# Retrospective Real-World Analysis of Patients with Geographic Atrophy (GA) Secondary to Age-Related Macular Degeneration Followed up for 3 Years

<u>Ehsan Rahimy</u>, M Ali Khan, Allen Ho, Ryan Corley, Mark Gallivan, Daniel Jones, Ramiro Ribeiro, Theodore Leng, Nancy Holekamp

July 13–16, 2022 40th Annual Meeting of the American Society of Retina Specialists, NY, USA



### Disclosures

- Ehsan Rahimy has the following financial interests or relationships to disclose:
  - Consultant and studies funded by Apellis Pharmaceuticals

### Introduction

- Longer-term outcomes regarding the progression of GA in patients with age-related macular degeneration (AMD) are lacking
- Here we report data from a retrospective analysis of clinical data from the American Academy of Ophthalmology IRIS<sup>®</sup> (Intelligent Research in Sight) Registry evaluating disease progression of patients with GA in one eye and either GA or choroidal neovascularization (CNV) in the other eye, over 3 years

AMD=age-related macular degeneration; CNV=choroidal neovascularization; GA=geographic atrophy.

### Patient disposition

ICD diagnosis code in at least one eye for GA between January 1st, 2016, to March 16, 2017, in the study eye	Presence of GA/CNV ICD diagnosis code in the fellow eye on the same day as a diagnosis code for GA in the study eye	Exclude patients with active or history of CNV in the study eye before the first GA record during the study period	Exclude adults <50 years of age and >110 years at index date, and patients without recorded birth sex	Exclude patients with missing baseline VA in the study eye & missing VA data for the study eye	Exclude patients with a history of any other retinal condition	Exclude patients with <3 years of follow-up. Visits will be defined as having an encounter, procedure, or diagnosis following the initial diagnosis of GA
(N=125,743)	(N=91,768)	(N=89,779)	(N=89,128)	(N=83,994)	(N=78,411)	(N=36,817)

### Patients with at least 3 years of follow-up (N=36,817)

CNV=choroidal neovascularization; GA=geographic atrophy; ICD=International Statistical Classification of Diseases and Related Health Problems; N=number; VA=visual acuity.

## **Cohorts for analysis**

COHORT 1 (n=21,	– GA:GA 789)	COHORT 2 – GA:CNV (n=14,976)		
Study eye: GA lesion location				
Cohort 1A	Cohort 1B	Cohort 2A	Cohort 2B	
Extrafoveal GA	Foveal GA	Extrafoveal GA	Foveal GA	
(n=10,720)	(n=11,069)	(n=7,263)	(n=7,713)	

- Patients excluded if they had a history of CNV or active CNV in the study eye, a history of any other retinal condition, or <36 months of follow-up
- Patients grouped by fellow eye status:
  - Cohort 1, GA:GA
  - Cohort 2, GA:CNV (study eye GA, fellow eye CNV)
- Subgroups were classified by lesion location: extrafoveal or foveal
- Main outcomes were study and fellow eye disease progression including VA over 3 years

CNV=choroidal neovascularization; GA=geographic atrophy; n=number; VA=visual acuity.

## Patient demographics (and treating provider)

	COHORT 1 – GA:GA (n=21,789)	COHORT 2 – GA:CNV (n=14,976)
Age (years) mean (SD)	80.1 (8.6)	81.8 (7.8)
Sex, n (%)		
Female	14,574 (66.9)	10,011 (66.9)
Male	7,215 (33.1)	4,965 (33.2)
Race, n (%)		
White or Caucasian	19,300 (88.6)	13,641 (91.1)
Black or African American	234 (1.1)	82 (0.6)
Asian	326 (1.5)	122 (0.8)
Other	148 (0.7)	102 (0.7)
Unknown	1,781 (8.2)	1,029 (6.9)
Treating provider, n (%)		
Retina specialist	14,471 (66.4)	13,638 (91.1)
General ophthalmologist	3,240 (14.9)	806 (5.4)
Non-retina specialist	3,083 (14.2)	353 (2.4)
Optometrist	906 (4.2)	111 (0.7)
Unknown	89 (0.4)	68 (0.5)

CNV=choroidal neovascularization; GA=geographic atrophy; n=number; SD=standard deviation.

## Study eye cumulative progression at Years 1, 2 and 3

#### Extrafoveal GA to foveal GA

Cohort 1A (n=10,720 eyes) and Cohort 2A (n=7,236)



- Progression to foveal GA is more common among bilateral GA eyes vs GA:CNV study eyes
  - 4.5–7.4% of GA:GA study eyes progressed each year, compared with 2.9–5.7% of GA:CNV study eyes
- Conversion to CNV is more common in GA:CNV study eyes
  - Over 30% of GA:CNV study eyes develop CNV at Year 3, compared with <15% of GA:GA study eyes</li>
  - Progression to CNV is slightly higher among eyes with extrafoveal GA vs foveal GA

#### **Extrafoveal GA to CNV**

Cohort 1A (n=10,720 eyes) and Cohort 2A (n=7,236)



#### Foveal GA to CNV



Cohort 1B (n=11,069 eyes) and Cohort 2B (n=7,713)

Overall population data included the total proportion of patients who progressed to foveal GA at any point during the follow-up period, which could extend beyond 3 years for select eyes. For this analysis, enrolled eyes with 3+ years of follow-up, but certain eyes may have >3 years of follow-up data. CNV=choroidal neovascularization; GA=geographic atrophy; n=number

## Median time to progression – study eye (1/2)

Study eye progression of patients with GA in both eyes (Cohort 1), patients with GA in one eye and CNV in the other eye (Cohort 2)

	COHORT 1A – GA:GA		COHORT 2A – GA:CNV	
	n	Median time to event, wks (IQRª)	n	Median time to event, wks (IQRª)
Study eye: Progression from extrafoveal GA to foveal GA				
Cases among incident patients	58	98.1 (52.0–144.5)	60	98.0 (33.1–125.7)
Cases among prevalent patients	2,280	79.1 (35.4–139.0)	1,018	71.0 (29.6–134.0)

Among the study eyes of patients with bilateral GA, median time to progression to foveal GA was ~98 weeks among newly
incident cases and ~79 weeks among prevalent GA cases

- Median time to foveal GA was nominally shorter (71 weeks) among prevalent patients with GA:CNV

Incident eyes defined as GA eyes with either early or intermediate AMD in year 2016 and an index date for GA in year 2017. alQR: 25th percentile–75th percentile. AMD=age-related macular degeneration; CNV=choroidal neovascularization; GA=geographic atrophy; IQR=interquartile range; n=number; wks=weeks.

## Median time to progression – study eye (2/2)

Study eye progression of patients with GA in both eyes (Cohort 1), patients with GA in one eye and CNV in the other eye (Cohort 2)

	COHORT 1 – GA:GA		COHORT 2 – GA:CNV	
	n	Median time to event, wks (IQRª)	n	Median time to event, wks (IQRª)
Study eye: Progression from extrafoveal GA to CNV				
Cases among incident patients	60	83.6 (44.7–151.1)	161	71.6 (38.0–123.9)
Cases among prevalent patients	1,850	89.0 (40.7–146.1)	2,587	78.6 (32.5–136.2)
Study eye: Progression from foveal GA to CNV				
Cases among incident patients	31	60.0 (49.7–106.9)	128	68.0 (26.0–113.0)
Cases among prevalent patients	1,614	85.7 (36.9–145.4)	2,531	68.4 (29.5–126.1)

• Time to progression from extrafoveal GA to CNV appeared to be faster among patients with fellow eye CNV

 Conversion occurred at a median of 72 weeks vs 84 weeks among newly incident cases, and 79 vs 89 weeks for prevalent cases among GA:CNV vs GA:GA study eyes, respectively

Incident eyes defined as GA eyes with either early or intermediate AMD in year 2016 and an index date for GA in year 2017. aIQR: 25th percentile–75th percentile. AMD=age-related macular degeneration; CNV=choroidal neovascularization; GA=geographic atrophy; IQR=interquartile range; n=number; wks=weeks.

## VA for study eye at index

	COHORT 1 – GA:GA (n=20,225)	COHORT 2 – GA:nAMD (n=13,926)
ETDRS Letters, mean (SD)	64.8 (20.0)	56.0 (25.8)
Proportion of patients, n (%)		
20/20 or better	2,742 (13.6)	1,145 (8.2)
<20/20 and ≥20/40	8,956 (44.3)	5,218 (37.5)
<20/40 and ≥20/100	5,460 (27.0)	3,574 (25.7)
<20/100 and ≥20/200	1,563 (7.7)	1,436 (10.3)
<20/200	1,504 (7.4)	2,553 (18.3)

• Among the patient eyes with available VA recordings (n=34,151), overall vision was worse among GA:nAMD study eyes than among GA:GA study eyes

 7.4% of GA:GA study eyes (Cohort 1) and 18.3% of GA:nAMD (Cohort 2) study eyes had worse than 20/200 vision at baseline

ETDRS=Early Treatment Diabetic Retinopathy Study; GA=geographic atrophy; n= number; nAMD=neovasular age-related macular degeneration; SD=standard deviation; VA=visual acuity. 10

## Study eye VA progression over 3 years

#### **Progression to severe blindness** (20/200 or worse) Bilateral GA and GA:CNV study eyes, VA: >20/200 and <20/40 at baseline



- Within 2 years, the rate increased to over a quarter of eyes and within 3 years, over a third of study eyes

### VA change at 12 months, 24 months, and 36 months



- Eyes with greater vision lose more letters in the first 3 years compared with eyes with poor vision at index
- Progression trends continue to occur into Year 3
  - Rate of change between Years 1 and 2 is similar rate of change from Years 2 and 3

### Conclusions

- GA caused substantial disease burden in this retrospective study of a large real-world database
- Eyes with GA lost significant vision over a 3-year period, consistent with trends in the previously reported 2-year analysis
- Eyes with good vision at baseline lost more letters over 3 years compared with eyes with poor vision
- Patients with GA are still at significant risk of being "lost-to-follow-up" as evidenced by more than 42,000 patients being excluded from this analysis with less than 3 years of follow-up data

AMD=age-related macular degeneration; CNV= choroidal neovascularization; GA=geographic atrophy.