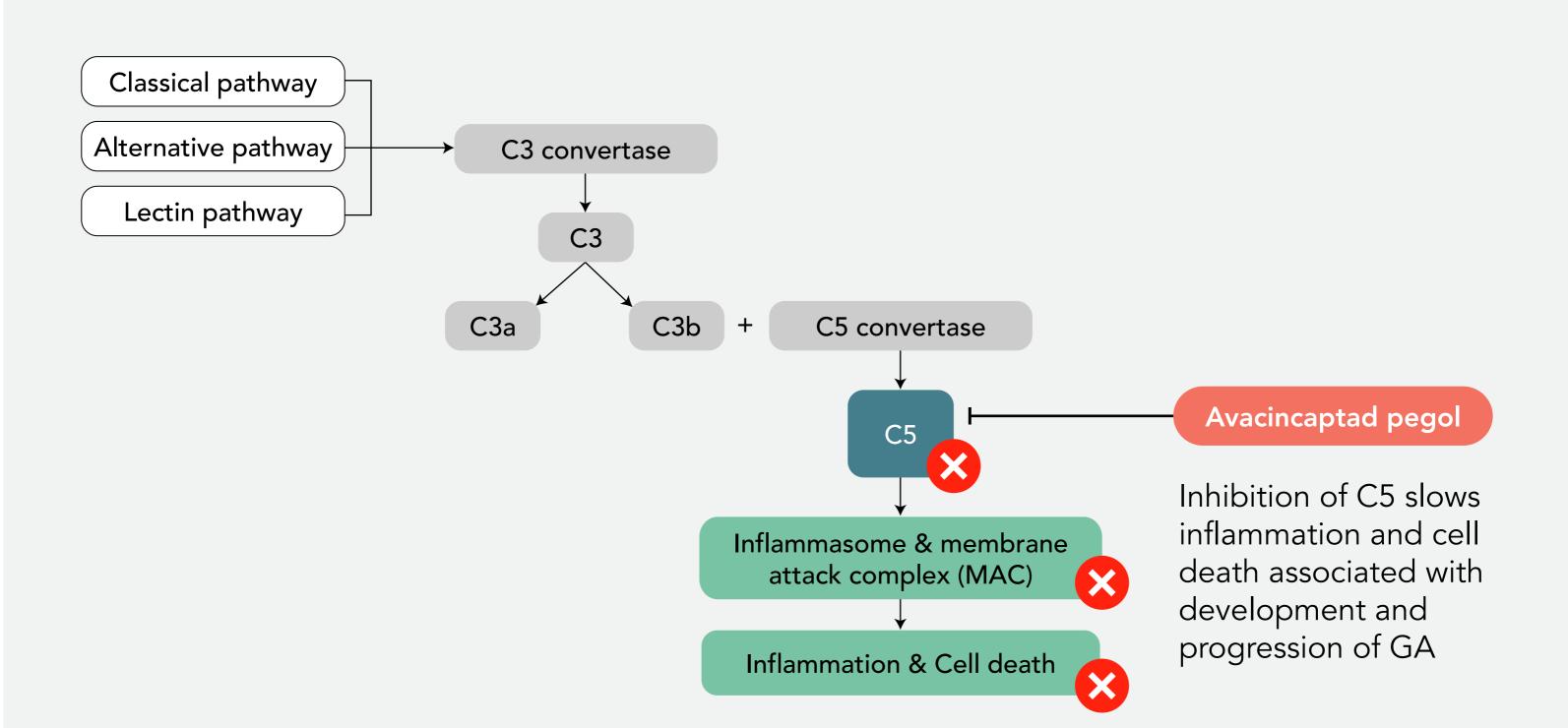
## The Efficacy of Avacincaptad Pegol in Geographic Atrophy: **GATHER1** and **GATHER2** Results

Avacincaptad pegol is a pegylated RNA aptamer designed to be a specific inhibitor of complement C5. The complement system plays a key role in the specialized immune defense mechanisms of the retina and the inhibition of C5 slows the inflammation and cell death associated with the development and progression of geographic atrophy (GA). GATHER1 is a Phase 2/3 study looking at the mean change in GA area from baseline over 12 months. GATHER2 is a Phase 3 study looking at the mean rate of growth (slope) in geographic atrophy area from baseline over 12 months.

Khanani AM, et al. Presented at The Retina Society 55<sup>th</sup> Annual Scientific Meeting; November 2-5, 2022; Pasadena, CA.

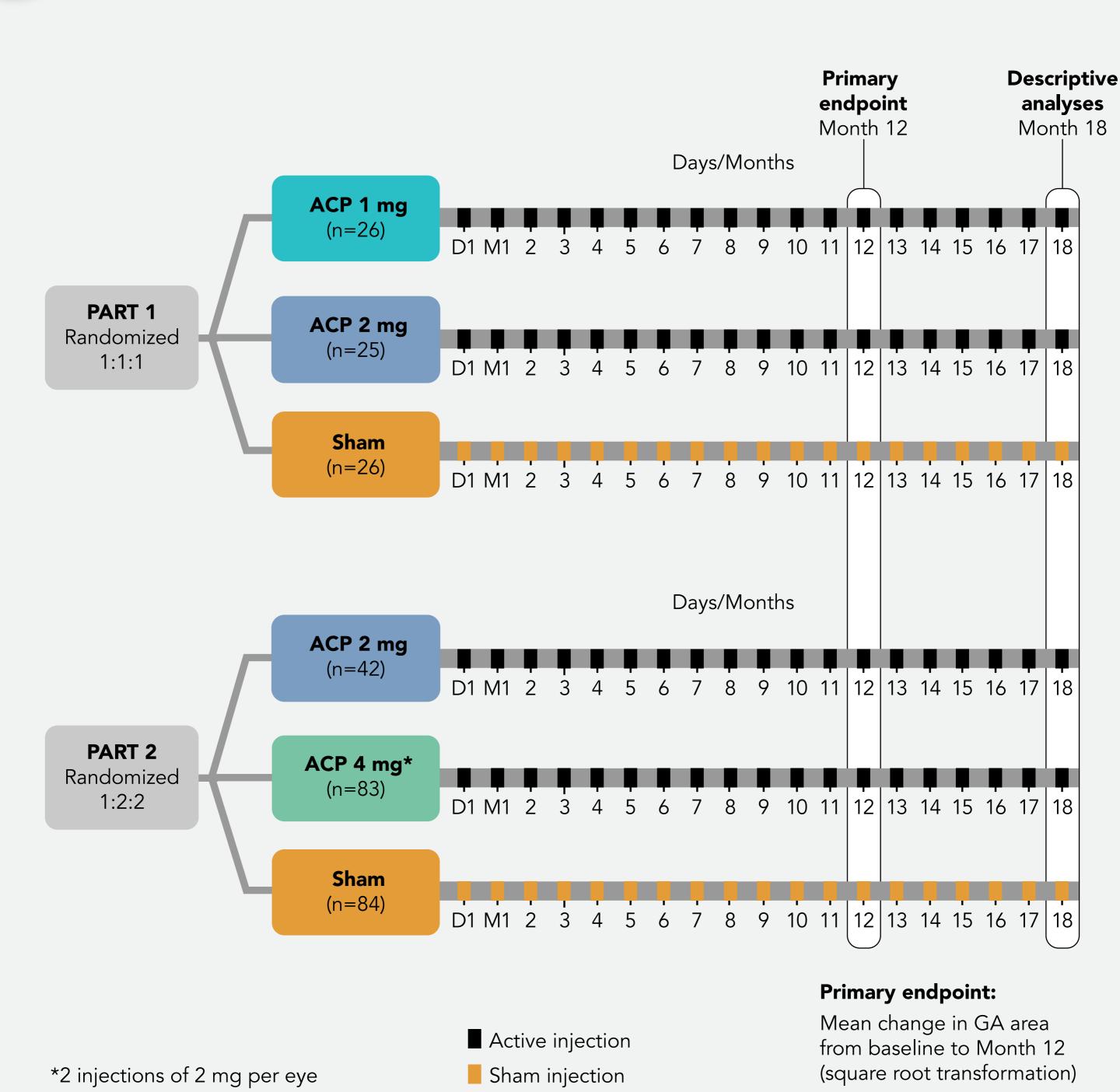


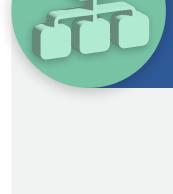
# C5 inhibition potentially preserves anti-inflammatory properties of C3a.





GATHER1 was a phase 2/3, international, prospective, randomized, double-masked, sham-controlled trial to analyze mean change in GA area from baseline to Month 12





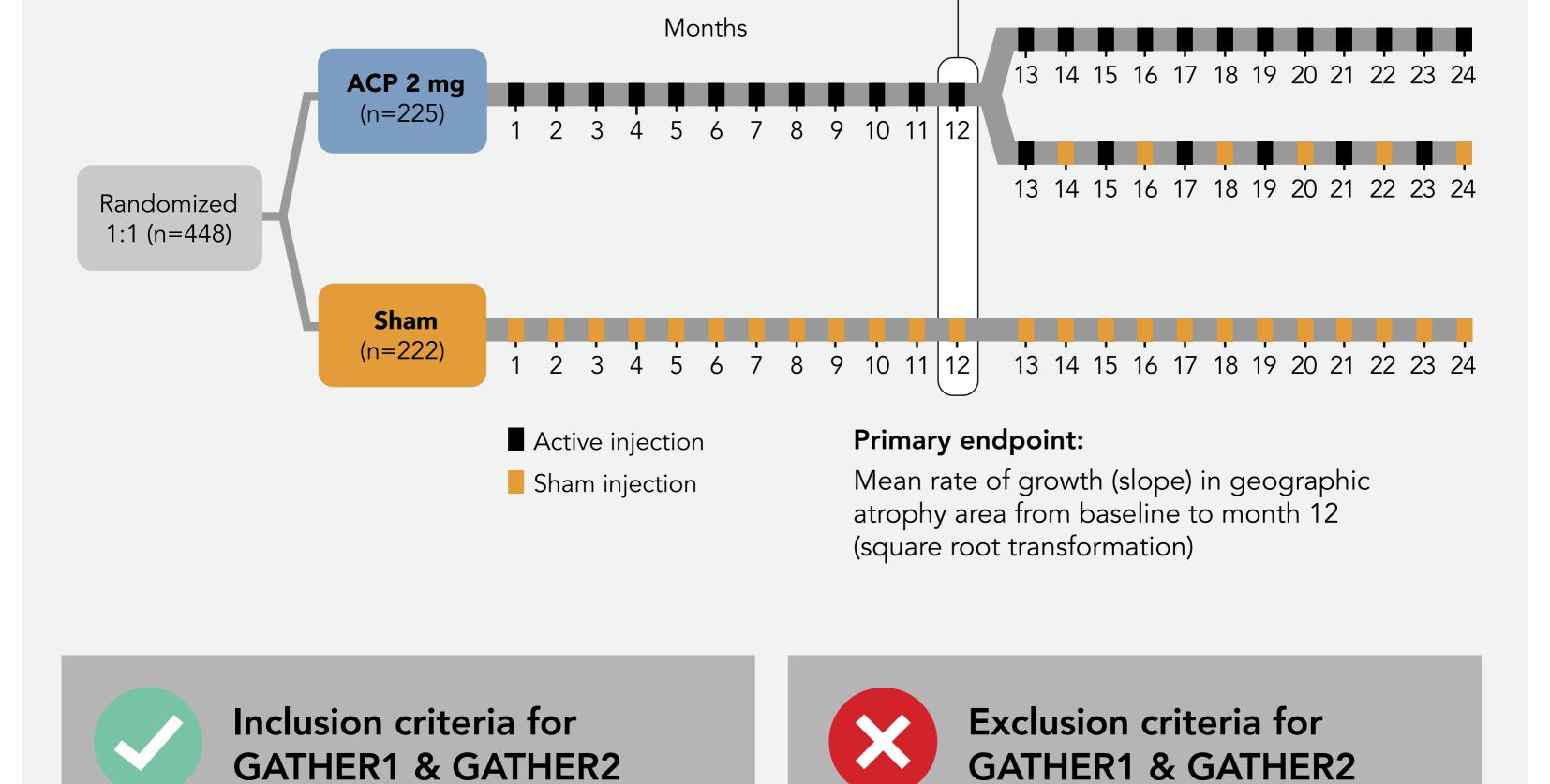
**Primary endpoint** 

Participants re-randomized 1:1

double-masked, sham-controlled study to analyze mean rate of growth (slope) in

GATHER2 was a phase 3, international, multicenter, prospective, randomized,

GA area from baseline to Month 12 (square root transformation)





### Non-center point involving ■ GA in part within 1500 µm from the

- foveal center • Total area between 2.5 mm<sup>2</sup> and 17.5 mm<sup>2</sup>
- (1-7 DA, respectively) • If multifocal lesions, at least 1 lesion had to be  $\geq 1.25 \text{ mm}^2 (0.5 \text{ DA})$

DA = disc area; GA = geographic atrophy.

GATHER2, respectively.

ACP 2 mg

# in either eye

#### Any prior treatment for AMD or any prior intravitreal treatment for any indication in either

GA secondary to any condition other than AMD

eye (except oral vitamin or mineral supplements) Any ocular condition in study eye that could progress during the study and potentially affect

• Evidence of CNV in either eye at baseline

factor • Any sign of diabetic retinopathy in either eye

central vision or otherwise act as a confounding

ACP 2 mg

Sham

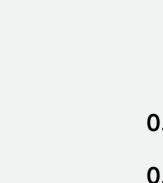
0.336

**GATHER1 GATHER2** 

0.292

Sham

AMD = age-related macular degeneration; BCVA = best-corrected visual acuity; CNV = choroidal neovascularization;



0.30

0.25

(n=222)(n=110)0.45 0.45 0.40 0.40 GA Growth (Slope) 27.4% reduction 14.3% reduction 0.402 0.392 0.35 0.35 vs. sham vs. sham

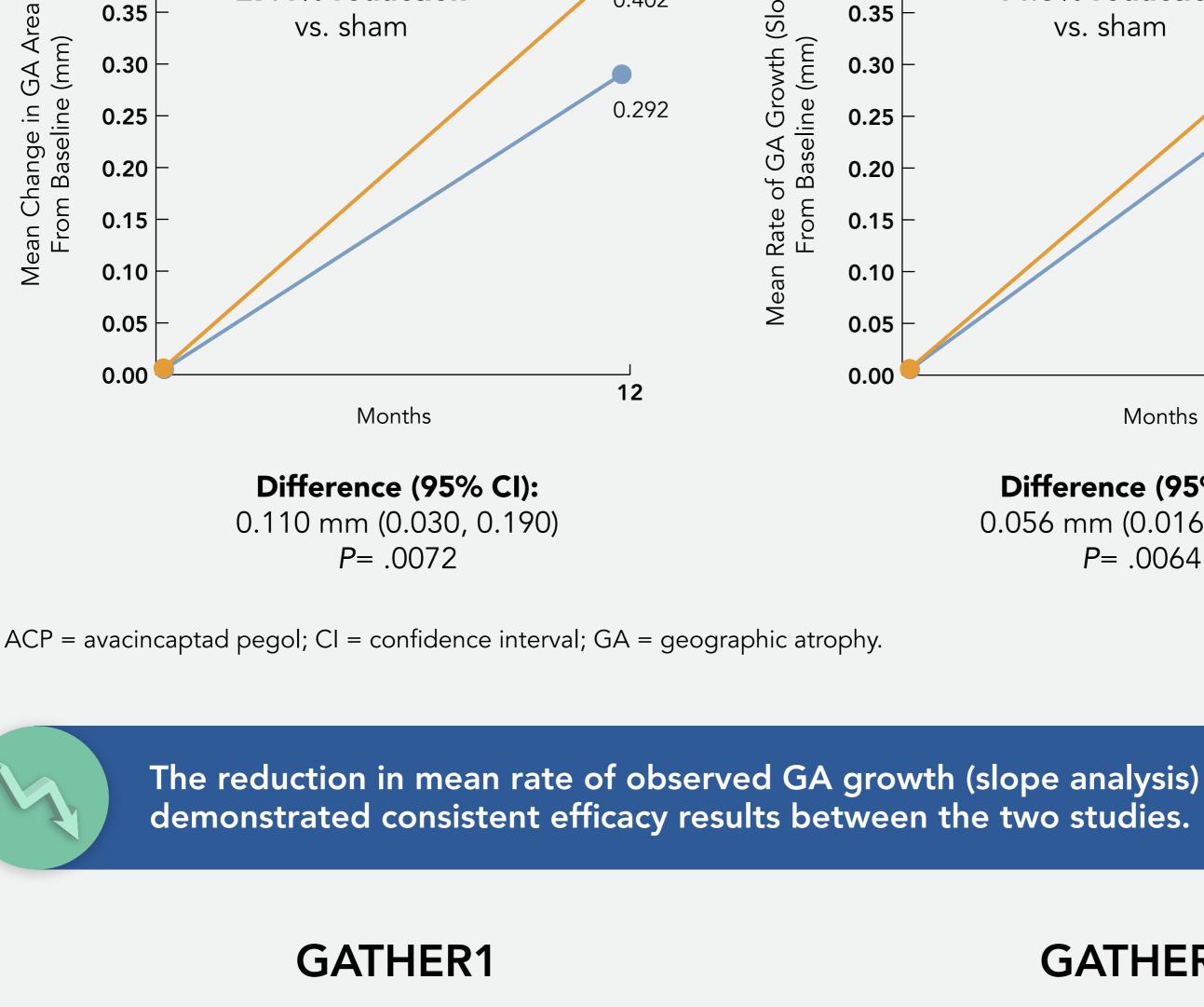
0.30

2.5

2.0

A significant reduction in the mean change in GA area and mean rate of GA growth

(slope analysis) from baseline to Month 12 was observed in both GATHER1 &

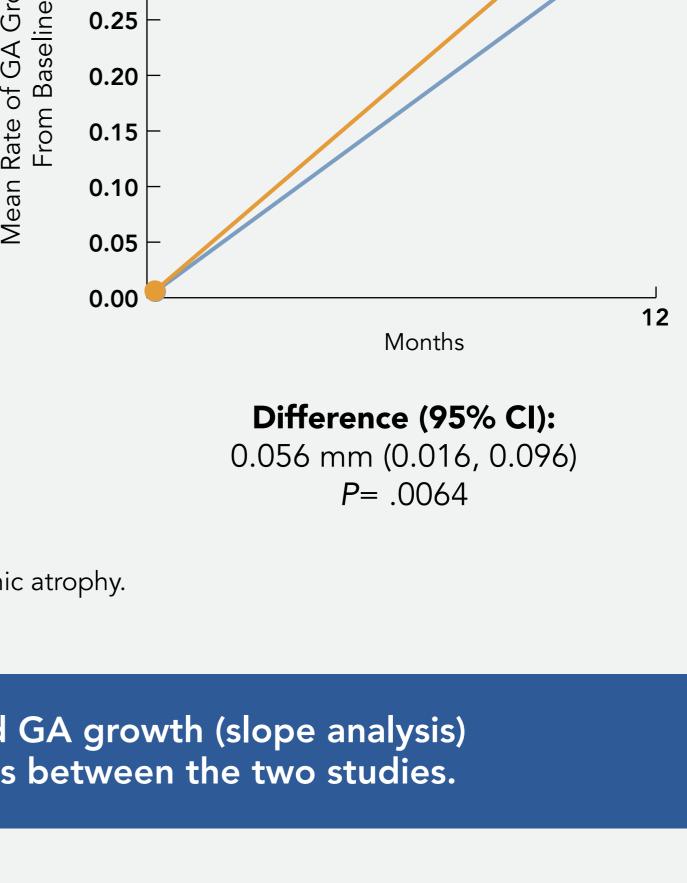


ACP 2 mg

(n=67)

35.4% reduction

ACP = avacincaptad pegol; CI = confidence interval; GA = geographic atrophy.



**GATHER2** 

Sham

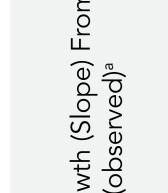
(n=222)

2.121

ACP 2 mg

(n=225)

17.7% reduction



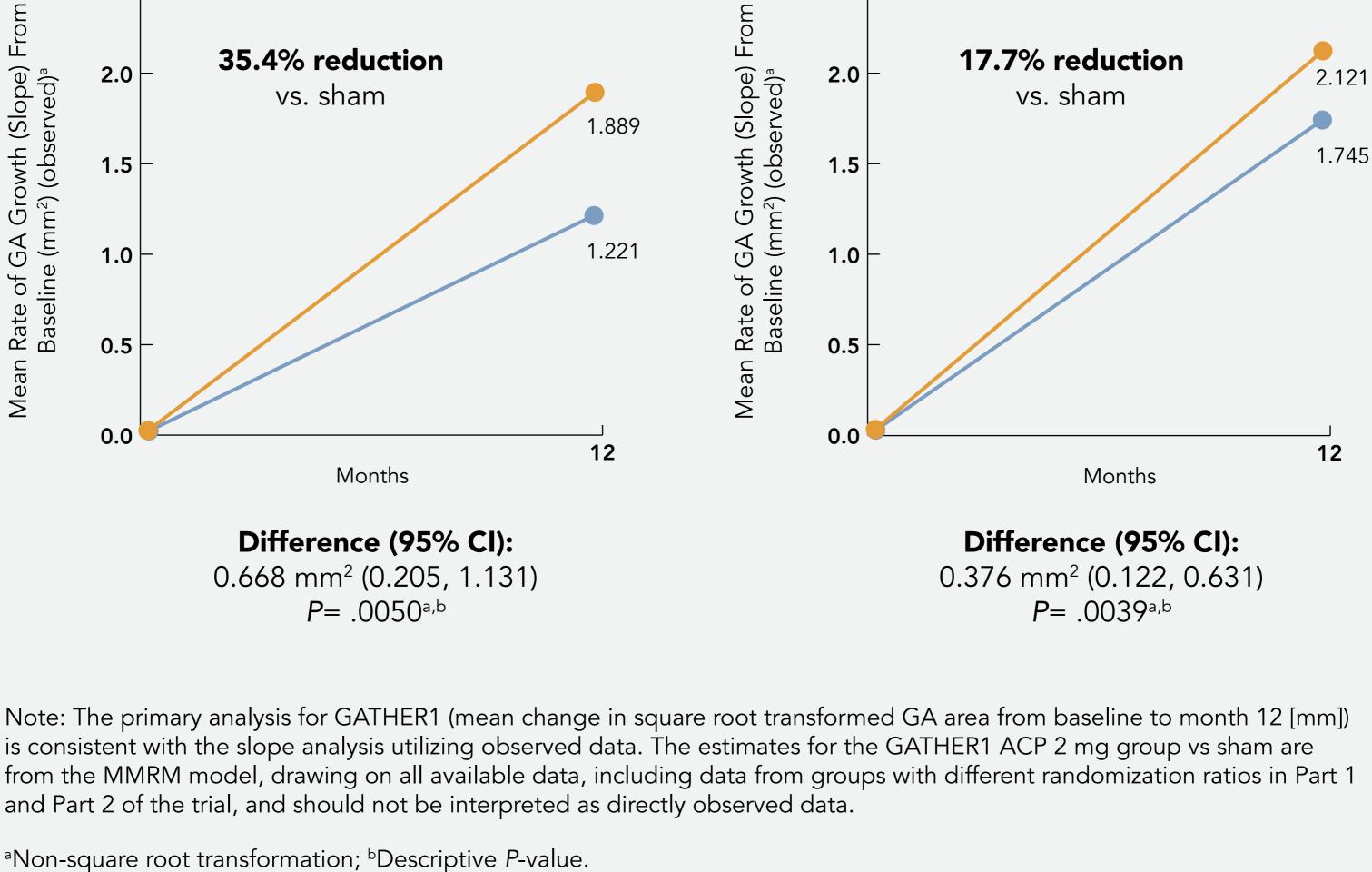
2.5

2.0

vs. sham vs. sham 1.889

Sham

(n=110)



Conclusion

Avacincaptad pegol is the first investigational therapy in GA to achieve the

12-month prespecified, primary endpoint, in two pivotal, phase 3 studies.