



**Rhopressa™ (ROCK-NET Inhibitor)
Rho Kinase Elevated Intraocular Pressure Treatment Trial
(Rocket 1) Phase 3 Topline Results**

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Any discussion of the potential use or expected success of our product candidates is subject to our product candidates being approved by regulatory authorities.

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- There are three Phase 3 registration trials for Rhopressa™, “Rocket 1,” a 90-day efficacy trial, the results of which are reported in this presentation, “Rocket 2,” a 12-month safety trial with a 90-day interim efficacy readout expected third quarter 2015, and “Rocket 3,” a safety-only study being conducted in Canada.
- Rocket 1 clinical trial evaluated
 - the ocular hypotensive efficacy of Rhopressa™ 0.02%, dosed QD, versus Timolol, 0.5%, dosed BID, in patients with elevated intraocular pressure (OAG and OHT) for 90 days
 - the ocular and systemic safety of Rhopressa™, 0.02% for 90 days

Rhopressa™ Rocket 1 Trial Design



Patients with open angle glaucoma (OAG) or ocular hypertension (OHT)
with IOP >20 mmHg and < 27 mmHg

N=411 randomized at 36 sites

(370 subjects per protocol)



Patients randomized
1:1

Rhopressa™
0.02% QD (PM)

Timolol 0.5%
BID



Primary endpoint: Mean IOP at
Weeks 2 and 6 and Day 90

Study Endpoints

Efficacy:

- The primary efficacy endpoint is the mean IOP at the following time points: 08:00, 10:00, and 16:00 at the Week 2, Week 6, and Day 90 visits
- Secondary efficacy endpoints include
 - IOP analysis stratified by baseline IOP above and below 24 mmHg
 - Mean change from baseline IOP
 - Mean percent change from diurnally adjusted baseline IOP
 - Mean diurnal and change from baseline diurnal IOP

Safety:

- Ocular and systemic safety measures

Trial Conduct

- Number of Subjects Randomized - 411
- Number of Early Terminations – 44 (Combined Rhopressa™ and Timolol)
 - 31 in Rhopressa™, 13 in Timolol
- Major Reasons for Early Termination (Combined Rhopressa™ and Timolol)
 - Adverse events (55%)
 - Protocol violation (18%)
 - Withdrawal of consent (11%)
 - Lack of efficacy (7%)
 - Investigator decision (4%)

Primary Endpoint: Topline Summary

- Per Protocol population (baseline IOP < 27 mmHg)
 - Rhopressa™ did not meet criteria for non-inferiority to Timolol
 - Rhopressa™ mean difference from Timolol ranged from -0.5 to +1.3 mmHg
 - Inferiority was driven by a subset of Rhopressa™ patients losing efficacy over time (~ 20%)

Rhopressa™ Rocket 1, Per Protocol

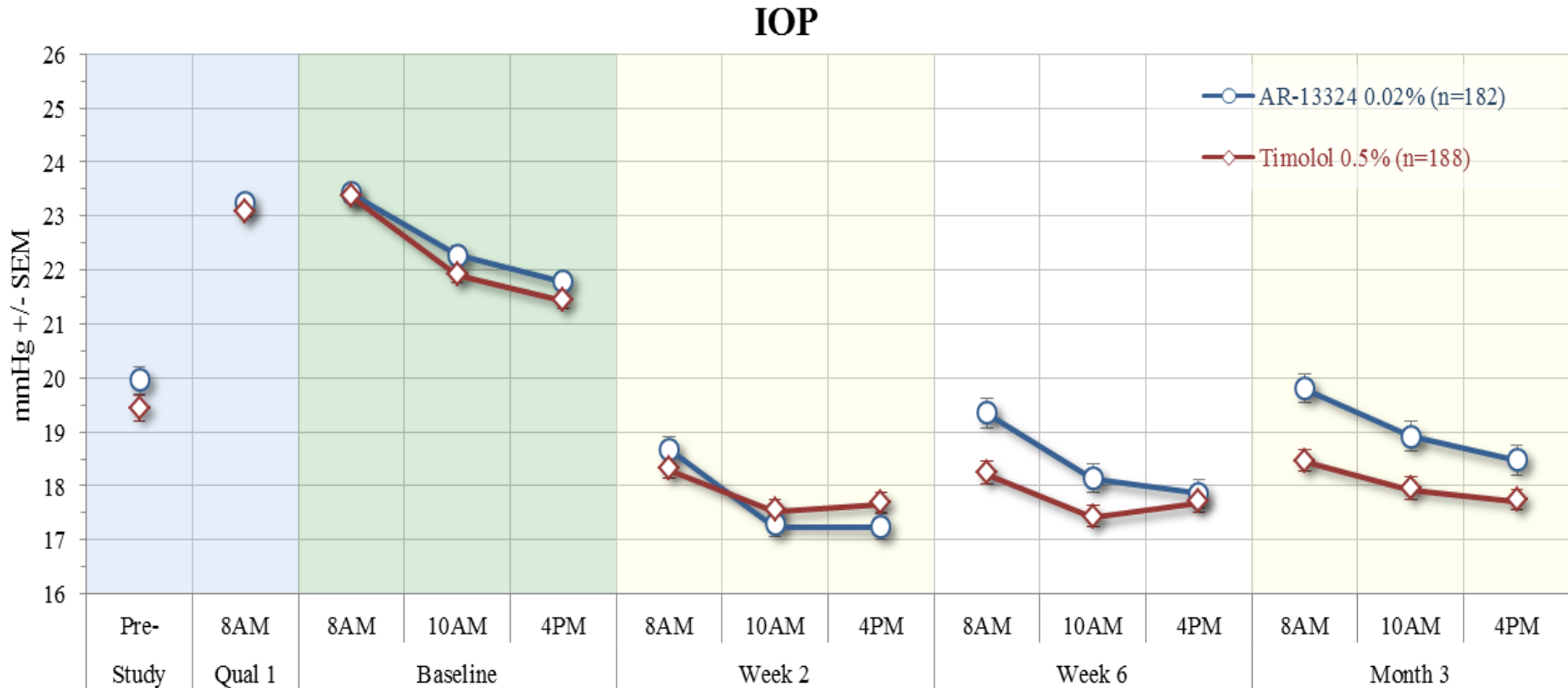


Baseline IOP < 27 mmHg at all time points

	Mean IOP		Rhopressa™– Timolol (95% CI)	
	Rhopressa™ N=182	Timolol N=188	Mean Difference	95% CI
Baseline				
8 AM	23.4	23.4		
10 AM	22.3	21.9		
4 PM	21.8	21.5		
Day 15				
8 AM	18.7	18.3	0.4	(-0.3, 1.0)
10 AM	17.3	17.6	-0.3	(-0.9, 0.4)
4 PM	17.2	17.7	-0.5	(-1.1, 0.2)
Day 43				
8 AM	19.4	18.2	1.1	(0.4, 1.8)*
10 AM	18.1	17.4	0.7	(0.0, 1.4)
4 PM	17.9	17.7	0.2	(-0.5, 0.8)
Day 90				
8 AM	19.8	18.5	1.3	(0.6, 2.0)*
10 AM	18.9	18.0	1.0	(0.3, 1.7)*
4 PM	18.5	17.7	0.7	(0.1, 1.4)

* Missed upper 95% CI criteria (<1.5 mmHg)

Mean Baseline IOP < 27 mmHg At All Time Points



Rhopressa™ Subjects With Decreasing Efficacy Over Time



(Decreasing efficacy over time defined as greater than 3 mmHg)

Correlation With Entry Baseline IOP

- <27 mmHg – 36 drifters (n=182, 19.8%)
- <26 mmHg – 18 drifters (n=133, 13.5%)
- <24 mmHg – 5 drifters (n=76, 6.6%)

Possible explanations for “drifter” subjects

- Severe trabecular meshwork damage (e.g. duration of disease)
- Dosing compliance
- Concomitant medications

n= number of subjects per protocol

Topline Summary (Lower Baseline IOP)

- Lower baseline populations (IOP < 26 mmHg and IOP < 24 mmHg)
 - IOP < 26 mmHg cohort (n=277), Rhopressa™ met non-inferiority criteria at all 9 time points and was numerically superior to Timolol at the majority of time points
 - IOP < 24 mmHg cohort (n=160), Rhopressa™ met non-inferiority criteria at all 9 time points and was numerically superior to Timolol at all 9 time points (Pre-specified analysis)
 - Subjects with loss of efficacy reduced by 50% in baseline IOP < 26 mmHg cohort and by 86% in baseline IOP < 24 mmHg

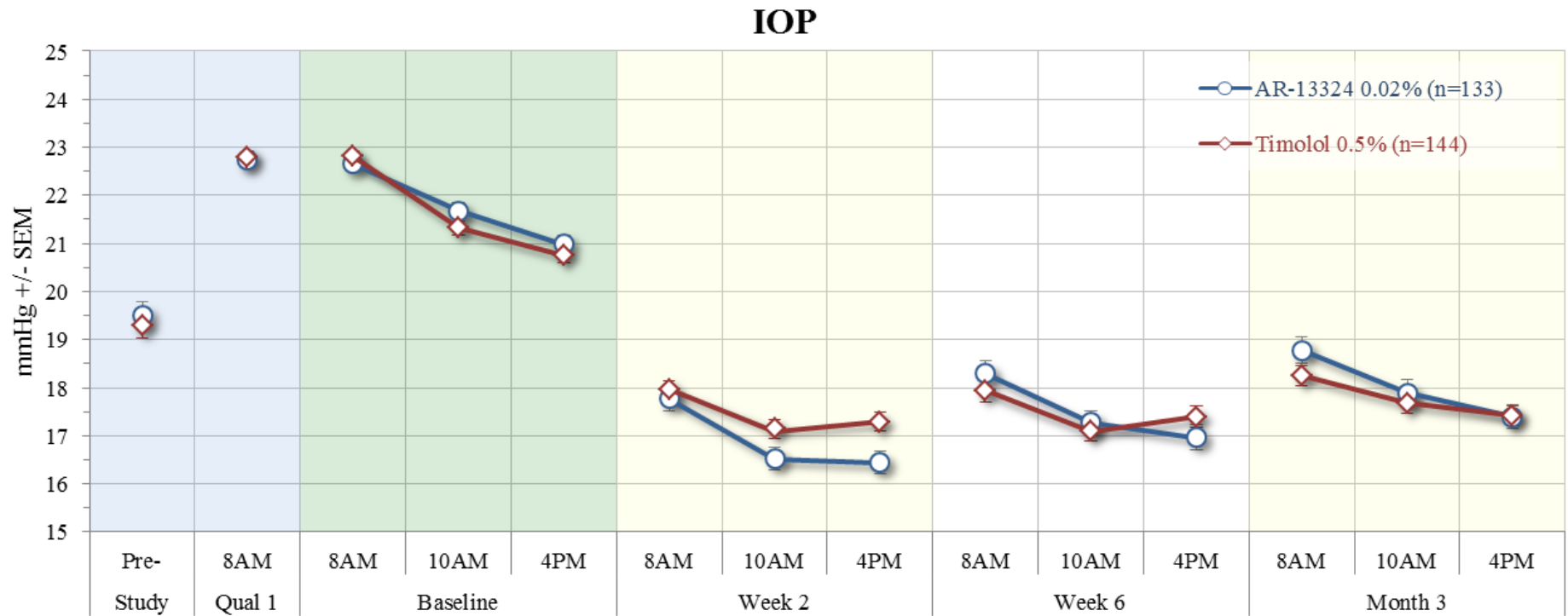
Rhopressa™ Rocket 1, Baseline IOP < 26 mmHg



Rhopressa™ met non-inferiority criteria at all time points and numerical superiority at majority of time points

	Mean IOP		Rhopressa™ – Timolol (95% CI)	
	Rhopressa™ N=133	Timolol N=144	Mean Difference	95% CI
Baseline				
8 AM	22.7	22.8		
10 AM	21.7	21.3		
4 PM	21.0	20.8		
Day 15				
8 AM	17.8	18.0	-0.2	(-0.8,0.4)
10 AM	16.5	17.1	-0.6	(-1.2,0.0)
4 PM	16.4	17.3	-0.9	(-1.5,-0.2)
Day 43				
8 AM	18.3	17.9	0.4	(-0.3,1.1)
10 AM	17.3	17.1	0.2	(-0.5,0.8)
4 PM	17.0	17.4	-0.4	(-1.1,0.3)
Day 90				
8 AM	18.8	18.3	0.5	(-0.2,1.3)
10 AM	17.9	17.7	0.2	(-0.5,0.9)
4 PM	17.4	17.4	-0.01	(-0.7,0.6)

Baseline IOP < 26 mmHg



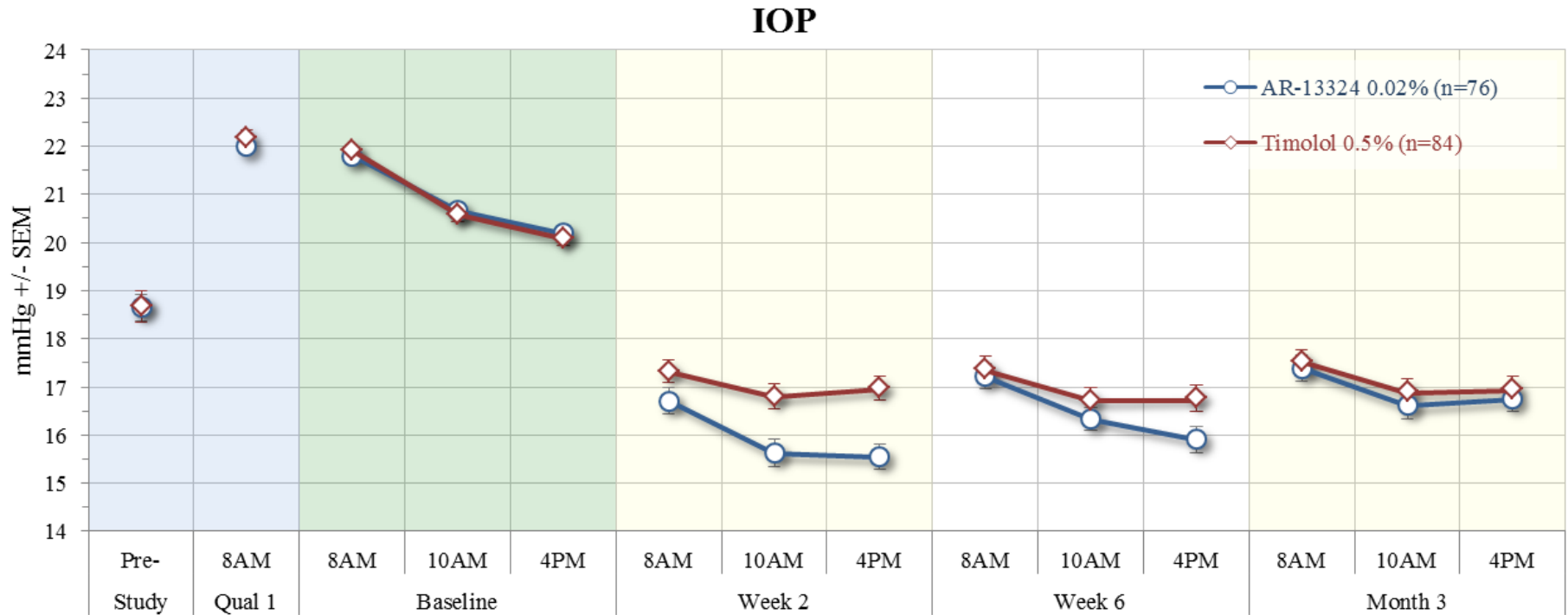
Rhopressa™ Rocket 1, Baseline IOP < 24 mmHg



Rhopressa™ met non-inferiority criteria at all 9 time points was numerically superior to Timolol at all 9 time points (Pre-specified analysis)

	Mean IOP		Rhopressa™– Timolol (95% CI)	
	Rhopressa™ N=76	Timolol N=84	Mean Difference	95% CI
Baseline				
8 AM	21.8	21.9		
10 AM	20.7	20.6		
4 PM	20.2	20.1		
Day 15				
8 AM	16.7	17.3	-0.6	(-1.3, 0.1)
10 AM	15.6	16.8	-1.2	(-1.9, -0.4)
4 PM	15.6	16.9	-1.4	(-2.1, -0.7)
Day 43				
8 AM	17.2	17.4	-0.2	(-0.9, 0.6)
10 AM	16.3	16.8	-0.4	(-1.2, 0.3)
4 PM	15.9	16.8	-0.9	(-1.7, -0.1)
Day 90				
8 AM	17.3	17.6	-0.3	(-1.0, 0.5)
10 AM	16.6	16.9	-0.3	(-1.1, 0.5)
4 PM	16.7	17.0	-0.3	(-1.1, 0.5)

Baseline IOP < 24 mmHg (Pre-specified analysis)



Safety/Tolerability Overview of Rhopressa™ (Days 15-90)



- There were no drug-related serious adverse events (SAEs)
- The most common adverse event was conjunctival hyperemia
 - Conjunctival hyperemia measured by biomicroscopy at 8am was ~35% of which 80% was mild
- Adverse events occurring in approximately 5-13% of the subjects receiving Rhopressa™ included: conjunctival hemorrhage, erythema of the eyelid, blurry vision and corneal deposits

Next Steps

- Rocket 2 efficacy results (expected in Q3 2015) important in determining the next steps, which may include the need for an additional Rhopressa™ registration trial
- Expect to file the Rhopressa™ NDA by the end of 2016, if additional trial is required
- Expect to commence Roclatan™ Phase 3 registration trials by the end of 2015, after review of Rocket 2 results
- Over \$179 million on the balance sheet as of the end of Q1 2015. We are well financed to execute our strategies