# Ocular Modifier Gene Therapy Program Update

A PHASE 1/2 STUDY TO ASSESS THE SAFETY AND EFFICACY OF OCU400 (MODIFIER GENE THERAPY) FOR RETINITIS PIGMENTOSA ASSOCIATED WITH *NR2E3* AND *RHO* MUTATIONS AND LEBER CONGENITAL AMAUROSIS WITH MUTATION(S) IN *CEP290* GENE

### Forward Looking Statements

This presentation contains forward-looking statements within the meaning of The Private Securities Litigation Reform Act of 1995, which are subject to risks and uncertainties, including, but not limited to, statements regarding the development of OCU400 and the interpretation of preliminary clinical trial results. We may, in some cases, use terms such as "predicts," "believes," "potential," "proposed," "continue," "estimates," "anticipates," "expects," "plans," "intends," "may," "could," "might," "will," "should," or other words that convey uncertainty of future events or outcomes to identify these forward-looking statements. Such statements are subject to numerous important factors, risks, and uncertainties that may cause actual events or results to differ materially from our current expectations, including, but not limited to, the risk that preliminary clinical data may not be indicative of final clinical data or data in later stage clinical trials. These and other risks and uncertainties are more fully described in our periodic filings with the Securities and Exchange Commission (SEC), including the risk factors described in the section entitled "Risk Factors" in the quarterly and annual reports that we file with the SEC. Any forward-looking statements that we make in this presentation speak only as of the date of this presentation. Except as required by law, we assume no obligation to update forward-looking statements contained in this presentation whether as a result of new information, future events, or otherwise, after the date of this presentation.



### Today's Agenda

Opening

Overview of Preliminary Safety and Efficacy Results

Closing

Shankar Musunuri, PhD, MBA Chairman, CEO and Co-founder, Ocugen

Arun Upadhyay, PhD Chief Scientific Officer, Head of Research, Development and Medical, Ocugen

Shankar Musunuri, PhD, MBA

#### Q&A

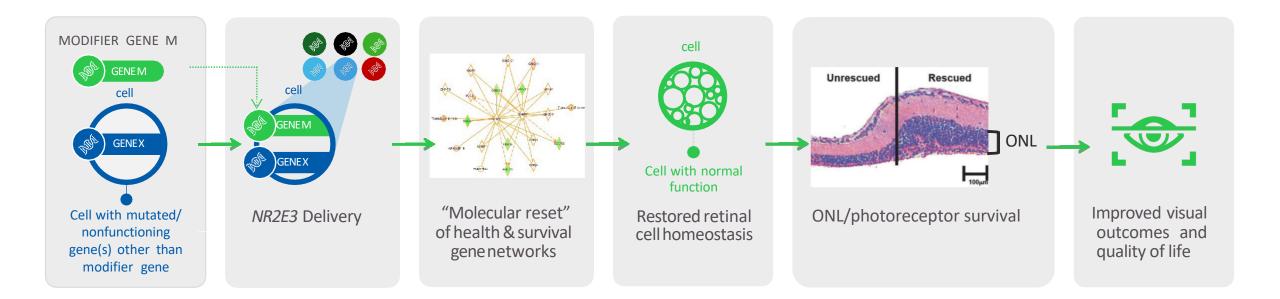
- Huma Qamar, MD, MPH, Head of Clinical Development and Medical Affairs, Ocugen
- David Birch, PhD, Scientific Director, Retina Foundation of the Southwest, Primary Investigator of the Study
- Neena B. Haider, PhD, Fellow of ARVO and Inventor of Modifier Gene Therapy



# Modifier Gene Therapy: A Broader Reach

Gene modifier therapy can potentially address multiple genetic defects with a single product utilizing a gene agnostic approach.

In patients with IRDs, this could mean:





# **Study Overview**

**Primary Endpoint: Safety** 

Safety of subretinal administration of OCU400

#### **Exploratory Endpoint: Efficacy**

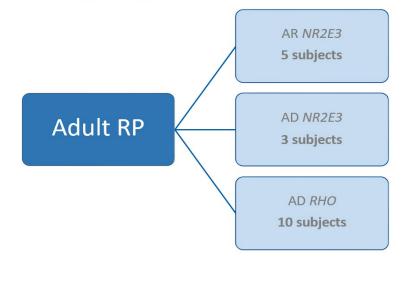
Multi-Luminance Mobility Test (MLMT)

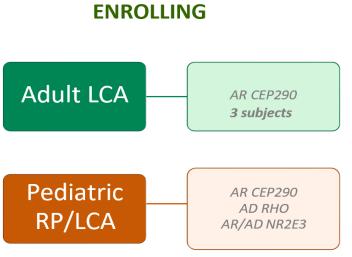
Best Corrected Visual Acuity (BCVA)

Clinical Trials.gov Identifier: NCT05203939

# **Enrollment Status**

#### COMPLETED







Multi-Luminance Mobility Test		
	Total Subjects for analyses (N=7) Subjects with 9-months follow-up : Cohort 1, N=3 Subjects with 6 months follow-up: N=1 from Cohor1 and N=3 from Cohort 2	
	Improvement ≥ 1 Lux	Improvement ≥ 2 Lux
Treated Eye	71.4%	28.6%
Untreated Eye	28.6%	0.0%

- 100% of treated eyes showed stability or improved MLMT scores
- 71.4% of treated eyes improved by at least 1 Lux Level vs ONLY 28.6% of untreated eyes
- 28.6 % of treated eyes improved by at least 2 Lux Level vs 0 % of untreated eyes

One subject had advanced RP at baseline with subsequent foveal detachment



#### MLMT is used as efficacy measure to assess visual function



Best Corrected Visual Acuity (BCVA) Score		
	Total Subjects for analyses (N=7) Subjects with 9-months follow-up : Cohort 1, N=3 Subjects with 6 months follow-up: N=1 from Cohor1 and N=3 from Cohort 2	
	Improvement ≥ 8 Letters	
Treated Eye	42.9%	
Untreated Eye	0.0%	



# OCU410: Dry Age-related Macular Degeneration (dAMD) and Stargardt Disease (STGD)

#### Dry AMD

#### Limited options, presenting significant unmet medical need

- US: 10M
- Worldwide: condition affects more than 266M people

#### Stargardt-an orphan disease

#### No treatment options exist

- US: 35,000
- Worldwide: condition affects approximately 800,000 people

# Recently approved therapy for geographic atrophy (GA)—advanced form of dAMD—has limitations

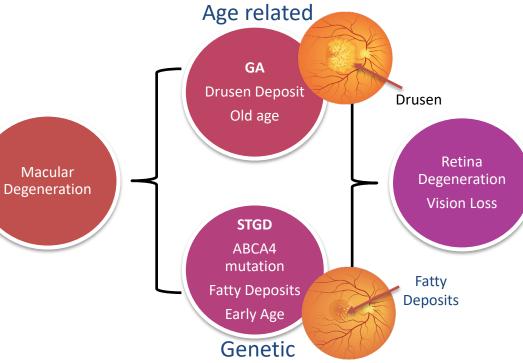
- Frequent intravitreal injections (N ~6-12 doses per year); Patient compliance
- Limited effect of GA lesion growth rate
- Approximately 12% of patients experience neovascular AMD when the drug is administered every month for two years

#### OCU410 addresses shortcomings of current approaches

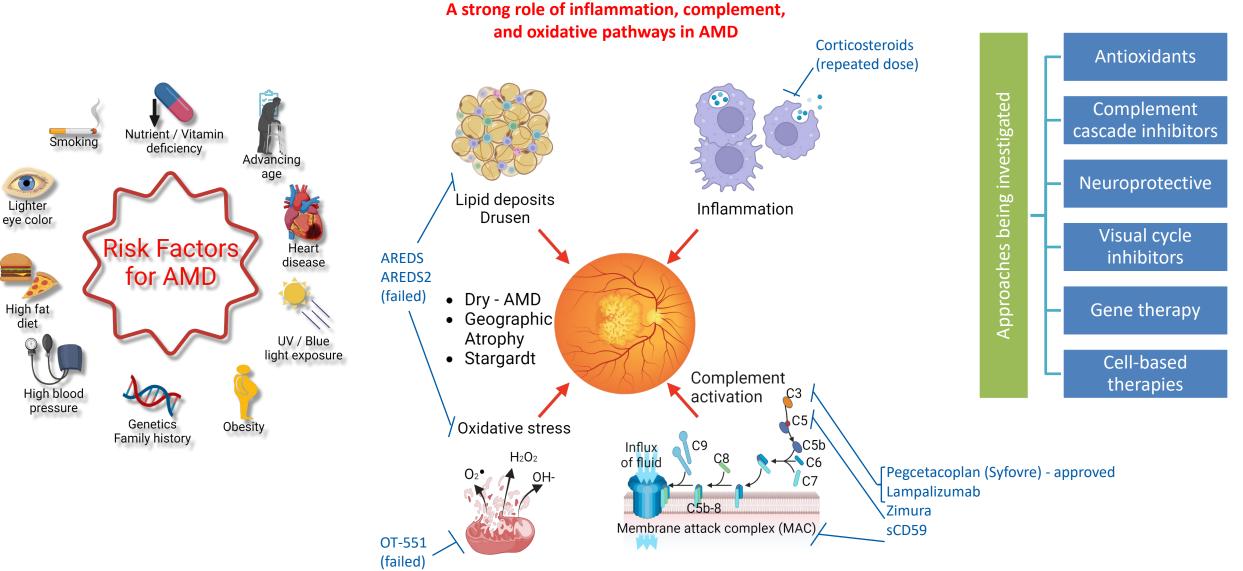
- Broad-spectrum, gene-agnostic approach
- Potential one-time, curative therapy with a single sub-retinal injection, using RORA

#### Plan to Initiate Phase 1/2 clinical trial in 2Q 2023





# AMD: Risk Factors, Treatment Options and Unmet Needs



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#### OCU410 (RORA): A Potential Modifier Therapeutic for Dry-AMD and STGD

7 Months RORA/GAPDH -1β /GAPI gene expi 200-Reduced cence (AU) 129S1 Drusen -<u>'</u> e 150-0.5 Rela 1.25 0.62 1.25 0.62 2.5 2.5 Autofluore 100-LPS Concentration [ in mg/mL] LPS Concentration [ in mg/mL] Untreated - ARP-19 Cells Abca4-/-IL-8 Lipid RORA-WT clone metabolism Blue /GAPDH Regulation of Inflammation OCU410 High Dose Months 1 7 1 7 1 7 Treated Abca4-/-Abca4<sup>-/-</sup> Abca4<sup>-/-</sup> 1.25 0.62 2.5 10 Control 0.62 1.25 2.5 10 0 5 OCU410 Uninjected LPS Concentration [ in mg/mL] LPS Concentration [ in mg/mL] **RORA / Macular Degeneration Model** NR1F1 Anti-complement: Increased anti-compliment (Cd59) protein Anti-oxidative: Improves ARPE19 cells survival 6 months **B6** Complement 90 ONL 0 activation RORA C3 IS/OS Norgmalized Cell Surgival Oxidative stress -------------------------------NR2E3 C5  $H_2O_2$ Untreated ------------------------------NR1D1 C9 C5b Influx of  $O_2^{\bullet}$ OHfluid Abca -/- mice OCU410-Treated Untre ONL IS/OS C5b-8 **CD59** Membrane attack complex (MAC) ONL IS/OS 50 1E3 1E4 1E5 C5b678 Cells 1E2 1e6 CD59 and DAPI

Anti-drusen activity and improves retinal function

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Anti-inflammatory: Suppresses inflammation in HMC3 cells

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Q&A

# Ocugen<sup>™</sup> Vision

Fully integrated, patient-centric biotech company focused on vaccines in support of public health and gene and cell therapies targeting unmet medical needs through **Courageous Innovation** 

