DEVELOPING TRANSFORMATIVE THERAPIES FOR RETINAL DISEASES

August 2021 NASDAQ: ISEE

Forward-looking statements

Any statements in this presentation about the Company's future expectations, plans and prospects constitute forward-looking statements for purposes of the safe harbor provisions under the Private Securities Litigation Reform Act of 1995. Forward-looking statements include any statements about the Company's strategy, future operations and future expectations and plans and prospects for the Company, and any other statements containing the words "anticipate," "believe," "estimate," "expect," "intend", "goal," "may", "might," "plan," "predict," "project," "seek," "target," "potential," "will," "would," "could," "continue," and similar expressions.

In this presentation, the Company's forward looking statements include statements about its expectations regarding availability of top-line data from and patient retention in its second Phase 3 trial (GATHER2) of Zimura in geographic atrophy secondary to AMD and use of its completed clinical trial of Zimura for the treatment of geographic atrophy secondary to AMD (GATHER1) as a Phase 3 trial, its development and regulatory strategy for Zimura and its other product candidates, including additional indications that the Company may pursue for the development of Zimura and IC-500, the Company's hypotheses regarding complement inhibition and HtrA1 inhibition as potential mechanisms of action for the treatment of retinal diseases, the implementation of its business and hiring plan, preliminary financial information, the timing, progress and results of clinical trials and other research and development activities, including regulatory submissions, the clinical meaningfulness of clinical trial results, the potential utility of its product candidates, estimates regarding the number of patients affected by the diseases and indications the Company's business development strategy.

Such forward-looking statements involve substantial risks and uncertainties that could cause the Company's research and 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, those related to the progression and duration of the COVID-19 pandemic and responsive measures thereto and related effects on the Company's research and development programs, operations and financial position, expectations for regulatory matters, the initiation and the progress of research and development programs and clinical trials, including enrollment and retention in clinical trials, availability of data from these programs, reliance on contract development and manufacturing organizations, contract research organizations and other third parties, establishment of manufacturing capabilities, developments from the Company's competitors and the marketplace for the Company's products, human capital matters, need for additional financing and other factors discussed in the "Risk Factors" section contained in the quarterly and annual reports that the Company files with the Securities and Exchange Commission.

Any forward-looking statements represent the Company's views only as of the date of this presentation. The Company anticipates that subsequent events and developments may cause its views to change. 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 except as required by law.

Diversified portfolio focused on retinal diseases

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Therapeutics for Age-Related Retinal Diseases (Large Market)

- Zimura (C5 inhibitor):
 - Positive data for the first of two Phase 3 trials (GATHER1)
 - Statistically significant 27% reduction in GA growth over 12 months (primary endpoint achieved)
 - Completed patient enrollment for second Phase 3 trial (GATHER2) in July 2021; topline data expected in 2H2022
 - Received Special Protocol Assessment (SPA) from FDA for GATHER2
 - Plan to file for NDA / MAA approvals following positive 12-month GATHER2 data
 - Plan to initiate clinical development in drusen with additional lifecycle initiatives ongoing
- IC-500 (HtrA1 Inhibitor): Complementary MOA adding to development stage AMD franchise

Cash Position

• Expected YE 2021 cash: \$215 million - \$225 million¹

¹ Estimate as of 8/4/21

Diversified portfolio focused on retinal diseases

Gene Therapy for Inherited Retinal Diseases (Orphan)

- Broad and diversified pipeline
 - Novel and cutting edge AAV gene therapy options
 - Five R&D programs in orphan inherited retinal diseases w/ no currently approved therapies in target diseases

Experienced Team with Extensive Drug Development Expertise in Retina

STRONG SENIOR TEAM WITH SIGNIFICANT OPHTHALMOLOGY EXPERIENCE



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Iveric Bio Pipeline

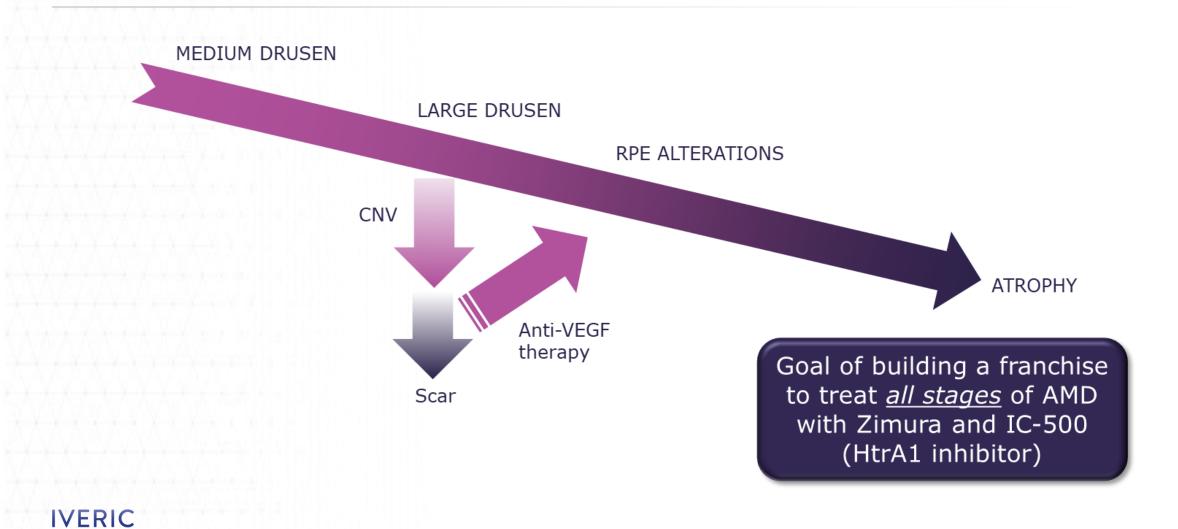
		Preclinical	Phase 1	Phase 2	Phase 3	
VA AVA	THERAPEUTICS PIPELINE					Status/Milestones
	Avacincaptad pegol (ZIMURA) GA secondary to AMD					GATHER1 (1 st Phase 3): Positive 12 & 18 month data reported GATHER2 (2 ND Phase 3): enrollment complete and SPA received; top-line data expected <u>2H 2022</u>
	Avacincaptad pegol (ZIMURA) Autosomal recessive Stargardt disease (STGD1)				•	Expanded enrollment (up to ~25 additional patients); trial on-going
	IC-500: HtrA1 inhibitor GA secondary to AMD					Plan to file IND in <u>2H 2022</u>
	AAV GENE THERAPY PIPELINE					
	IC-200: BEST1-Related IRDs					Plan to initiate Phase 1/2 in autosomal recessive bestrophinopathy in ${\bf Q42021}$
	IC-100: RHO-adRP Rhodopsin-mediated autosomal- dominant RP					Further development under evaluation
	mini-CEP290: LCA10 Leber congenital amaurosis type 10					Lead construct identified; planning development
	mini-ABCA4: STGD1* Autosomal recessive Stargardt disease (STGD1)					Evaluating preclinical data
	mini-USH2A: USH2A-Related IRDs*					Preliminary results expected 1H 2022

AGE-RELATED MACULAR DEGENERATION (AMD)

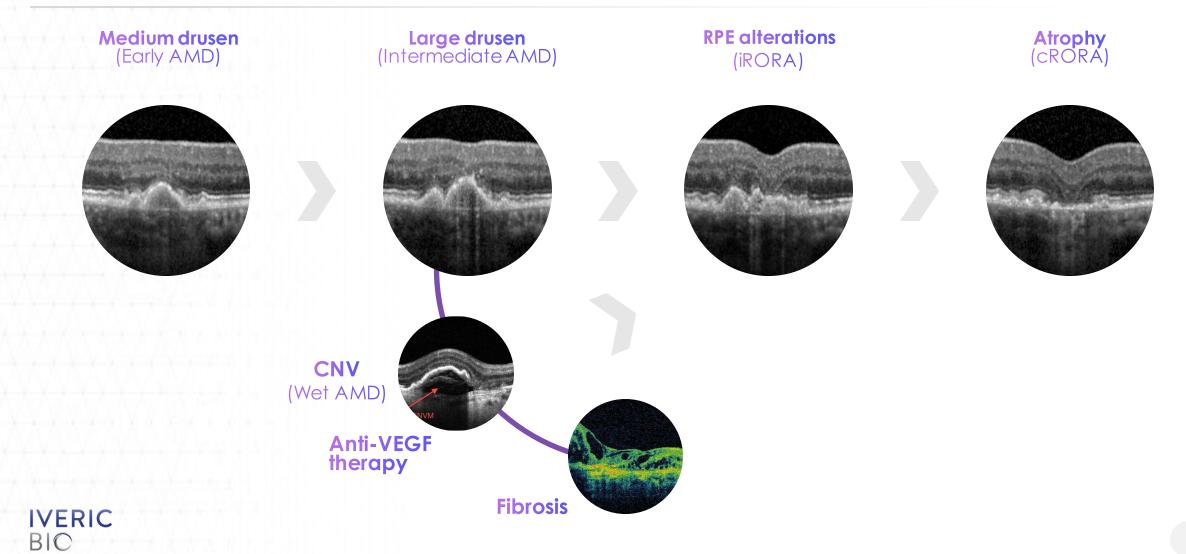
Disease Overview & Market Size

Pathway of AMD disease progression

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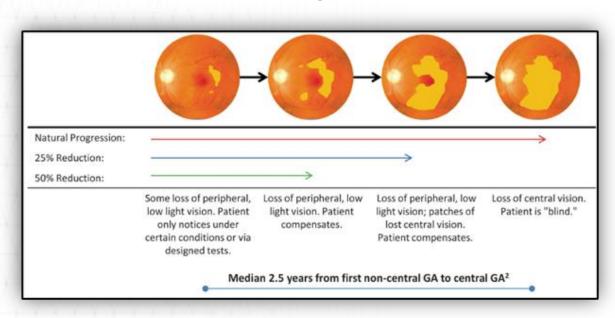


Pathway of AMD disease progression



Growth rate and loss of vision depend on GA location

Geographic Atrophy: loss of photoreceptors over time

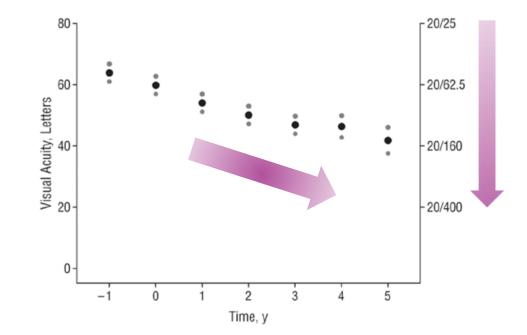


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Increase In Area of Degeneration Over Time

Loss of Vision Over Time



GA severely impacts vision in ~1.5 million patients in the US alone

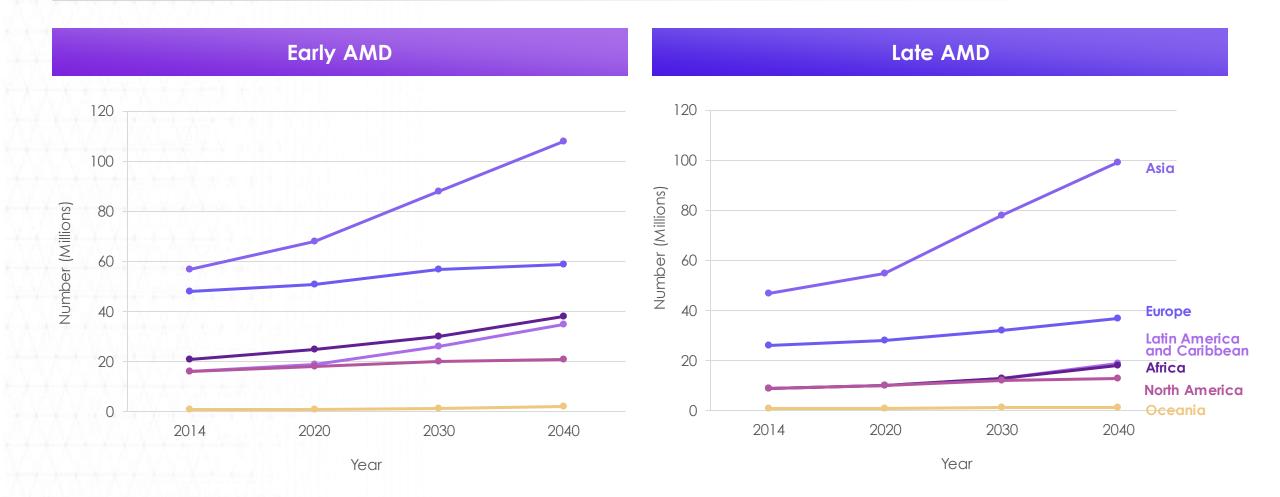
Leading cause of central vision loss in individuals over 50 years old in developed countries¹ Severely affects vision and often threatens complete vision loss in an estimated **1.5 million** individuals in the United States and **5 million** individuals worldwide² Early signs of retinal changes are seen in individuals as young as **30–40 years old**³ Studies show **GA** severity increases with age¹

One-third of the population is affected by GA by the time individuals are 80 years old³

IVERIC BIC 1. Ferris FL, Wilkinson CP, Bird A, Chakravarthy U, Chew E, Csaky K, et al. Clinical Classification of Age-related Macular Degeneration. Ophthalmology. 2013;120(4):844-851. 2. Boyer DS, Schmidt-Erfurth U, van Lookeren Campagne M, Henry EC, Brittain C. The pathophysiology of geographic atrophy secondary to age-related macular degeneration and the complement pathway as a therapeutic target. *Retina*. 2017;37(5):819-835. 3. Ratnayaka JA, Lotery AJ. Challenges in studying geographic atrophy (GA) age-related macular degeneration: the potential of a new mouse model with GA-like features. *Neural Regen Res*. 2020;15(5):863-864. doi:10.4103/1673-5374.268972.

AMD is projected to increase in global prevalence

Projected number of individuals with AMD by region¹



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COMPLEMENT ACTIVATION IN GA

What We Know About the Role of Complement in the Pathogenesis of GA

Genetic link: Complement & AMD

Complement Factor H Polymorphism in Age-Related Macular

Degeneration

Robert J. Klein¹, Caroline Zeiss^{2,*}, Emily Y. Chew^{3,*}, Jen-Yue Tsai^{4,*}, Richard S. Sackler¹, Chad Haynes¹, Alice K. Henning⁵, John Paul SanGiovanni³, Shrikant M. Mane⁶, Susan T. Mayne⁷, Michael B. Bracken⁷, Frederick L. Ferris³, Jurg Ott¹, Colin Barnstable², and Josephine Hoh.^{7,†}

"In individuals homozygous for the risk allele, the likelihood of AMD is increased by a factor of 7.4" *

THE PATHOPHYSIOLOGY OF GEOGRAPHIC ATROPHY SECONDARY TO AGE-RELATED MACULAR DEGENERATION AND THE COMPLEMENT PATHWAY AS A THERAPEUTIC TARGET

DAVID S. BOYER, MD,* URSULA SCHMIDT-ERFURTH, MD,† MENNO VAN LOOKEREN CAMPAGNE, PHD,‡ ERIN C. HENRY, PHD,‡ CHRISTOPHER BRITTAIN, MBBS§

Complement System in Pathogenesis of AMD: Dual Player in Degeneration and Protection of Retinal Tissue

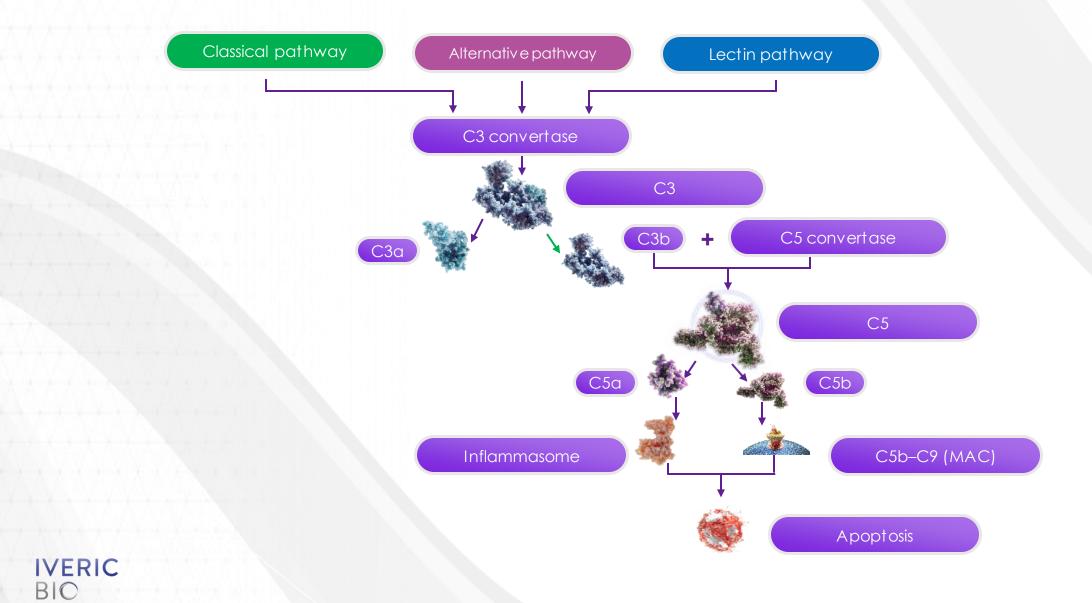
Milosz P. Kawa,¹ Anna Machalinska,^{2,3} Dorota Roginska,¹ and Boguslaw Machalinski¹

Complement Activation Levels Are Related to Disease Stage in AMD

Thomas J. Heesterbeek,¹ Yara T. E. Lechanteur,¹ Laura Lorés-Motta,^{1,2} Tina Schick,³ Mohamed R. Daha,⁴ Lebriz Altay,³ Sandra Liakopoulos,³ Dzenita Smailhodzic,¹ Anneke I. den Hollander,^{1,2} Carel B. Hoyng,¹ Eiko K. de Jong,¹ and B. Jeroen Klevering¹

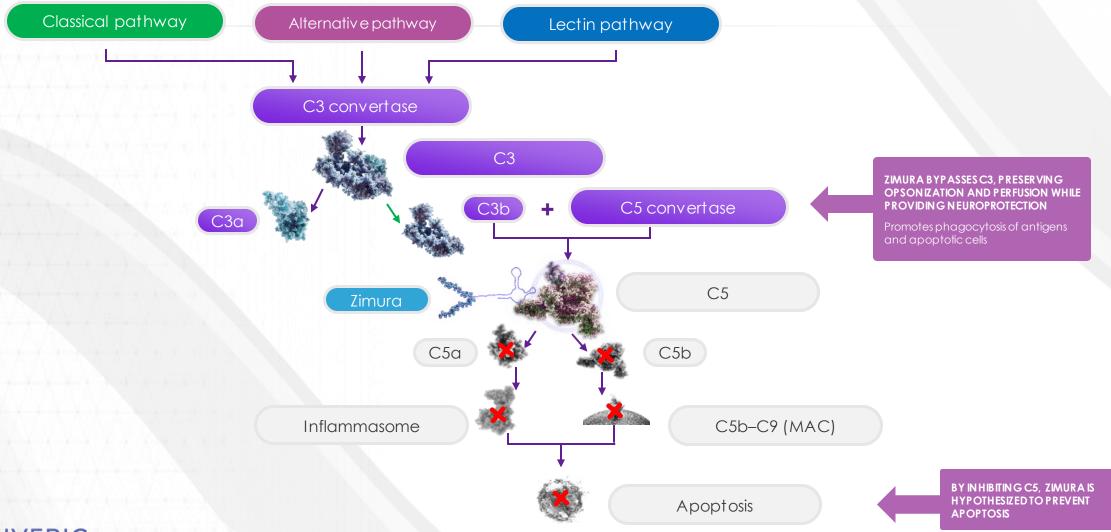
* Quotation from: Klein, et al. Complement Factor H Polymorphism in Age-Related Macular Degeneration. *Science*. 2005 April 15; 308(5720): 385-389.

Activated complement leads to inflammation and cell death



WHY IS ZIMURA® IMPORTANT?

Zimura targets C5, inhibiting the 2 triggers of cell death, preserving the remainder of the pathway



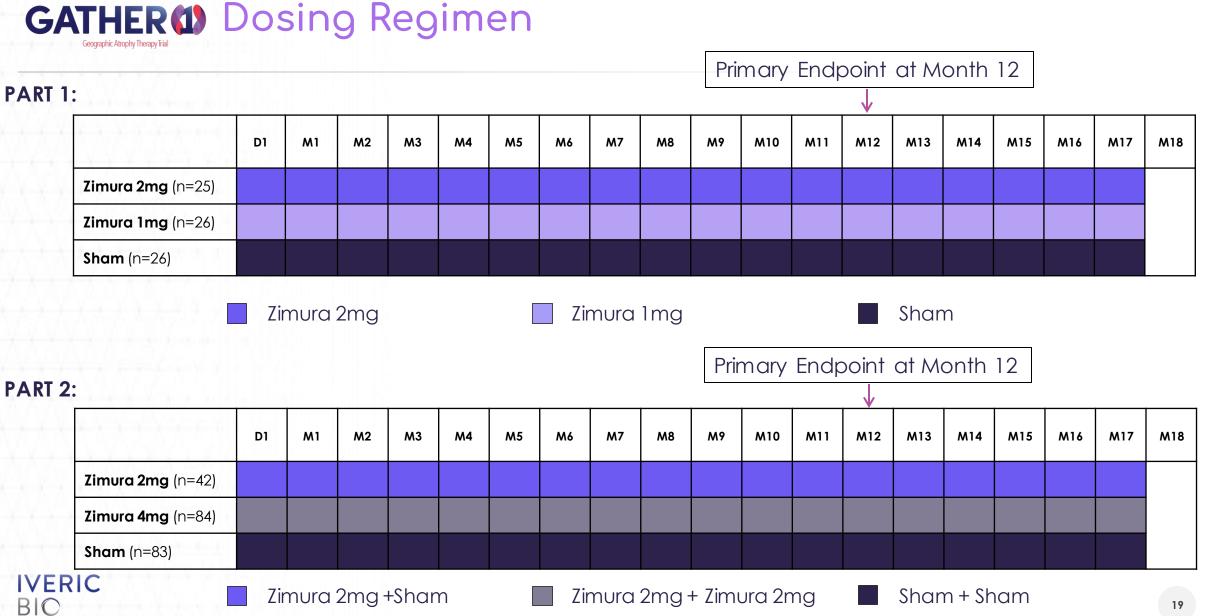
IVERIC BIC

Xu H, Chen M. Targeting the complement system for the management of retinal inflammatory and degenerative diseases. *European Journal of Pharmacology*. 2016;787:94-104.

ZIMURA® PHASE 3 PROGRAM IN GA SECONDARY TO AMD



(Geographic Atrophy Therapy Trials)



GATHER1: Key Inclusion Criteria

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Non-foveal GA secondary to dry AMD

- Total GA area \geq 2.5 and \leq 17.5 mm² (1 and 7 disk areas [DA] respectively), determined by screening images of FAF
- If GA is multifocal, at least one focal lesion should measure \geq 1.25 mm² (0.5 DA)
- GA in part within 1500 microns from the foveal center
- The atrophic lesion must be able to be photographed in its entirety
- Best corrected visual acuity in the SE between 20/25 20/320, inclusive

GATHER1: Primary efficacy endpoint achieved

Mean Rate of Change in Geographic Atrophy (GA) Area from Baseline to Month 12 (MRM Analysis) (Square Root Transformation, ITT Population)

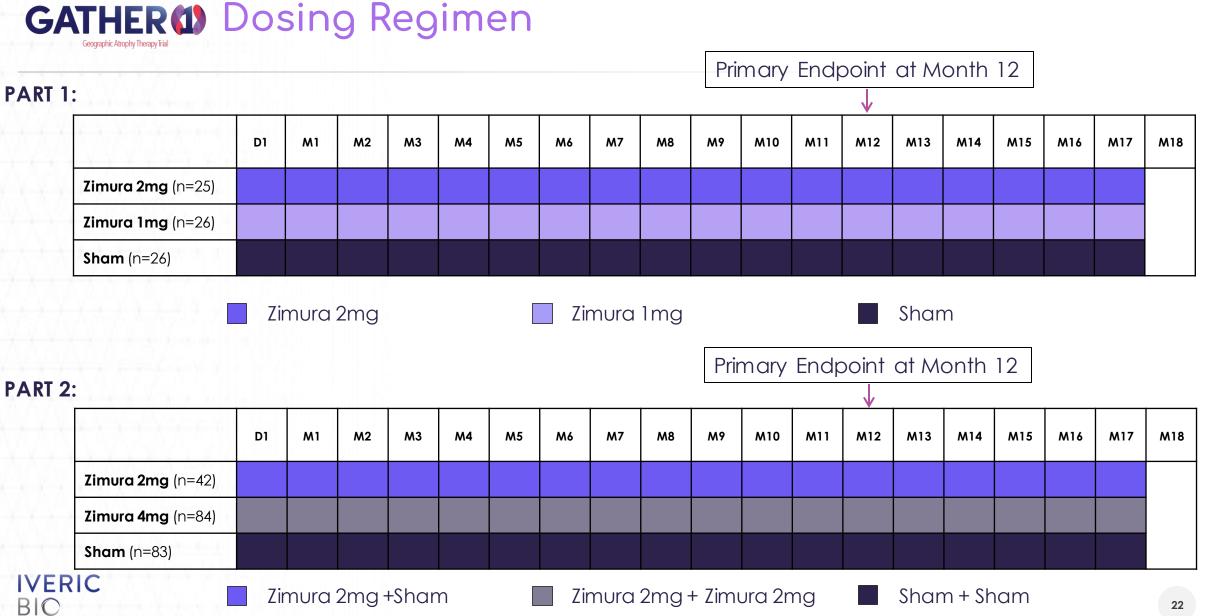
Cohort	Zimura 2mg (N=67)	Sham 2mg (N=110)	Difference	P-value	% Difference
Mean Change in GA ^(a)	0.292 ^(c)	0.402 ^(c)	0.110	0.0072 ^(b)	27.38%
Cohort	Zimura 4mg (N=83)	Sham 4mg (N=84)	Difference	P-value	% Difference
Mean Change in GA ^(a)	0.321	0.444	0.124	0.0051 ^(b)	27.81%



(a) = mm, based on the least squares means from the MRM model

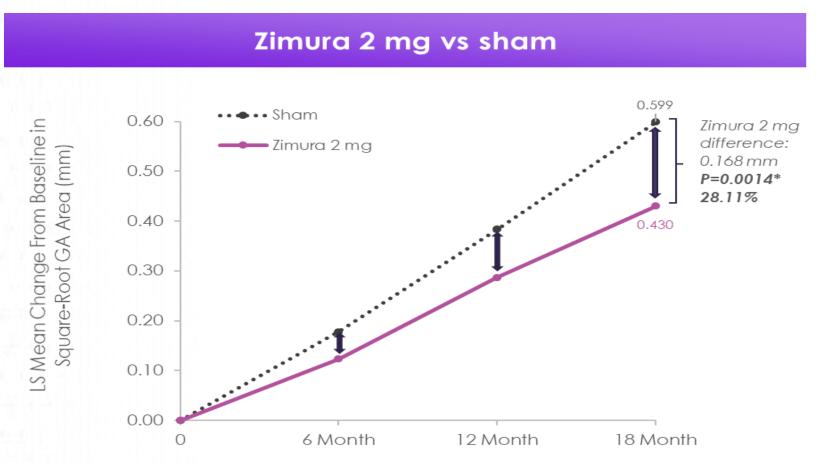
(b) = reflects statistically significant p-value; Hochberg procedure was used for significance testing

(c) = these least squares means are estimates of the MRM model, drawing on all available data, including data from groups with different randomization ratios in Part 1 and Part 2, and should not be interpreted as directly observed data



GATHER1: Decrease in GA growth over 18 months Zimura 2 mg vs. Sham (square root transformation)

MEAN RATE OF GROWTH IN GA AREA AS MEASURED BY SQUARE ROOT TRANSFORMATION OVER 18 MONTHS

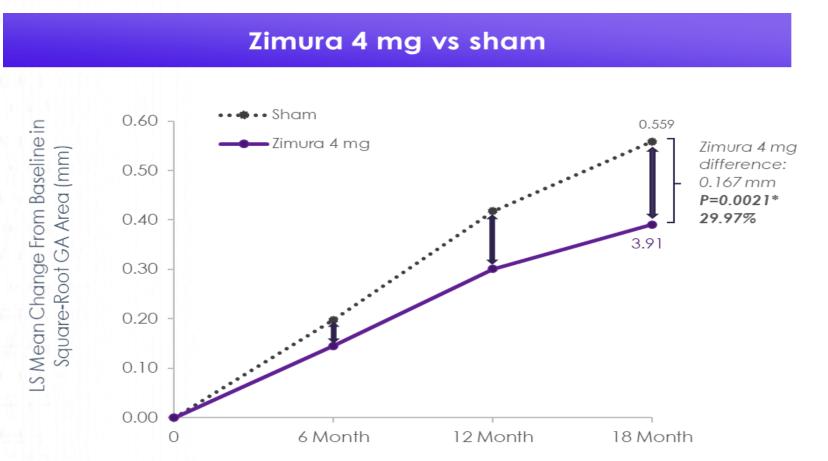


IVERIC BIC

Based on LSMEANS from MRM model; ITT population Hochberg procedure was used for significance testing; prespecified and descriptive analysis. These least squares means are estimates of the MRM model, drawing on all available data, including data from groups with different randomization ratios in Part 1 and Part 2, and should not be interpreted as directly observed data. *18-month P values are descriptive in nature.

GATHER1: Decrease in GA growth over 18 months Zimura 4 mg vs. Sham (square root transformation)

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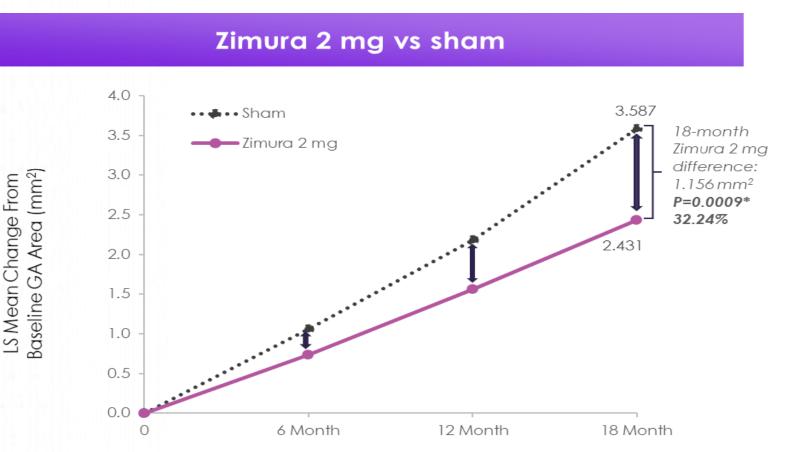


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GATHER1: Decrease in GA growth over 18 months Zimura 2 mg vs. Sham (non-square root transformation)

MEAN RATE OF GROWTHIN GA AREA AS MEASURED BY NON-SQUARE-ROOT GA LESION AREA OVER 18 MONTHS

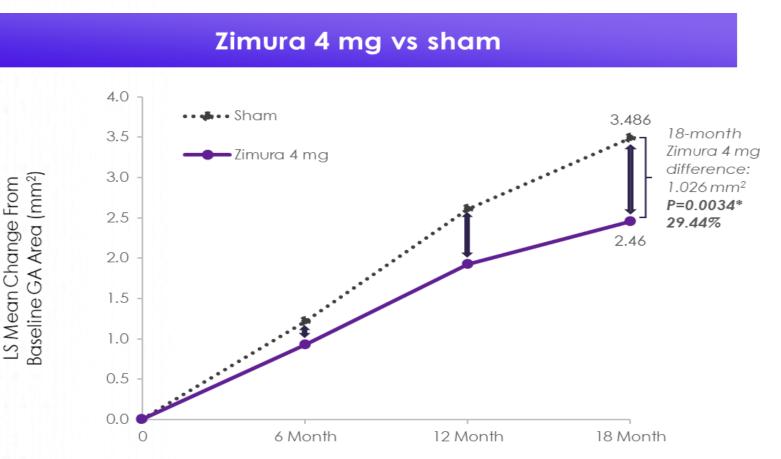


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GATHER1: Decrease in GA growth over 18 months Zimura 4 mg vs. Sham (non-square root transformation)

MEAN RATE OF GROWTH IN GA AREA AS MEASURED BY NON-SQUARE-ROOT GA LESION AREA OVER 18 MONTHS



IVERIC BIC

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Zimura was generally well tolerated over 18 months



Zimura was generally well tolerated after 18 months of continuous administration

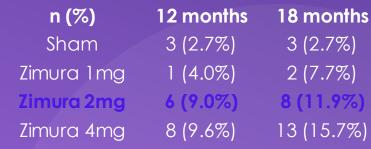


No reported Zimura-related inflammation

 \checkmark

The most frequently reported ocular adverse events were related to the injection procedure

Incidence of study eye CNV:

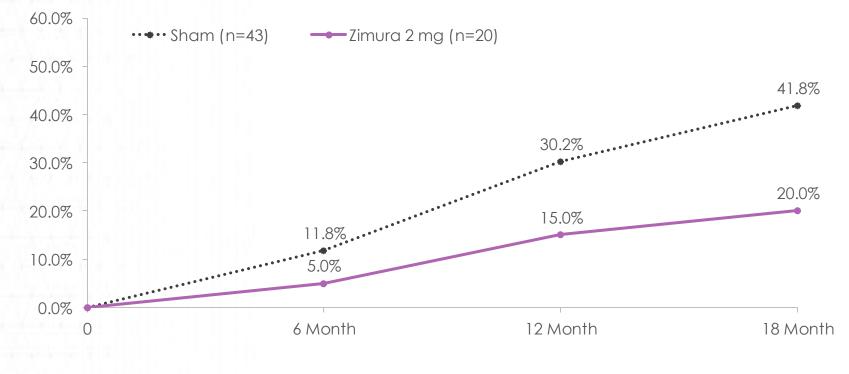


IVERICBICBased on investigator-reported safety events.

GATHER Progression of iRORA to cRORA

Proportion of patients that progress from iRORA to cRORA (Zimura 2 mg vs. Sham)

(post-hoc analysis)



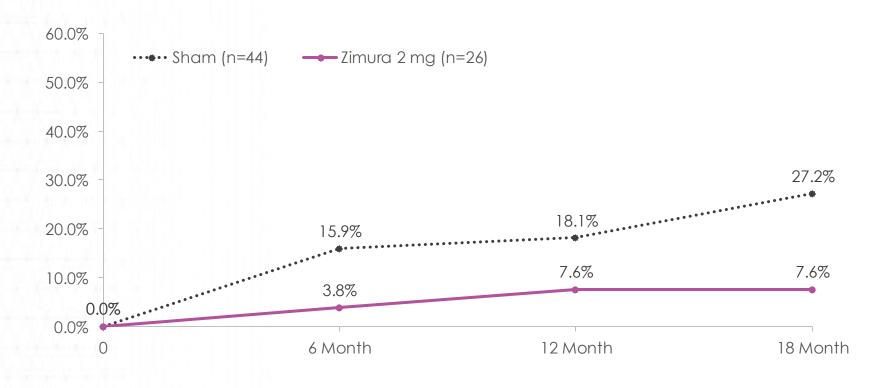
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*iRORA: Incomplete RPE + Outer Retinal Atrophy; cRORA: Complete RPE + Outer Retinal Atrophy GA is a subset of cRORA (excludes region of CNV)

GATHER (1) Progression of Drusen to iRORA/cRORA

Proportion of patients that progress from drusen to iRORA or cRORA (Zimura 2 mg vs. Sham)

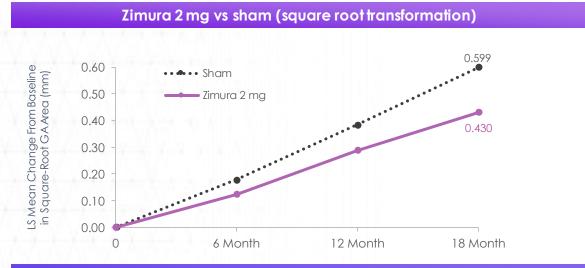
(post-hoc analysis)



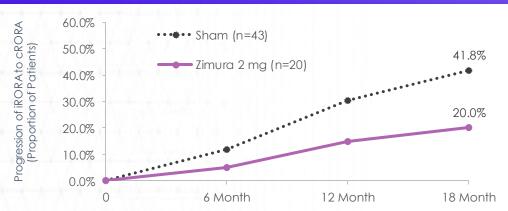
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*iRORA: Incomplete RPE + Outer Retinal Atrophy; cRORA: Complete RPE + Outer Retinal Atrophy GA is a subset of cRORA (excludes region of CNV)

Potential to alter natural history of disease



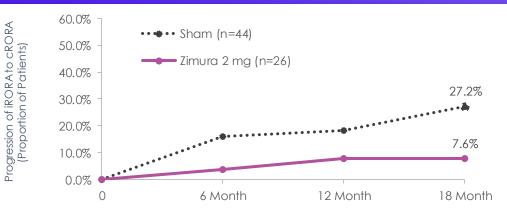
Progression of iRORA to cRORA (post-hoc analysis)



Zimura 2 mg vs sham (non-square root transformation)



Progression of drusen to iRORA/cRORA (post-hoc analysis)



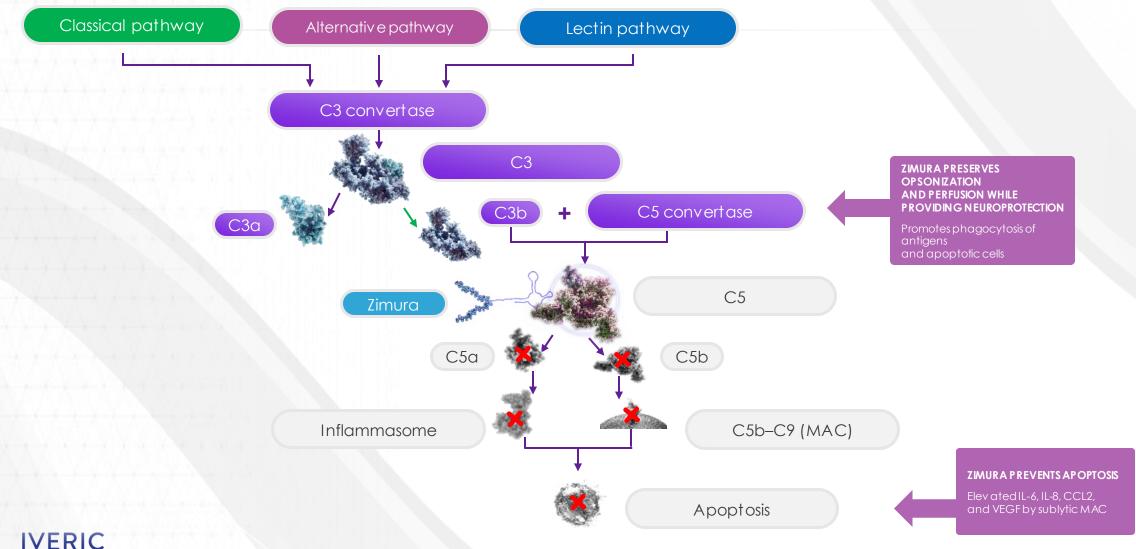
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INHIBITION OF C5

What Are The Potential Advantages of Inhibiting the Complement System at C5?

Inhibiting the 2 triggers of cell death, preserving the remainder of the pathway



BIO Xu H, Chen M. Targeting the co and degenerative diseases. Eu

Xu H, Chen M. Targeting the complement system for the management of retinal inflammatory and degenerative diseases. *European Journal of Pharmacology*. 2016;787:94-104.

C5 inhibition: Potential safety advantages

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Complement C3a receptors play roles in endotoxemia, ischemiareperfusion, neurotrauma and ALS models



C3aR is protective in these models (knockout worsens disease)



C3-CR3 is also protective in a retinal degeneration model



Global blockade of C3, as opposed to C5, may prevent the beneficial activities of C3a, while also increasing infection risk

Source: J. Exp. Med. 2019 Vol. 216 No. 8 1925–1943. J Immunol 2006; 176:4315-4322. J Immunol 2015;194: 3542–3548 Wu et al., 2013, PNAS; Brennan et al., 2019, JCI Insight. Woodruff unpublished data.

C5 inhibition: Potential safety advantages

"Deficiency of C3 or CR3 decreased microglial phagocytosis of apoptotic photoreceptors and increased microglial neurotoxicity to photoreceptors,..."

C3- and CR3-dependent microglial clearance protects photoreceptors in retinitis pigmentosa

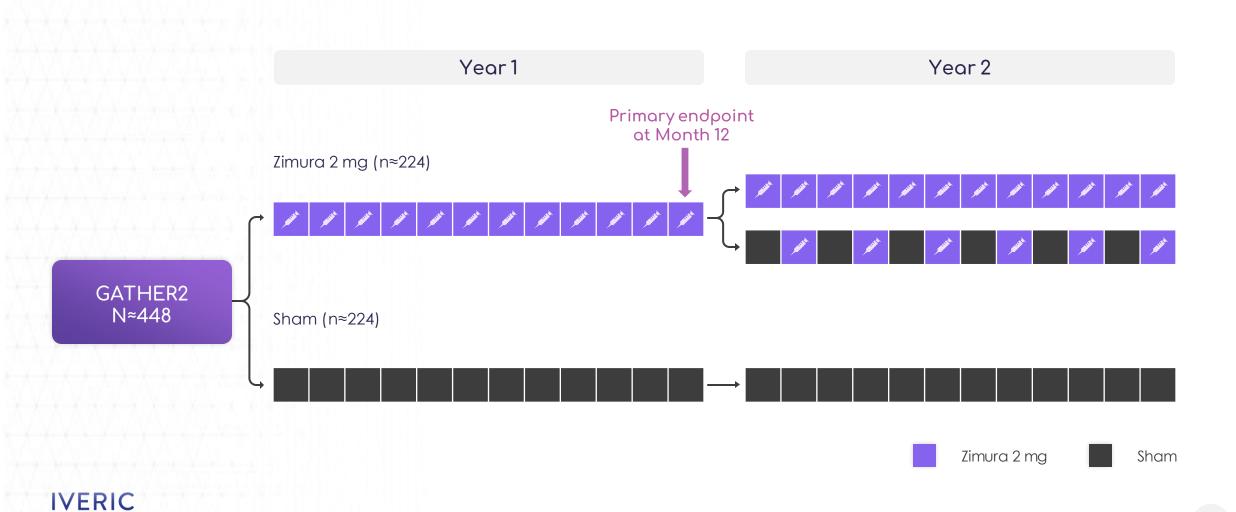
Sean M. Silverman, Wenxin Ma[®], Xu Wang, Lian Zhao[®], and Wai T. Wong[®]

Complement activation has been implicated as contributing to neurodegeneration in retinal and brain pathologies, but its role in retinitis pigmentosa (RP), an inherited and largely incurable photoreceptor degenerative disease, is unclear. We found that multiple complement components were markedly up-regulated in retinas with human RP and the rd10 mouse model, coinciding spatiotemporally with photoreceptor degeneration, with increased C3 expression and activation localizing to activated retinal microglia. Genetic ablation of C3 accelerated structural and functional photoreceptor degeneration and altered retinal inflammatory gene expression. These phenotypes were recapitulated by genetic deletion of CR3, a microgliaexpressed receptor for the C3 activation product iC3b, implicating C3-CR3 signaling as a regulator of microglia-photoreceptor interactions. Deficiency of C3 or CR3 decreased microglial phagocytosis of apoptotic photoreceptors and increased microglial neurotoxicity to photoreceptors, demonstrating a novel adaptive role for complement-mediated microglial clearance of apoptotic photoreceptors in RP. These homeostatic neuroinflammatory mechanisms are relevant to the design and interpretation of immunomodulatory therapeutic approaches to retinal degenerative disease.



Second Pivotal Clinical Trial of Zimura in GA

GATHER(2) Primary endpoint at Month 12



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BUILDING AN AMD FRANCHISE

Evidence for the role of HtrA1 in AMD pathogenesis

Target backed by strong human genetic and pre-clinical/clinical evidence



Strong human genetic evidence associates ocular HtrA1 overexpression with geographic atrophy and all neovascular forms of AMD



Compelling preclinical and clinical evidence for role of HtrA1 in AMD

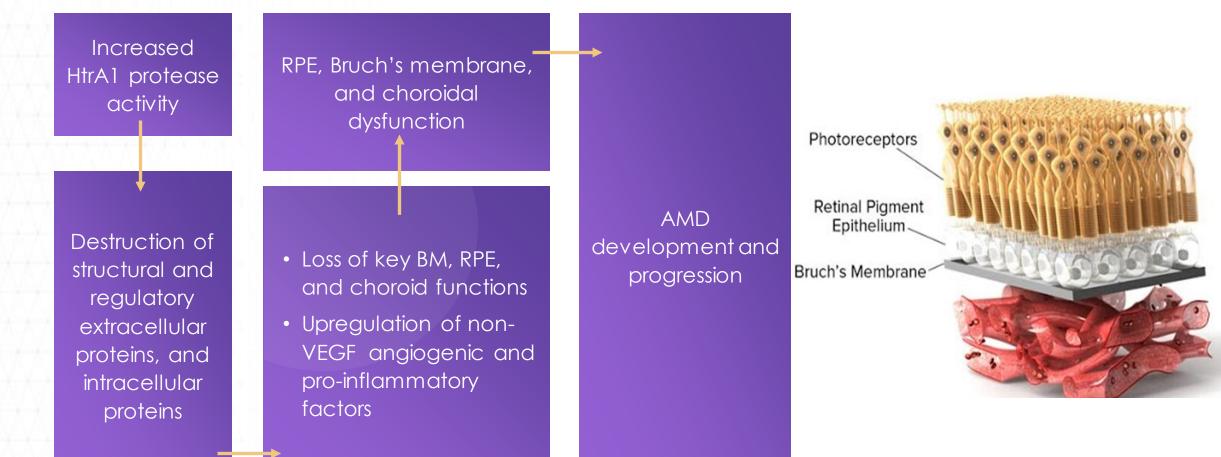


HtrA1 is non-overlapping and could augment the effects of targeting other AMD treatment pathways



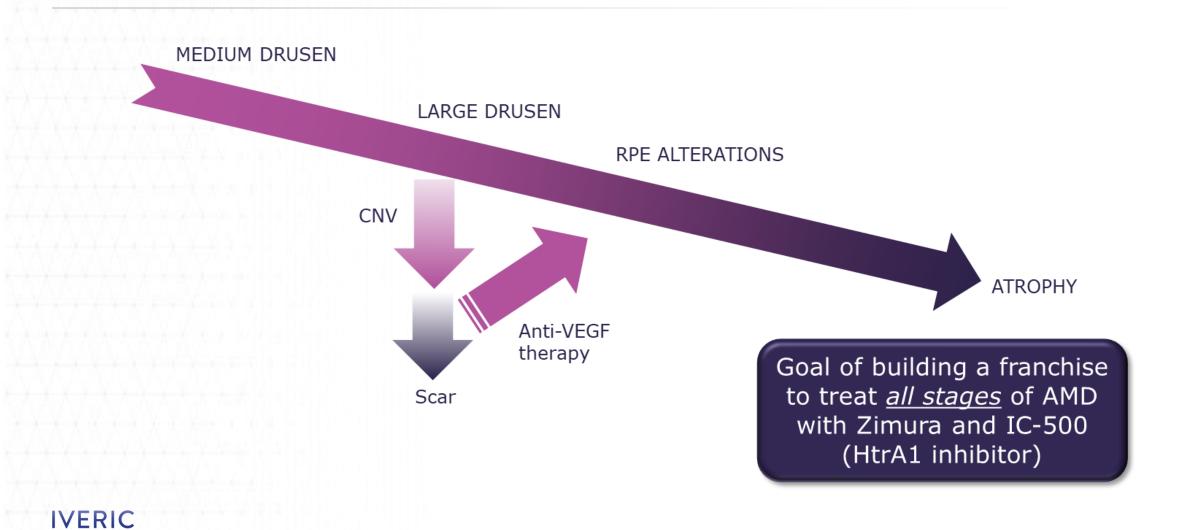
Proposed mechanism of HtrA1 activity in AMD

Destruction of extracellular matrix proteins leads to epithelium dysfunction



Multiple shots on goal in AMD

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EXECUTION AND REGULATORY CLARITY

GATHER(2) Enrollment remained strong throughout the pandemic

Time to Complete Enrollment was Four Months Ahead of Original Timeline

Press Release

Iveric Bio Completes Patient Enrollment of GATHER2 Pivotal Clinical Trial of Zimura® Ahead of Schedule

07.26.2021

- Topline Data Expected in 2H 2022; if Positive, New Drug Application Expected -

Injection fidelity is the most meaningful marker of patient retention



12-Month Injection Fidelity Rate

87%

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Target 12- Month Injection Fidelity Rate

> 90%	
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Injection Fidelity Calculation:

Total Number of Injections or Sham Administered

Total Number of expected injections or Sham

GATHER⁽²⁾ Regulatory path

First Known Special Protocol Assessment in GA

Press Release

Iveric Bio Receives FDA Agreement Under Special Protocol Assessment (SPA) for GATHER2 Phase 3 Clinical Trial of Zimura® in Geographic Atrophy Secondary to Age-Related Macular Degeneration

07.06.2021

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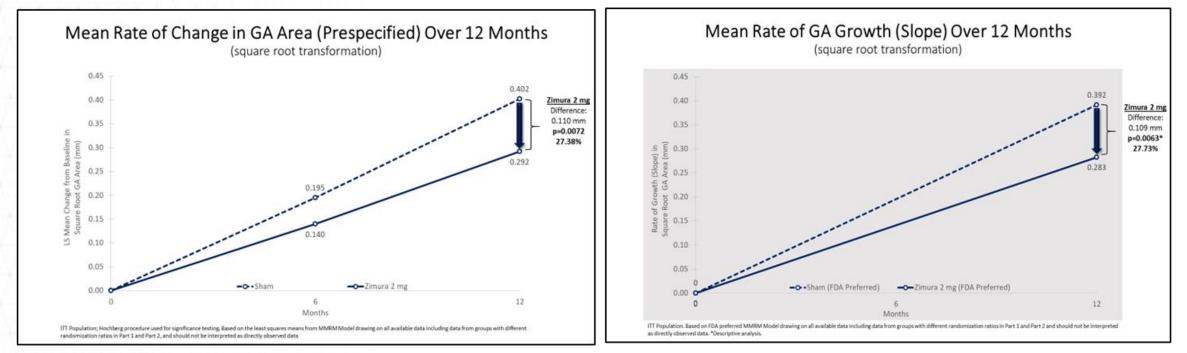
- GATHER2 Enrollment and Retention Continue to Exceed Expectations; Completion of Enrollment Expected Late July of this Year and Topline Data Expected Second Half of 2022 -

GATHER1: 2mg vs. sham mean rate of change in GA area (prespecified) and mean rate of GA growth (slope) (post-hoc)

FDA Preferred Analysis Supports Prespecified Analysis

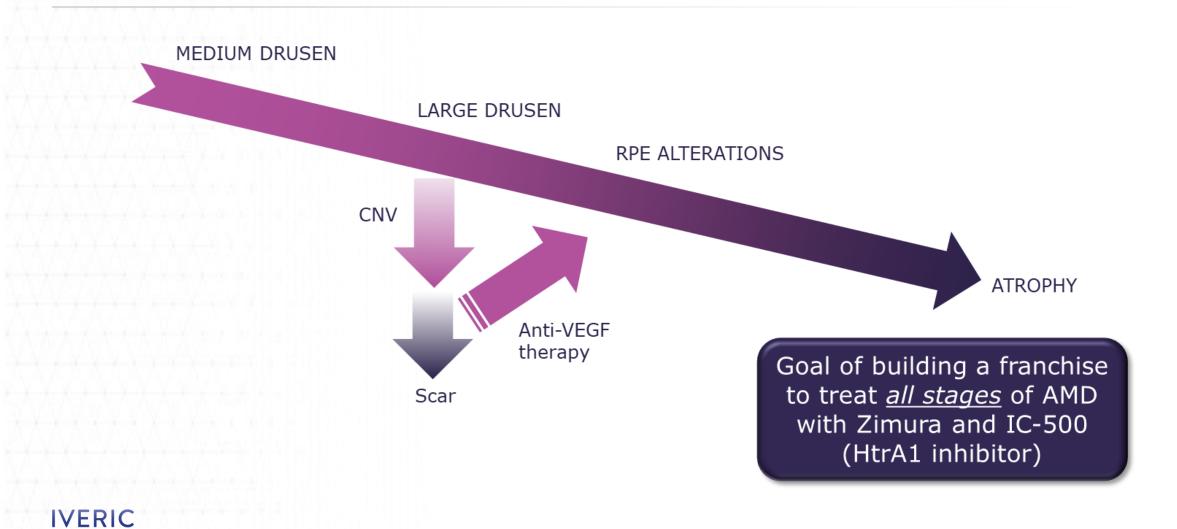
Prespecified Analysis

FDA preferred Analysis



Pathway of AMD disease progression

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Recent and planned milestones

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GATHER2 enrollment completed (July 2021)



Hired Chief Commercial Officer (August 2021)



Initiate Phase 1/2 trial for IC-200 in patients with AR BEST Disease (Q4 2021)



Initiate clinical trial of Zimura in drusen (2022)

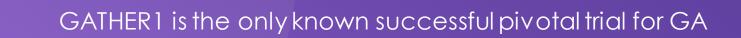


GATHER2 topline data readout (2H 2022)



Summary

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If positive, we expect GATHER2 will be the final pivotal trial required for FDA and EMA approval for GA

Zimura has the potential to impact earlier stages of AMD

We believe we are well positioned to expand Zimura's
indications, build an AMD franchise and, subject to regulatory approval, commercialize Zimura for GA