



# Virtual KOL/IR Event

November 14, 2020

# Forward-looking Statements

Statements contained in this press release regarding events or results that may occur in the future are “forward-looking statements” within the meaning of the Private Securities Litigation Reform Act of 1995. Such statements include, but are not limited to, statements regarding: the potential for ADV-022 in treating patients with wet AMD and DME; the expected growth of the incidence of new cases of wet AMD in the U.S. as its population ages; Adverum’s expectations that it will present longer-term data from the OPTIC Phase 1 trial for ADV-022 in wet AMD in the first half of 2021 and data from the INFINITY Phase 2 trial for ADV-022 in DME in the second half of 2021; Adverum’s plans to accelerate the development and future commercial launch plans for ADV-022; and Adverum’s expectations as to its plans to advance ADV-022 in wet AMD by initiating a pivotal trial mid-2021. All of these statements are based on certain assumptions made by Adverum on current conditions, expected future developments and other factors Adverum believes are appropriate in the circumstances. Adverum may not achieve any of these in a timely manner, or at all, or otherwise carry out the intentions or meet the expectations disclosed in its forward-looking statements, and you should not place undue reliance on these forward-looking statements. Actual results and the timing of events could differ materially from those anticipated in such forward-looking statements as a result of various risks and uncertainties, which include risks inherent to, without limitation: Adverum’s novel technology, which makes it difficult to predict the time and cost of product candidate development and obtaining regulatory approval; the results of early clinical trials not always being predictive of future results; the potential for preliminary or interim results of clinical trials to change as the clinical trial continues or in connection with the preparation and analysis of final results; the potential for future complications or side effects in connection with use of ADV-022; obtaining regulatory approval for gene therapy product candidates; enrolling patients in clinical trials; reliance on third parties for conducting the OPTIC and INFINITY trials and vector production; the effects of the COVID-19 pandemic on the company’s operations and on the company’s ongoing clinical trials; and ability to fund operations through completion of the OPTIC and INFINITY trials and thereafter. Risks and uncertainties facing Adverum are described more fully in Adverum’s Form 10-Q filed with the SEC on November 5, 2020 under the heading “Risk Factors.” All forward-looking statements contained in this press release speak only as of the date on which they were made. Adverum undertakes no obligation to update such statements to reflect events that occur or circumstances that exist after the date on which they were made.

# Agenda for OPTIC Data Virtual KOL/IR Event Saturday, Nov 14, 2020 7:30-9am PT

Time	Presentation	Speaker
5 Minutes	Opening Remarks	<b>Laurent Fischer, M.D.</b> CEO, Adverum Biotechnologies
25 Minutes	<b>OPTIC Phase 1 Presentation</b>	<b>Carl D. Regillo, M.D., F.A.C.S</b> Director, Wills Eye Hospital Retina Service Investigator in OPTIC Phase 1 Trial
15 Minutes	<b>Ocular Inflammation Overview</b>	<b>Steven Yeh, M.D.</b> Associate Professor, Director, Section of Uveitis and Ocular Immunology, Emory Eye Center
15 Minutes	<b>Fireside Chat – Dr. Boyer and Dr. Fischer</b>	<b>David S. Boyer, M.D.</b> Senior Partner, Retina-Vitreous Associates Medical Group and Adjunct Clinical Professor of Ophthalmology, University of Southern California/ Keck School of Medicine, Los Angeles Investigator in OPTIC Phase 1 Trial
30 Minutes	Q&A	Laurent Fischer, M.D. Aaron Osborne, M.B.B.S. Leone Patterson David S. Boyer, M.D. Carl D. Regillo, M.D., F.A.C.S. Steven Yeh, M.D.
	Closing Remarks	<b>Laurent Fischer, M.D.</b>

# Phase 1 Study of Intravitreal Gene Therapy with ADVIM-022 for Neovascular Age-related Macular Degeneration (OPTIC Trial Cohorts 1–4)

Carl D. Regillo, M.D., F.A.C.S

Director of the Wills Eye Hospital Retina Service

*(on behalf of the OPTIC investigators)*



November 14, 2020

# Disclosures

- **Grant Support:** Genentech, Regeneron, Novartis, Allergan, Astellis, Notal, Chengdu Kanghong, Opthea, Iveric, Adverum, RegenXBio, Kodiak, Graybug
- **Consultant:** Genentech, Novartis, Allergan, Notal, Takeda, Kodiak, Graybug, Lineage, Opthea, Eyepoint, Iveric, Aldeyra, Merck, Adverum, Chengdu Kanghong

# Key Takeaways for ADVIM-022 (OPTIC Trial)

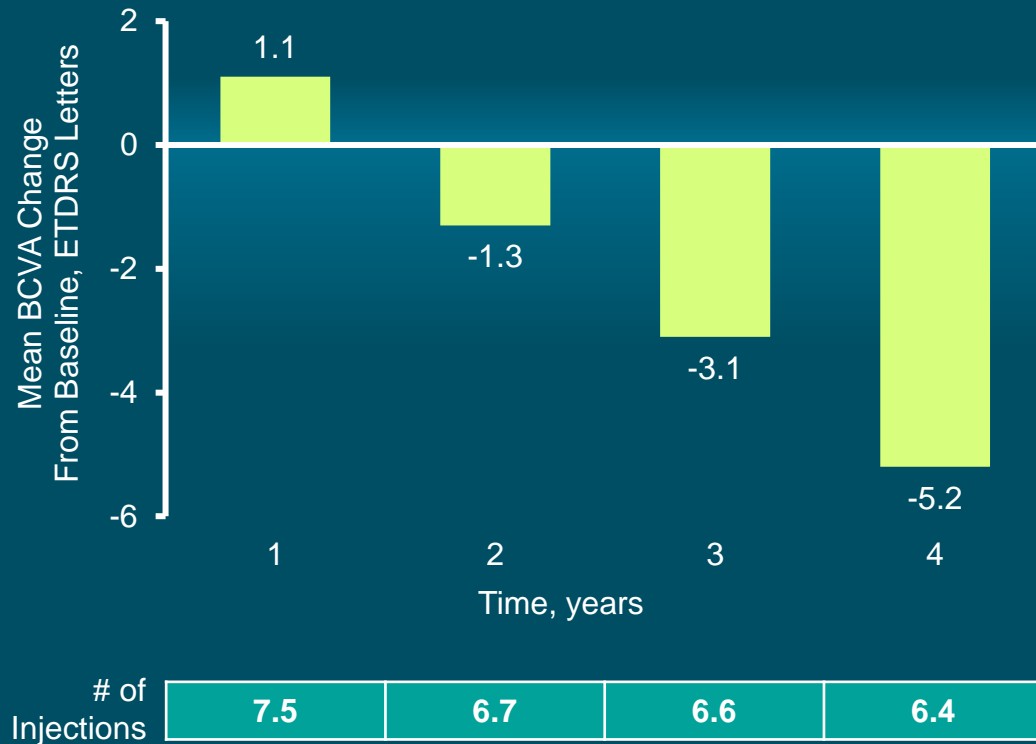
- Continues to be well tolerated with a favorable safety profile at both high and low doses
- Show robust and sustained efficacy at both high and low doses
- Durability out to 92 weeks from a single IVT injection with zero supplemental injections in Cohort 1
- Robust aqueous anti-VEGF protein expression observed at 18 months in Cohort 1
- Substantial reduction in annualized injection frequency following ADVIM-022
- Most patients are supplemental injection free in OPTIC
- Warrant further investigation in larger studies

# Real-world anti-VEGF Patient Outcomes

*Under treatment leads to vision loss over time*

98,821 Eyes from 79,885 US Patients  
Receiving Routine Intravitreal anti-VEGF Therapy

Development Approach to Deliver  
Long-Term Efficacy

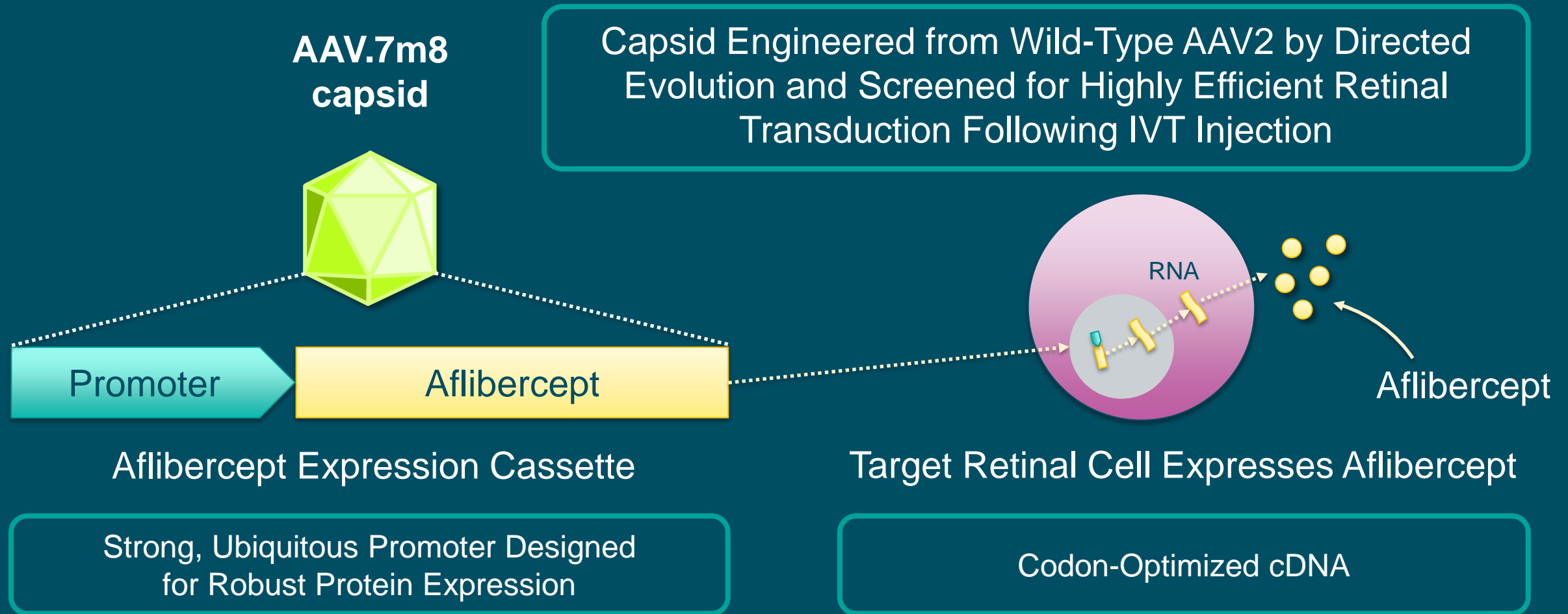


## Gene Therapy

In-Office Intravitreal Injection  
to Establish an Intraocular  
anti-VEGF Biofactory

# ADVM-022: Adeno-Associated Virus Gene Therapy Vector

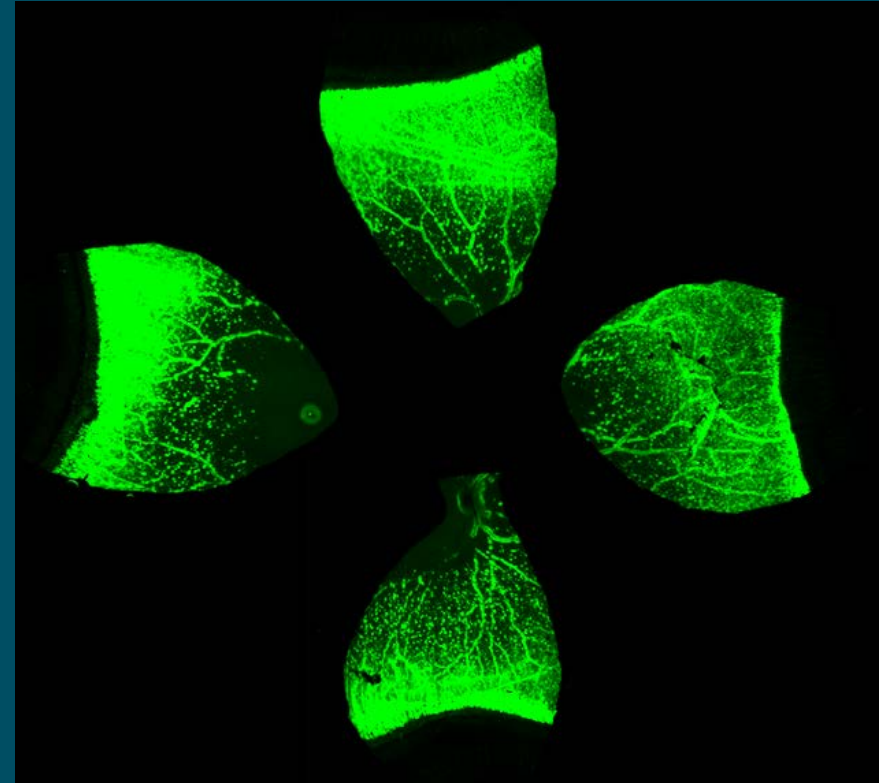
*Designed for continuous delivery of aflibercept by intravitreal injection*





# Intravitreal Injection of AAV.7m8 Results in Robust Cellular Transduction and Protein Expression in the Eye

- Advanced AAV.7m8 vector developed using directed evolution to:
  - Enable efficient intravitreal delivery<sup>1,3</sup>
  - Increase transduction of retinal cells<sup>1,3</sup>
  - Increase protein expression<sup>1</sup>
- Protein expression in NHPs:
  - Photoreceptors, ganglion cells<sup>1–3</sup>
  - Bipolar cells, Müller cells, optic nerve<sup>2</sup>
  - Ciliary epithelium, iris pigment epithelium<sup>2</sup>



Green Fluorescent Protein Expression In  
Non-Human Primate Retina<sup>1</sup>

# Preclinical NHP Data Demonstrate Long-Term Sustained Aflibercept Levels Comparable to Aflibercept Bolus Injection



\*Time after IVT injection of bolus aflibercept protein (1.2 mg/eye; separate study)  
when similar aflibercept levels were observed in NHPs  
IVT, intravitreal therapy; NHP, non-human primate

1. Kiss, S. Ann Meeting of the Am Soc Gene Cell Ther; 2019, Washington, DC  
2. Grishanin, R Ann Congress Eur Soc Gene Cell Ther; 2018, Lausanne, Switzerland

# OPTIC: Phase 1, Two-Year Multicenter Dose-Ranging Study of ADVM-022 in Neovascular AMD

- **Primary Objective**

- Assess the safety and tolerability of a single IVT injection of ADVM-022

- **Secondary Objective**

- Evaluate vision (BCVA)
- Evaluate anatomy (SD-OCT)
- Assess the need for rescue therapy



Oral steroid prophylaxis\*: Cohort 1 ( $6 \times 10^{11}$  vg/eye, n=6) and Cohort 2 ( $2 \times 10^{11}$  vg/eye, n=6)

Steroid eye drops prophylaxis\*\*: Cohort 3 ( $2 \times 10^{11}$  vg/eye, n=9) and Cohort 4 ( $6 \times 10^{11}$  vg/eye, n=9)

**Patients Receive Rescue Aflibercept (2 mg IVT) if *any* of the Following Criteria are Met:**

1. Loss of  $\geq 10$  letters in BCVA from baseline that is attributed to intraretinal or subretinal fluid observed by the investigator
2. Increase in central subfield thickness  $> 75 \mu\text{m}$  from baseline
3. Presence of vision-threatening hemorrhage due to AMD

\*Subjects received prophylaxis of 60 mg oral prednisone for 6 days starting at Day -3 followed by 7-day taper.

\*\*Subjects receive prophylaxis of QID difluprednate eye drops for 3 weeks starting at Day 1 followed by a 3-week taper.

BCVA, best-corrected visual acuity; IVT, intravitreal therapy; SD-OCT, spectral domain optical coherence tomography; QID, 4x/day

# OPTIC Update for Cohorts 1-4 as of October 15, 2020

	Cohort 1 (N=6)	Cohort 2 (N=6)	Cohort 3 (N=9)	Cohort 4* (N=9)
<b>ADVM-022 Dose, vg/eye</b>	<b>High Dose</b> 6×10 <sup>11</sup>	<b>Low Dose</b> 2×10 <sup>11</sup>	<b>Low Dose</b> 2×10 <sup>11</sup>	<b>High Dose</b> 6×10 <sup>11</sup>
<b>Steroid Prophylaxis</b>	<b>Oral</b> 13-day course	<b>Oral</b> 13-day course	<b>Eye drops</b> 6-week course	<b>Eye drops</b> 6-week course
<b>Follow-Up, Weeks</b>	64–92 weeks (median 86)	64–68 weeks (median 64)	32–48 weeks (median 48)	12–24 weeks (median 16)
<b>Subject Disposition</b>	No discontinuations, some visits missed due to COVID-19 concerns	No discontinuations	No discontinuations, some visits missed due to COVID-19 concerns	No discontinuations
<b>Baseline Characteristics</b>	✓	✓	✓	✓
<b>Safety Data</b>	✓	✓	✓	✓
<b>Efficacy Data<sup>†</sup></b>	✓	✓	✓	N/A
<b>Aqueous anti-VEGF Protein Expression Data</b>	N=2 at week 76	N/A	N/A	N/A

\*Cohort 4 has less than 6 months of follow-up

<sup>†</sup>Includes BCVA and CST outcomes

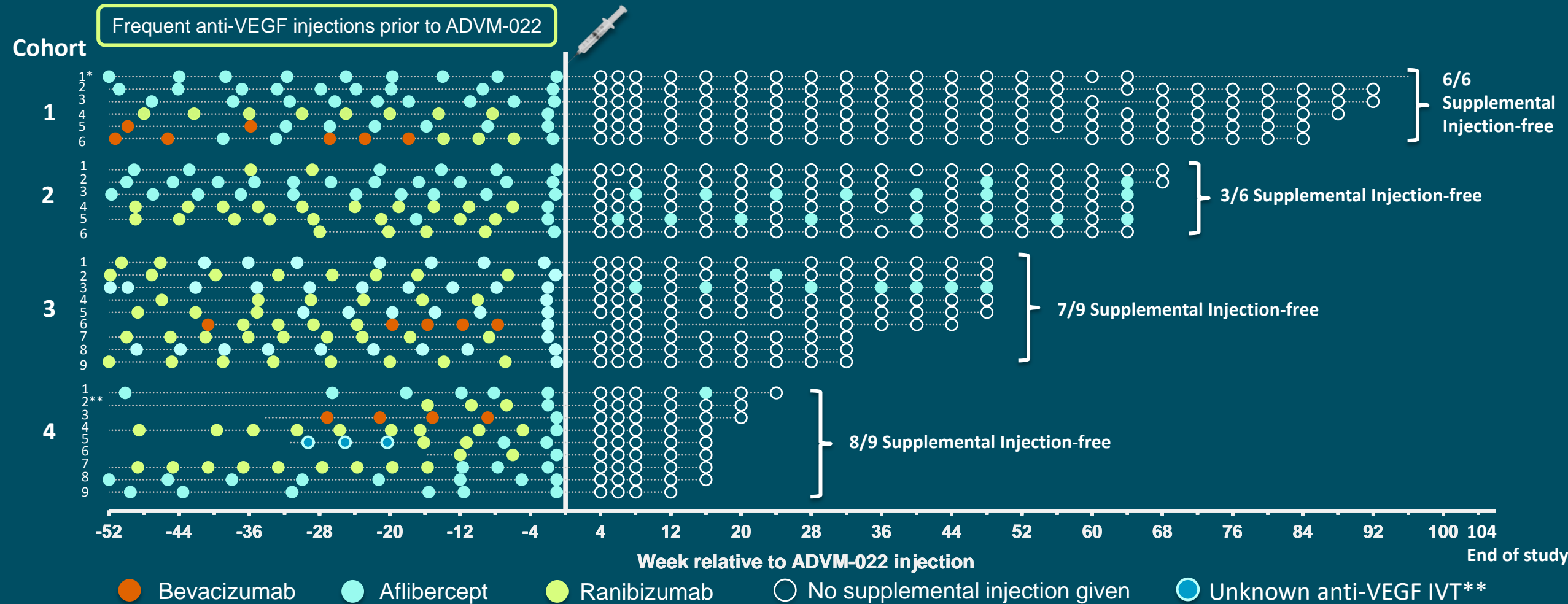
# Neovascular AMD Study Population Previously Required Frequent Injections to Maintain Vision

Baseline Characteristics	Cohort 1 (N=6)	Cohort 2 (N=6)	Cohort 3 (N=9)	Cohort 4 (N=9)
Mean (range) Age, Years	79.0 (62–88)	79.8 (74–90)	77.4 (65–90)	79.9 (68–88)
Mean (range) Years Since nAMD Diagnosis	4.5 (0.9–10.6)	4.1 (0.5–6.8)	3.3 (0.7–8.0)	3.2 (0.2–8.0)
Mean (range) Number anti-VEGF Injections Since Initial Diagnosis*	38.2 (7–109)	34.0 (4–69)	24.8 (9–70)	28.5 (2–58)**
Mean (range) Number anti-VEGF Injections in 12 Months Prior to ADVM-022	9.2 (8–11)	9.2 (5–11)	9.1 (7–10)	7.1 (3–12)**
Mean (range) BCVA, ETDRS Letters Approximate Snellen Equivalent	65.8 (57–77) 20/50	64.7 (53–72) 20/50	65.9 (53–75) 20/50	65.0 (54–77) 20/50
Mean (range) CST, $\mu$ m	369.2 (293–561)	307.7 (235–339)	473.4 (301–857)	398.6 (255–538)

\*Not including the mandated aflibercept at Screening; \*\*Excluding Patient #2 with incomplete prior anti-VEGF data.

BCVA, best corrected visual acuity; CST, central subfield thickness; ETDRS, Early Treatment Diabetic Retinopathy Study  
nAMD, neovascular age-related macular degeneration; VEGF, vascular endothelial growth factor

# Substantial Reduction in anti-VEGF Treatments Following a Single IVT Injection of ADVM-022



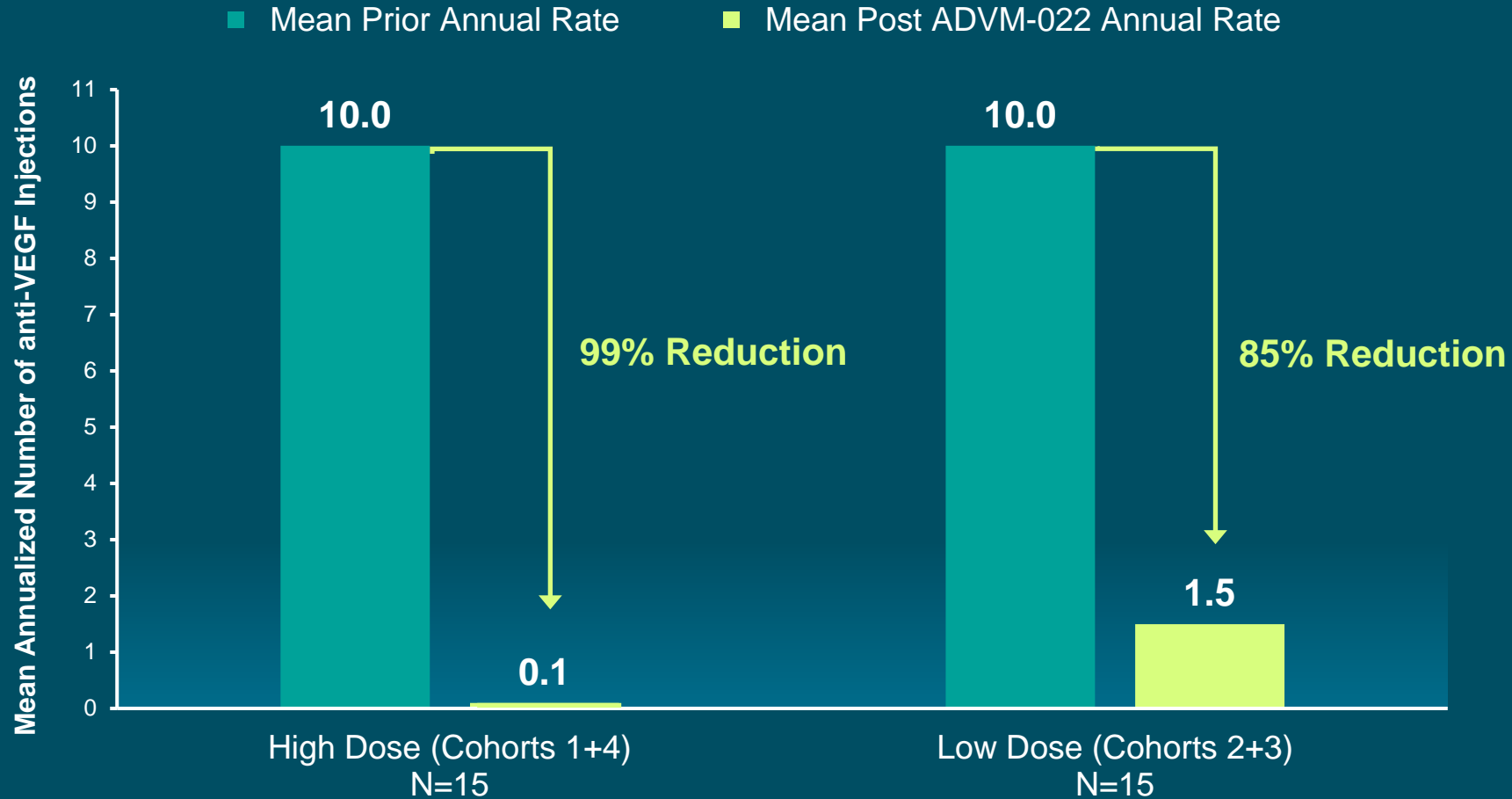
Five patients were diagnosed <1 year prior to ADVM-022 injection: one each in Cohorts 2 and 3, three in Cohort 4.

\*Cohort 1, Patient 1 remains on study but have missed visits post Week 64; \*\*Incomplete prior data for Cohort 4, Patient 2;

†Received in a clinical trial not yet unmasked (NCT04049266).

Data cut: October 15, 2020

# Substantial Reduction in Annualized anti-VEGF Injection Frequency Following ADVIM-022



*Annualized rate (Prior) = (number of IVTs in 12 months prior to ADVIM-022) / (days from the first IVT in the past 12 months to ADVIM-022 / 365.25).*

*Annualized rate (Post) = (numbers of aflibercept IVTs since ADVIM-022) / (days from ADVIM-022 to the last study follow-up / 365.25).*

*Data cut: October 15, 2020*



# Safety Summary Across Cohorts through October 15, 2020

- No ADVIM-022-related non-ocular adverse events
  - No deaths or discontinuations in OPTIC
- When observed, inflammation has been responsive to and manageable with steroid eye drops
- No clinical or fluorescein\* evidence of posterior inflammation
  - No vasculitis, retinitis, choroiditis, vascular occlusions or endophthalmitis
- All ADVIM-022-related ocular AEs were mild (78%) to moderate (22%)
  - One AE of special interest of moderate recurrent uveitis deemed to be related to ADVIM-022 was responsive to steroid eye drops (Cohort 1)
- One unrelated ocular SAE of retinal detachment surgically repaired and resolved (Cohort 1)
- Two patients had mild AEs of IOP elevation that resolved
  - One patient had two mild IOP elevations (highest 24 mmHg) that were both treated with Combigan® eye drops
  - One case in a patient on Combigan® for ocular hypertension at baseline which resolved with no change to treatment

\*Fluorescein angiography of posterior pole

IOP, intraocular pressure; AEs, adverse events; SAEs, serious AEs



# Adverse Events Across Cohorts as of October 15, 2020

*ADVM-022 related events were mild (78%) or moderate (22%)*

Adverse Events		Cohort 1 (N=6)		Cohort 2 (N=6)		Cohort 3 (N=9)		Cohort 4 (N=9)	
		6×10 <sup>11</sup> vg/eye Oral steroids 13-day prophylaxis		2×10 <sup>11</sup> vg/eye Oral steroids 13-day prophylaxis		2×10 <sup>11</sup> vg/eye Steroid eye drops 6-week prophylaxis		6×10 <sup>11</sup> vg/eye Steroid eye drops 6-week prophylaxis	
		Subjects	Events	Subjects	Events	Subjects	Events	Subjects	Events
Ocular	Serious	2	2*	0	0	0	0	0	0
	ADVM-022 Related**	6	31	5	21	5	15	6	19
	Total Ocular	6	54	5	34	8	31	8	23
Non-Ocular†	Serious ‡	1	1	0	0	2	2	0	0
	Total Non-Ocular†	5	18	6	7	5	10	2	2

\* Retinal detachment (unrelated to ADVM-022) and recurrent moderate uveitis (likely related to ADVM-022)

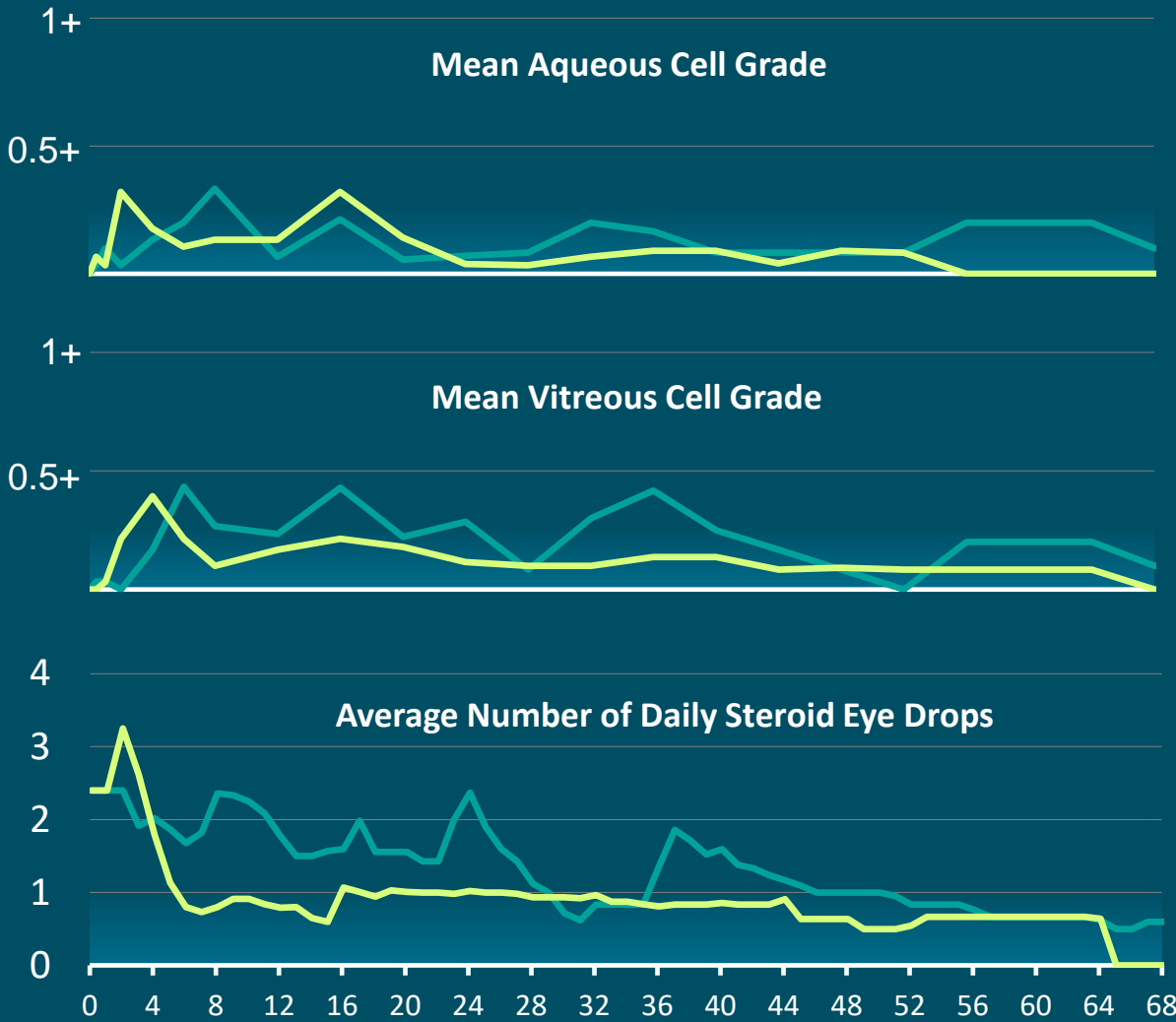
\*\* ADVM-022 related ocular events were mild (78%) or moderate (22%)

† None of the non-ocular AEs were ADVM-022 related

‡ Serious non-ocular AEs included degenerative intervertebral disc disease (1) in Cohort 1; and COPD exacerbation (1), and stable angina pectoris (1) in Cohort 3

# Ocular Cell Grade and Steroid Eye Drop Use Decreases over Time

High Dose (6x10<sup>11</sup> vg/eye)      Low Dose (2x10<sup>11</sup> vg/eye)



Decreasing trend over time for:

- Average aqueous cell grade
- Average vitreous cell grade
- Average steroid eye drop use

Cell grades as assessed by slit lamp  
Grade categories are based on the Standardization of Uveitis Nomenclature (SUN) criteria for aqueous cells and National Institutes of Health (NIH) guidelines for vitreous cells.  
Aqueous cells: 0.5+: 1-5 cells 1+: 6-15 cells 2+: 16-25 cells 3+: 26-50 cells 4+: >50 cells  
Vitreous cells: 0.5+: 1-10 cells 1+: 11-20 cells 2+: 21-30 cells 3+: 31-100 cells 4+: >100 cells;  
rare cells are captured as 0.5+ for this analysis

# Ocular Cellular Inflammation & Topical Steroid Eye Drop Overview

*Latest Outcomes as of October 15, 2020*

Dose	Cohort 1 High Dose (N=6)	Cohort 2 Low Dose (N=6)	Cohort 3 Low Dose (N=9)	Cohort 4 High Dose (N=9)
Follow-Up	64–92 weeks (median 86)	64–68 weeks (median 64)	32–48 weeks (median 48)	12–24 weeks (median 16)
Average Aqueous Cell Grade	0.08	0.00	0.06	0.11
Average Vitreous Cell Grade	0.17	0.00	0.06	0.11
% with any cellular inflammation	33%	0%	11%	22%
Average # of daily drops	1.2	0.5	0.8	1.9

At the most recent visit:

- Low average cell grades
- Low average number of daily drops
- Cohort 4 still in early follow-up
- Slow tapering implemented

*High Dose -  $6 \times 10^{11}$  vg/eye*

*Low Dose -  $2 \times 10^{11}$  vg/eye*

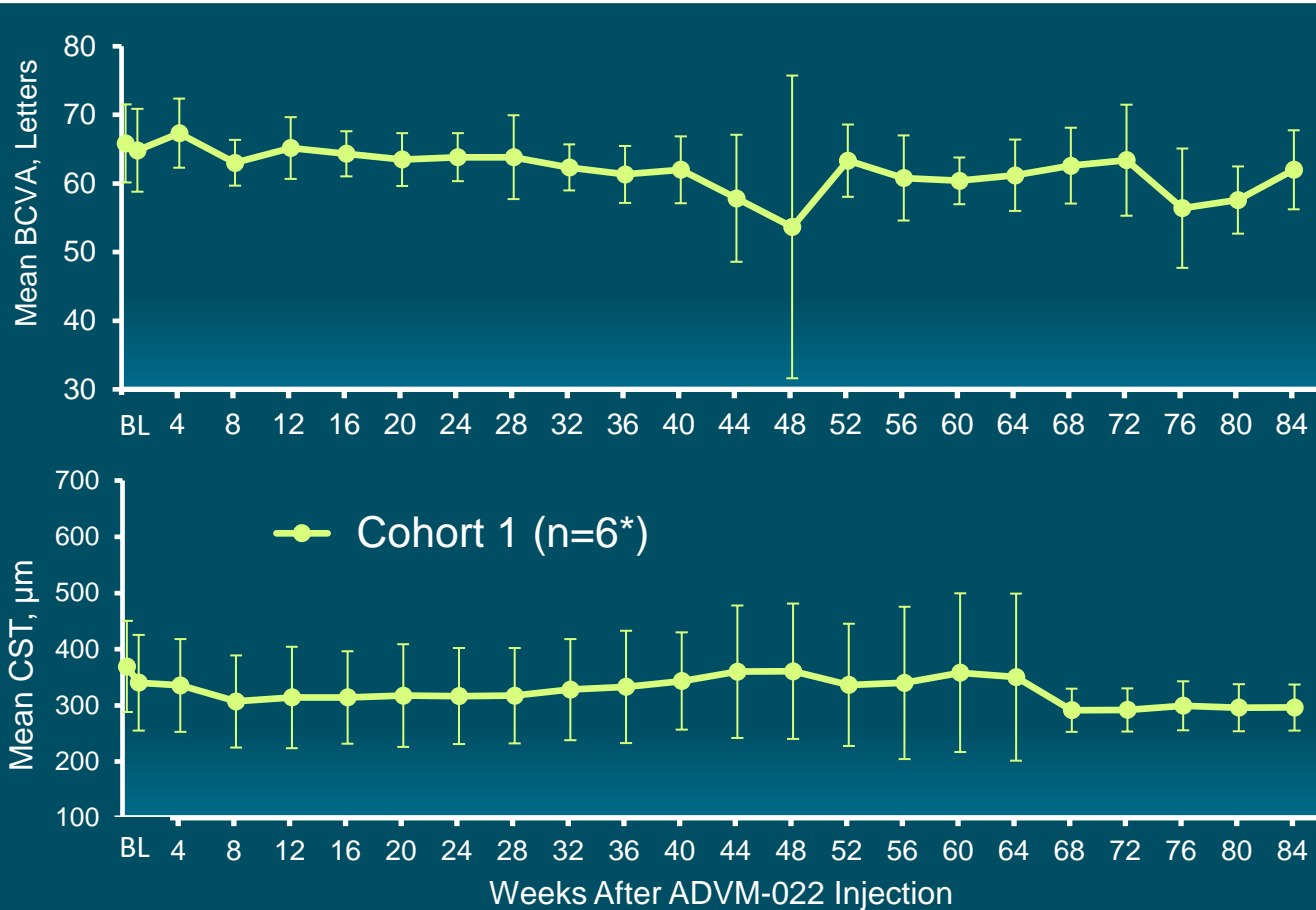
*Averages calculated across entire cohort*

# Cohort 1: BCVA and CST Stable, Zero Supplemental Injections

Robust anti-VEGF Protein Expression observed at 18 months



Mean (90% CI) by Visit Through Week 84



Latest Outcomes as of Oct. 15, 2020

Follow-Up	64–92 weeks (median 86)
Rescue-Free Patients	100% (6/6)
Mean BCVA Change from Baseline	
All Patients	–2.5 Letters
Mean CST Change from Baseline	
All Patients	–19.7 μm

Mean Aqueous anti-VEGF Protein level\*\*

Week 76 (n=2)	1840 ng/mL
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\*One patient had low BCVA and no CST values at 44 and 48 weeks due to retinal detachment; N=5 from Week 56 to 84

Aflibercept 2 mg IVT administered at baseline, 7–15 days prior to ADVIM-022 IVT (Day 1);

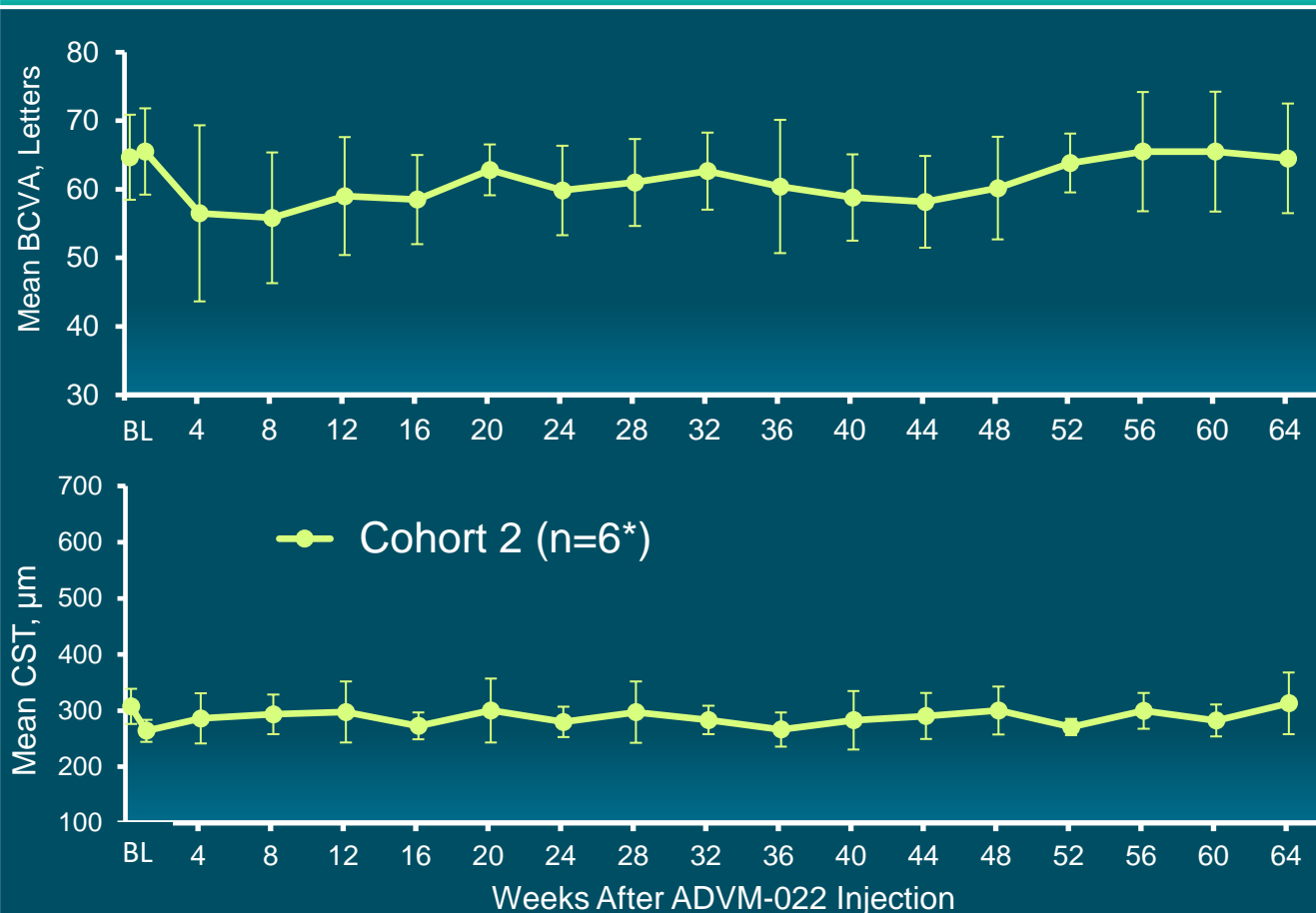
BCVA, best corrected visual acuity; CST, central subfield thickness; BL, baseline; D, day; W, week

Error bars are 90% CIs of the mean absolute BCVA and CST using T-distribution

\*\* Available aqueous humor aflibercept protein samples from Cohort 1 subjects enrolled in optional aqueous humor sampling study

# Cohort 2: BCVA and CST Maintained Over Time

Mean (90% CI) by Visit Through Week 64



Latest Outcomes as of Oct. 15, 2020

Follow-Up	64–68 weeks (median 64)
Rescue-Free Patients	50% (3/6)
Mean BCVA Change from Baseline	
All Patients	+0.2 Letters
Rescue-Free Patients	+1.0 Letters
Mean CST Change from Baseline	
All Patients	–1.0 μm
Rescue-Free Patients	–23.7 μm

\* N=5 for Week 36 and 40 visits

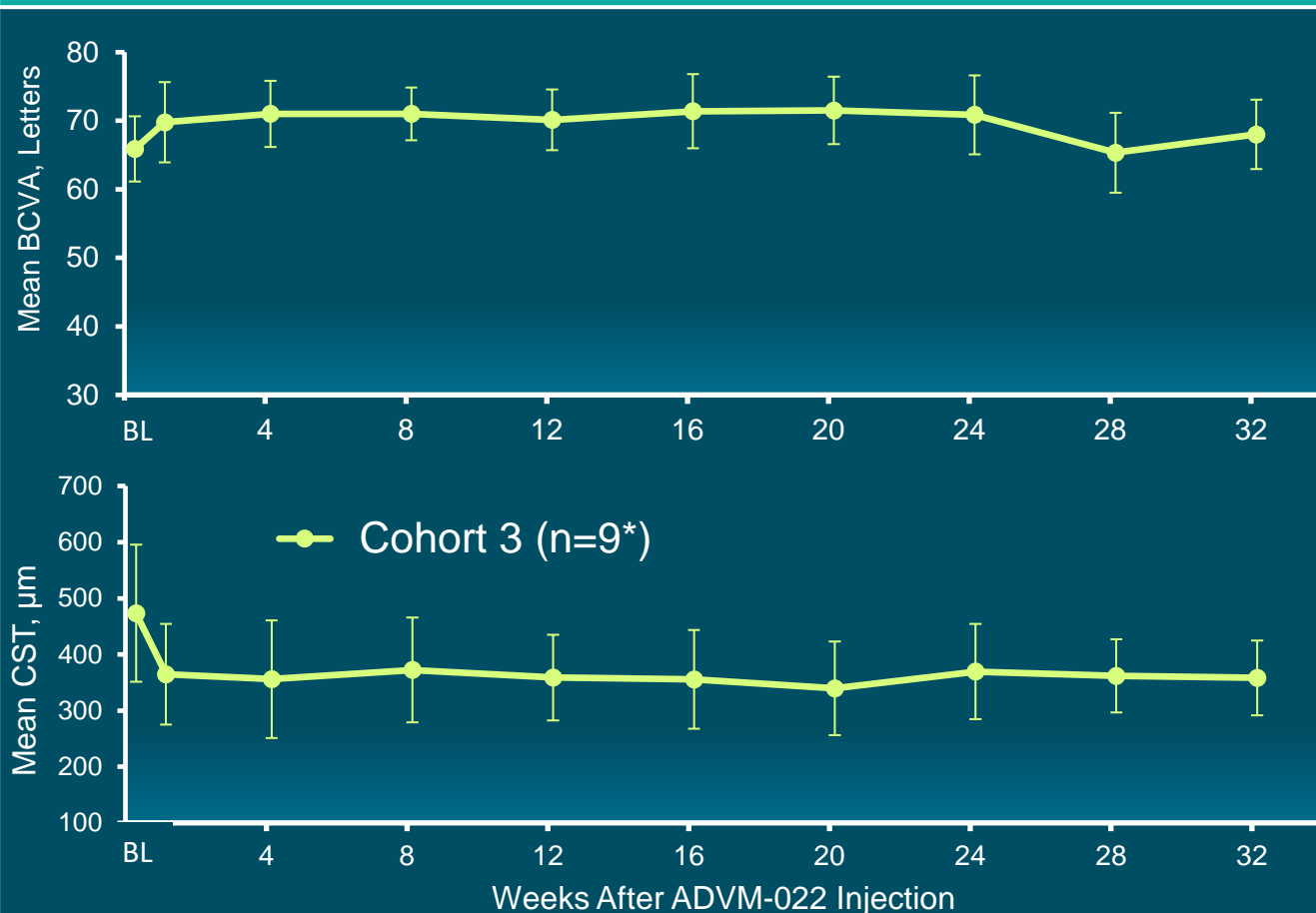
Aflibercept 2 mg IVT administered at baseline, 7–15 days prior to ADV-022 IVT (Day 1).

BCVA, best corrected visual acuity; CST, central subfield thickness; BL, baseline; D, day; W, week

Error bars are 90% CIs of the mean absolute BCVA and CST using T-distribution

# Cohort 3: BCVA Maintained and CST Improved

Mean (90% CI) by Visit Through Week 32



Latest Outcomes as of Oct. 15, 2020

Follow-Up	32–48 weeks (median 48)
Rescue-Free Patients	78% (7/9)
Mean BCVA Change from Baseline	
All Patients	–0.9 Letters
Rescue-Free Patients	+4.1 Letters
Mean CST Change from Baseline	
All Patients	–113.4 μm
Rescue-Free Patients	–132.7 μm

\*N=8 for Week 4, 16 and 20; N=7 at Week 24

Aflibercept 2 mg IVT administered at baseline, 7–15 days prior to ADV-022 IVT (Day 1)

BCVA, best corrected visual acuity; CST, central subfield thickness; BL, baseline; D, day; W, week

Error bars are 90% CIs of the mean absolute BCVA and CST using T-distribution

# Case Study: Cohort 3, Subject 5

## *Persistent fluid despite frequent anti-VEGF injections*

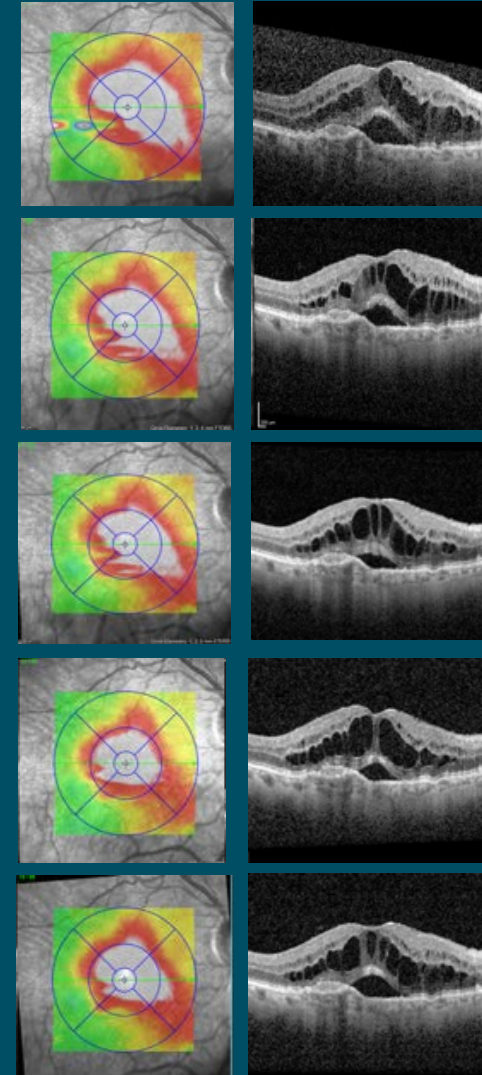
OCT scans and treatment intervals from most recent 5 anti-VEGF injections visits prior to OPTIC

82 Year Old Male	
Previous IVT, n*	19
IVT in Last 12 Months, n	9



Aflibercept injections

\* Excluding the aflibercept injection received at the Screening visit  
IVT, intravitreal therapy; OCT, optical coherence tomography;  
VEGF, vascular endothelial growth factor



Weeks Prior to  
ADVM-022



–30 weeks



–25 weeks



–20 weeks



–15 weeks



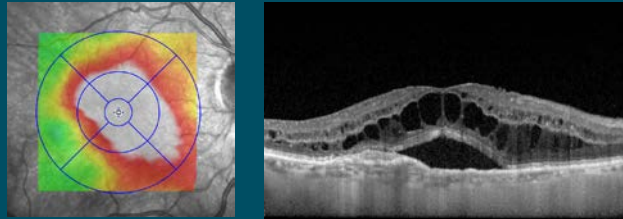
–10 weeks




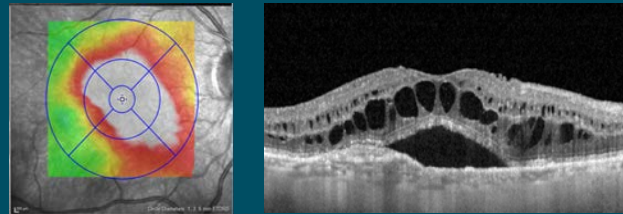
# Case Study: Cohort 3, Subject 5


## *Rapid and sustained anatomical improvements*

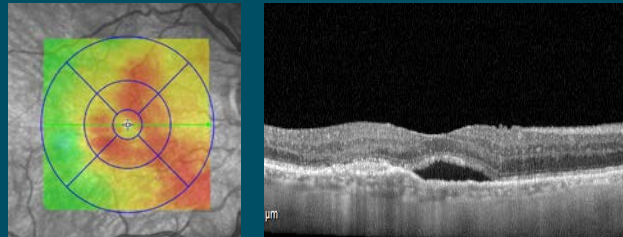
–3 weeks  
Screening  
BCVA: 77 letters  
CST: 678  $\mu$ m



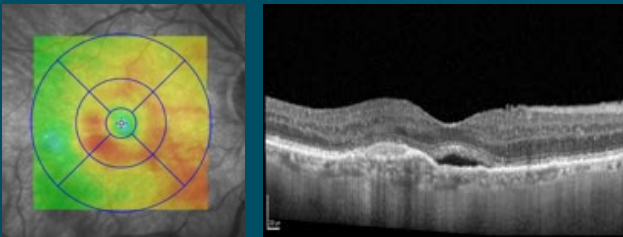
 Aflibercept IVT  
–2 weeks  
BCVA: 75 letters  
CST: 664  $\mu$ m



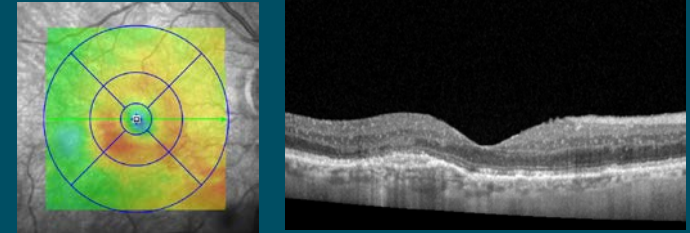
 ADVIM-022  
0 weeks  
BCVA: 82 letters  
CST: 355  $\mu$ m



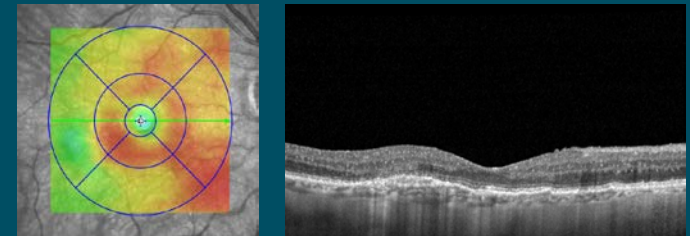
+1 week  
BCVA: 80 letters  
CST: 338  $\mu$ m



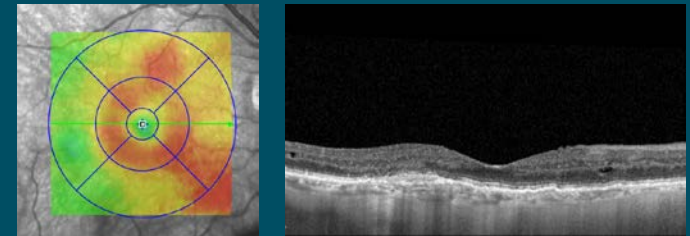
+12 weeks  
BCVA: 81 letters  
CST: 257  $\mu$ m



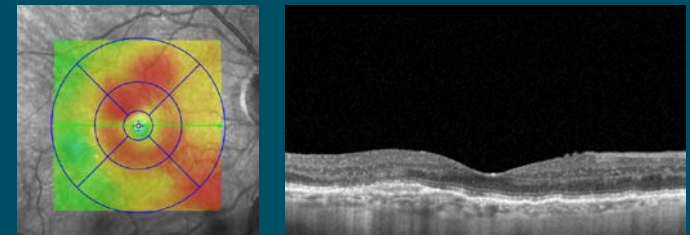
+24 weeks  
BCVA: 83 letters  
CST: 272  $\mu$ m



+36 weeks  
BCVA: 83 letters  
CST: 286  $\mu$ m



+48 weeks  
BCVA: 83 letters  
CST: 300  $\mu$ m





# ADVM-022 Greatly Reduced anti-VEGF Injection Burden in wet AMD – Warrants Further Investigation in Larger Studies



- ADVM-022 continues to be well tolerated with a favorable safety profile at both high and low doses (n=30)
  - All ADVM-022-related ocular adverse events were mild (78%) to moderate (22%)
  - Ocular inflammation, when observed, has been responsive to steroid eye drops
- ADVM-022 continues to show robust and sustained efficacy at both high and low doses
  - Mean BCVA maintained
  - Mean CST maintained to improved
- Durability out to 92 weeks from a single IVT injection with zero supplemental injections in Cohort 1
- Robust aqueous anti-VEGF protein expression observed at 18 months in Cohort 1
- Substantial reduction in annualized anti-VEGF injection frequency following ADVM-022 in patients who previously required frequent injections to maintain vision:
  - High dose: 99% reduction
  - Low dose: 85% reduction
- Most patients are supplemental anti-VEGF injection free in OPTIC:
  - High dose: 14/15 patients injection free
  - Low dose: 10/15 patients injection free

# ADVM-022 Acknowledgments

## Investigators, Study Teams and Participants

- David Boyer MD
- Brandon Busbee MD
- Brian Joondeph MD
- Arshad Khanani MD
- James Major MD
- Dante Pieramici MD
- Carl Regillo MD
- Charles Wykoff MD, PhD
- Mehdi Gasmi PhD
- Szilard Kiss MD
- Aaron Osborne MBBS
- Carol Hoang PharmD
- Adam Turpcu PhD
- Carol Chung PhD



# Thank you



# Fireside Chat with Dr. David Boyer

## **David S. Boyer, M.D.**

Senior Partner, Retina-Vitreous Associates Medical Group and Adjunct Clinical Professor of Ophthalmology, University of Southern California/ Keck School of Medicine, Los Angeles  
Investigator in OPTIC Phase 1 Trial



# OPTIC and INFINITY Investigator

- Dr. David Boyer, Retina Specialist and Clinical Trial Investigator
- Senior Partner, Retina-Vitreous Associates Medical Group (LA Retina) servicing greater Los Angeles area
- All retina practice with 11 retina specialists
- Involved in extensive clinical research
- Enrolled a third of patients into OPTIC
- Enrolled the first and last patients into OPTIC



Retina-Vitreous Associates Medical Group  
Diseases | Surgery of the Retina and Vitreous

# Fireside Chat with Dr. David Boyer

## **David S. Boyer, M.D.**

Senior Partner, Retina-Vitreous Associates Medical Group and Adjunct Clinical Professor of Ophthalmology, University of Southern California/ Keck School of Medicine, Los Angeles  
Investigator in OPTIC Phase 1 Trial

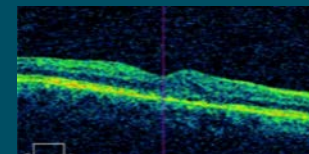
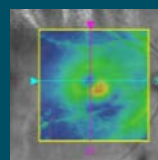


# Case Study: Cohort 1, Subject 5

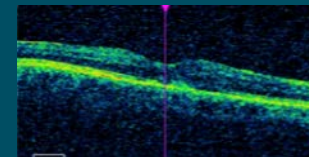
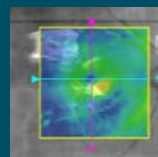
*Frequent anti-VEGF injections for persistent retinal fluid prior to ADVIM-022*

88 Year Old Male	
Study Eye	Right
Lens status	Pseudophakic
Previous IVT, n*	7
IVT in Last 12 Months, n	8

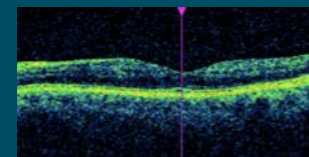
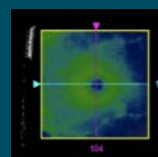
Weeks Prior to  
ADVIM-022



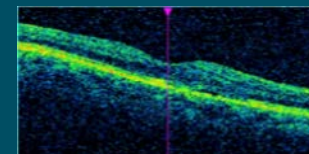
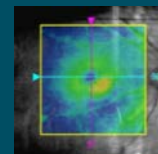
–32 weeks



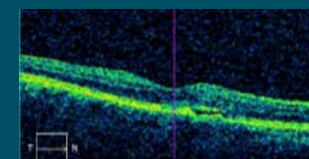
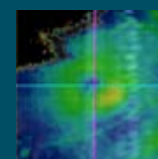
–27 weeks



–22 weeks



–16 weeks



–9 weeks

 Aflibercept injections

\* Excluding the aflibercept injection received at the Screening visit  
IVT, intravitreal therapy; OCT, optical coherence tomography;  
VEGF, vascular endothelial growth factor

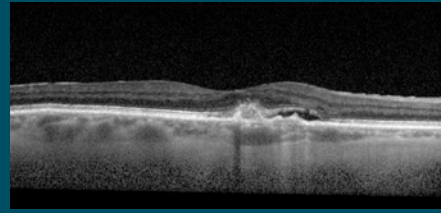
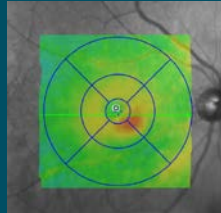


# Case Study: Cohort 1, Subject 5

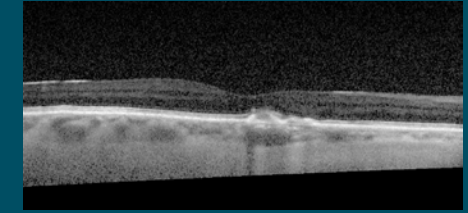
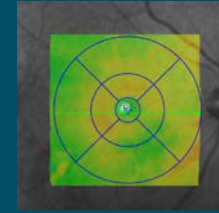
*Zero anti-VEGF Injections with resolved retinal fluid through 18 months after ADVN-022*



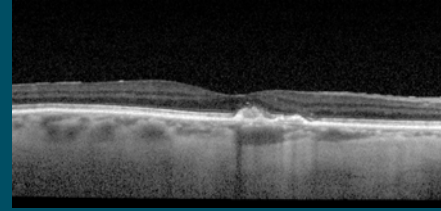
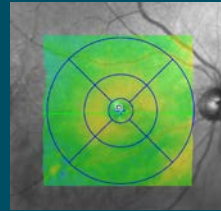
Aflibercept IVT  
-2 weeks  
BCVA: 64 letters  
CST: 293  $\mu\text{m}$



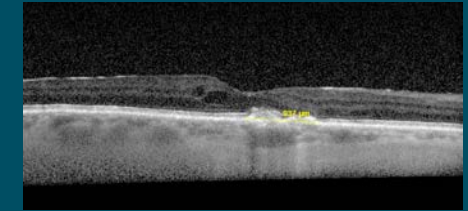
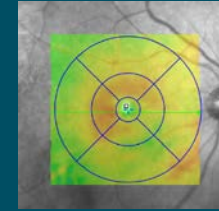
+24 week  
BCVA: 62 letters  
CST: 268  $\mu\text{m}$



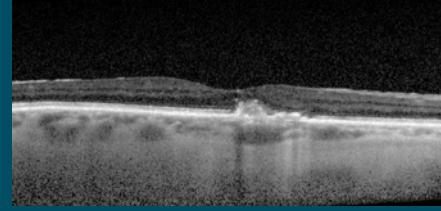
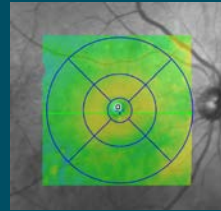
ADV-022  
0 weeks  
BCVA: 65 letters  
CST: 262  $\mu\text{m}$



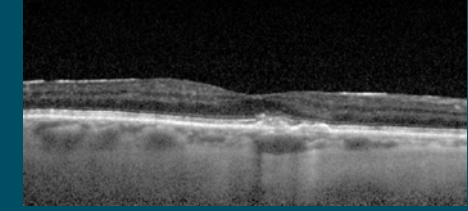
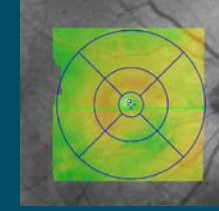
+36 weeks  
BCVA: 61 letters  
CST: 294  $\mu\text{m}$



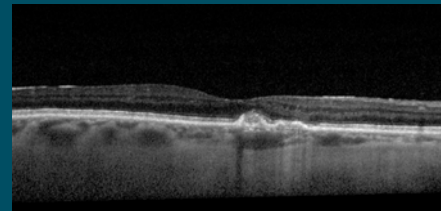
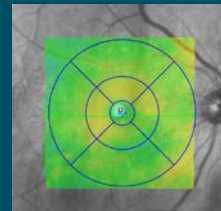
+4 week  
BCVA: 61 letters  
CST: 259  $\mu\text{m}$



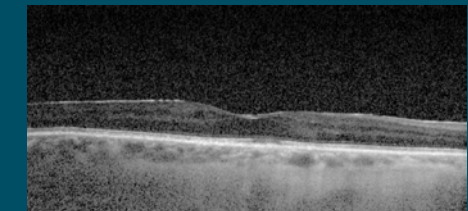
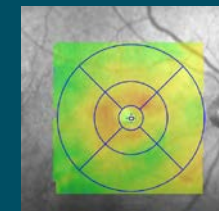
+52 weeks  
BCVA: 65 letters  
CST: 293  $\mu\text{m}$



+8 week  
BCVA: 65 letters  
CST: 254  $\mu\text{m}$



+84 weeks  
BCVA: 64 letters  
CST: 313  $\mu\text{m}$





# Fireside Chat with Dr. David Boyer

## **David S. Boyer, M.D.**

Senior Partner, Retina-Vitreous Associates Medical Group and Adjunct Clinical Professor of Ophthalmology, University of Southern California/ Keck School of Medicine, Los Angeles  
Investigator in OPTIC Phase 1 Trial



# Q&A

# Execution and Anticipated Milestones

## ADVM-022 for Wet AMD OPTIC Phase 1 Clinical Trial

- ☒ Presented additional data from Cohort 1 at Macula 20/20 in January 2020
- ☒ Presented new interim data from Cohorts 1 and 2 at Angiogenesis in February 2020
- ☒ Completed patient dosing in Cohort 3
- ☒ Presented new interim data from Cohorts 1-3 in May 2020
- ☒ Completed patient dosing in Cohort 4 in July 2020
- ☒ Presented new interim data from Cohorts 1-4 in August 2020
- ☒ Partial clinical hold removed by FDA
- ☒ Present additional data from Cohorts 1-4 in November 2020
- ☐ Present longer-term data, including additional anti-VEGF protein expression data, **1H2021**
- ☐ Initiate first pivotal trial **mid-2021**

## ADVM-022 for DME INFINITY Phase 2 Clinical Trial

- ☒ Submitted IND in DR to FDA
- ☒ Randomized first patient in July 2020
- ☐ Present data from INFINITY clinical trial (24-week primary endpoint assessment) **2H2021**

ADVERUM  
BIOTECHNOLOGIES