## EYP-1901 in wet AMD DAVIO 2 Phase 2 Clinical Trial Topline Data

December 4, 2023



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Committed to developing therapeutics to improve the lives of patients with serious retinal diseases

## Pipeline represents potentially substantial opportunities using our bioerodible Durasert E<sup>™</sup> IVT delivery technology

- **EYP-1901** vorolanib, a selective and patented TKI
  - Positive topline Phase 2 data in wet AMD
  - Phase 3 trials in wet AMD planned to initiate in 2H 2024
  - Topline Phase 2 data in NPDR anticipated in Q2 2024
  - Phase 2 trial in DME planned to commence in Q1 2024
- **EYP-2301** *razuprotafib,* a patented TIE-2 agonist for serious retinal diseases

#### Durasert<sup>®</sup> - proven, safe IVT drug delivery technology

- Routine in-office IVT injection
- Bioerodible and non-erodible formulations
- Safely administered to thousands of patient eyes across four FDA approved products with non-erodible formulations

#### **Strong Balance Sheet**

- \$136.0M of cash and investments on September 30, 2023
- Cash runway into 2025

IVT, intravitreal injection



EYP-1901 (vorolanib in Durasert E) a potential treatment for wet AMD featuring sustained delivery for 6-months or longer





There is a Significant Need for More Durable Therapies in Wet AMD



#### Many patients with wet AMD are chronically undertreated

 >80% of Retina Specialists say undertreatment is due to patient noncompliance, scheduling limitations or provider preference for less frequent dosing<sup>1</sup>



## Current "treat and extend" protocol still places significant burden on physicians and patients

 Chronic disease treated with short acting anti-VEGF biologics



#### A delay in care/missed visit can result in vision loss

A delay in treatment of only 5.34 weeks resulted in vision loss<sup>2</sup>

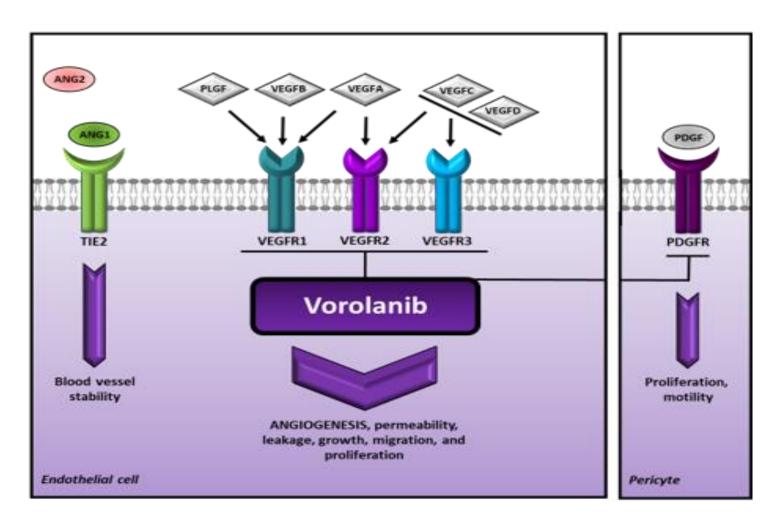


- An aging population means significantly more injections in a patient's lifetime
  - Current anti-VEGF treatments are dosed on average every two months in the United States<sup>3</sup>

1. 2022 PAT Survey; 2. American Academy of Ophthalmology, *The Effect of Delay in Care Among Patients Requiring Intravitreal Injections*, Welin Song, BS et al; 3. NIH *Current and Upcoming Anti-VEGF Therapies and Dosing Strategies for the treatment of neovascular AMD: a comparative review*, Saira Khanna et al, Dec. 2019

## Vorolanib Brings a Potential New MOA to the Treatment of VEGF-Mediated Retinal Diseases by Blocking all Isoforms of VEFG and PDGF

- Potent and selective pan–VEGF receptor inhibition
- Composition of matter patent into 2037
- Demonstrated neuroprotection in a validated retinal detachment animal model
- Blocks PDGF which may lead to antifibrotic benefit
- Reduced off-target binding and does not inhibit TIE-2 at clinically relevant doses



SoC, standard of care; ANG, angiopoietin; PDGF(R), platelet-derived growth factor (receptor); PLGF, placental growth factor; TIE-2, tyrosine-protein kinase receptor TIE-2; VEGF(R), vascular endothelial growth factor (receptor).



### TECHNOLOGY DURASERT®



#### Safe Sustained IVT Drug Delivery

- Delivered by a single in-office IVT injection
- Continuous, stable release of drug
- Zero-order kinetics

#### Durasert E<sup>™</sup>: bioerodible

- Insert consists of drug embedded within a bioerodible matrix
- Designed to deplete drug load before matrix fully erodes
- ILUVIEN<sup>®1</sup>

YUTIQ<sup>®1</sup>

- RETISERT<sup>®2</sup>
- VITRASERT<sup>®2</sup>

Durasert<sup>®</sup>: non-erodible

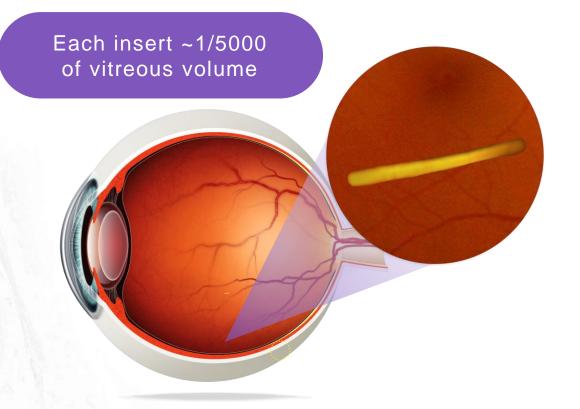
bioerodible matrix

polyimide shell:

• Drug embedded within a

coated with non-erodible

## EYP-1901: Receptor Binding Vorolanib In Bioerodible Durasert E<sup>™</sup>



- Delivered in the physician office via standard intravitreal injection technique
- Immediately bioavailable featuring an initial burst of drug followed by zero order kinetics release for ~9 months
- Positive safety and efficacy data in wet AMD from Phase 1 DAVIO clinical trial
- Continued positive safety data in ongoing Phase 2 clinical trials with all patients at least six months post injection
- Shipped and stored at ambient temperature





## The DAVIO 2 Clinical Trial – Background

A non-inferiority trial evaluating two doses of EYP-1901 against an aflibercept control in wet AMD The DAVIO 2 clinical trial was designed to evaluate EYP-1901 in wet AMD and support Phase 3 clinical trials based on a Type C meeting with FDA

**Design:** Multi-center, randomized, double-masked trial in patients with previously treated wet AMD

#### **Anti-VEGF supplement criteria:**

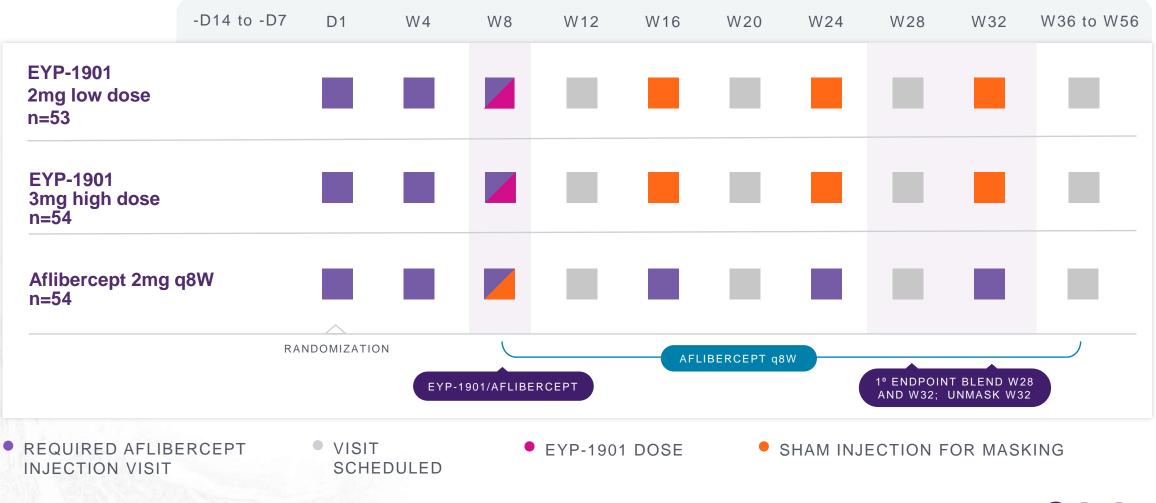
- 5 letter loss with 75 microns of new fluid
- Other criteria
  - 10 letter loss due to wet AMD
  - 100 microns new fluid x 2 visits
  - New retinal hemorrhage from wet AMD
  - Investigator discretion

**Primary outcome:** difference in mean change in BCVA from Day 1 to Week 28 and 32 (blended)

**Key secondary endpoints:** safety, reduction in treatment burden, percent of eyes supplement-free up to six months and anatomical results



# DAVIO 2 Clinical Trial is Randomized, Double-Masked, Aflibercept Controlled\* with a Single EYP-1901 Treatment at Two Doses



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\*Aflibercept on-label control required by FDA



## DAVIO 2 Baseline Characteristics Well Balanced Across Arms

	Aflibercept 2mg q8W (n=54)	EYP-1901 2mg (n=50)	EYP-1901 3mg (n=52)
Mean age, years (range)	75.9 (52-93)	76.4 (61-93)	75.4 (56-89)
Female, %	53.7%	64.0%	67.3%
Mean BCVA, ETDRS letters (range)	73.4 (41-85)	73.9 (52-84)	74.9 (46-85)
Mean CST, µm (range)	265.7 (178-348)	267.0 (192-400)	262.9 (186-345)
Median length of time for wet AMD diagnosis prior to screening, months (range)	28.1 (2.4-273.6)	24.2 (2.4-168.0)	28.1 (2.4-145.2)
Mean # of injections normalized to 12 months prior to screening (range)	9.5 (1.0-12.0)	10.1 (2.0-13.0)	10.0 (2.0-13.3)



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## Phase 2 DAVIO 2 Clinical Trial Topline Results

A NON-INFERIORITY TRIAL VERSUS AN AFLIBERCEPT CONTROL





## EYP-1901 Phase 2 DAVIO 2 Clinical Trial Met All Objectives

#### MET PRIMARY ENDPOINT

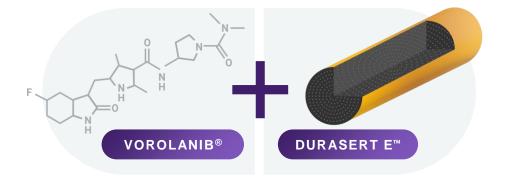
 Statistically non-inferior change in BCVA versus aflibercept control with a numerical difference of only -0.3 and -0.4 letters, respectively for the 2 mg and 3 mg dose

#### ✓ Continued favorable safety profile:

- No EYP-1901-related SAEs reported
- Ocular AEs majority are mild in severity and expected with mode of administration

#### MET ALL SECONDARY ENDPOINTS

- ✓ ~80% reduction in treatment burden at 6-months
- Nearly two-thirds of eyes supplement-free up to sixmonths
- ✓ Strong anatomical control in both EYP-1901 arms





DAVIO 2 Achieved Primary Endpoint of Non-Inferiority Change in Visual Acuity in Both EYP-1901 Arms; Statistically and Numerically Non-Inferior to Control (95% CI)

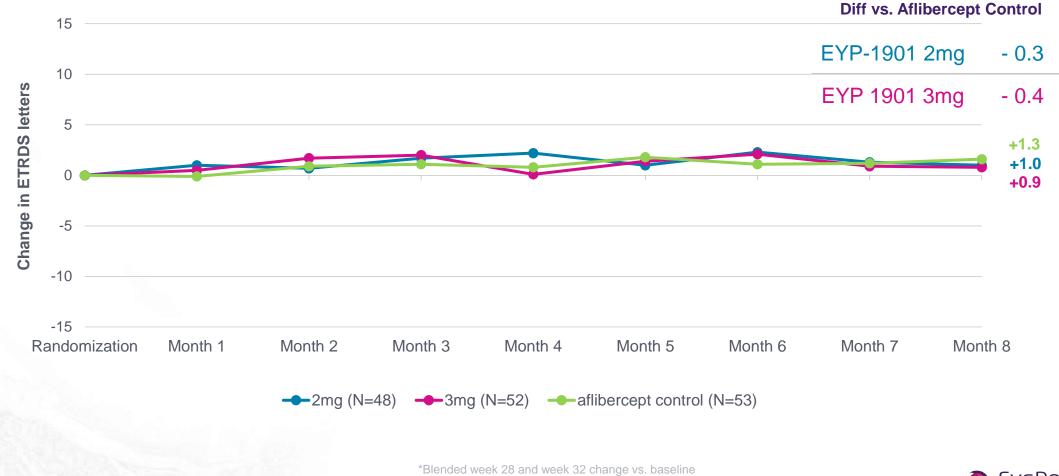
	EYP-1901 2mg	EYP-1901 3mg	Aflibercept 2mg q8W
Change in BCVA* vs. Baseline	+ 1.0 letters	+ 0.9 letters	+ 1.3 letters
Difference vs. Aflibercept Control	- 0.3 letters	- 0.4 letters	NA
Statistics	NI (95% CI)	NI (95% CI)	NA

#### Non-inferiority Margin = - 4.5 letters per FDA guidance

\*Blended week 28 and week 32 NI, Non-inferior; CI, Confidence Interval PRELIMINARY DATA – PENDING FINAL ANALYSIS



# EYP-1901 was Statistically Non-Inferior in Change in BCVA Compared to the Aflibercept Control (95% CI)



MEAN CHANGE IN BCVA FROM BASELINE

\*Blended week 28 and week 32 change vs. baseline CI, Confidence Interval PRELIMINARY DATA – PENDING FINAL ANALYSIS



# EYP-1901 Demonstrated a Favorable Safety Profile in the Phase 2 DAVIO 2 Trial (Data Cut as of November 7, 2023)

- No reported EYP-1901-related ocular SAEs
  - Four ocular SAEs reported in a study eye none deemed related to EYP-1901<sup>1</sup>
- No reported EYP-1901-related systemic SAEs
- AEs reported were generally mild and expected with IVT<sup>2</sup>
- No cases of:
  - Insert migration into the anterior chamber
  - Retinal occlusive vasculitis
- Low patient discontinuation rate of 4% up to week 32
  - No discontinuations were related to AEs or EYP-1901 treatment

 1- As determined by investigator
 2- Further details to be provided following completion of internal review SAE, serious adverse event; AE, adverse event; IVT, intravitreal injection
 PRELIMINARY DATA – PENDING FINAL ANALYSIS



### Clinically Meaningful Reduction in Treatment Burden Supports EYP-1901 as a Maintenance Treatment For Wet AMD

	EYP-1901 2mg	EYP-1901 3mg
Mean number of injections week 8 through week 32	0.55	0.71
Mean number of injections 6 months prior to screening*	5.07	4.98
Reduction in treatment burden vs. 6 months prior (%)	89%	85%



# EYP-1901 Demonstrated a Significant Reduction in Treatment Burden vs. the Aflibercept Control Arm

	EYP-1901 2mg	EYP-1901 3mg	Aflibercept 2mg q8W
Mean number of injections week 8 through week 32	0.55	0.71	3.32
Reduction in treatment burden vs. aflibercept control (%)	83%	79%	NA



## EYP-1901 Demonstrated Clinically Meaningful Supplement-Free Rates

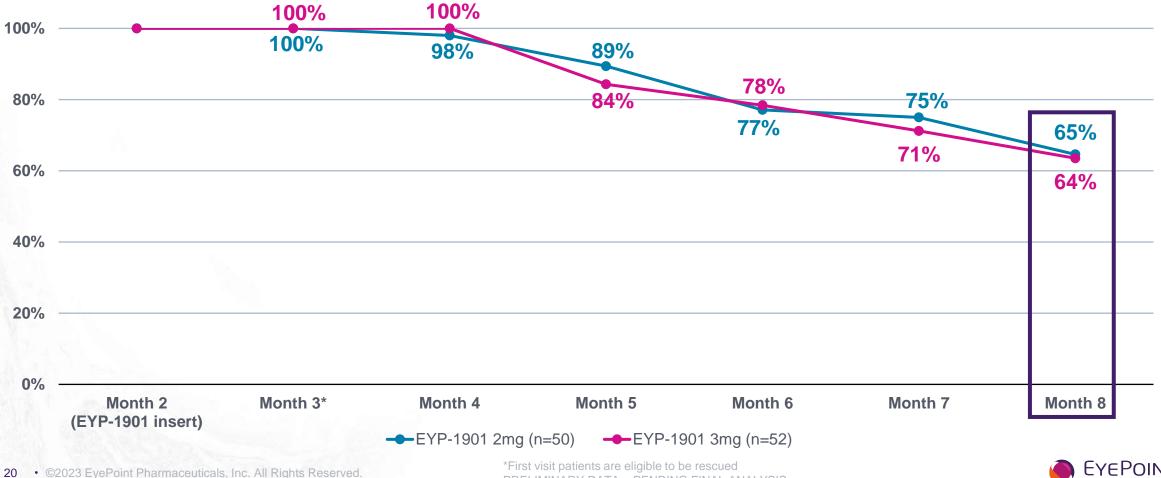
#### PERCENT OF PATIENTS SUPPLEMENT-FREE UP TO SIX MONTHS AFTER ADMINISTRATION OF EYP-1901

	EYP-1901 2MG	EYP-1901 3mg
Supplement-Free Rates	65%	64%



Nearly Two-Thirds of Eyes Treated with EYP-1901 were Supplement-Free up to Six Months After a Single Injection

#### SUMMARY OF SUPPLEMENT-FREE RATES BY WEEK



PRELIMINARY DATA - PENDING FINAL ANALYSIS

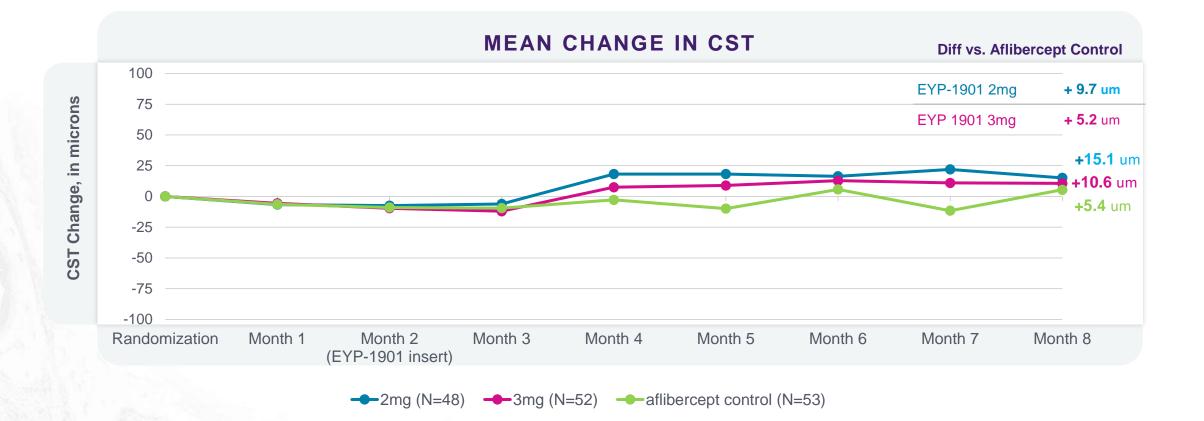
PHARMACEUTICALS

### Stable Anatomy via OCT in EYP-1901 Arms

	EYP-1901 2mg	EYP-1901 3mg	Aflibercept 2mg q8W
Baseline CST (mean)	262.9	267.0	265.7
Change in OCT at Week 32 vs. Baseline	+ 15.1 microns	+ 10.6 microns	+ 5.4 microns
Difference vs. Aflibercept Control	+ 9.7 microns	+ 5.2 microns	NA



### Data from DAVIO 2 Suggests Strong Anatomic Control with OCT Change Below 10 microns at Week 32 Compared to the Aflibercept Control



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## EYP-1901 Phase 2 DAVIO 2 Clinical Trial Met All Objectives

#### MET PRIMARY ENDPOINT

 Statistically non-inferior change in BCVA versus aflibercept control with a numerical difference of only -0.3 and -0.4 letters, respectively for the 2 mg and 3 mg dose

#### ✓ Continued favorable safety profile:

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#### MET ALL SECONDARY ENDPOINTS

- ✓ ~80% reduction in treatment burden at 6-months
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# EYP-1901 Phase 2 DAVIO 2 Clinical Trial Met All Primary and Secondary Endpoints

Endpoint	DAVIO 2 Topline	DAVIO 2 Lower Limit (as included in November deck)
Mean change in BCVA vs. aflibercept control	<ul> <li>- 0.3 letters (EYP-1901 2mg)</li> <li>- 0.4 letters (EYP-1901 3mg)</li> <li>Statistically non-inferior (CI 95%)</li> </ul>	<ul><li>&lt; - 3.0 letters</li><li>Potentially underpowered</li></ul>
Safety	<ul> <li>No reported EYP-1901-related ocular SAEs</li> <li>No reported EYP-1901- related systemic SAEs</li> </ul>	Favorable safety profile
Reduction in treatment burden	<ul> <li>89% (EYP-1901 2mg)*</li> <li>85% (EYP-1901 3mg)*</li> </ul>	50% or better
Supplement-free rate	<ul> <li>65% (EYP-1901 2mg)</li> <li>64% (EYP-1901 3mg)</li> </ul>	50% or better
Mean change in CST on OCT	<ul> <li>+ 15.1 microns (EYP-1901 2mg)</li> <li>+ 10.6 microns (EYP-1901 3mg)</li> </ul>	Within ~30 microns



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