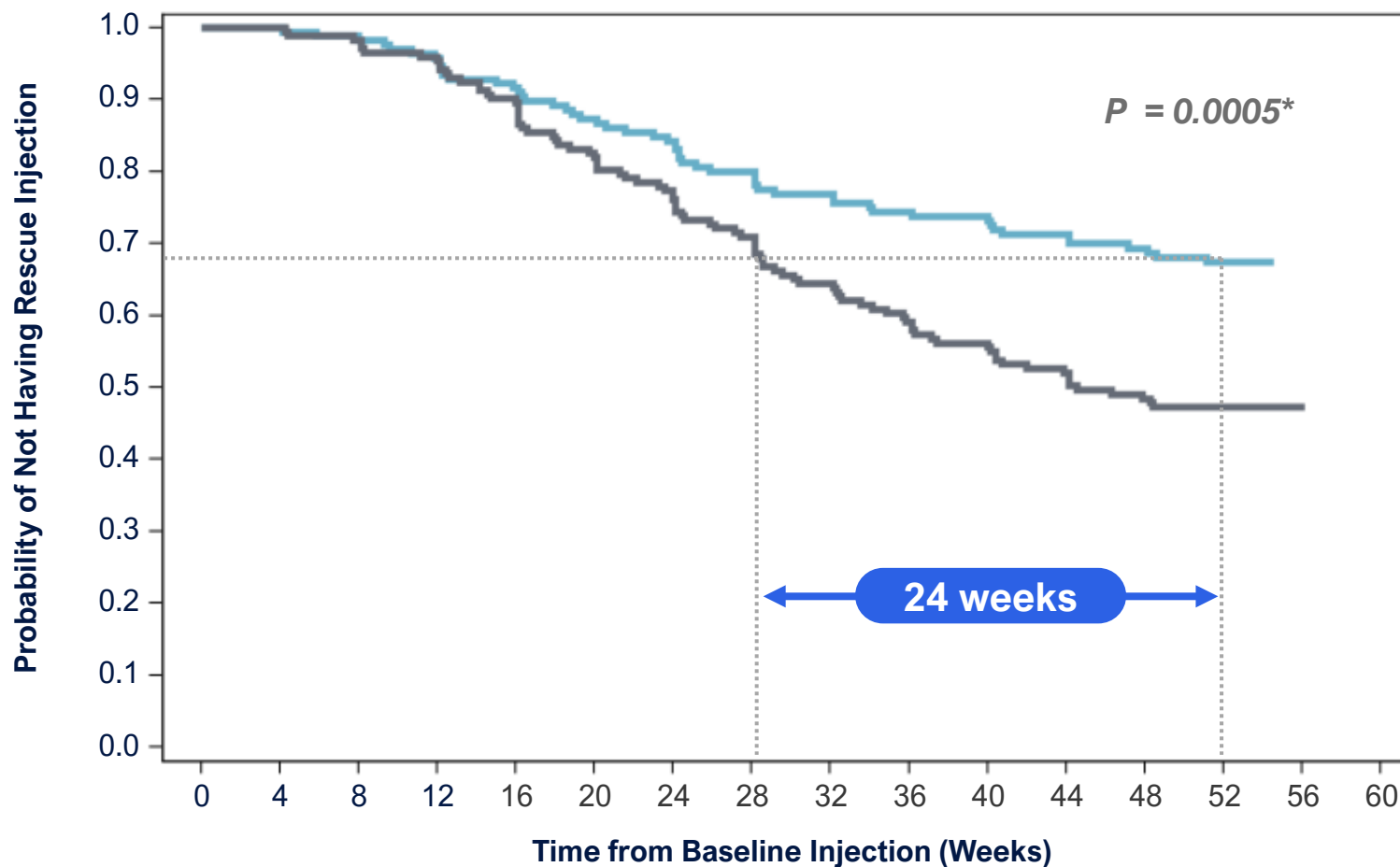
SOL-1

Post Hoc Analysis: Proportion of Rescue-Free Subjects



OTX-TKI Extended Durability, Significantly Delaying First Rescue Injection

Secondary Endpoint: Time to First Rescue Treatment

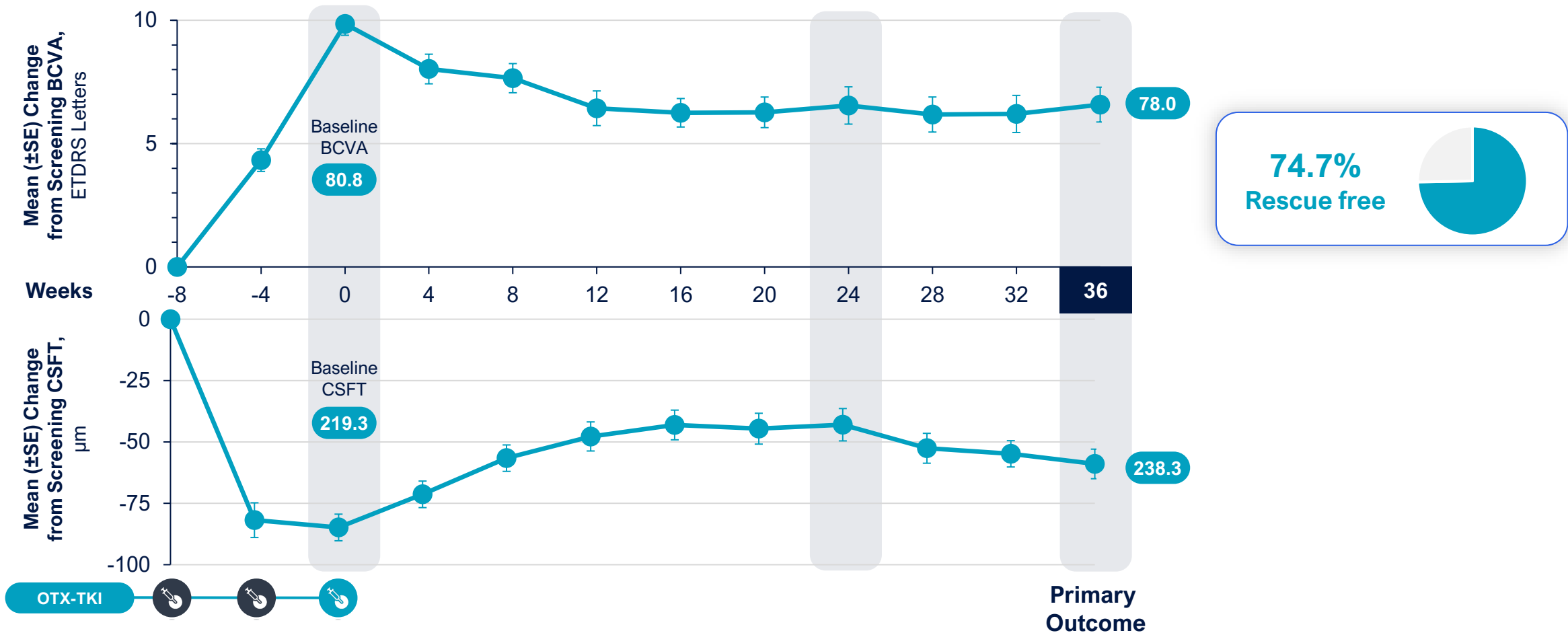


— OTX-TKI 0.45 mg (n=170)
— Aflibercept 2 mg (n=172)

On Average, OTX-TKI Subjects Maintained Vision and Anatomy, with Most Remaining Rescue-Free

SOL-1

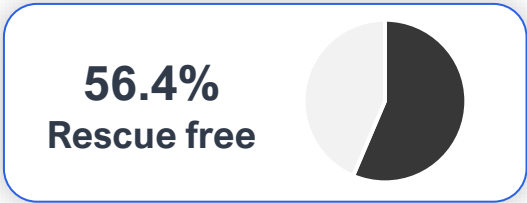
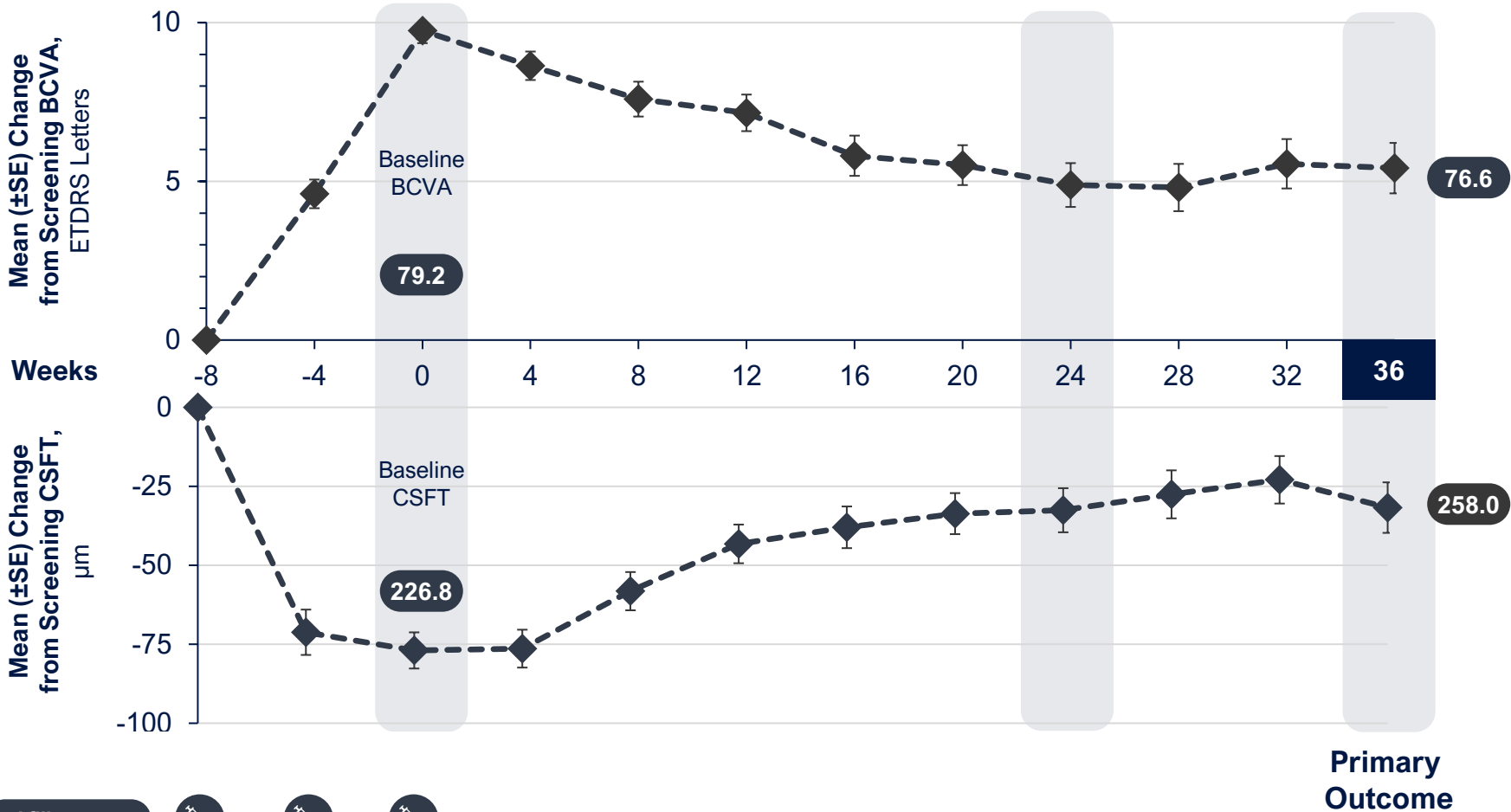
Post Hoc Analysis: Mean Change in BCVA and CSFT from Screening



On Average, OTX-TKI Subjects Maintained Vision and Anatomy, with Most Remaining Rescue-Free



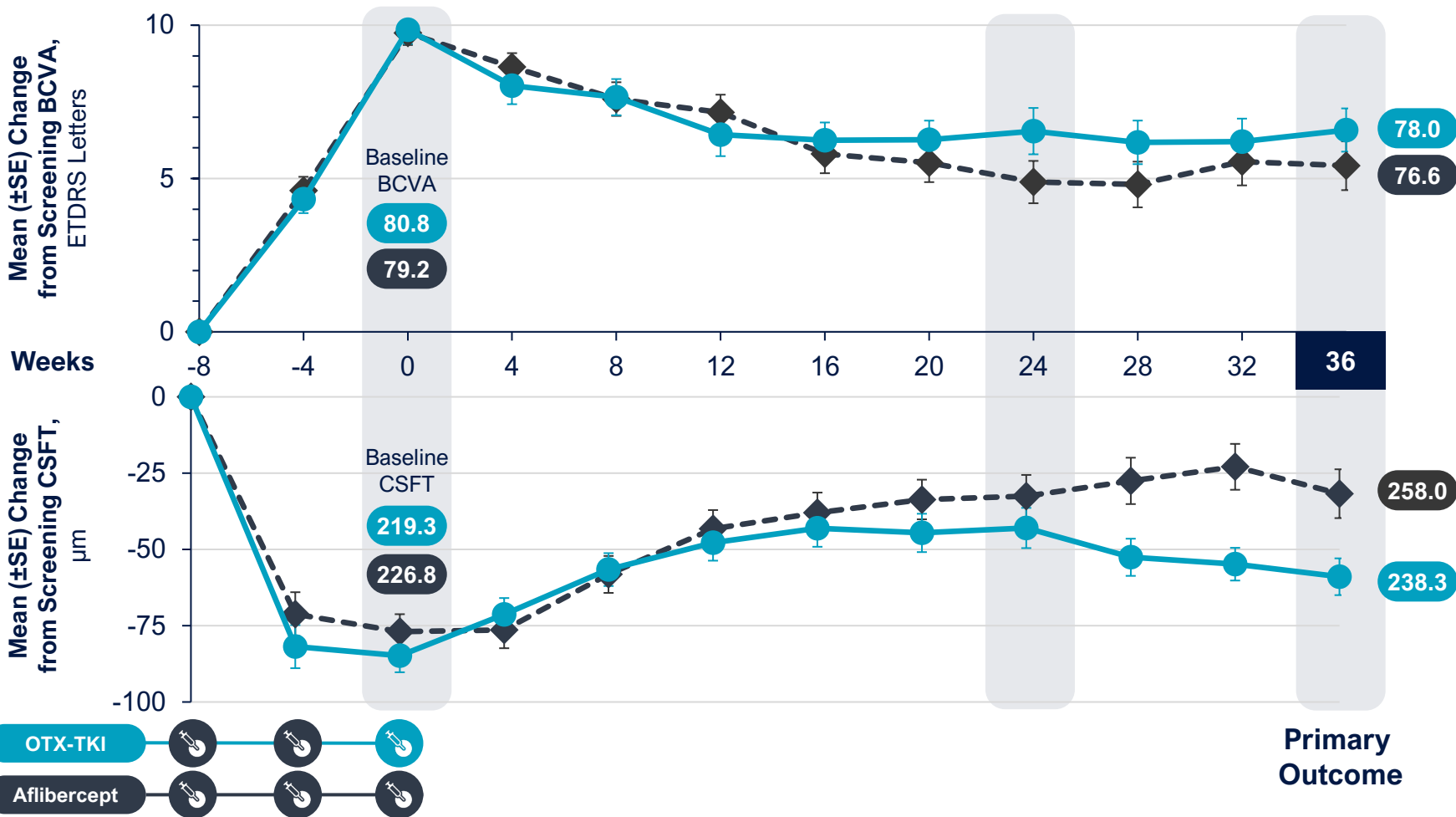
Post Hoc Analysis: Mean Change in BCVA and CSFT from Screening



On Average, OTX-TKI Subjects Maintained Vision and Anatomy, with Most Remaining Rescue-Free

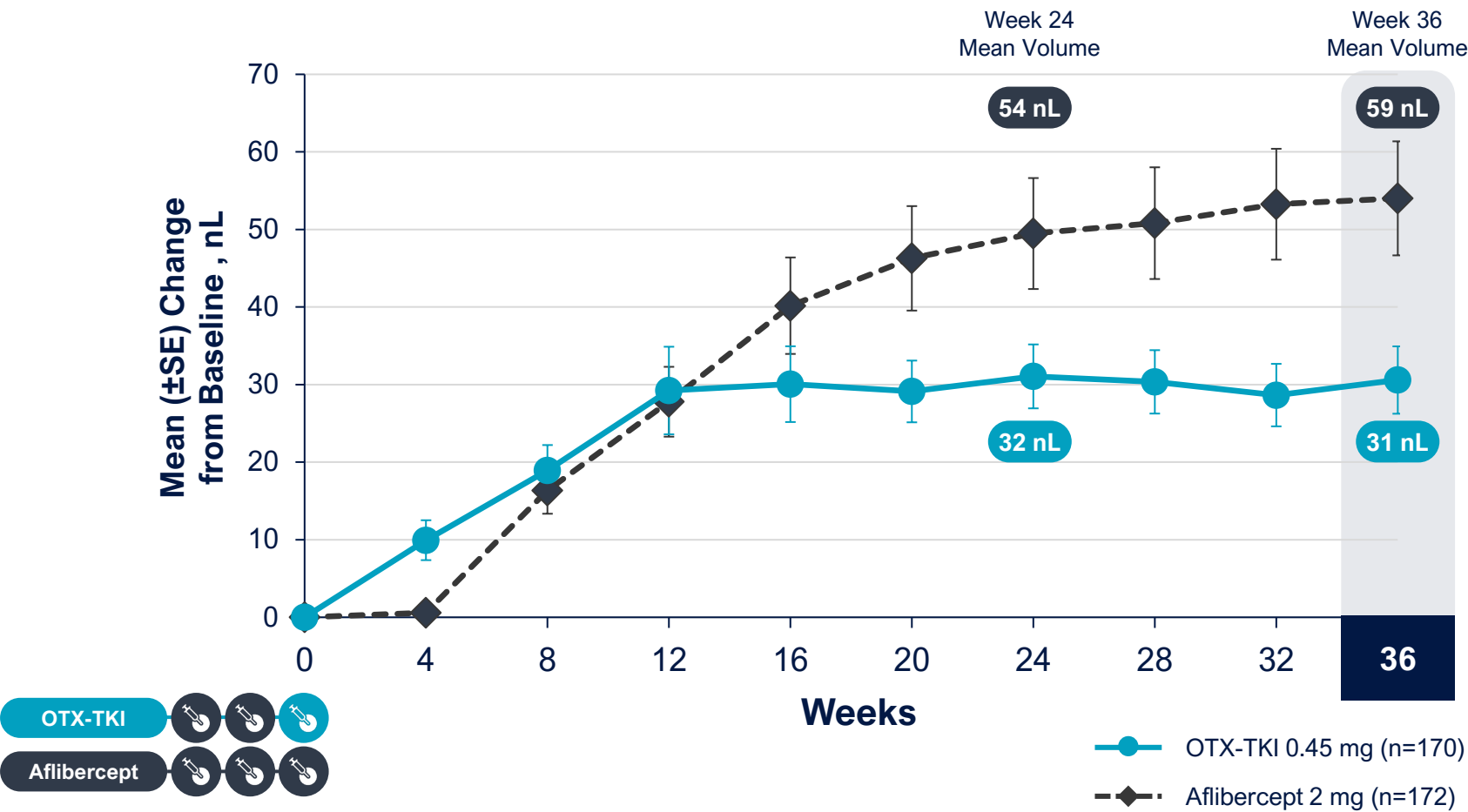


Post Hoc Analysis: Mean Change in BCVA and CSFT from Screening

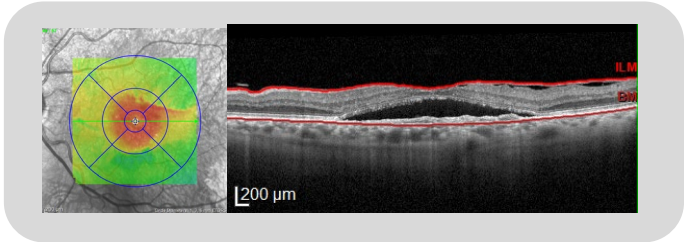


OTX-TKI Delivered Sustained Fluid Control

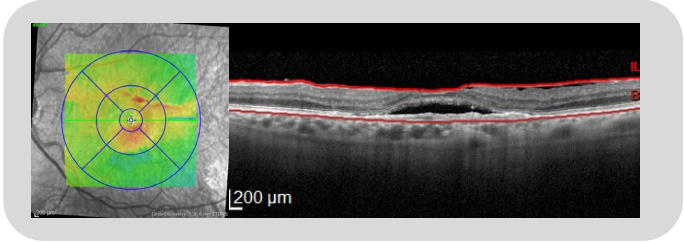
Prespecified Exploratory Analysis: Mean Change in Intra-Retinal and Sub-Retinal Fluid Volume



Total Fluid Volume: 60 nL



Total Fluid Volume: 30 nL



No Treatment or Procedure Related SAEs

Non-Ocular and Ocular Adverse Events in the Study Eye

Subjects with AEs Through Week 52, n (%)	OTX-TKI 0.45 mg n = 170	Aflibercept 2 mg n = 172
Non-ocular AEs		
≥ 1 AE	84 (49.4)	73 (42.4)
≥ 1 SAE	19 (11.2)	21 (12.2)
≥ 1 treatment-related AE	0	0
≥ 1 treatment-related SAE	0	0
≥ 1 study procedure-related AE	1 (0.6)	0
≥ 1 study procedure-related SAE	0	0
AE leading to study discontinuation	3 (1.8)	3 (1.7)
AE leading to death*	2 (1.2)	1 (0.6)
Ocular AEs in the study eye		
≥ 1 AE	90 (52.9)	58 (33.7)
≥ 1 SAE†	1 (0.6)	0
≥ 1 treatment-related AE	15 (8.8)	1 (0.6)
≥ 1 treatment-related SAE	0	0
≥ 1 study procedure-related AE	15 (8.8)	7 (4.1)
≥ 1 study procedure-related SAE	0	0
Ocular AE leading to study discontinuation	0	0

OTX-TKI was Generally Well Tolerated

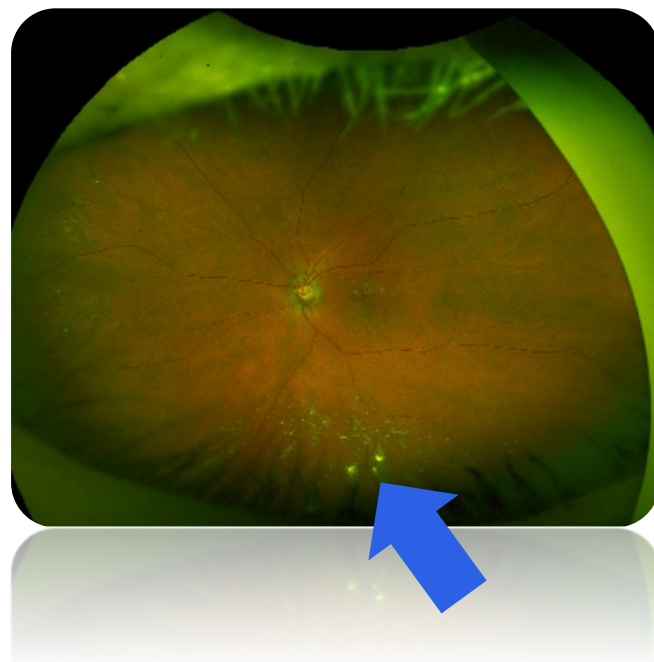
Ocular Adverse Events in the Study Eye

Subjects with Ocular AEs (> 2%) Through Week 52, n (%)	OTX-TKI 0.45 mg n = 170	Aflibercept 2 mg n = 172
Vitreous floaters	21 (12.4)	2 (1.2)
Cataract	12 (7.1)	5 (2.9)
Conjunctival hemorrhage	11 (6.5)	5 (2.9)
Retinal hemorrhage	10 (5.9)	17 (9.9)
Dry eye	7 (4.1)	2 (1.2)
Vitreous detachment	7 (4.1)	3 (1.7)
Punctate keratitis	6 (3.5)	0
Vitreous opacities	6 (3.5)	0
Eye pain	5 (2.9)	1 (0.6)
Anterior chamber opacity	4 (2.4)	0
Posterior capsule opacification	4 (2.4)	6 (3.5)

- Nine IOI events observed in study eyes of 7 subjects in the OTX-TKI arm
 - All cases were mild or moderate in severity and resolved
- **There were no cases of endophthalmitis, occlusive or non-occlusive retinal vasculitis**

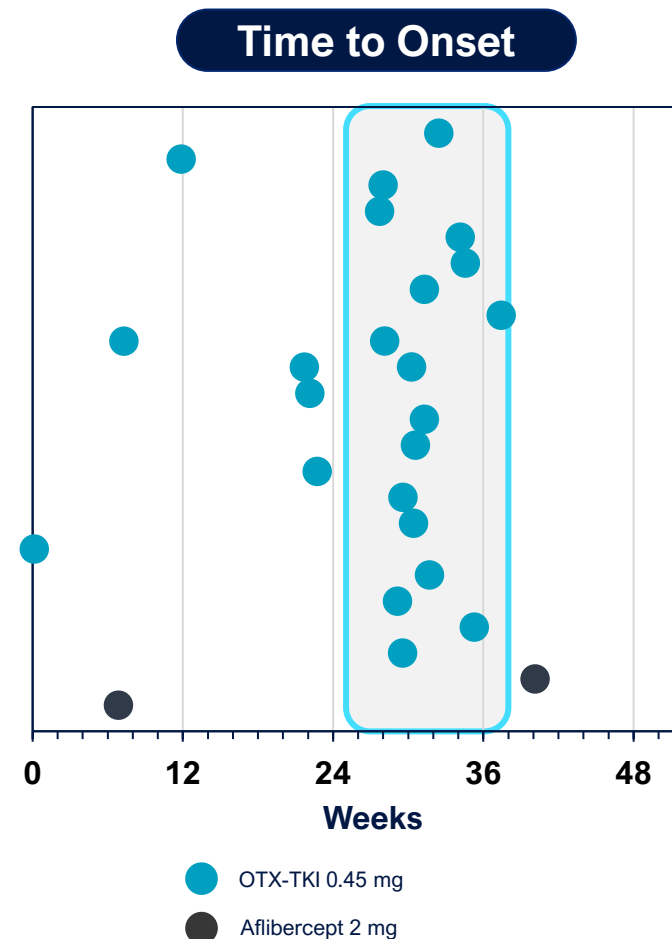
Appearance of Floaters May Correspond to Hydrogel Dissolution and Drug Elution

Evaluation of Vitreous Floaters



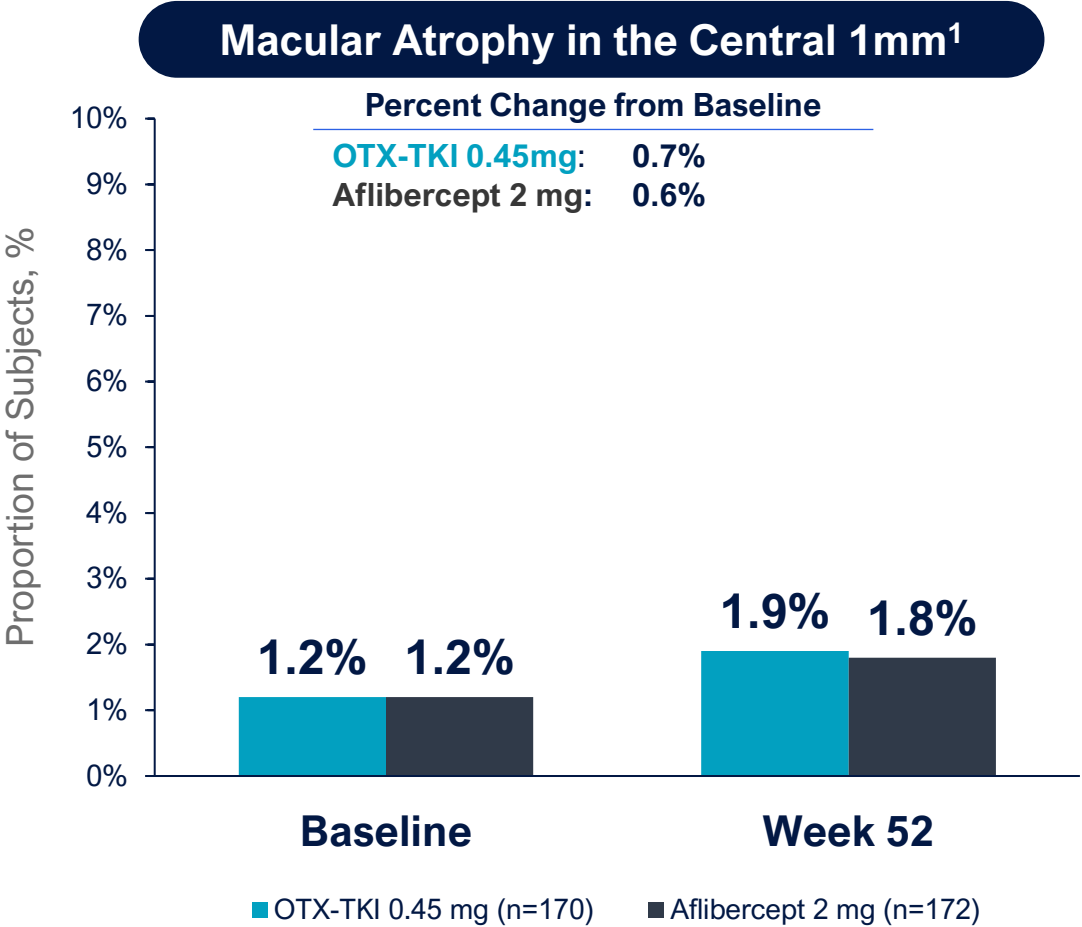
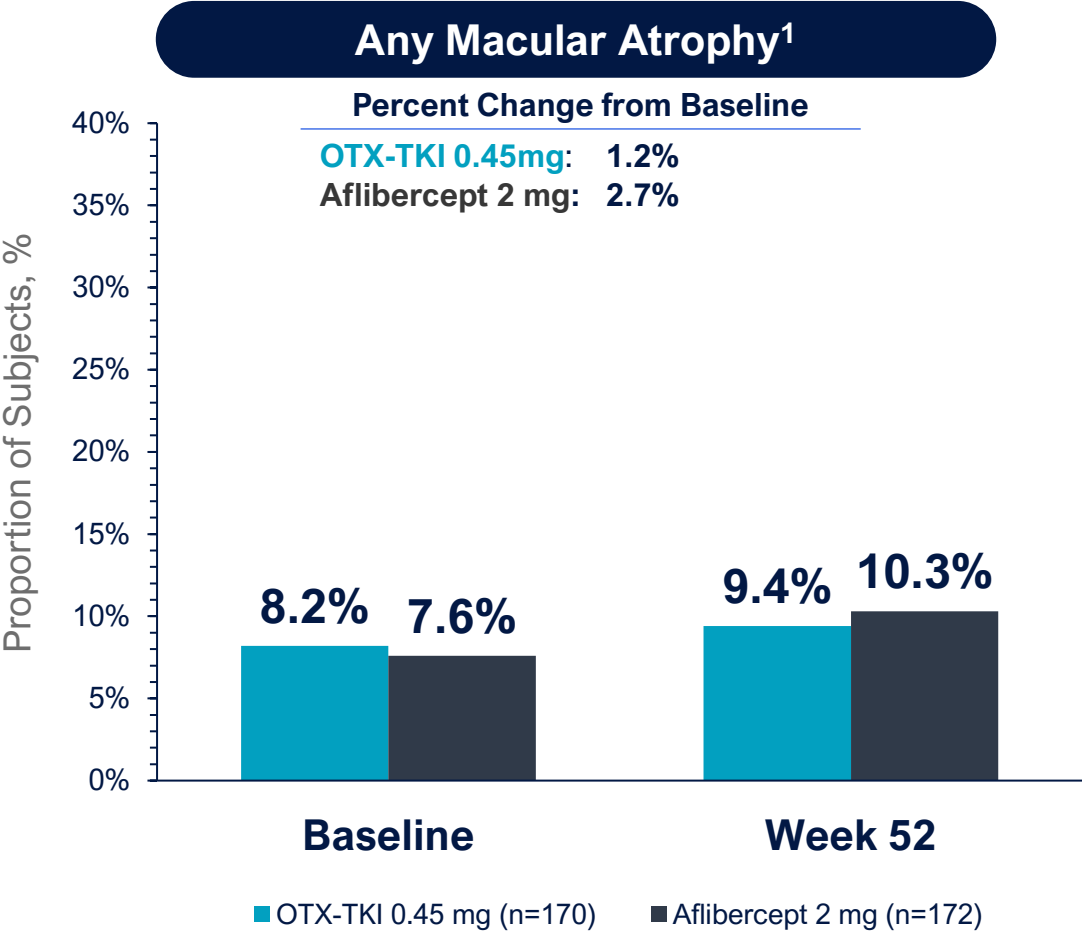
	OTX-TKI 0.45 mg n = 23*	Aflibercept 2 mg n = 2
Time to onset, days		
Mean (SD)	187.9 (64.0)	164.5 (164.8)
Median	207.0	164.5
BCVA change from prior visit before onset of floaters, ETDRS letters		
Mean change (SD)	-0.4 (5.0)	-5.5 (6.4)
Median change	0.0	-5.5

- All cases were mild to moderate in severity
- No evidence of IOI associated with floaters
- Onset of vitreous floaters AE coincides with drug elution stage of OTX-TKI hydrogel bioresorption process
- **For all subjects with vitreous floaters AE reported, drug particles are no longer visible**



Incidence of Macular Atrophy in SOL-1

Safety Analysis



¹cRORA measured by the reading center
cRORA, complete retinal pigment epithelium and outer retinal atrophy

Superiority Primary Endpoint Successfully Met

OTX-TKI Demonstrated Superior Maintenance of Vision Compared to Aflibercept 2mg at Week 36*

First trial to demonstrate durability ≥ 9 months

In a post hoc analysis, 74.7% of subjects treated with OTX-TKI did not receive rescue treatments by Week 36

Unmatched Durability

Visual and anatomic stability with OTX-TKI

In a post hoc analysis, OTX-TKI subjects maintained visual and anatomical outcomes, with most remaining rescue-free through Week 36

Sustained Disease Control

OTX-TKI demonstrated a well-tolerated safety profile through Week 52

There were no cases of endophthalmitis, occlusive or non-occlusive retinal vasculitis

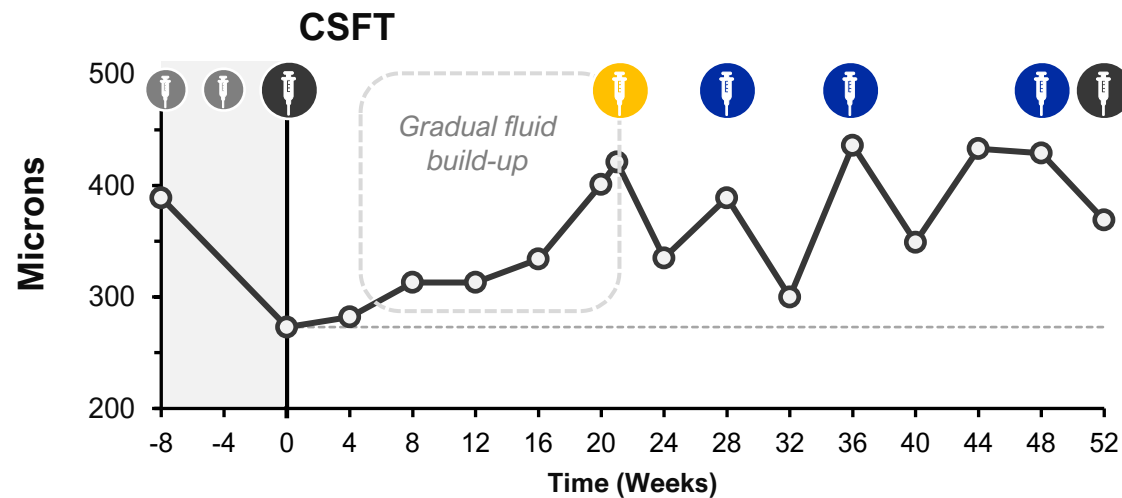
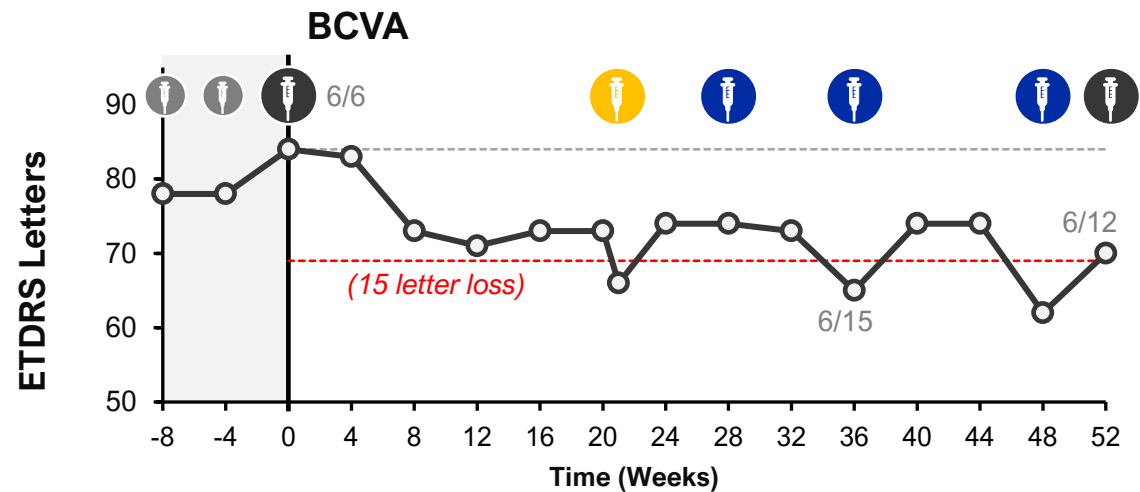
Well-Tolerated

OTX-TKI is an investigational product candidate which has not been approved by the FDA or any regulatory authority.

Clinical Evidence

Adrian Fung, MD

Aflibercept Case 1



Screening Aflibercept (2mg)

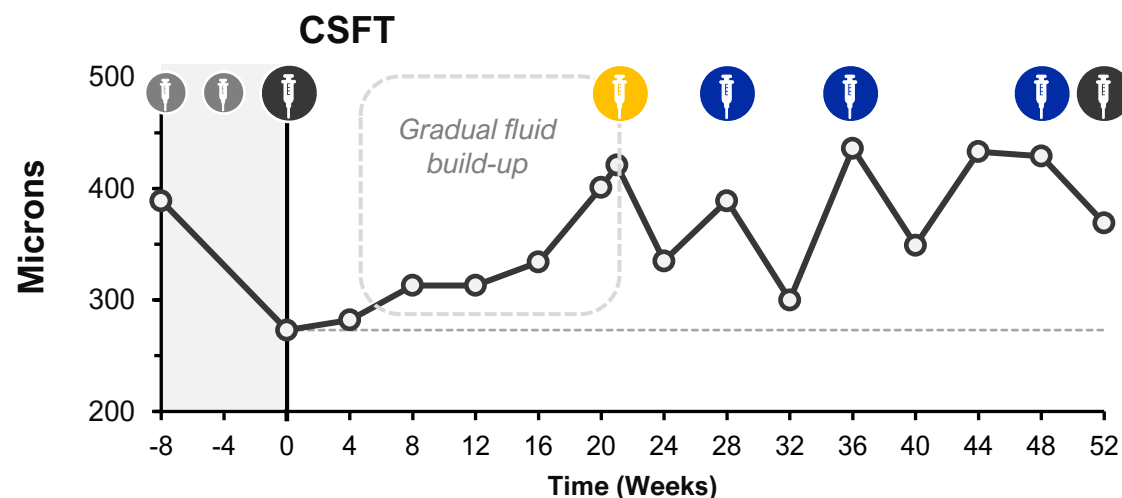
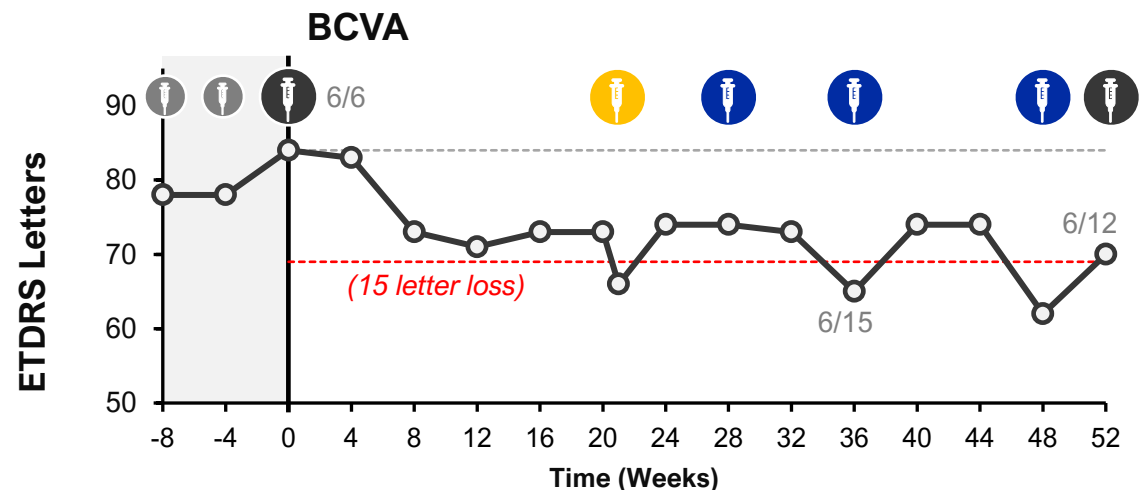
Study Aflibercept (2 mg)

Per-Protocol Rescue Aflibercept (2mg)

Investigator Discretion Rescue

-8 Weeks	BCVA: 78 Ltrs CSFT: 389 μ m	
Baseline	BCVA: 84 Ltrs Metric Snellen: 6/6 CSFT: 273 μ m	
Week 4	BCVA (Δ): 83 (-1) Ltrs CSFT (Δ): 282 (+9) μ m	
Week 8	BCVA (Δ): 73 (-11) Ltrs CSFT (Δ): 313 (+40) μ m	
Week 12	BCVA (Δ): 71 (-13) Ltrs CSFT (Δ): 313 (+40) μ m	
Week 16	BCVA (Δ): 73 (-11) Ltrs CSFT (Δ): 334 (+61) μ m	
Week 21 unscheduled	BCVA (Δ): 66 (-18) Ltrs CSFT (Δ): 421 (+148) μ m	

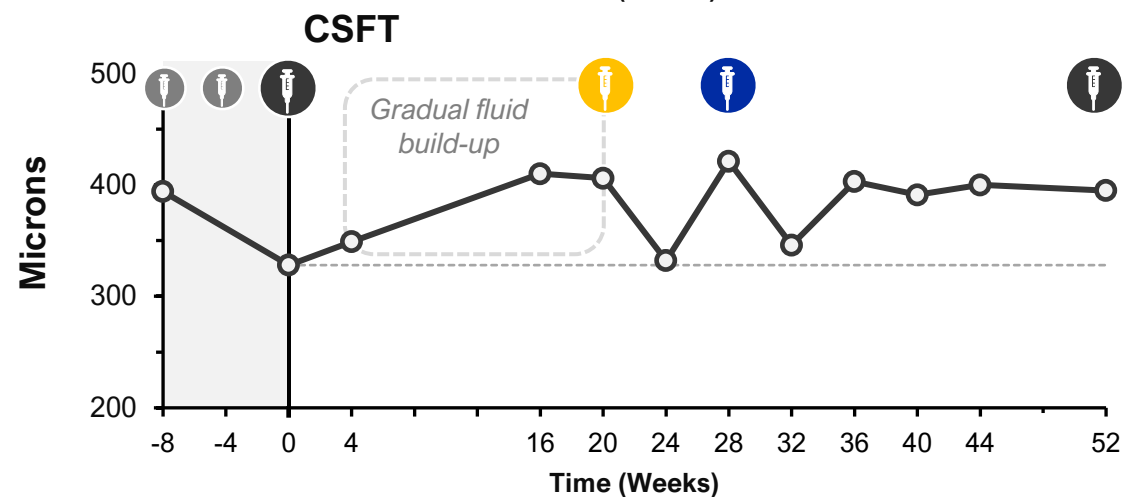
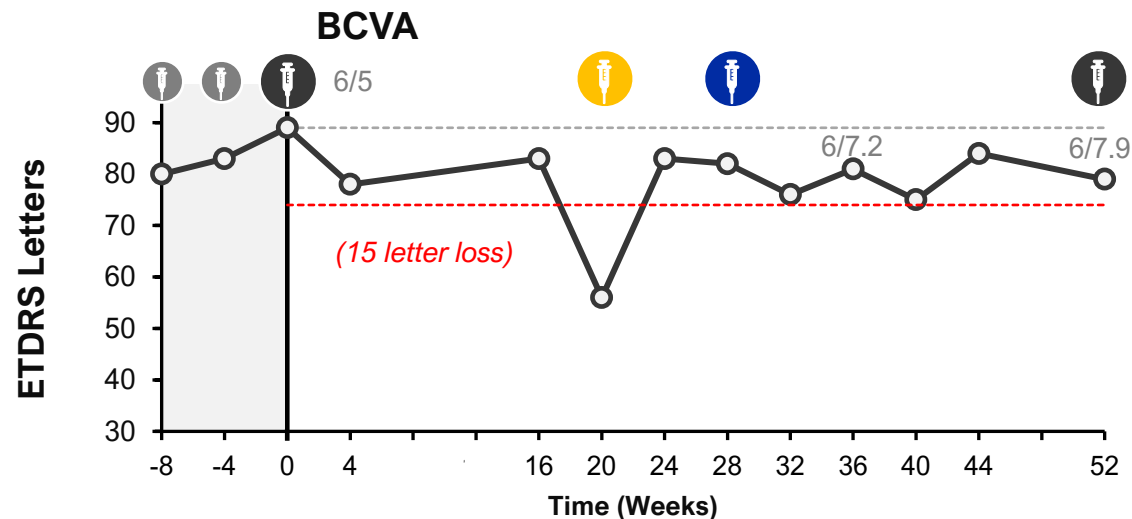
Aflibercept Case 1



Screening Aflibercept (2mg) Study Aflibercept (2 mg) Per-Protocol Rescue Aflibercept (2mg) Investigator Discretion Rescue

Week 24	BCVA (Δ): 74 (-10) Ltrs CSFT (Δ): 335 (+62) μ m		
Week 28	BCVA (Δ): 74 (-10) Ltrs CSFT (Δ): 389 (+116) μ m		
Week 32	BCVA (Δ): 73 (-11) Ltrs CSFT (Δ): 300 (+27) μ m		
Week 36	BCVA (Δ): 65 (-19) Ltrs Metric Snellen: 6/15 CSFT (Δ): 436 (+163) μ m		
Week 48	BCVA (Δ): 62 (-22) Ltrs CSFT (Δ): 429 (+156) μ m		
Week 52	BCVA (Δ): 70 (-14) Ltrs Metric Snellen: 6/12 CSFT (Δ): 369 (+96) μ m		

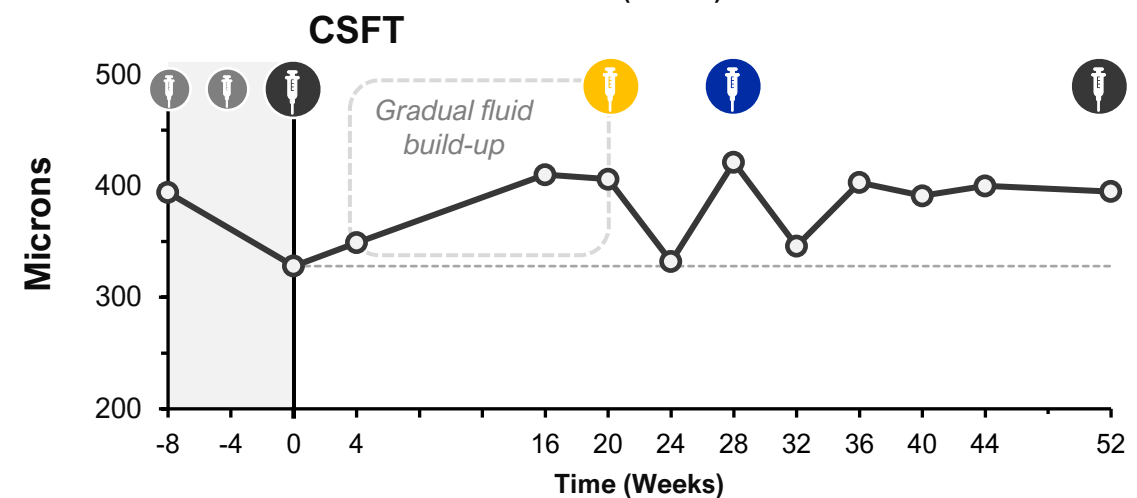
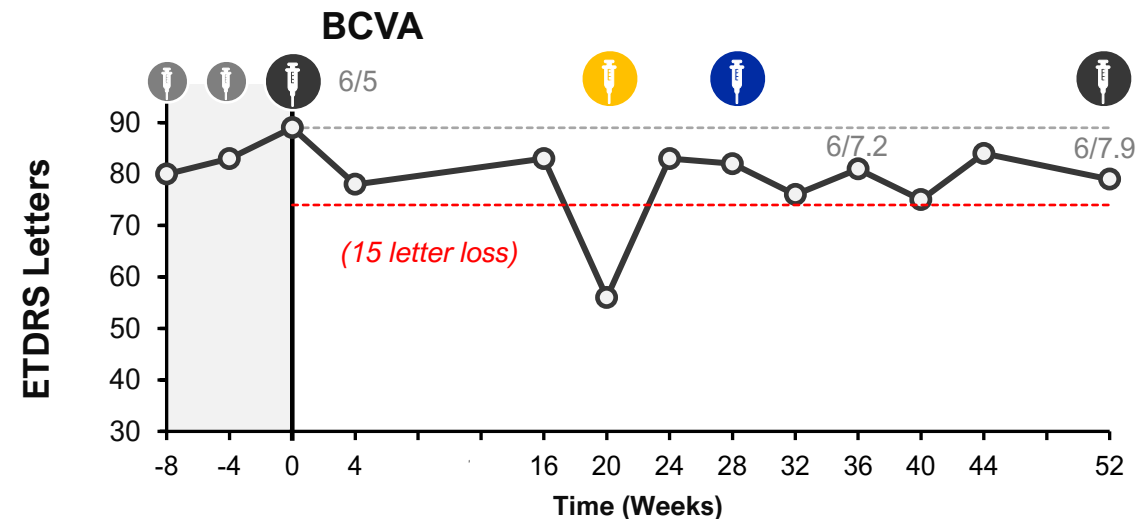
Aflibercept Case 2



Screening Aflibercept (2mg)
 Study Aflibercept (2 mg)
 Per-Protocol Rescue Aflibercept (2mg)
 Investigator Discretion Rescue

-8 Weeks	BCVA: 80 Ltrs CSFT: 394 μ m	
Baseline	BCVA: 89 Ltrs Metric Snellen: 6/5 CSFT: 328 μ m	
Week 4	BCVA (Δ): 78 (-11) Ltrs CSFT (Δ): 349 (+21) μ m	
Week 16	BCVA (Δ): 83 (-6) Ltrs CSFT (Δ): 410 (+82) μ m	
Week 20	BCVA (Δ): 56 (-33) Ltrs CSFT (Δ): 406 (+78) μ m	
Week 24	BCVA (Δ): 83 (-6) Ltrs CSFT (Δ): 332 (+4) μ m	
Week 28	BCVA (Δ): 82 (-7) Ltrs CSFT (Δ): 421 (+93) μ m	
Week 36	BCVA (Δ): 81(-8) Ltrs Metric Snellen: 6/7.2 CSFT (Δ): 403 (+75) μ m	

Aflibercept Case 2

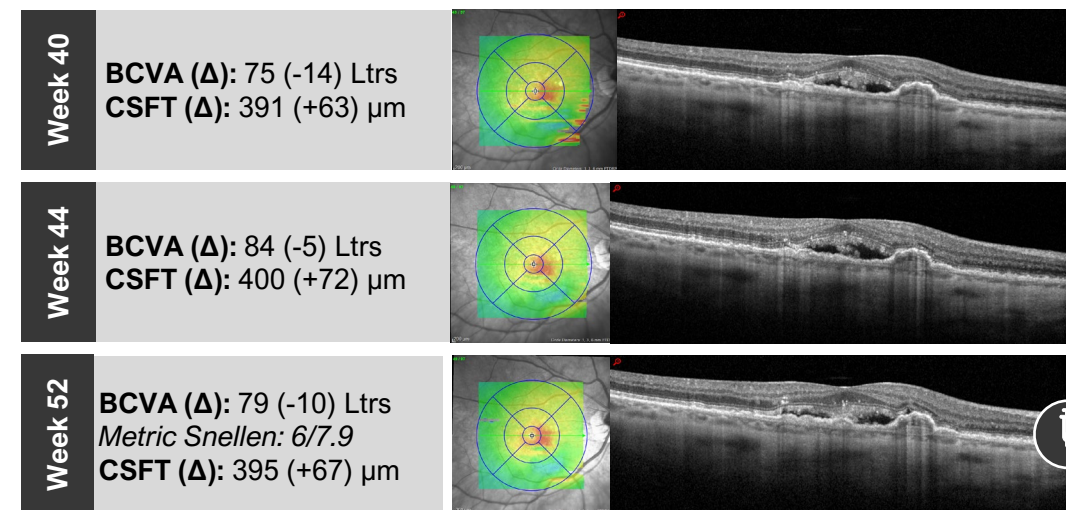


Screening Aflibercept (2mg)

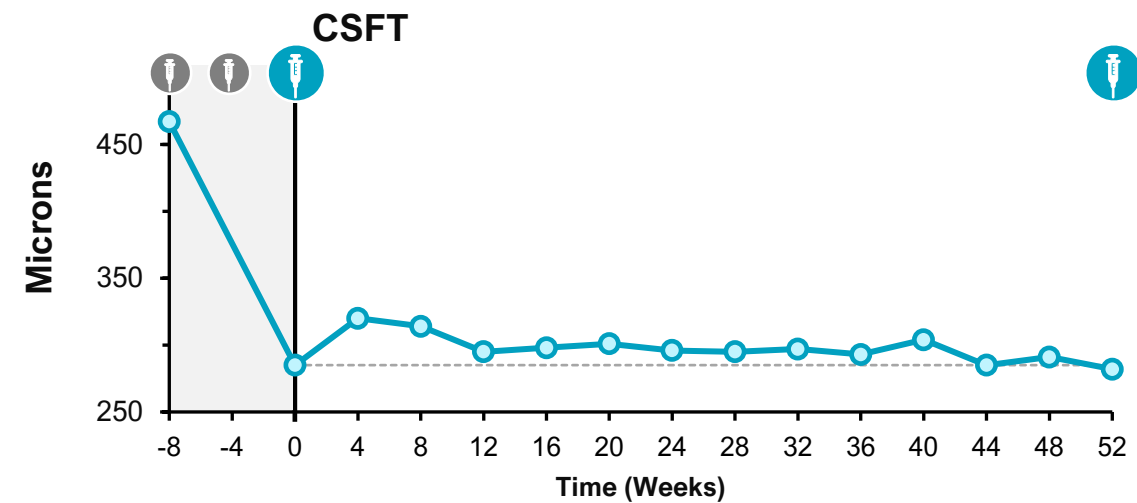
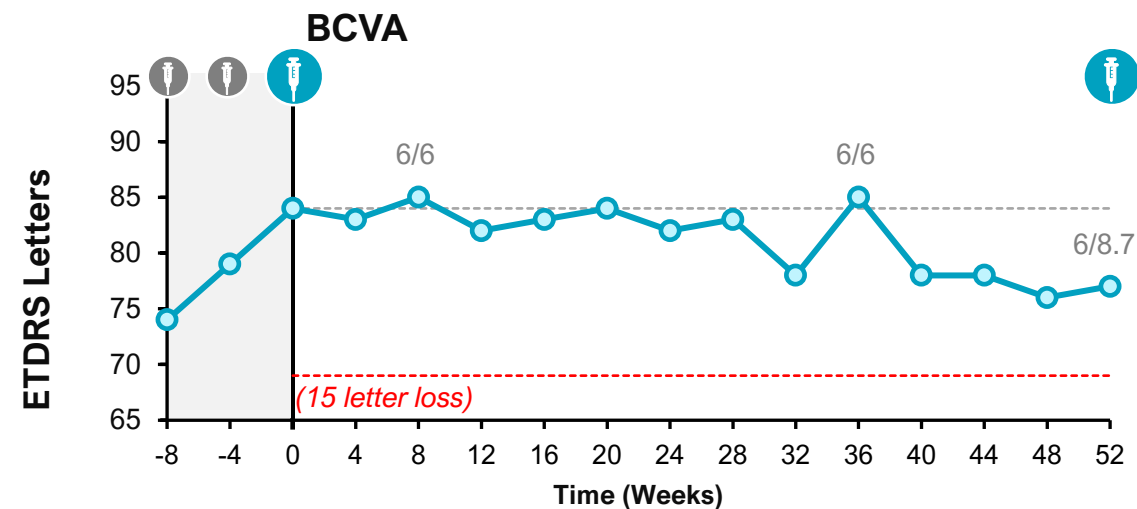
Study Aflibercept (2 mg)

Per-Protocol Rescue Aflibercept (2mg)

Investigator Discretion Rescue



OTX-TKI Case 1



Screening
Aflibercept (2mg)

OTX-TKI
(0.45mg)

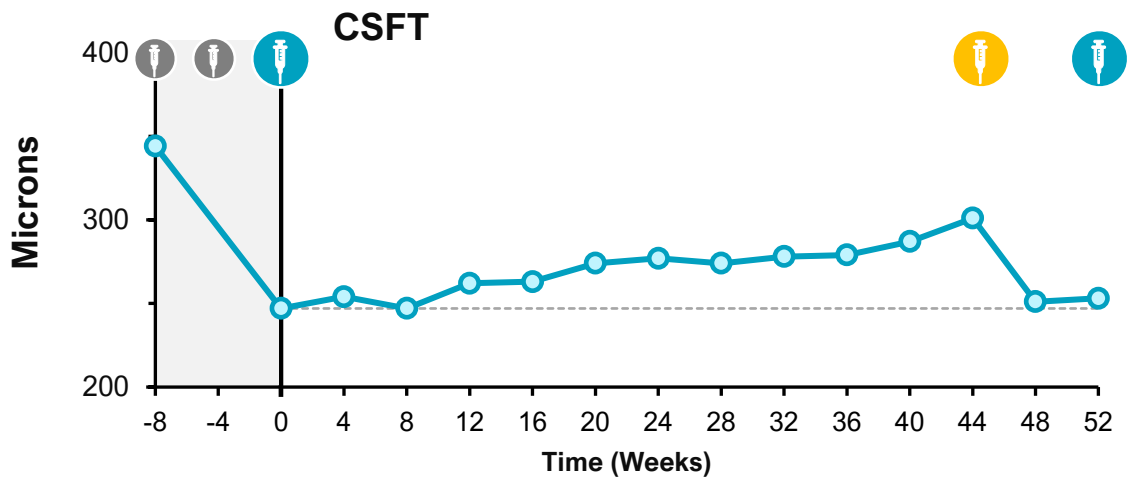
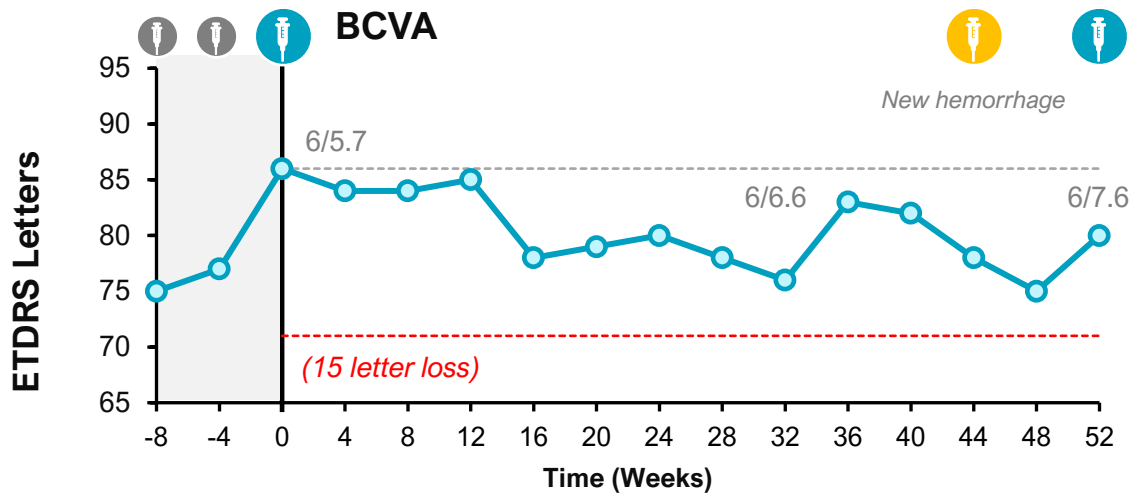


Retina
experience
redefined

Case study of one subject; individual results may vary
BCVA, best-corrected visual acuity; CSFT, central subfield thickness; ETDRS, Early Treatment Diabetic Retinopathy Study; Ltrs, letters

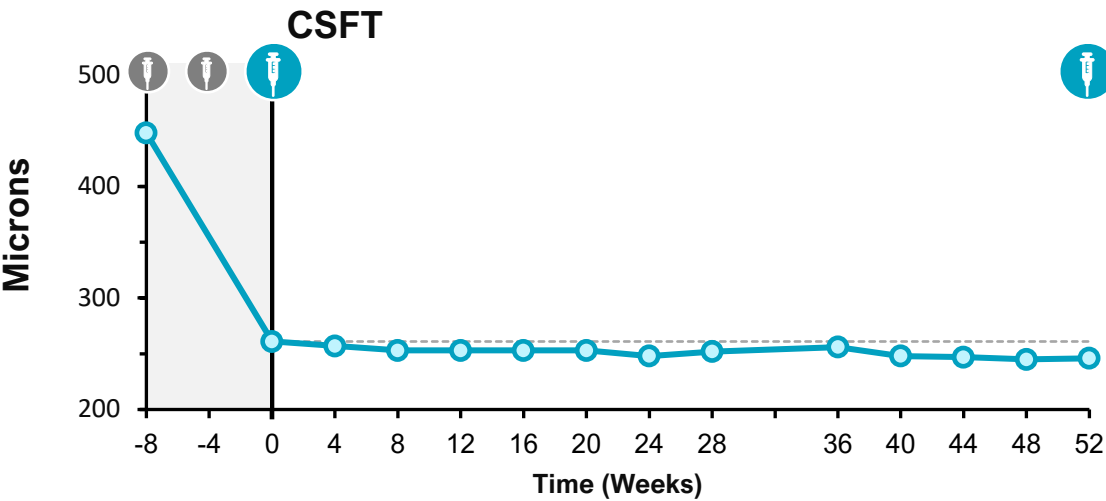
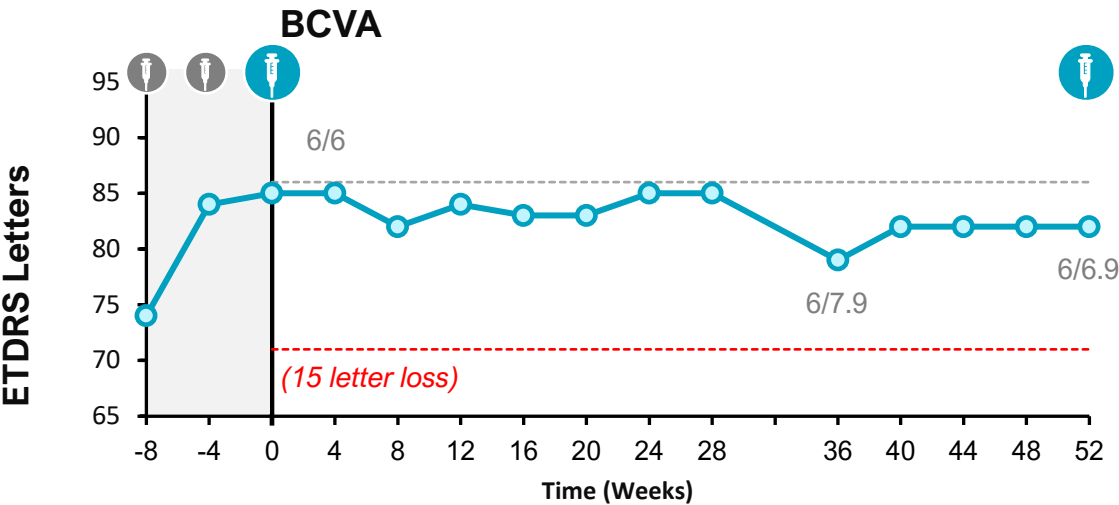
-8 Weeks	BCVA: 74 Ltrs (-10) CSFT: 467 μ m	
Baseline	BCVA: 84 Ltrs Metric Snellen: 6/6 CSFT: 285 μ m	
Week 4	BCVA (Δ): 83 (-1) Ltrs CSFT (Δ): 320 (+35) μ m	
Week 12	BCVA (Δ): 82 (-2) Ltrs CSFT (Δ): 295 (+10) μ m	
Week 24	BCVA (Δ): 82 (-2) Ltrs CSFT (Δ): 296 (+11) μ m	
Week 36	BCVA (Δ): 85 (+1) Ltrs Metric Snellen: 6/6 CSFT (Δ): 293 (+8) μ m	
Week 44	BCVA (Δ): 78 (-6) Ltrs CSFT (Δ): 285 (0) μ m	
Week 52	BCVA (Δ): 77 (-7) Ltrs Metric Snellen: 6/8.7 CSFT (Δ): 282 (-3) μ m	

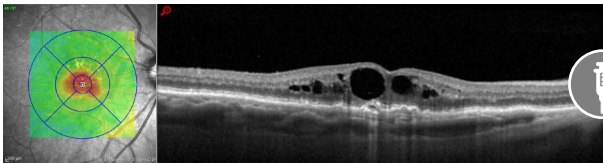
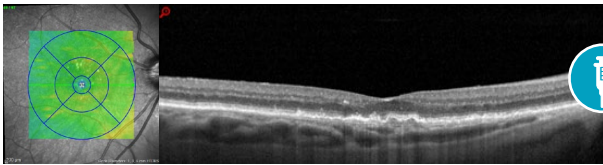
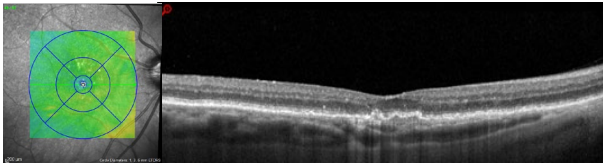
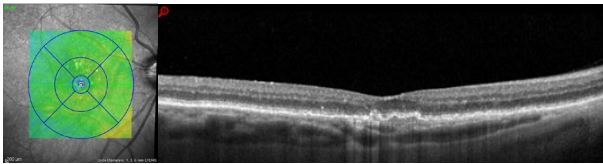
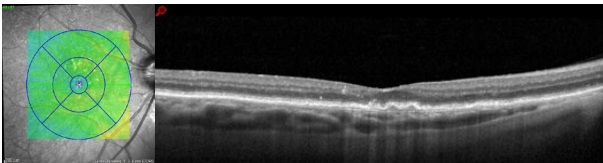
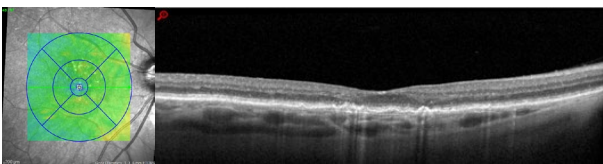
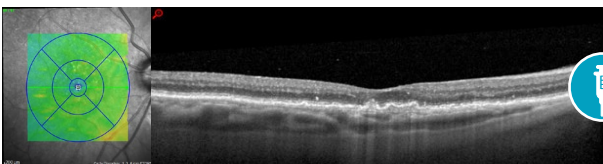
OTX-TKI Case 2



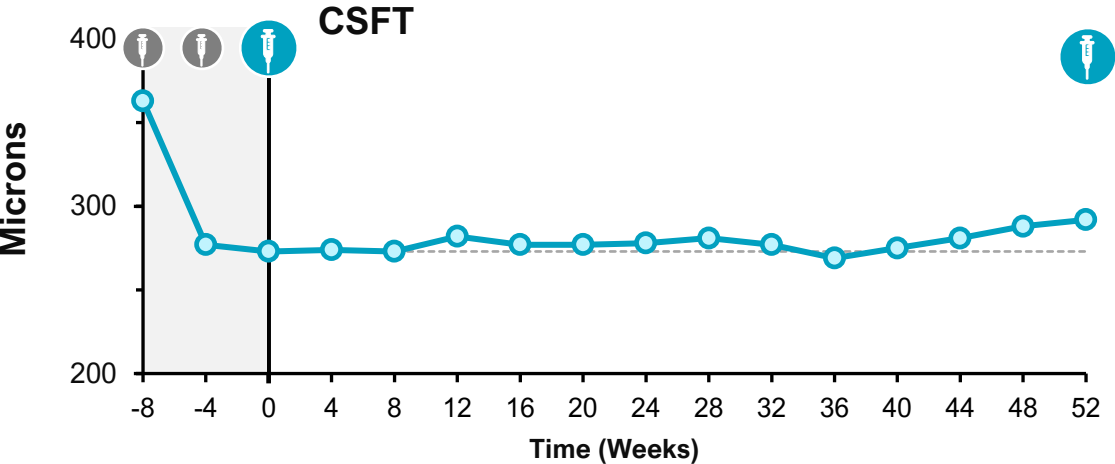
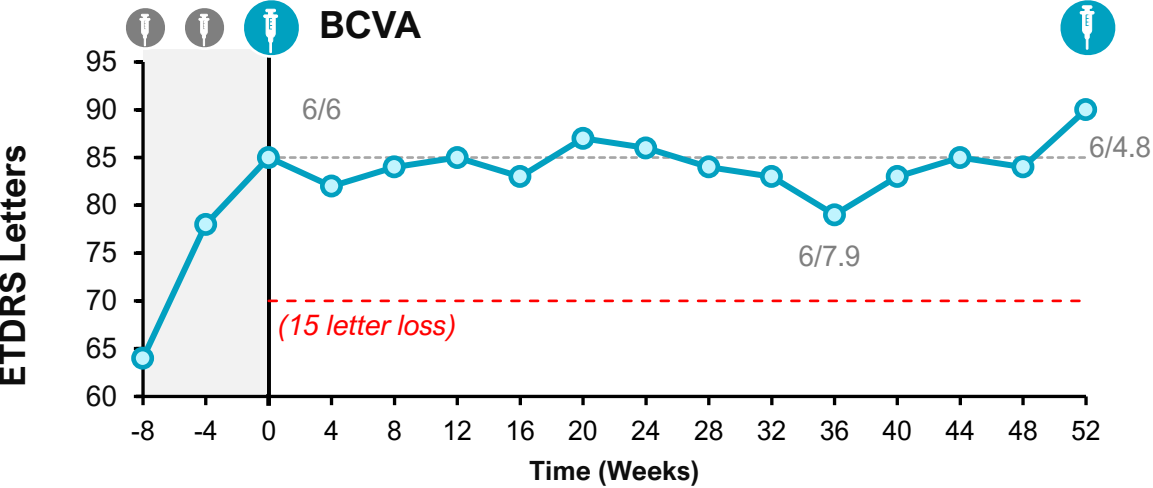
-8 Weeks	BCVA: 75 Ltrs CSFT: 344 µm	
Baseline	BCVA: 86 Ltrs Metric Snellen: 6/5.7 CSFT: 247 µm	
Week 4	BCVA (Δ): 84 (-2) Ltrs CSFT (Δ): 254 (+7) µm	
Week 12	BCVA (Δ): 85 (-1) Ltrs CSFT (Δ): 262 (+15) µm	
Week 24	BCVA (Δ): 80 (-6) Ltrs CSFT (Δ): 277 (+30) µm	
Week 36	BCVA (Δ): 83 (-3) Ltrs Metric Snellen: 6/6.6 CSFT (Δ): 279 (+32) µm	
Week 44	BCVA (Δ): 78 (-8) Ltrs CSFT (Δ): 301 (+54) µm	
Week 52	BCVA (Δ): 80 (-6) Ltrs Metric Snellen: 6/7.6 CSFT (Δ): 253 (+6) µm	

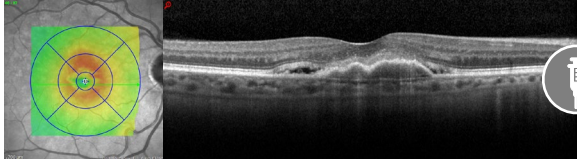
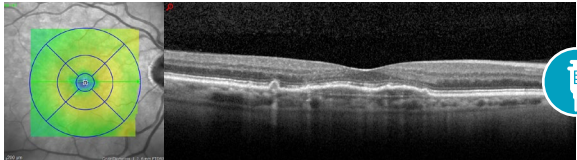
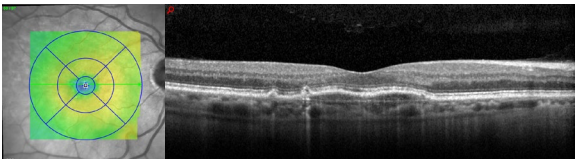
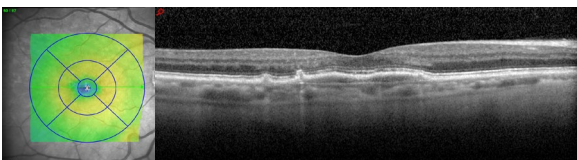
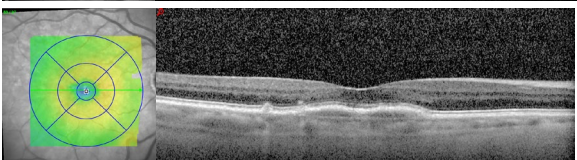
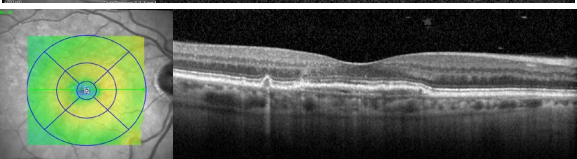
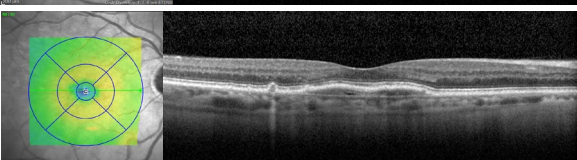
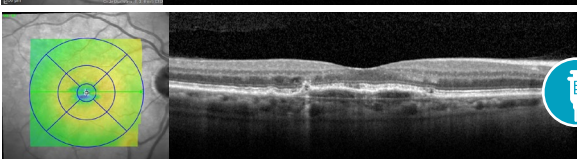
OTX-TKI Case 3



-8 Weeks	BCVA: 74 Ltrs CSFT: 448 µm	
Baseline	BCVA: 85 Ltrs Metric Snellen: 6/6 CSFT: 261 µm	
Week 4	BCVA (Δ): 85 (0) Ltrs CSFT (Δ): 257 (-4) µm	
Week 12	BCVA (Δ): 84 (-1) Ltrs CSFT (Δ): 253 (-8) µm	
Week 24	BCVA (Δ): 85 (0) Ltrs CSFT (Δ): 248 (-13) µm	
Week 36	BCVA (Δ): 79 (-6) Ltrs Metric Snellen: 6/7.9 CSFT (Δ): 256 (-5) µm	
Week 52	BCVA (Δ): 82 (-3) Ltrs Metric Snellen: 6/6.9 CSFT (Δ): 246 (-15) µm	

OTX-TKI Case 4

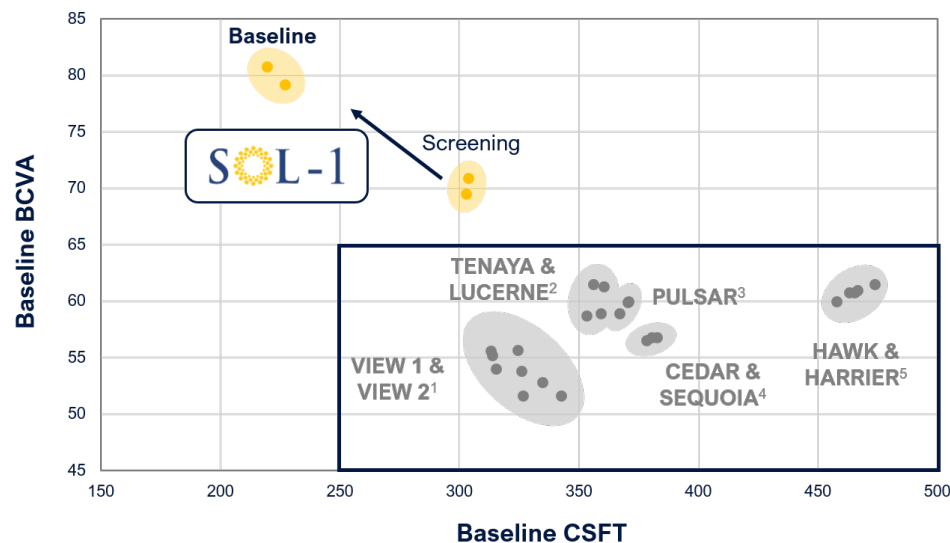


-8 Weeks	BCVA: 64 Ltrs CSFT: 363 μ m	
Baseline	BCVA: 85 Ltrs Metric Snellen: 6/6 CSFT: 273 μ m	
Week 4	BCVA (Δ): 82 (-3) Ltrs CSFT (Δ): 274 (+1) μ m	
Week 16	BCVA (Δ): 83 (-2) Ltrs CSFT (Δ): 277 (+4) μ m	
Week 24	BCVA (Δ): 86 (+1) Ltrs CSFT (Δ): 278 (+5) μ m	
Week 36	BCVA (Δ): 79 (-6) Ltrs Metric Snellen: 6/7.9 CSFT (Δ): 269 (-4) μ m	
Week 44	BCVA (Δ): 85 (0) Ltrs CSFT (Δ): 281 (+8) μ m	
Week 52	BCVA (Δ): 90 (+5) Ltrs Metric Snellen: 6/4.8 CSFT (Δ): 292 (+19) μ m	

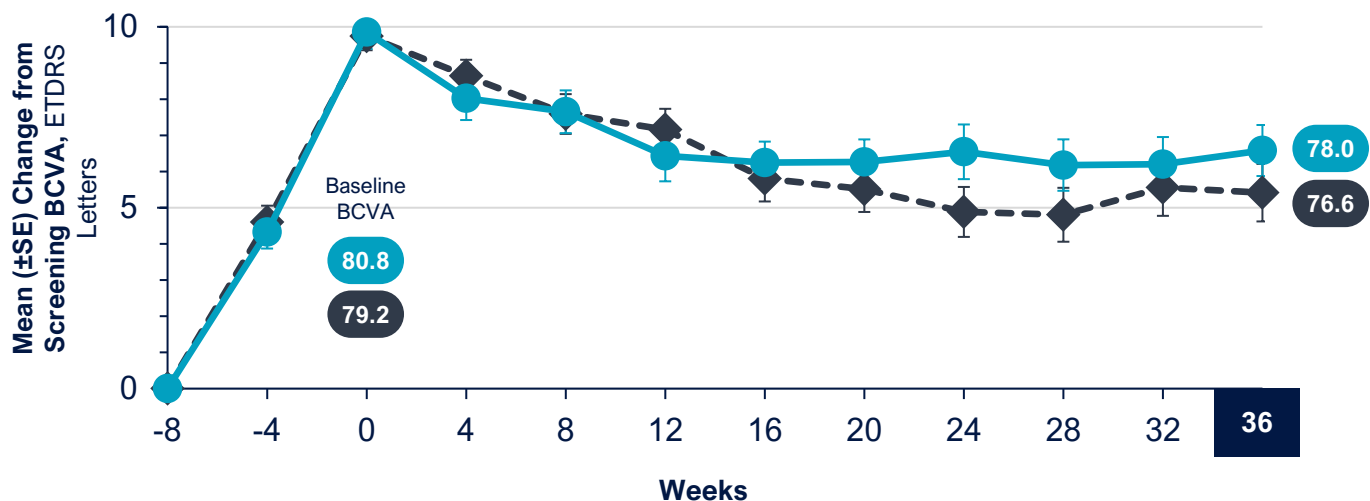
Key Insights

Adnan Tufail, MD

Baseline BCVA Across nAMD Trials



SOL-1 : Mean Change in BCVA from Screening
Post Hoc Analysis

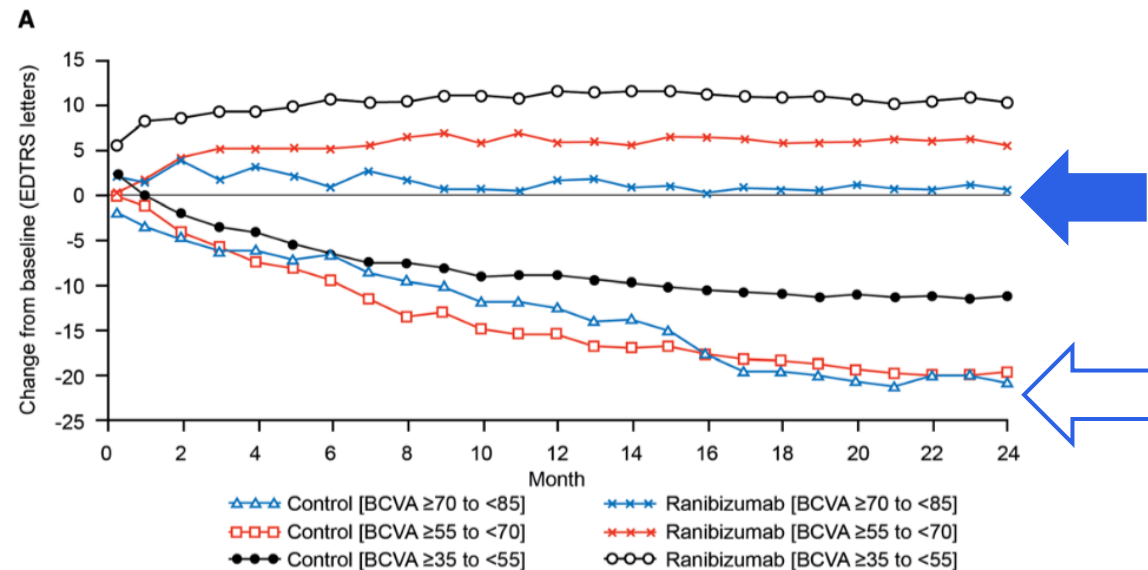


Does Reduced Potential for Visual Acuity Gain Occur in Pivotal Trials?

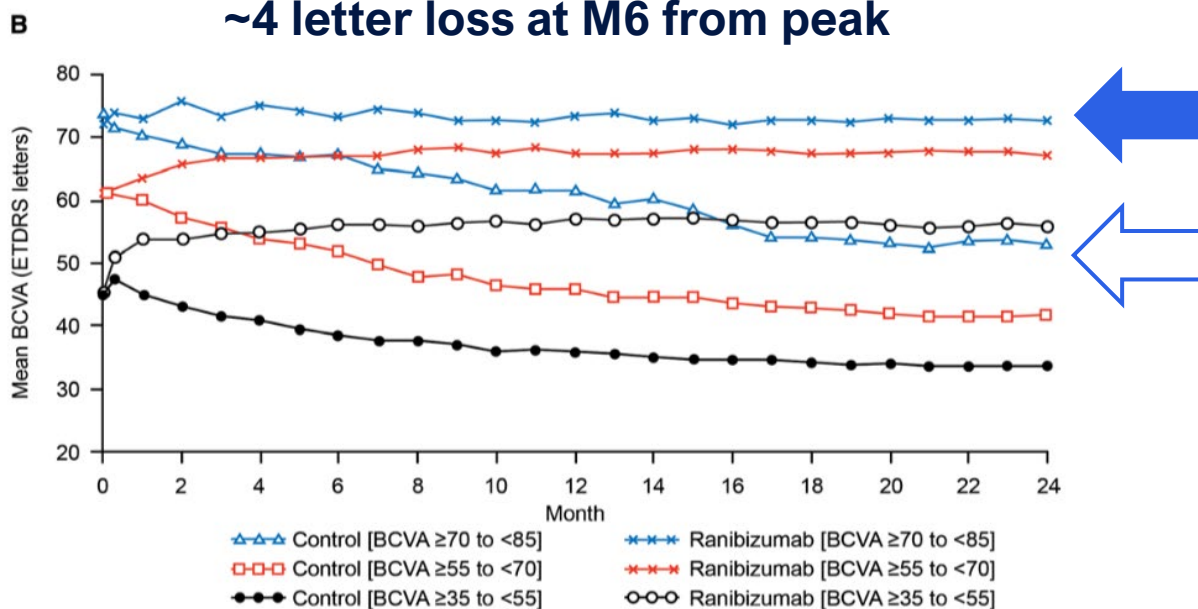
Sub-Analysis of ANCHOR and MARINA

Visual benefit versus visual gain: what is the effect of baseline covariants in the treatment arm relative to the control arm? A pooled analysis of ANCHOR and MARINA

Adnan Tufail¹, Philippe Margaron², Tadhg Guerin³, Michael Larsen⁴



In good VA group (baseline 70-85 letters),
~4 letter loss at M6 from peak

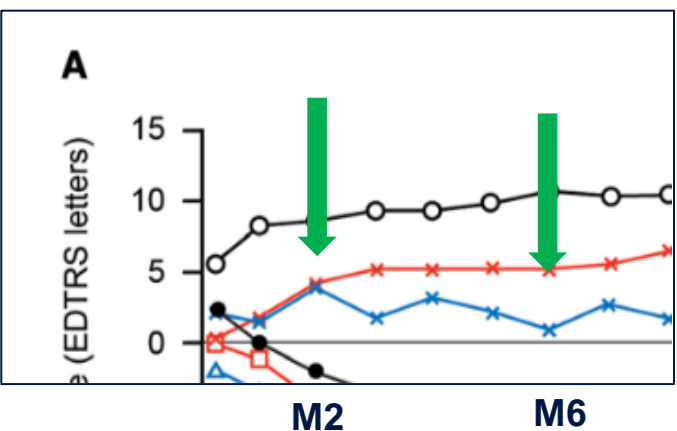
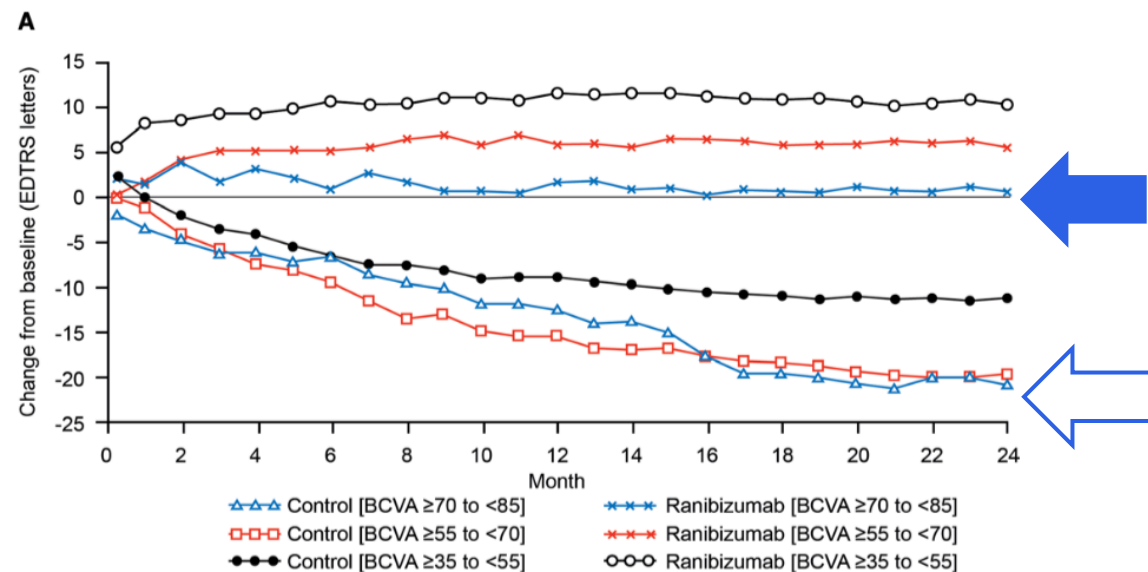


Does Reduced Potential for Visual Acuity Gain Occur in Pivotal Trials?

Sub-Analysis of ANCHOR and MARINA

Visual benefit versus visual gain: what is the effect of baseline covariants in the treatment arm relative to the control arm? A pooled analysis of ANCHOR and MARINA

Adnan Tufail ¹, Philippe Margaron, ² Tadhg Guerin, ³ Michael Larsen ⁴

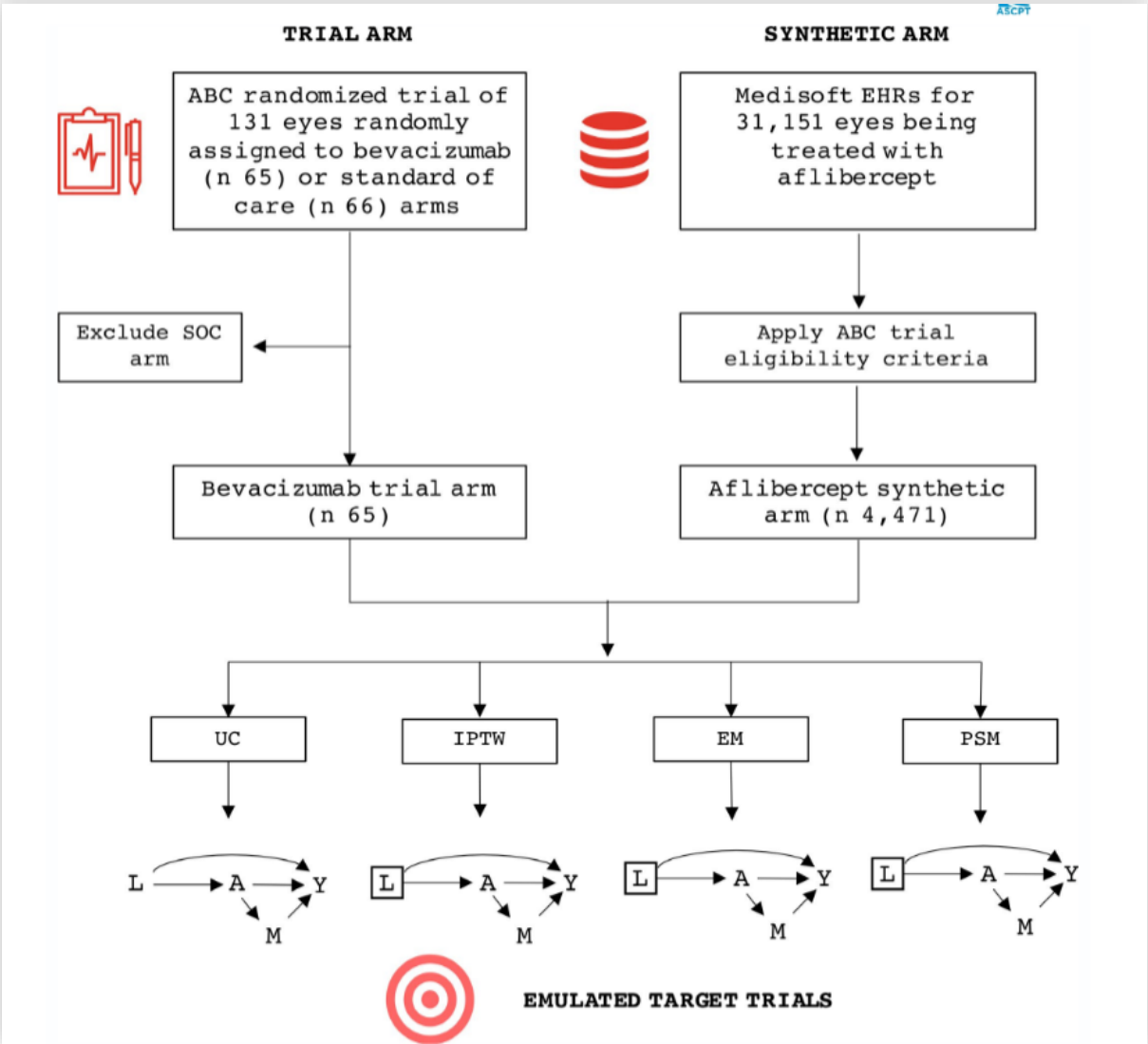


ARTICLE

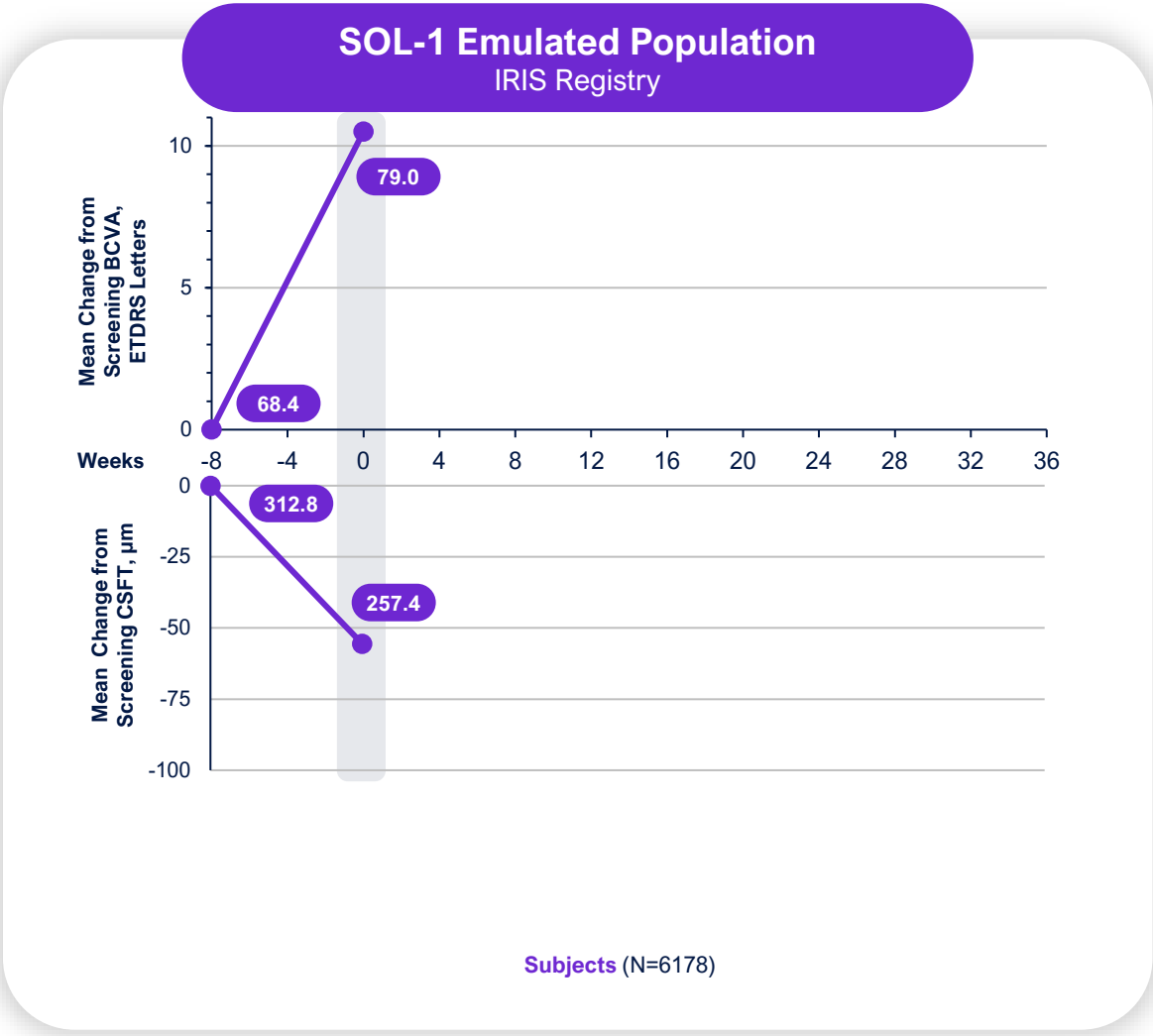
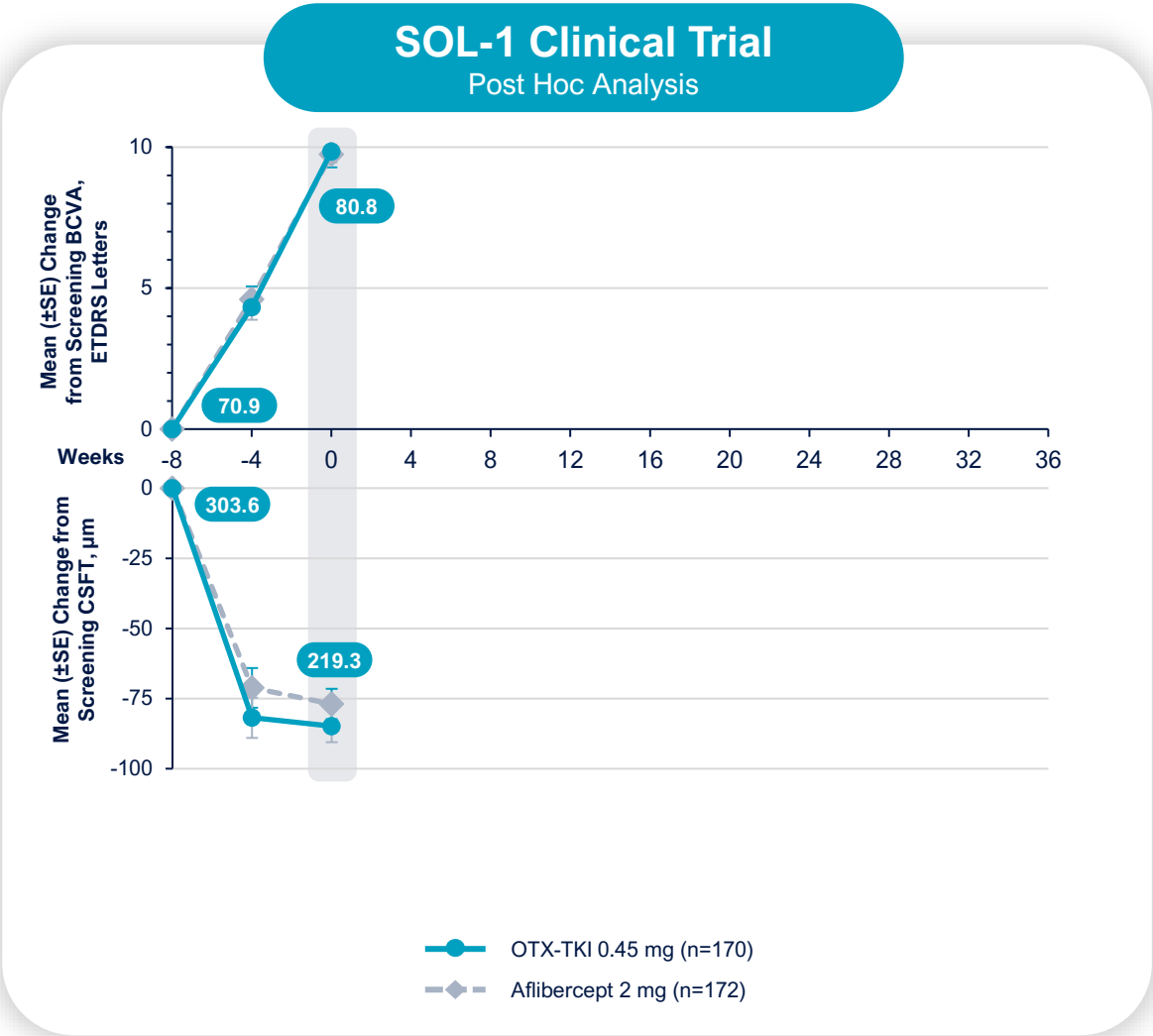
Contextualizing single-arm trials with real-world data: An emulated target trial comparing therapies for neovascular age-related macular degeneration

Darren S. Thomas¹ | Aaron Y. Lee² | Philipp L. Müller^{3,4,5} | Roy Schwartz^{4,5,6} | Abraham Olvera-Barrios^{4,5} | Alasdair N. Warwick^{4,7} | Praven J. Patel⁶ | Tjebo F.C. Heeren^{4,5} | Catherine Egan^{4,5,6} | Paul Taylor¹ | Adnan Tufail^{4,5,6} | on behalf of the UK AMD EMR Users Group⁷

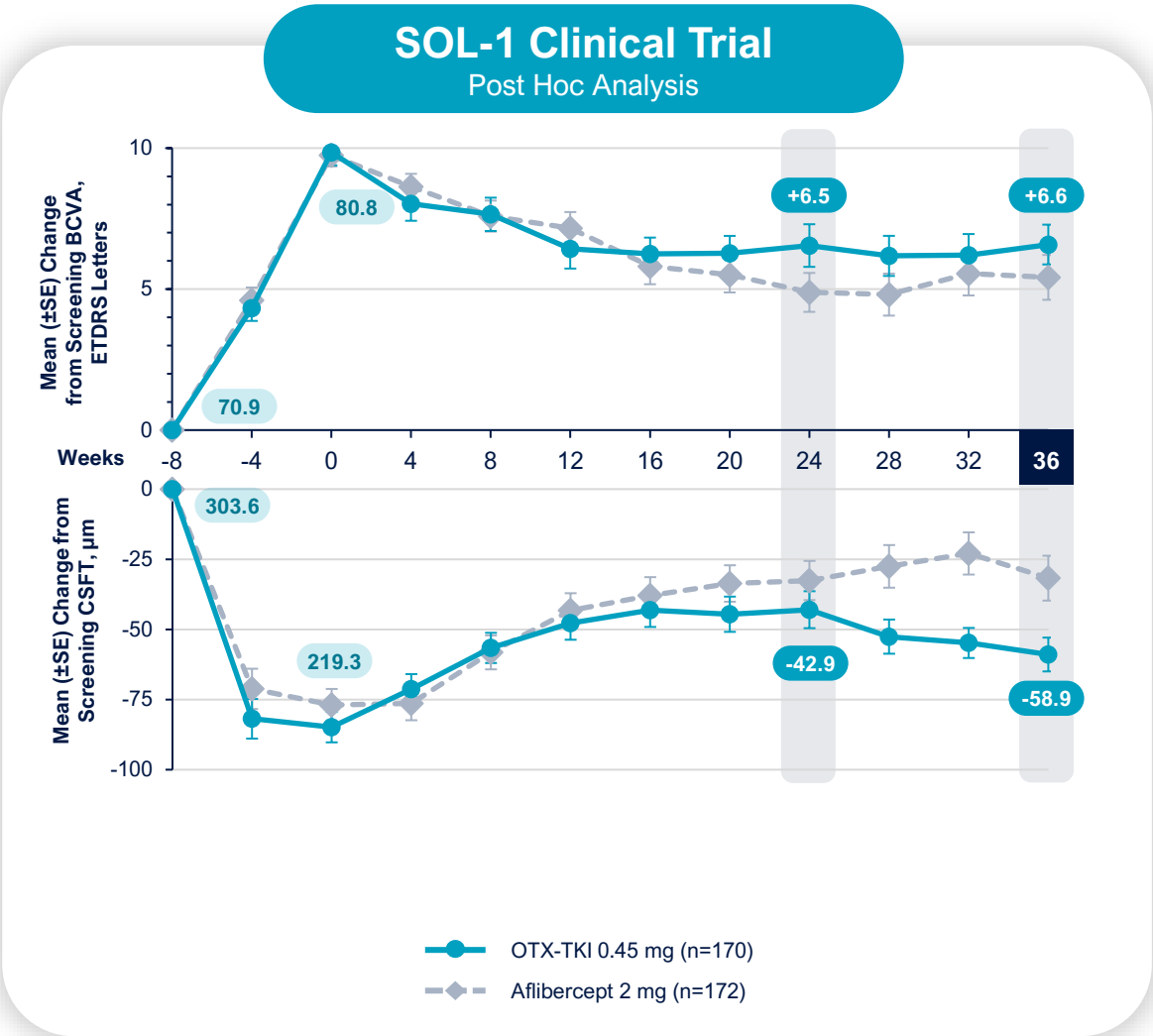
Using a similar methodology, we emulated SOL-1 entry criteria to the IRIS Registry



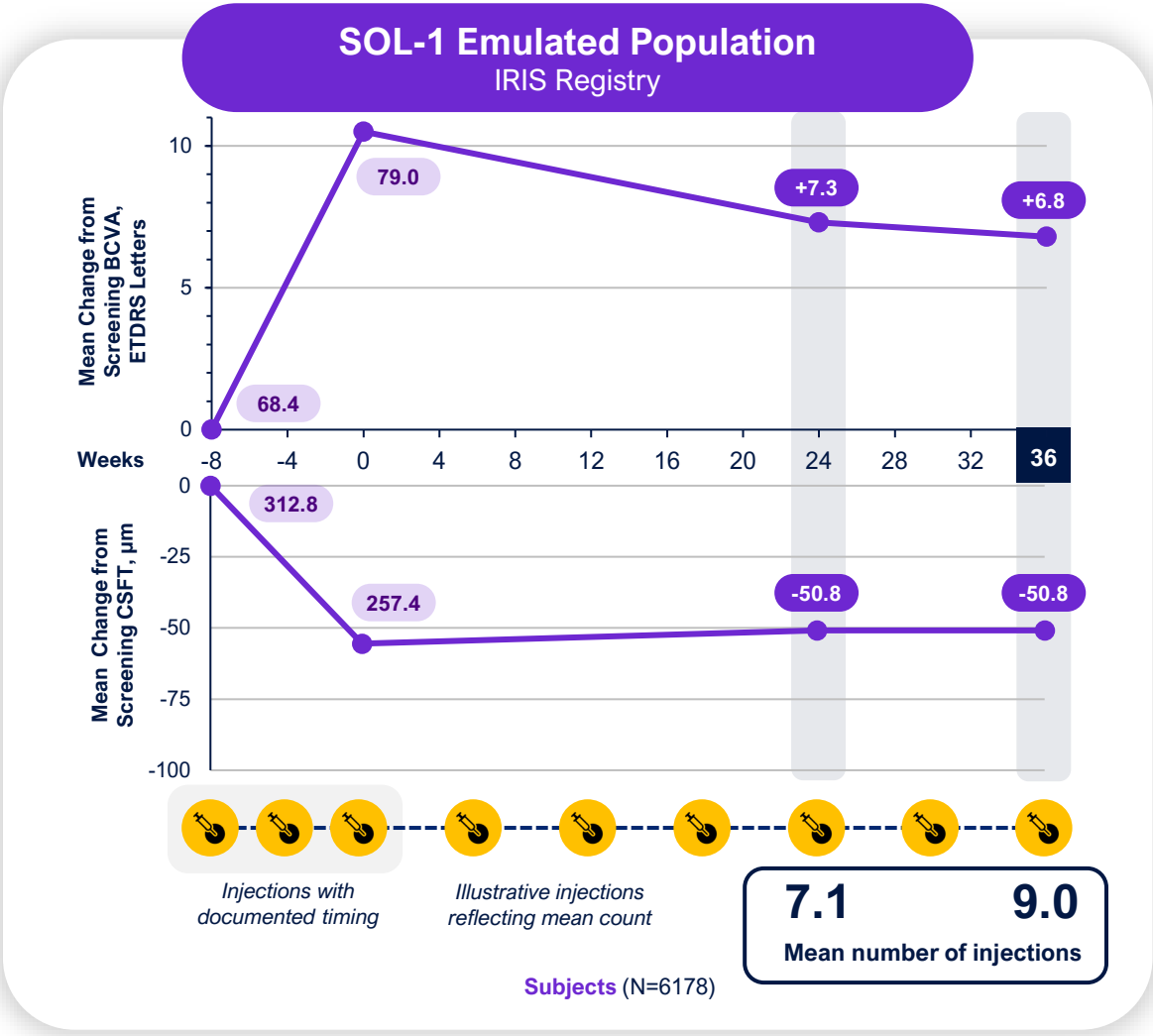
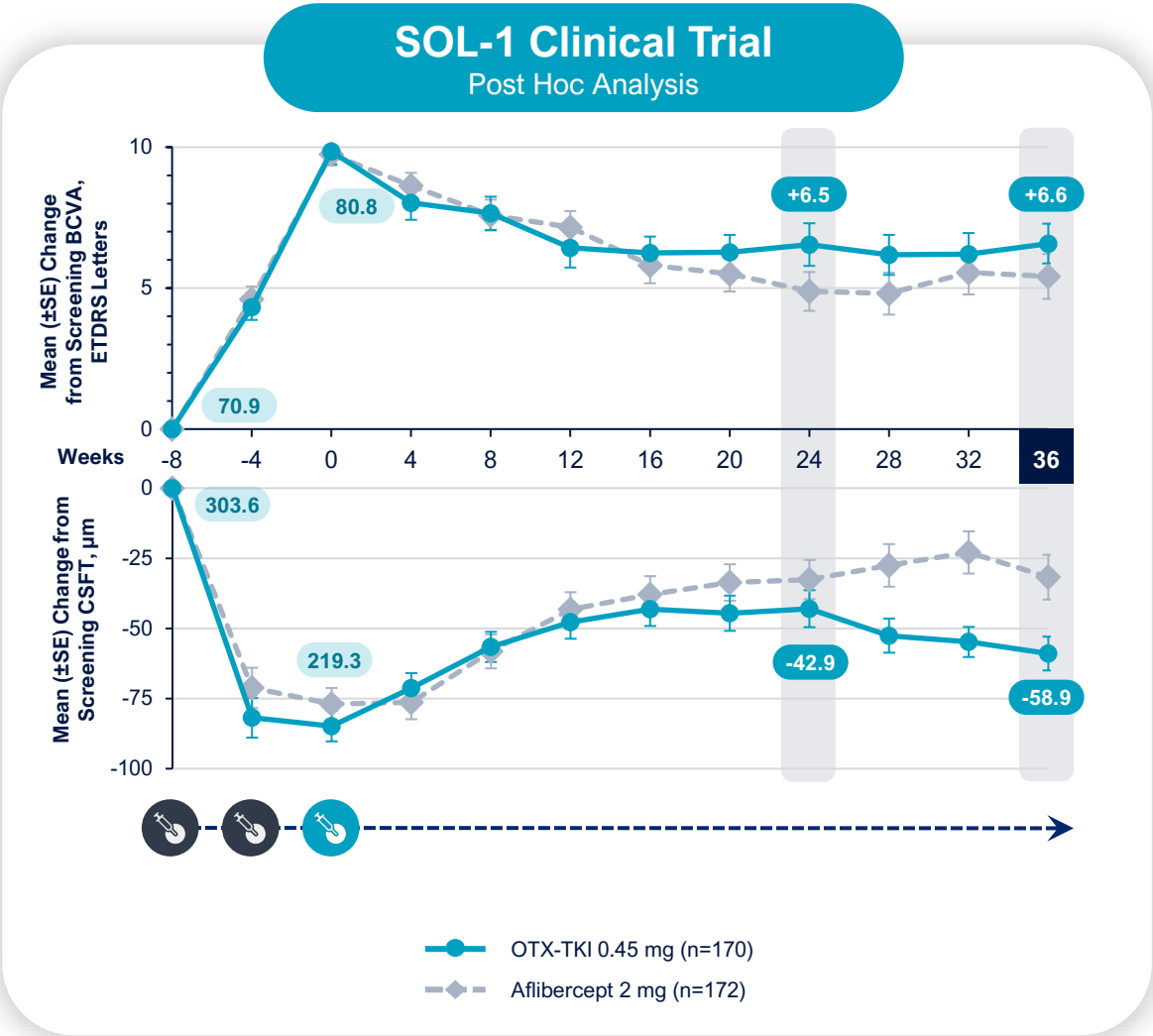
How Does SOL-1 Data Align with Real World Outcomes?



How Does SOL-1 Data Align with Real World Outcomes?

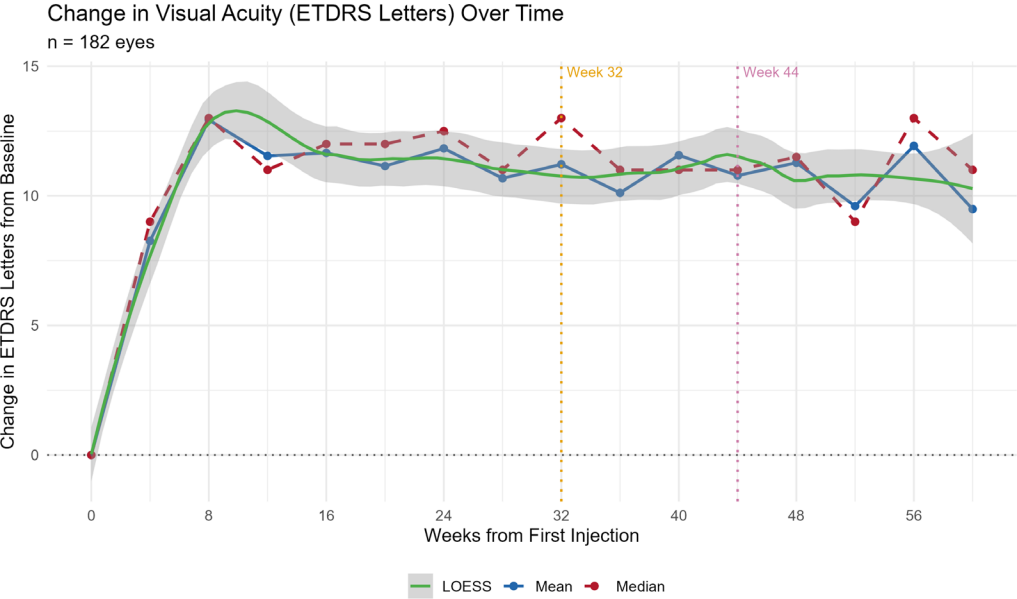


How Does SOL-1 Data Align with Real World Outcomes?



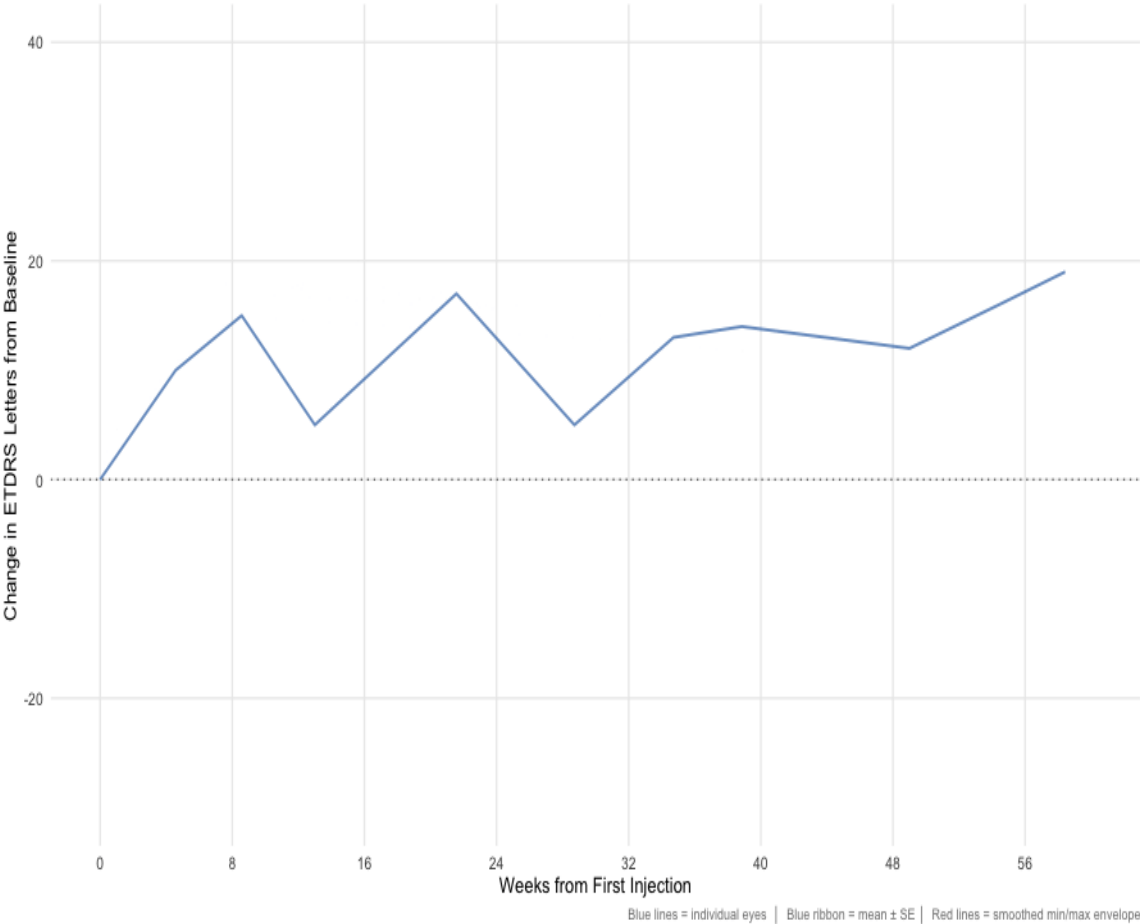
How Does SOL-1 Data Align with Real World Outcomes?

UK Dataset (2022 -2025)



Change in Visual Acuity (ETDRS Letters) Over Time

Eyes: 1 / 182

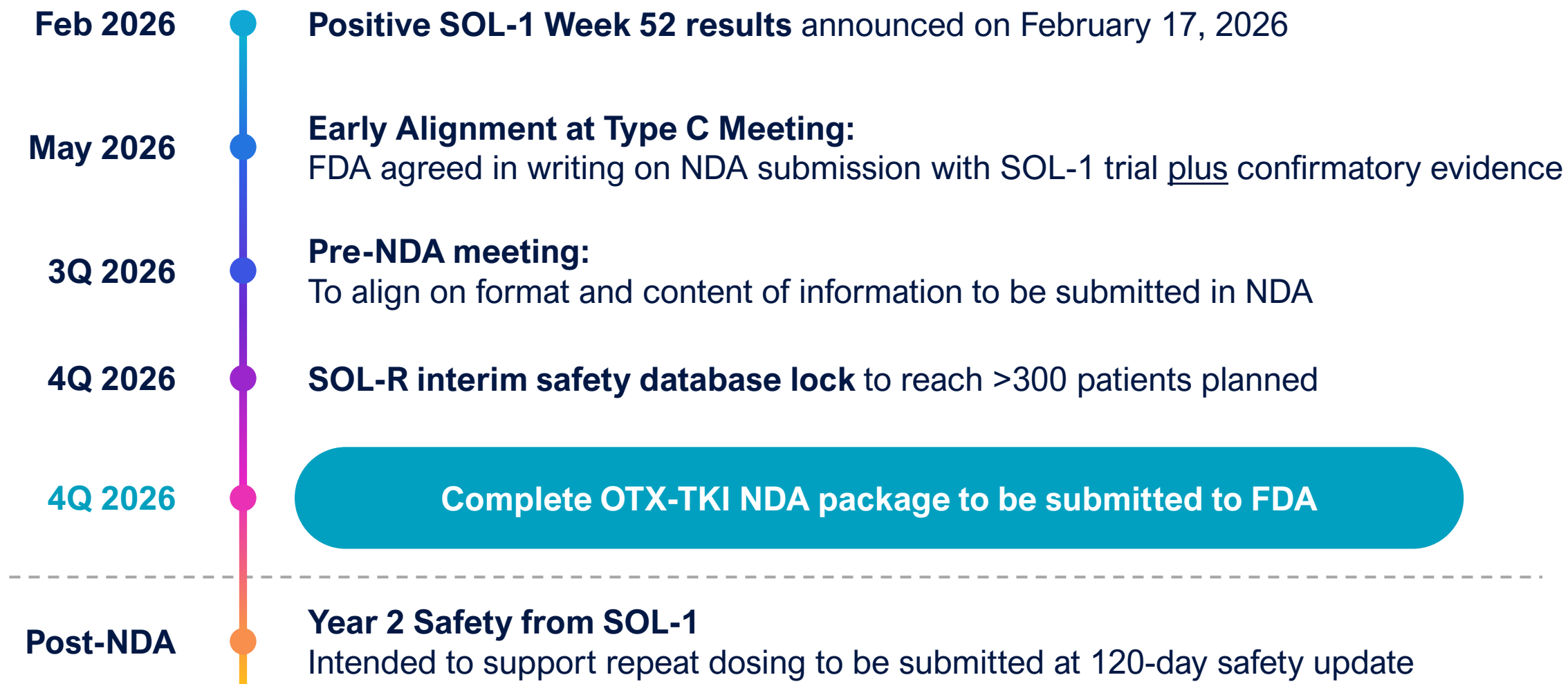


Age, mean = 78.9 years
Screen VA, mean = 65 letters
Baseline VA, mean = 78 letters

Clinical Program Update

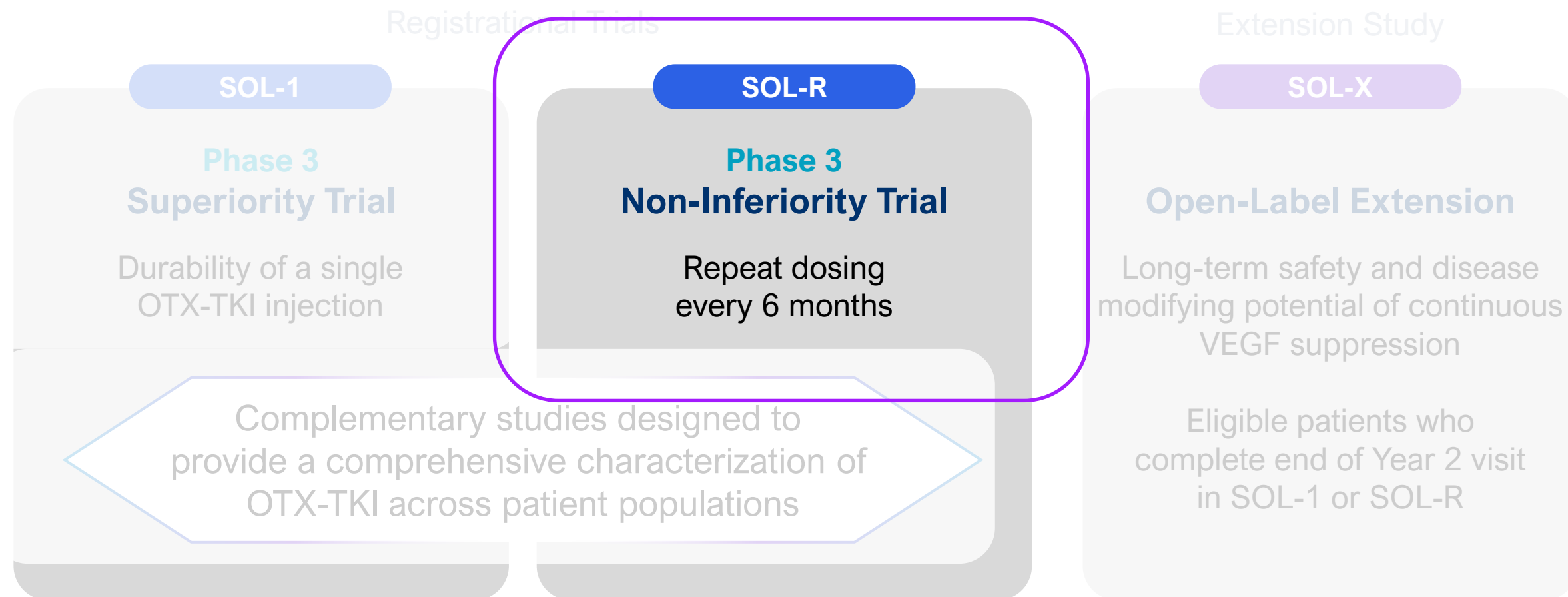
Jeffrey Heier, MD

Timeline to OTX-TKI NDA Submission Package for nAMD

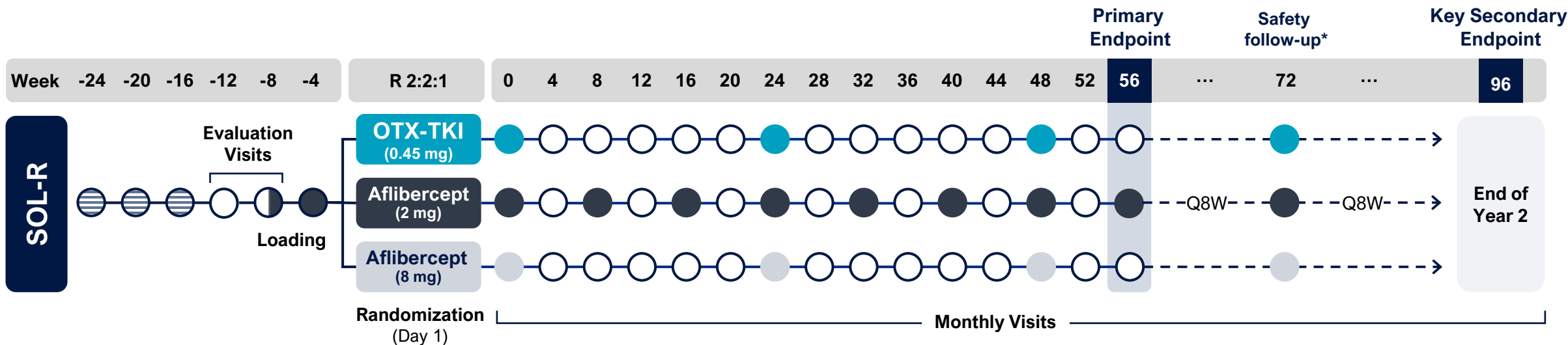


SOL: OTX-TKI Phase 3 Clinical Program in nAMD

Evaluating the efficacy, durability and safety of OTX-TKI in nAMD



SOL-R: Evaluation of with OTX-TKI relative to aflibercept 2 mg and 8 mg



Aflibercept 8mg dosing identical with no sham injections

Primary analysis:

OTX-TKI dosed Q24W vs.
aflibercept 2mg Q8W

Key Secondary Endpoint:

Superiority comparison at Year 2
vs. aflibercept 8mg

Secondary Endpoint:

Evaluating prevention of fibrosis
& atrophy vs. aflibercept 2mg

SOL: OTX-TKI Phase 3 Clinical Program in nAMD

Evaluating the efficacy, durability and safety of OTX-TKI in nAMD

Registrational Trials

SOL-1

Phase 3 Superiority Trial

Durability of a single
OTX-TKI injection

SOL-R

Phase 3 Non-Inferiority Trial

Repeat dosing
every 6 months

Complementary studies designed to
provide a comprehensive characterization of
OTX-TKI across patient populations

Extension Study

SOL-X

Open-Label Extension

Long-term safety and disease
modifying potential of continuous
VEGF suppression

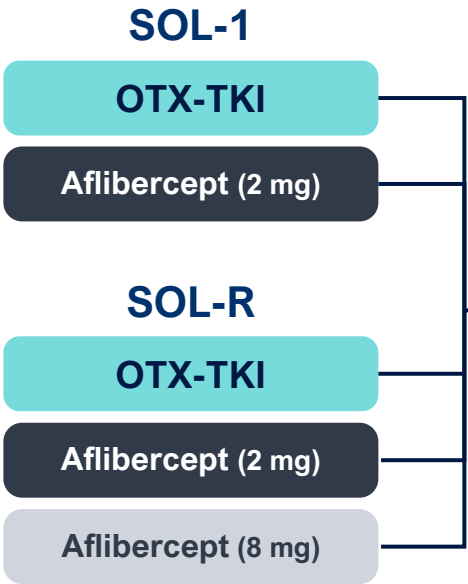
Eligible patients who
complete end of Year 2 visit
in SOL-1 or SOL-R

SOL-X: Unlocking Long-Term Value with OTX-TKI

Designed to evaluate safety and disease modifying potential of OTX-TKI

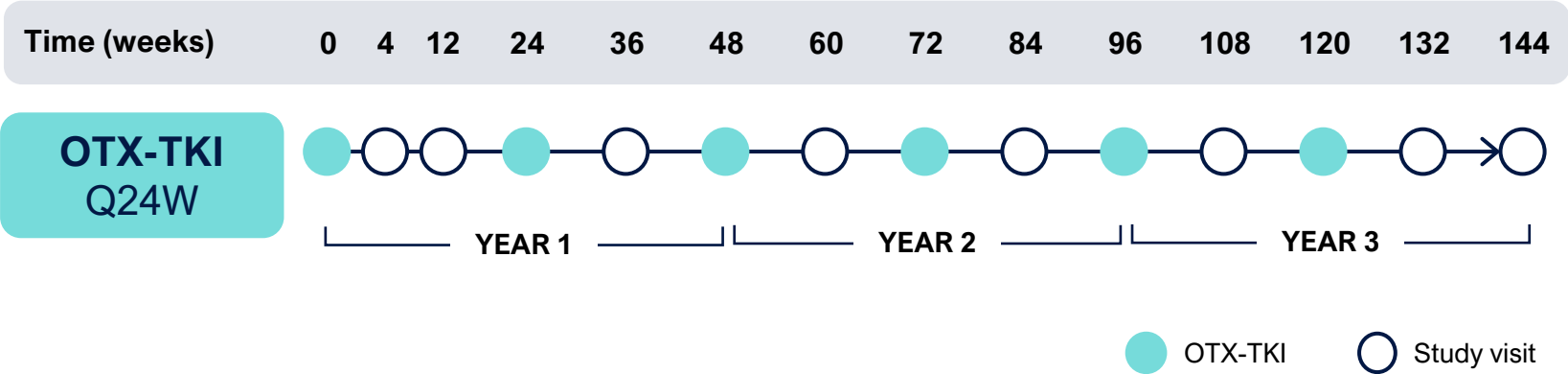
Primary Trials

Initial Treatment Arms



SOL-X

Three-year additional follow-up

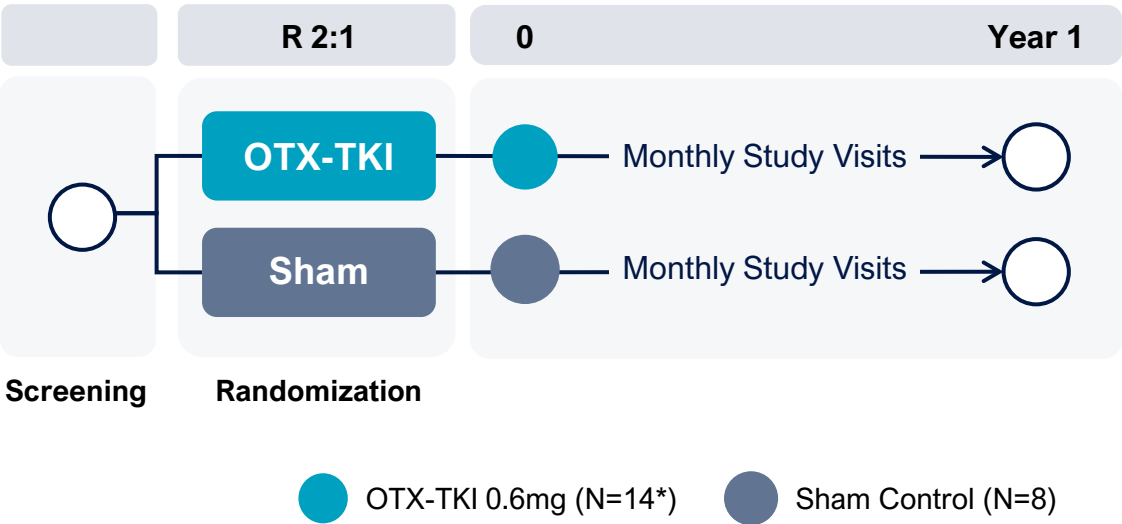


HELIOS-1 Study of OTX-TKI in NPDR

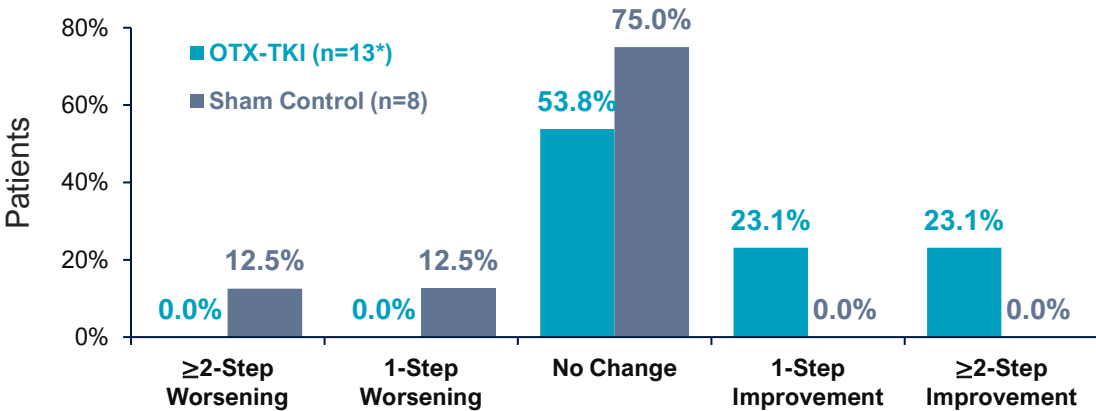


Design

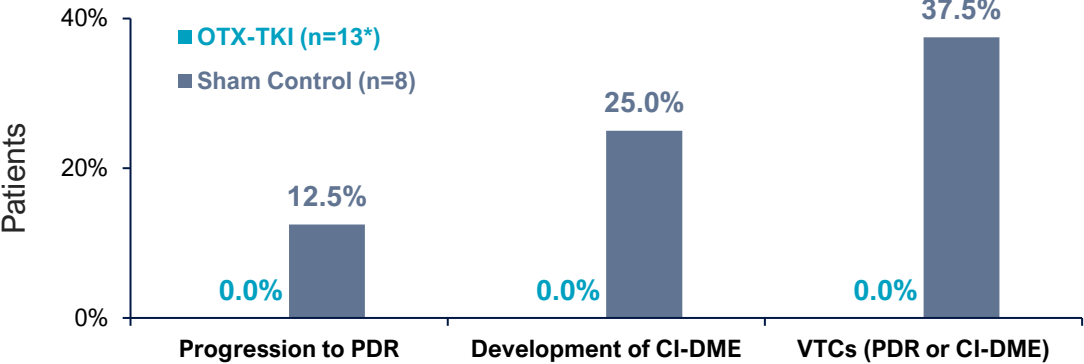
Multi-center, double-masked, randomized, parallel group study of OTX-TKI in moderately-severe to severe NPDR without CI-DME



Change in DRSS from Baseline to Week 48¹



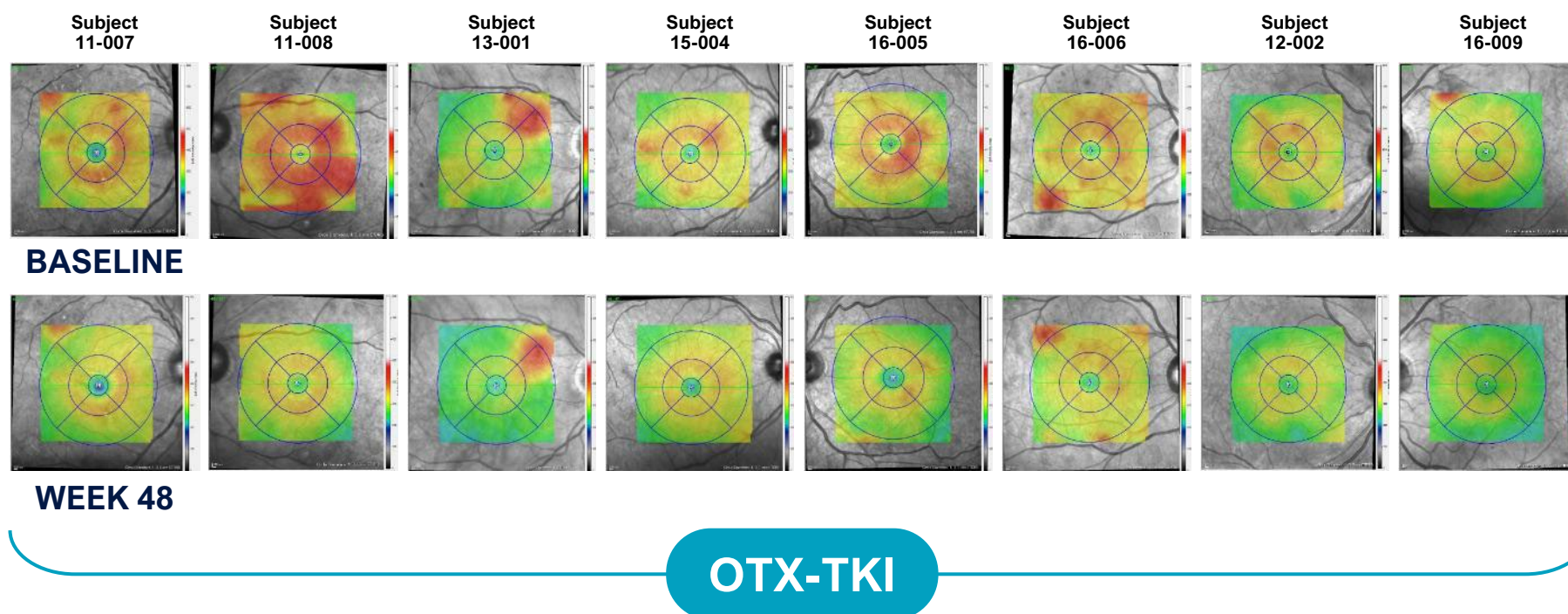
Progression to VTCs at Week 48²



¹14 subjects enrolled in OTX-TKI treatment arm, with one subject death unrelated to treatment.
DRSS, diabetic retinopathy severity scale; NPDR, non-proliferative diabetic retinopathy; CI-DME, center-involved diabetic macular edema; PDR, proliferative diabetic retinopathy; VTC, vision-threatening complication

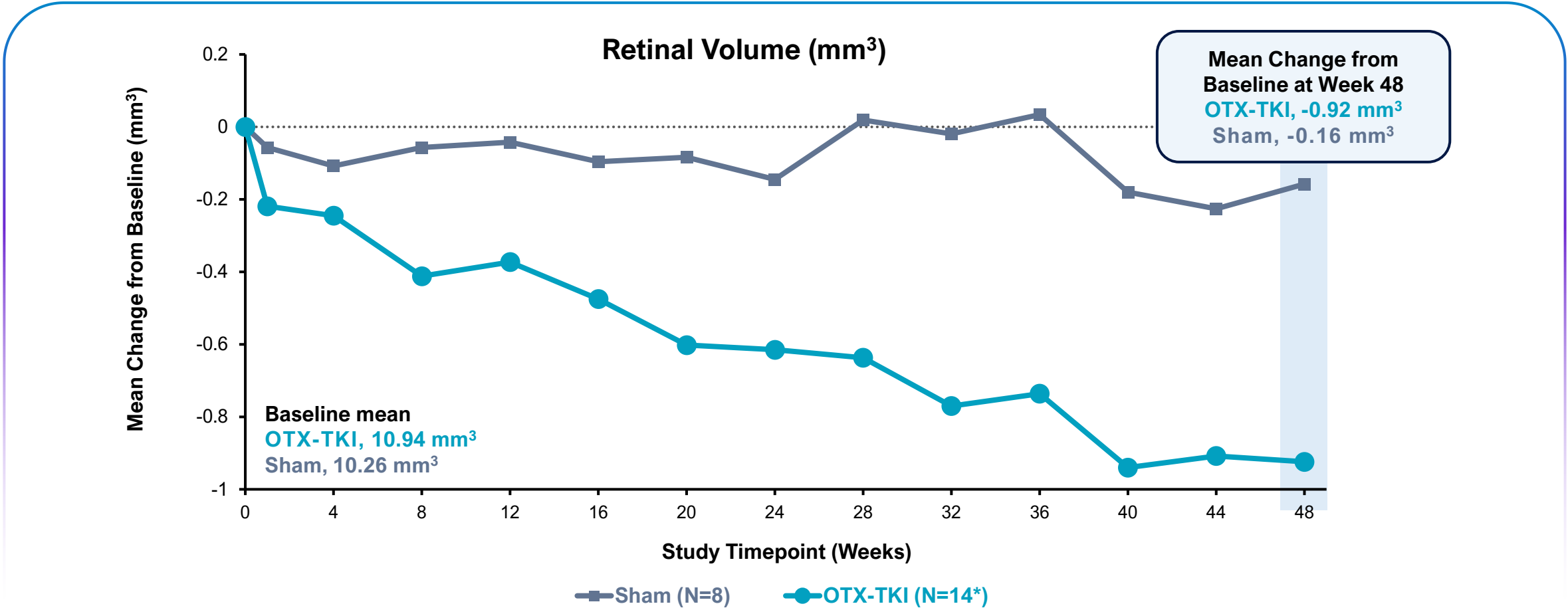
HELIOS-1: DME Improvement in All Subjects with OTX-TKI

HELIOS-1 OTX-TKI Subjects with Non-CI DME (N=8)



OTX-TKI Subjects Consistently Had Greater Reductions from Baseline in Retinal Volume Compared to Sham

Post Hoc Analysis

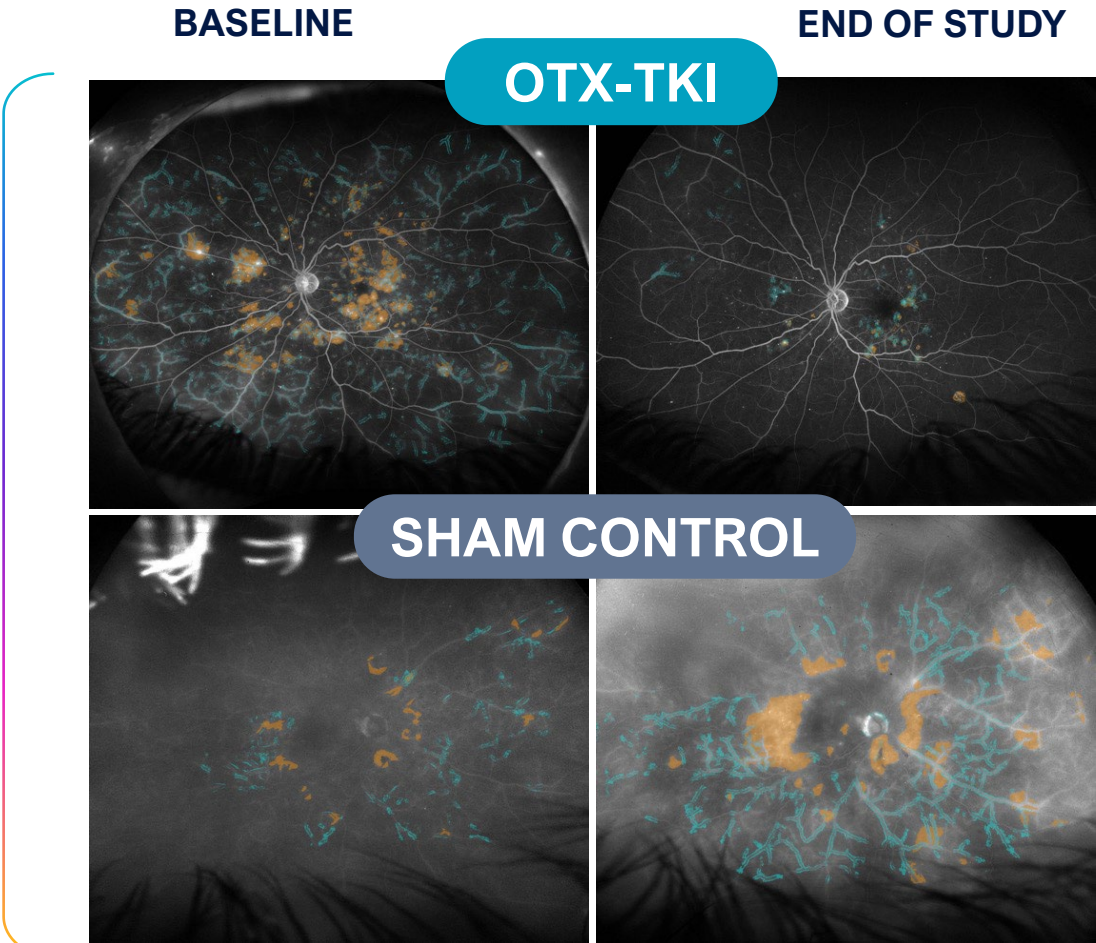


HELIOS-1: Total Retinal Vascular Leakage

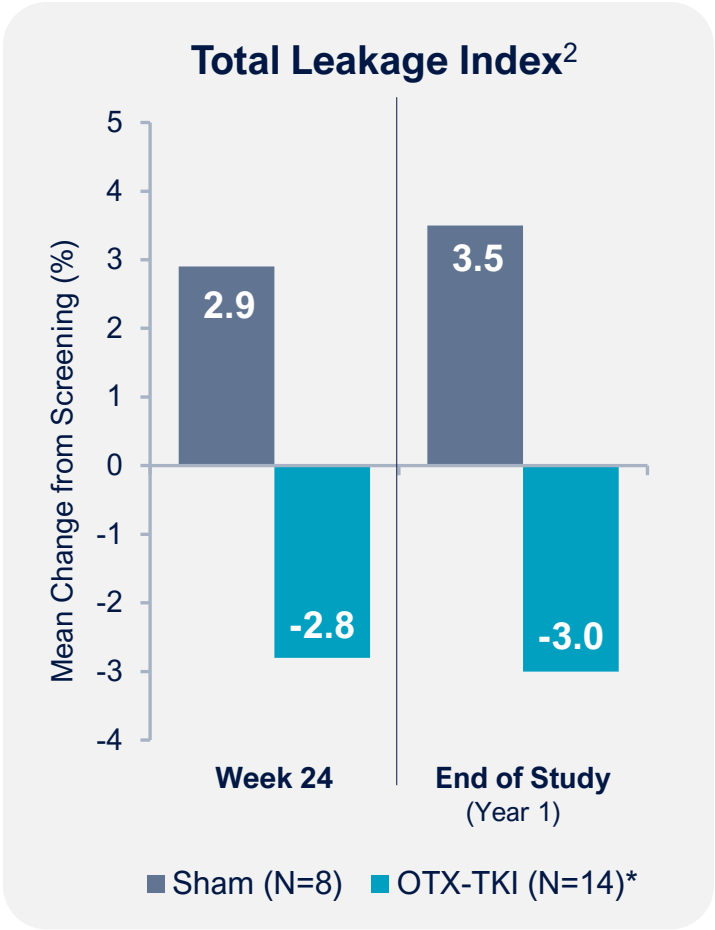
Post Hoc Analysis



- Higher levels of total retinal vascular leakage associated with¹:
- More advanced DR
 - Greater risk of disease progression
 - Worse visual outcomes



Clinical images of ultrawidefield fluorescein angiography (UWF-FA) leakage of one patient. Individual results may vary.



OTX-TKI is an investigational product candidate which has not been approved by the FDA or any regulatory authority.

*14 patients enrolled in OTX-TKI treatment arm, with one patient death unrelated to treatment.
1. Ehlers J, et al. *Ophthalmol Retina*. 2025;9(3):243-252. 2. Figures from Ehlers J, et al. Volumetric Macular Fluid Analysis of the Impact of a Single Axitinib Intravitreal Implant (OTX-TKI) from the HELIOS Clinical Trial for Diabetic Retinopathy. Presented at Angiogenesis. Virtual. February 8, 2025.
DR, diabetic retinopathy

Diabetic Retinopathy Program

HELIOS-3 Study Design



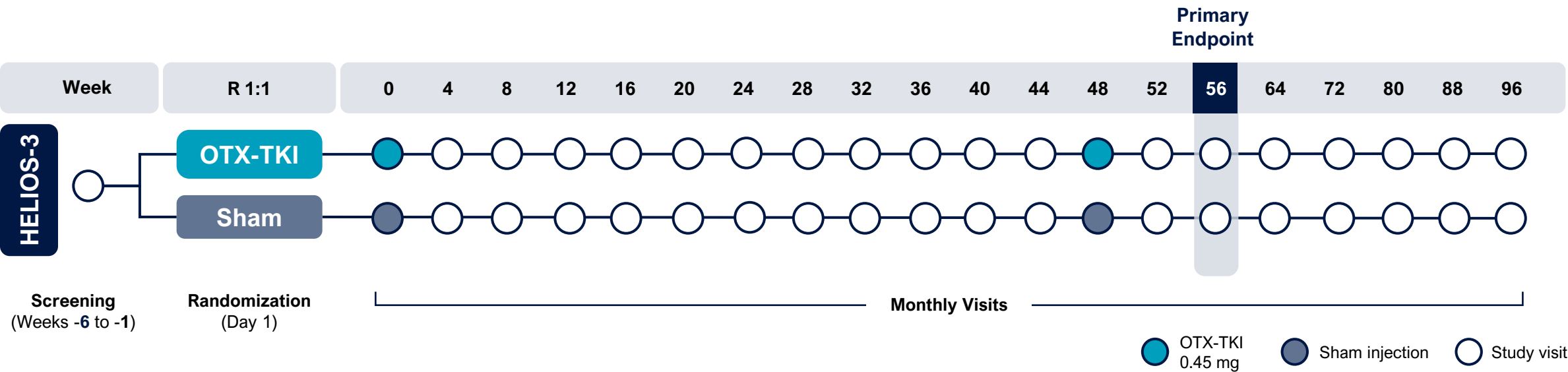
Evaluation of only Q12M OTX-TKI regimen vs Sham

Design

Superiority study of OTX-TKI in ~620 subjects with moderately-severe to severe NPDR without CI-DME

Primary Endpoint (Week 56)

Ordinal DRSS 2-step change status at **Week 56** from baseline
(≥ 2 step improvement, ≥ 2 -step worsening, < 2 -step change in either direction)



OTX-TKI Development Program Summary

OTX-TKI

OTX-TKI demonstrated superior maintenance of vision* to aflibercept 2 mg at Week 36 in nAMD



nAMD Program

PHASE 3

Superiority Trial

Durability of a single OTX-TKI injection



Primary endpoint met

SOL-1

PHASE 3

Non-Inferiority Trial

Repeat dosing every 6 months

Randomization complete - topline data expected in Q1 '28

SOL-R

DR Program

PHASE 3

Superiority Trial

Evaluating yearly dosing

Enrollment in HELIOS-3 is underway

HELIOS-3

Redefining the Management of nAMD

The Impact of Durability and Sustained Disease Control

August 29, 2026

Asia-Pacific Vitreo-retina Society Congress | Gold Coast, Australia

