



Corporate Review

August 2018

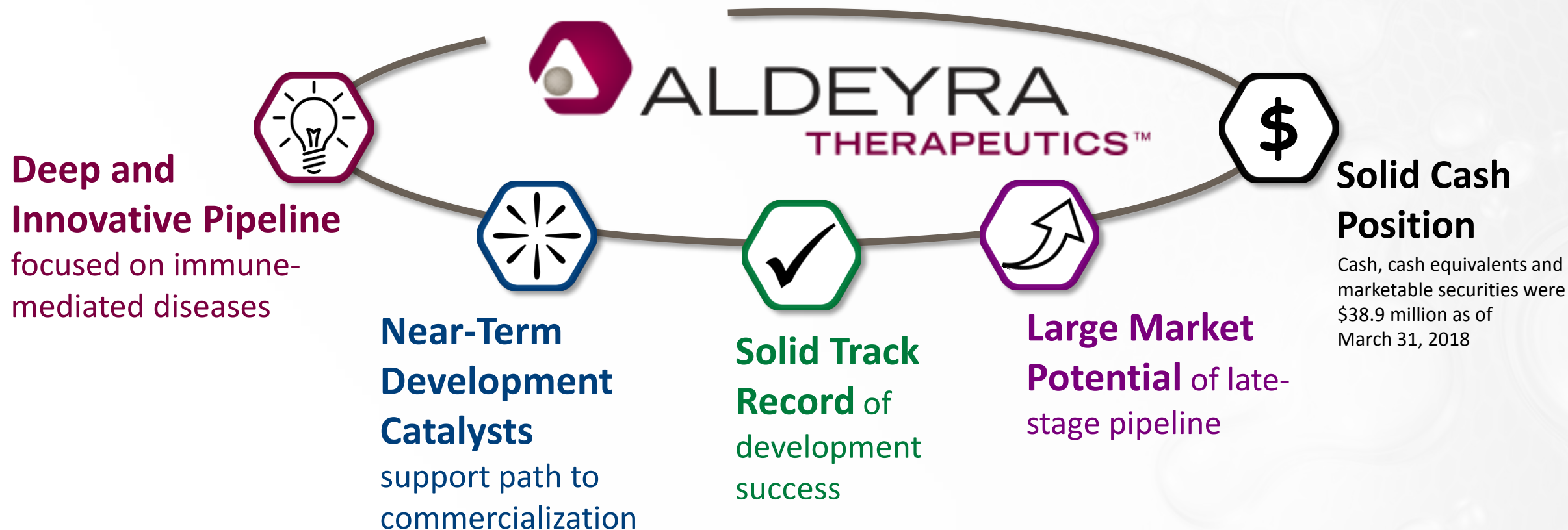
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Developing Next-Generation Medicines to Improve the Lives of Patients with Immune-Mediated Diseases



Our Mission

Developing Next-Generation Medicines to Improve the Lives of Patients with Immune-Mediated Diseases



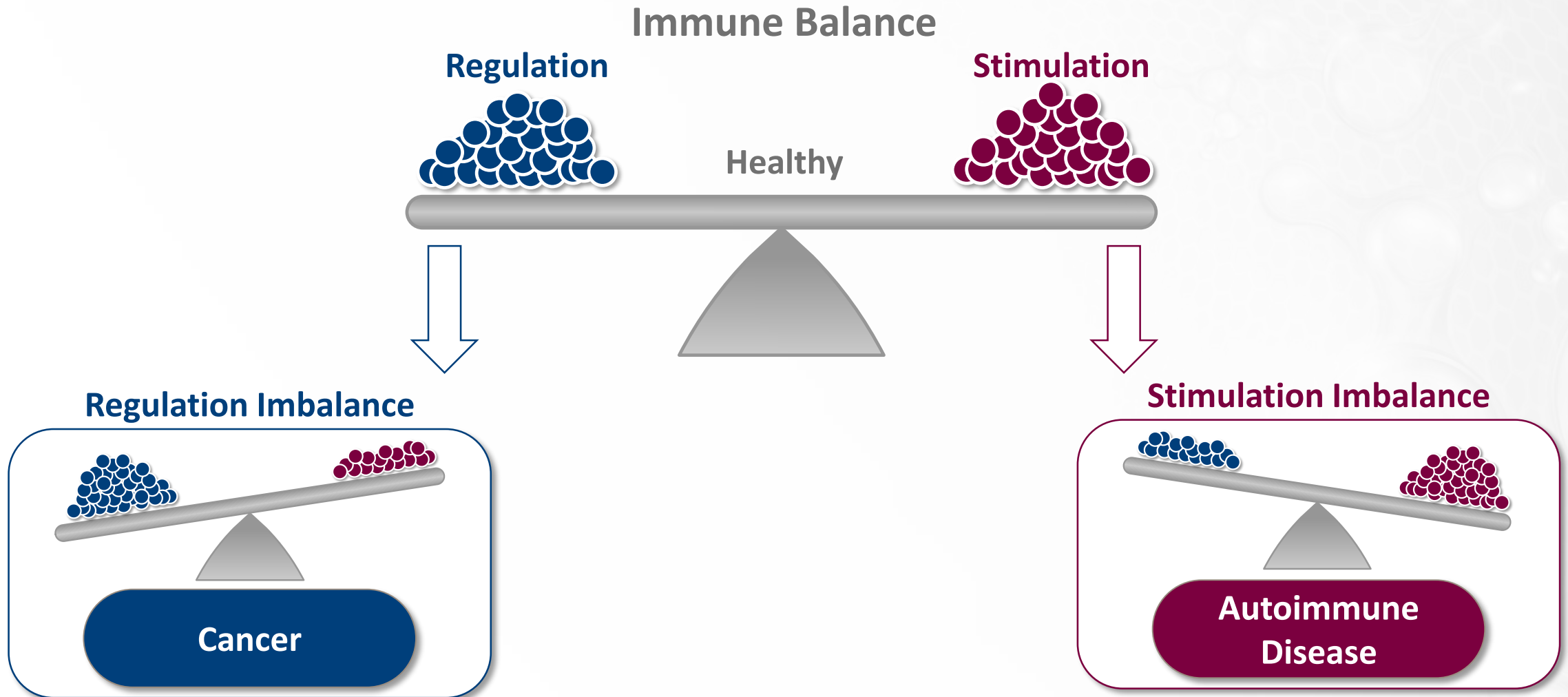
Suffer from some form of **immune-mediated disease**



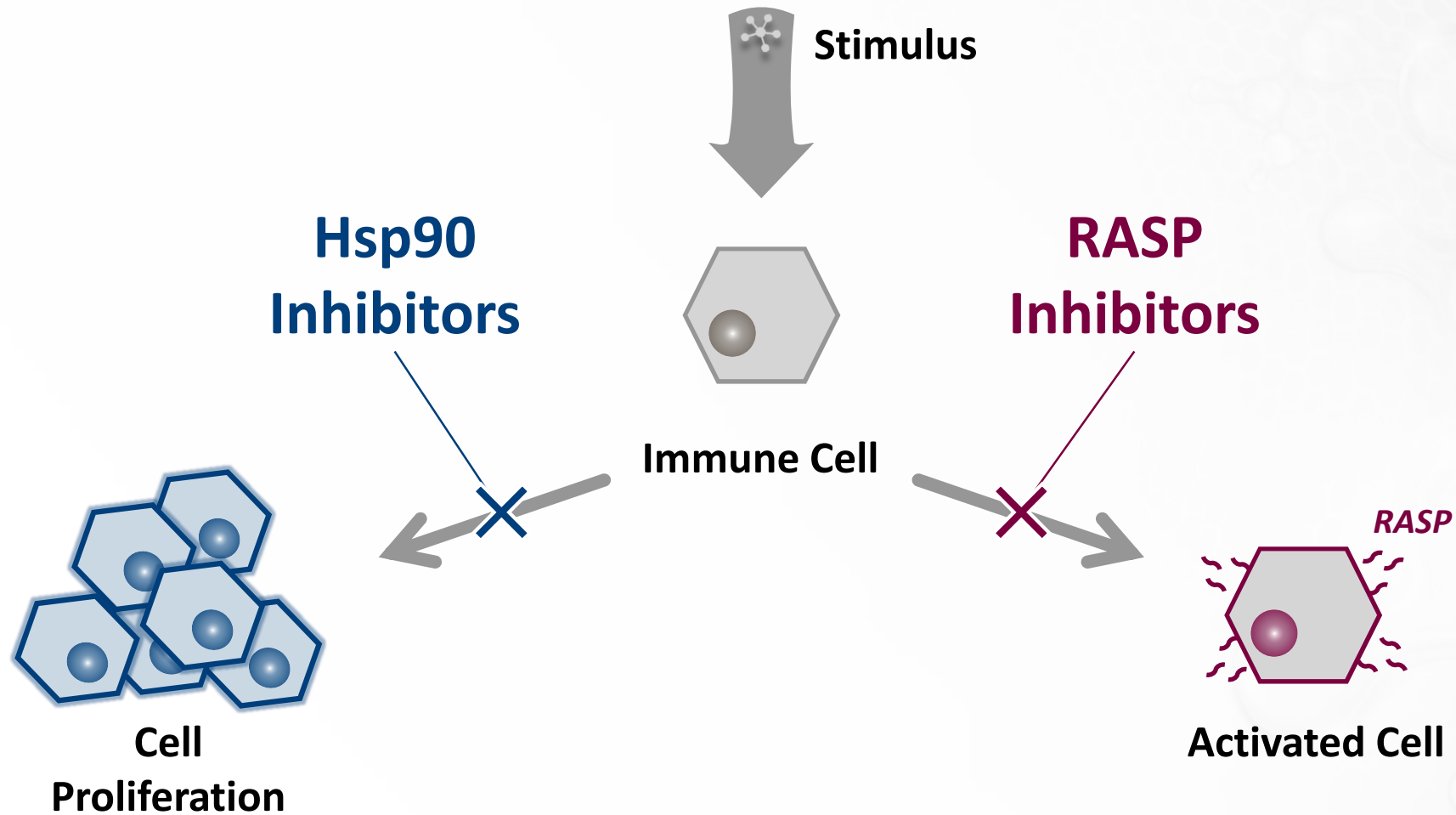
Disease control elusive despite existing therapies, and thus **novel approaches are needed**

Source: Shurin and Smolkin, Advances in Experimental Medicines and Biology 601:3-12, 2007;
Kuek et al, Postgraduate Medical Journal 83(978): 251-260, 2007.

Immune System Imbalance Leads to Disease




Novel Approaches to Address Immune-Mediated Disease



RASP = Reactive Aldehydes Species

Deep and Innovative Pipeline

Mechanism	Compound	Indication	Preclinical	Phase 1	Phase 2	Phase 3	Next Expected Milestone
RASP Inhibitors	Reproxalap Ocular	Dry Eye Disease					Phase 2b results H2-2018
		Allergic Conjunctivitis					Phase 3 results H2-2018 / 2019
		Noninfectious Anterior Uveitis					Phase 3 results 2019
	Reproxalap Dermal	Sjögren-Larsson Syndrome					Phase 3, Part 1 results 2019
	ADX-629 Systemic	Autoimmune Disease					
	ADX-103	Retinal Disease					
	Not Disclosed	Systemic Inflammatory Disease				<i>Research Collaboration</i> 	
Hsp90 Inhibitors	ADX-1612	PTLD					
		Ovarian Cancer				<i>Investigator Sponsored Trial</i>	
		Mesothelioma				<i>Investigator Sponsored Trial</i>	Phase 2 results H2-2018
	ADX-1615	Autoimmune Disease					
		Cancer					
Anti-Inflammatory	Not Disclosed	Ocular Inflammation					



RASP = Reactive Aldehydes Species

PTLD = Post-Transplant Lymphoproliferative Disorder

✓ = Positive Phase 2 clinical data reported in 2016 – 2017



Reproxalap: Our Lead Product Candidate

Potential Benefits Over Standard of Care Across Four Indications

		Reproxalap Development Stage	Current Standard of Care	Potential Reproxalap Competitive Advantages†
 Ocular Inflammation	Dry Eye Disease	Phase 2b	Xiidra®, Restasis®	Rapid onset, broader activity
	Allergic Conjunctivitis	Phase 3	Antihistamines	Non-drying, durable activity; Responder superiority vs. vehicle
	Noninfectious Anterior Uveitis	Phase 3	Corticosteroids	No expected risk of glaucoma or other corticosteroid toxicities
 Inborn Errors of Metabolism	Sjögren-Larsson Syndrome	Phase 3	Bathing, Moisturizers	Clinically demonstrated efficacy; Currently no FDA or EMA approved therapy

† Pending clinical data, regulatory discussions, payor negotiations, competition, potential legislative changes, and other factors, which may not be in Aldeyra's control. Preliminary assumptions are subject to change.

Reproxalap: Target Therapies with Significant Market Potential

		Market and Commercialization Potential			
 Ocular Inflammation		Estimated U.S. Population*	Healthcare Providers	Commercial Build-out	Pricing Benchmarks [†]
	Dry Eye Disease	20 million	Ophthalmologists and Optometrists	Internal Sales Force or Partner	↑ \$500 or greater per course ↓
	Allergic Conjunctivitis	30 million	Ophthalmologists and Optometrists	Internal Sales Force or Partner	
	Noninfectious Anterior Uveitis	150,000	Anterior Segment Ophthalmologists (~30 Centers)	Internal Sales Force or Partner	
 Inborn Errors of Metabolism	Sjögren-Larsson Syndrome	1,000 [‡]	Pediatric Geneticists, Tertiary Care Dermatologists	Internal Sales Force or Partner	\$200,000 - \$400,000 per year

*Aldeyra estimates based on internal market research and publicly available information.

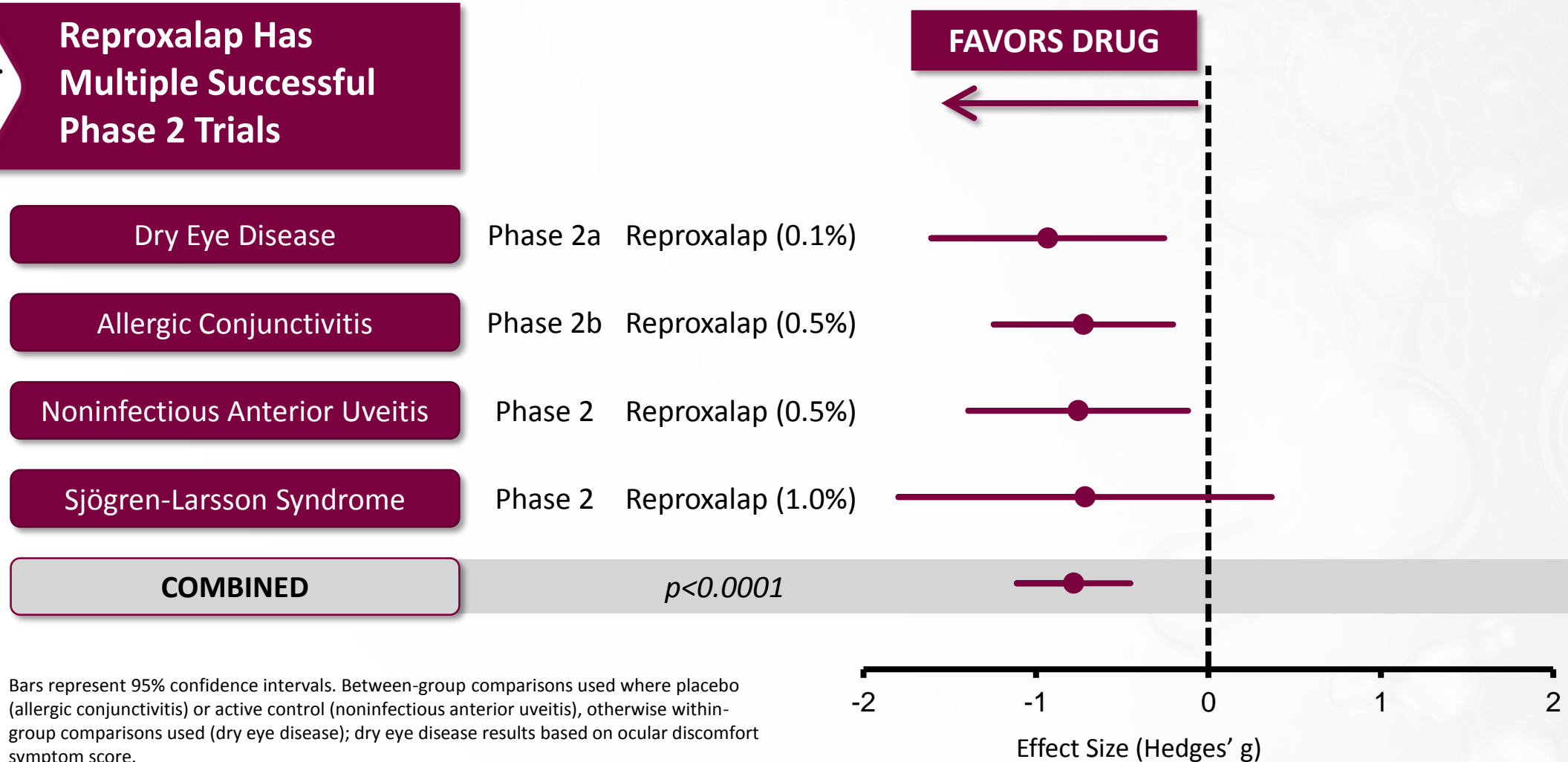
[†]Pending clinical data, regulatory discussions, payor negotiations, competition, potential legislative changes, and other factors, which may not be in Aldeyra's control. Preliminary assumptions are subject to change.

[‡]Extrapolated from a Swedish estimate and a U.S. genetic mutation analysis, it is generally assumed that there are approximately 1,000 Sjögren-Larsson Syndrome (SLS) patients in the United States and a greater number of SLS patients in Europe.

Reproxalap: Meta-Analysis Strongly Supports Drug Activity



**Reproxalap Has
Multiple Successful
Phase 2 Trials**



Source: Aldeyra analysis of Phase 2 clinical trial data on file.



Reproxalap: Ocular Inflammation

- Dry Eye Disease
- Allergic Conjunctivitis
- Noninfectious Anterior Uveitis

Dry Eye Disease: A Chronic Disease with Inadequate Therapy

Large Disease Burden

20
million

of **adults in the U.S.** estimated to suffer from Dry Eye Disease (DED)



vs



Women are twice as likely to suffer from DED than men

Age 50+

>3x

DED **increases with age**, with those over age 50 three times more likely to suffer from DED



DED can significantly effect vision-related **quality of life**

Inadequate Current Therapy

Restasis®

2017 Sales: \$1.5 billion

- Only a subset of patients respond favorably
- May take up to six weeks or longer to have an effect

Xiidra® (launched 2016)

2017 Sales: \$259 million

- Up to 25% of users experience eye irritation or discomfort and an associated bad taste

A Unique Opportunity

Reproxalap

- A **novel and differentiated approach** to treat DED
- **Rapid Improvement of multiple signs and symptoms** observed in patients with DED in a Phase 2a clinical trial
- Phase 2b **results expected H2 2018**

Sources: "Dry Eyes" by R. M. Shtein, MD; www.uptodate.com, May 2018; Farrand et al; American Journal of Ophthalmology 90:98, 2017; Allergan 10K and Shire 10K; Aldeyra research.

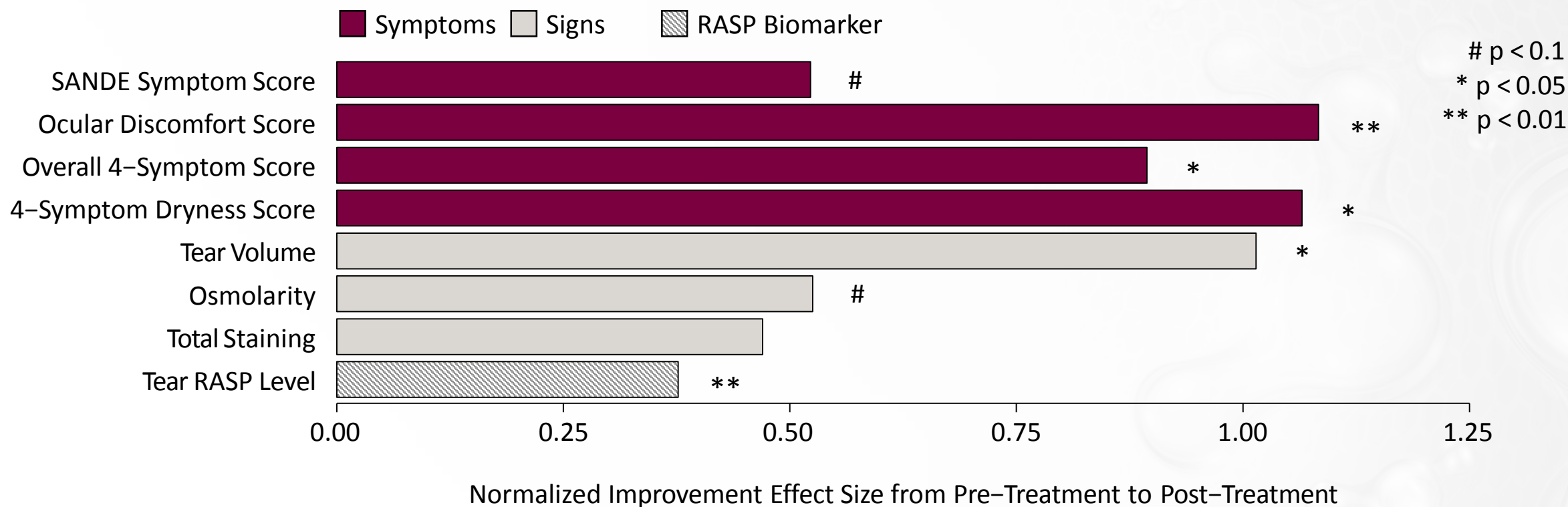
Reproxalap Improved Numerous Dry Eye Disease Signs and Symptoms in Phase 2a Clinical Trial

Endpoint (Pooled Data)		Pre-Treatment*	Post-Treatment*	p value
Symptoms	Symptom Assessment in Dry Eye (SANDE) Score (0-100)	61	52	0.003
	Ocular Discomfort Score (0-4)	2.3	1.5	0.00002
	Overall 4-Symptom Score (0-4)	2.6	2.0	0.0004
Signs	Tear Volume (Schirmer Test)	5.6 mm	8.3 mm	0.008
	Osmolarity	304 mOsm/L	294 mOsm/L	0.003
	Total Staining (Lissamine Green) (0-20)	5.2	4.3	0.002

After one month of therapy, multiple signs and symptoms of dry eye disease improved, a broad and rapid therapeutic response.

Improvement Effect Sizes Were Robust and Statistically Significant in Phase 2a Clinical Trial

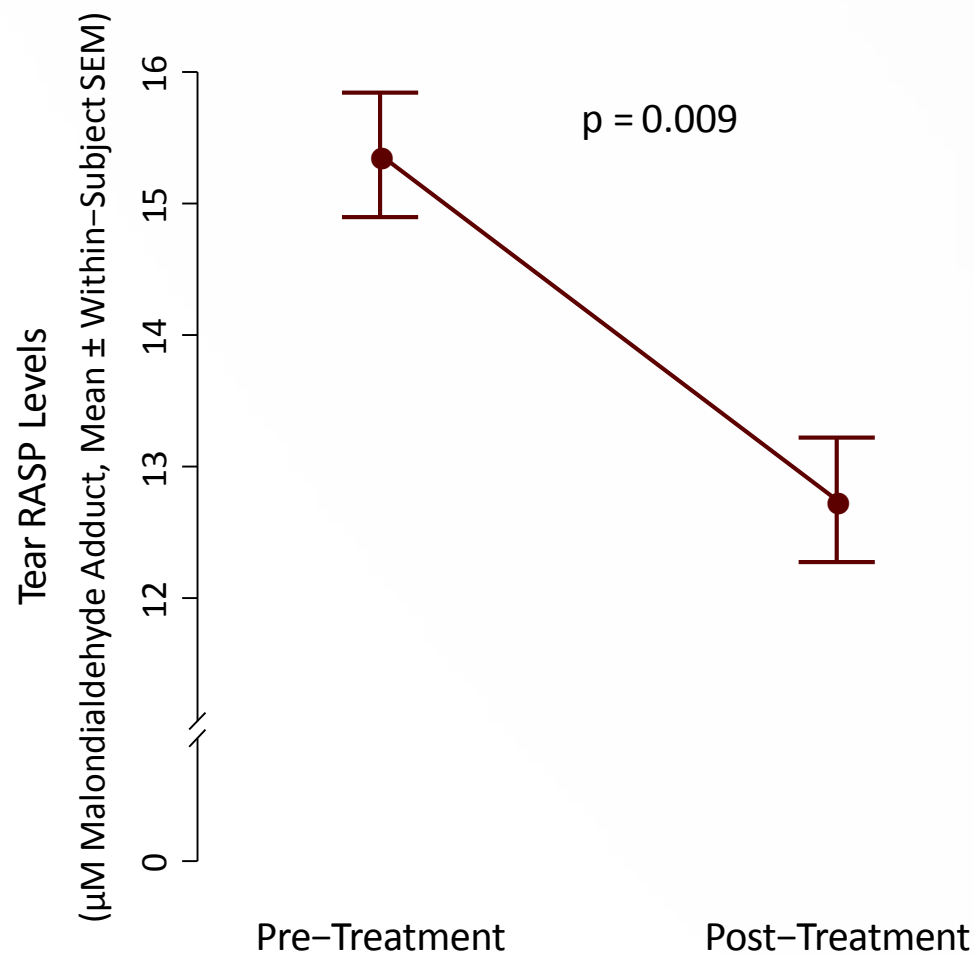
0.1% Reproxalap Improvement Effect Size Across Dry Eye Disease Signs and Symptoms



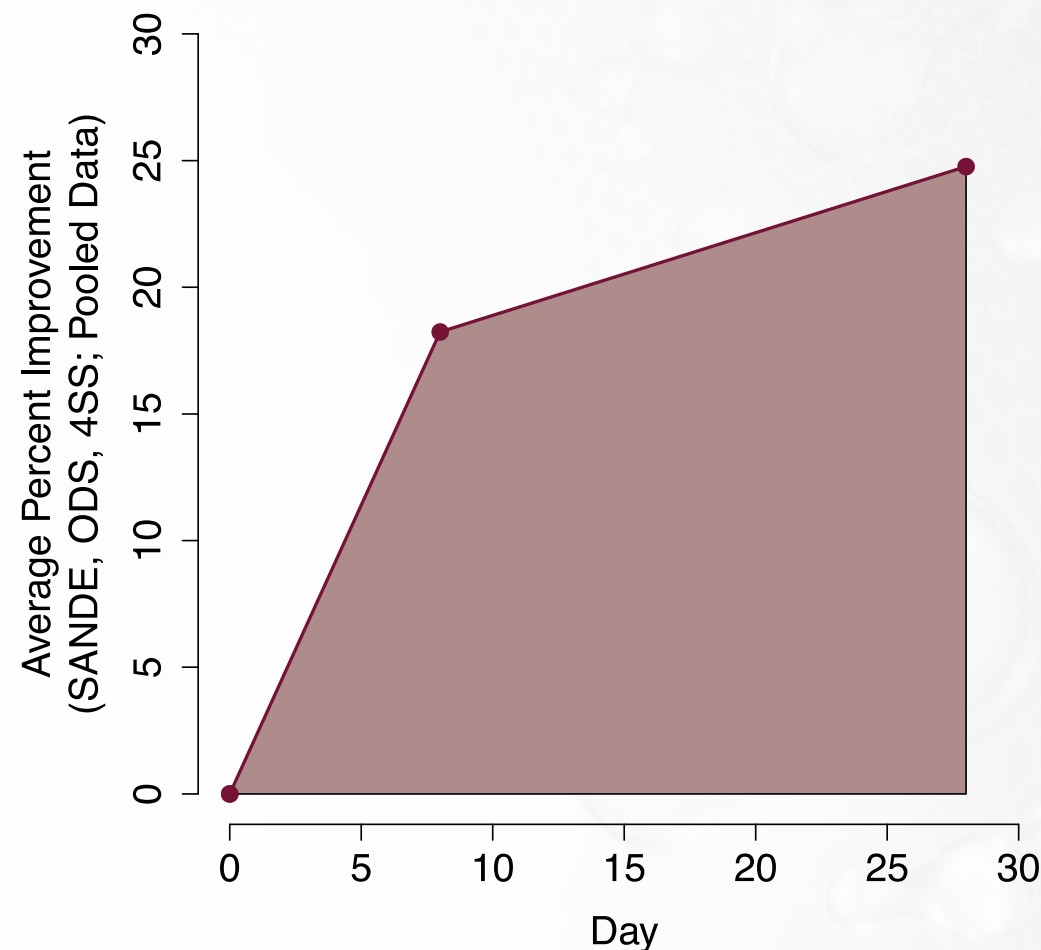
RASP = Reactive Aldehydes Species

Effect size = Mean difference from Day 0 to Day 28 / Standard Deviation of Day 0.

Drug Activity in Phase 2a Clinical Trial Supported by Biomarker Reduction and Increased Efficacy Over Time



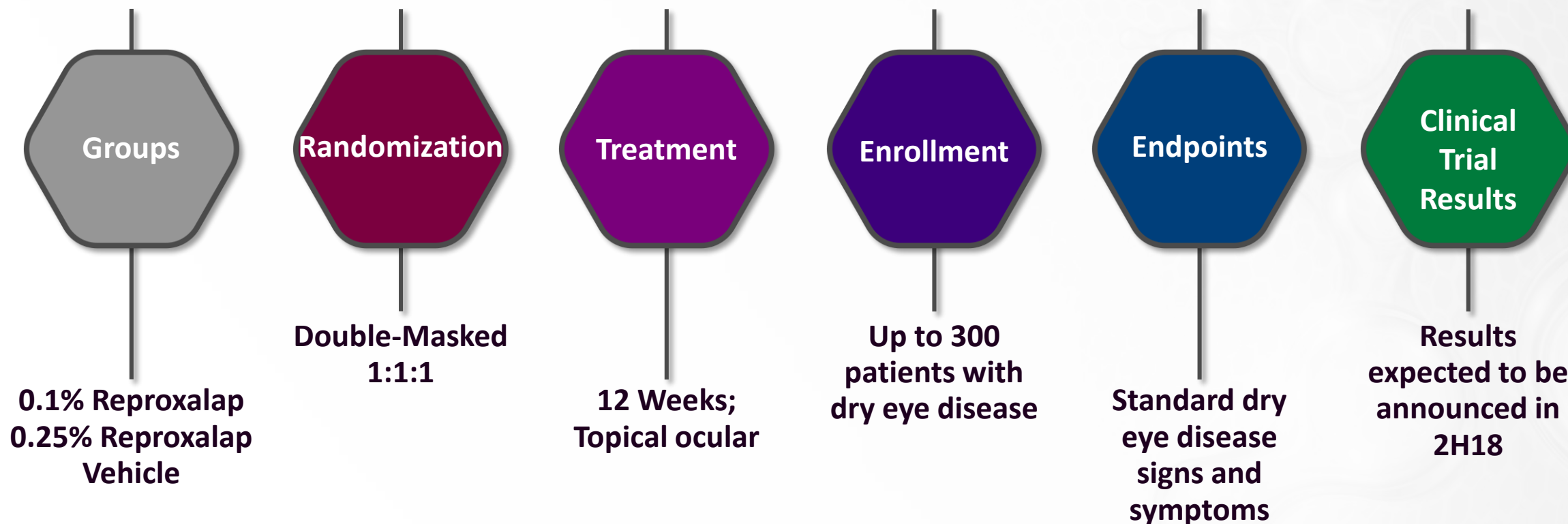
RASP = Reactive Aldehydes Species
Pre-Treatment = Day 0, Post-Treatment = Day 28.



SANDE=Symptom Assessment in Dry Eye Score, ODS=Ocular Discomfort Score,
4SS=Overall 4-Symptom Score

Dry Eye Disease Phase 2b Clinical Trial Design

Initiated January 2018



Further information can be found on www.clinicaltrials.gov: Trial #NCT03404115.

Allergic Conjunctivitis: A Common Disease with Unmet Medical Need

Large Disease Burden

20%
globally

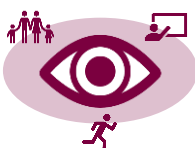
20% or more of people globally suffer from allergic conjunctivitis (AC) annually, and prevalence is increasing



AC can cause persistently disturbing symptoms **acutely, seasonally, and perennially**



Comorbidities with AC are common, including ocular conditions such as dry eye disease



AC may limit patient **quality of life**, affecting daily activities and psychosocial relations

Unmet Medical Need

24%

- Antihistamines are not effective in an estimated 24% of treated AC patients

2%

- Approximately 2% of AC patients have **severe conditions** and may be **steroid-dependent**

A Unique Opportunity

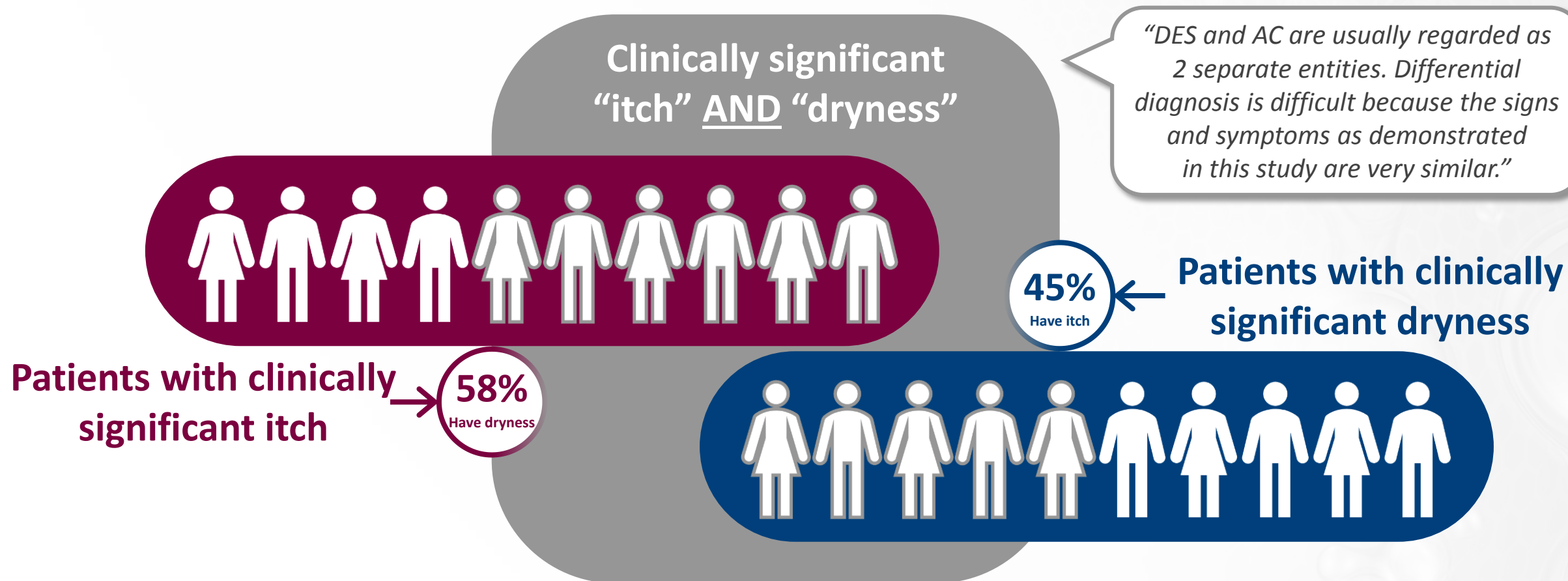
Reproxalap

- A **novel and differentiated approach** to treat AC
- **Mitigated post-histaminic allergy** at levels statistically superior to control in two Phase 2 clinical trials
- Phase 3 **results expected H2 2018 or early 2019**

Sources: "Allergic Conjunctivitis" by Hemran et al; www.uptodate.com, Dec. 2017; Sanchez et al; J Investig Allergol Clin Immunol Suppl. 2: 1-19, 2011; Leonardi et al, Clinical & Experimental Allergy, 45, 1118, 2015; Abelson et al, Allergy Clin Immunol 115:118, 2005; Aldeyra 2017 US physician market research.

Allergic Conjunctivitis and Dry Eye Disease are Related, and Comorbidity is Common

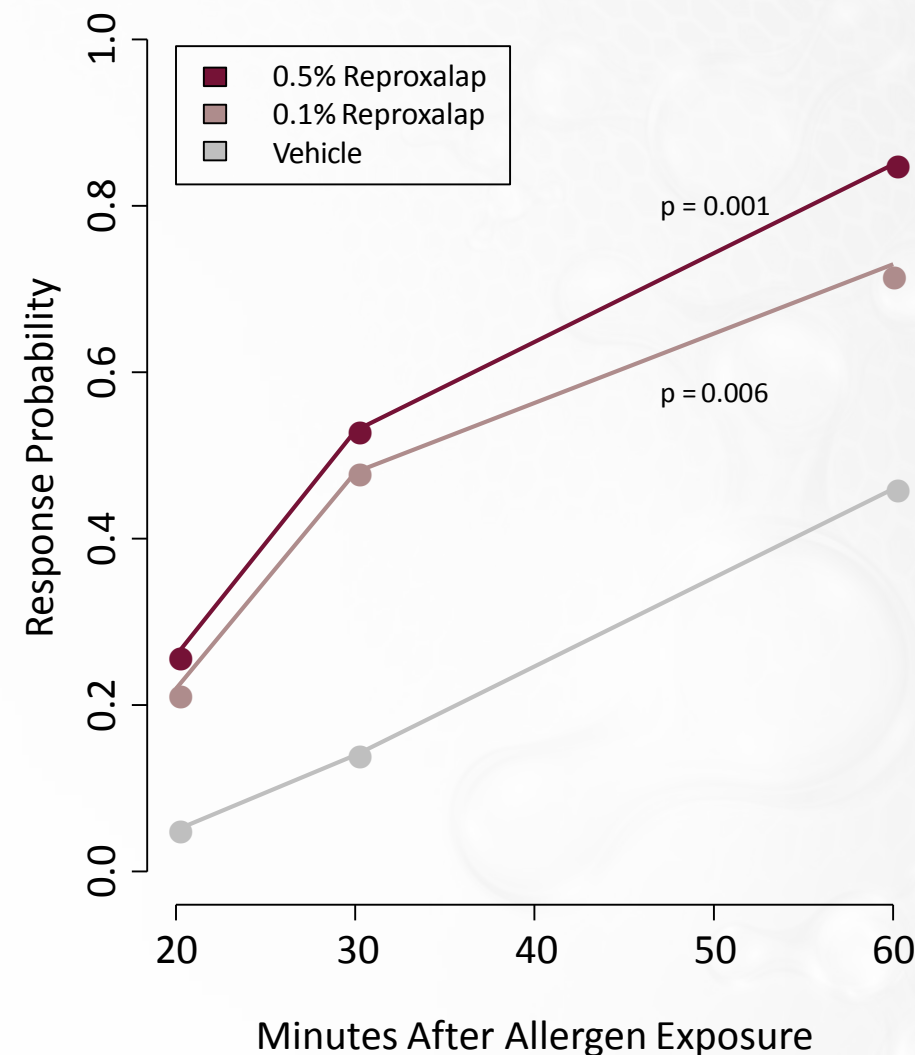
2011 Study of Allergic Conjunctivitis and Dry Eye Syndrome



Source: M.M. Hom et al. / Ann Allergy Asthma Immunol 108 (2012) 163–166.

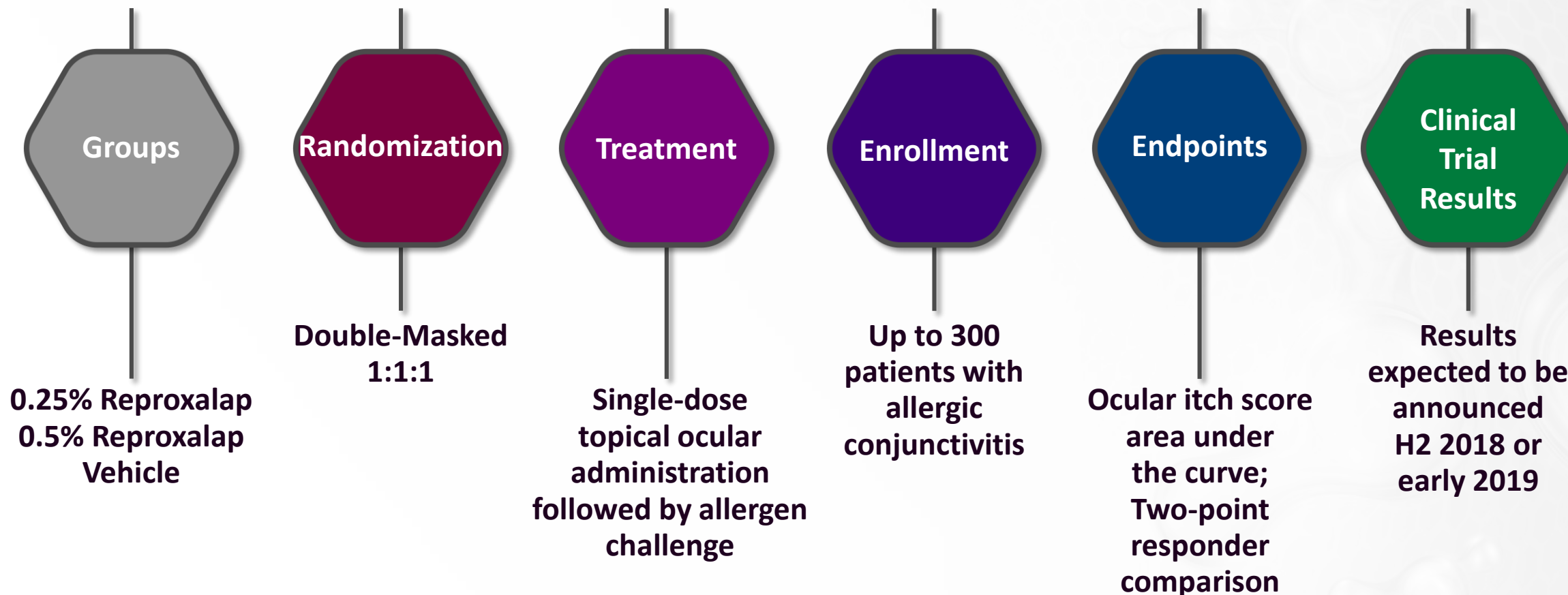
Reproxalap Groups Showed Higher and More Durable Clinical Responses vs. Vehicle Group in Phase 2b Clinical Trial

- On the ocular itch scale (range 0 – 4), clinical response was defined as improvement of two points from peak itch score.
- Probability of response statistically higher in 0.1% and 0.5% reproxalap groups vs. vehicle ($p=0.006$ and $p=0.001$, respectively).



ALLEVIATE Trial Design in Allergic Conjunctivitis

Phase 3 Clinical Trial Initiated April 2018



Further information can be found on www.clinicaltrials.gov: Trial #NCT03494504.

Noninfectious Anterior Uveitis: A Serious Disease That Can Cause Loss of Vision

Serious Inflammatory Disease



Noninfectious Anterior Uveitis (NAU) is a severe **autoimmune acute ocular inflammation**



Inflammatory cells in front of eye cause **pain, photophobia, and loss of vision**

150K
annually

NAU is a **rare disease** with an estimated 150,000 U.S. patients per year



NAU has a big impact on **quality of life**, leading to loss of work and significant economic burden

Inadequate Current Therapy

Steroids

- Currently treated with **corticosteroids**, which may lead to **cataracts and glaucoma**

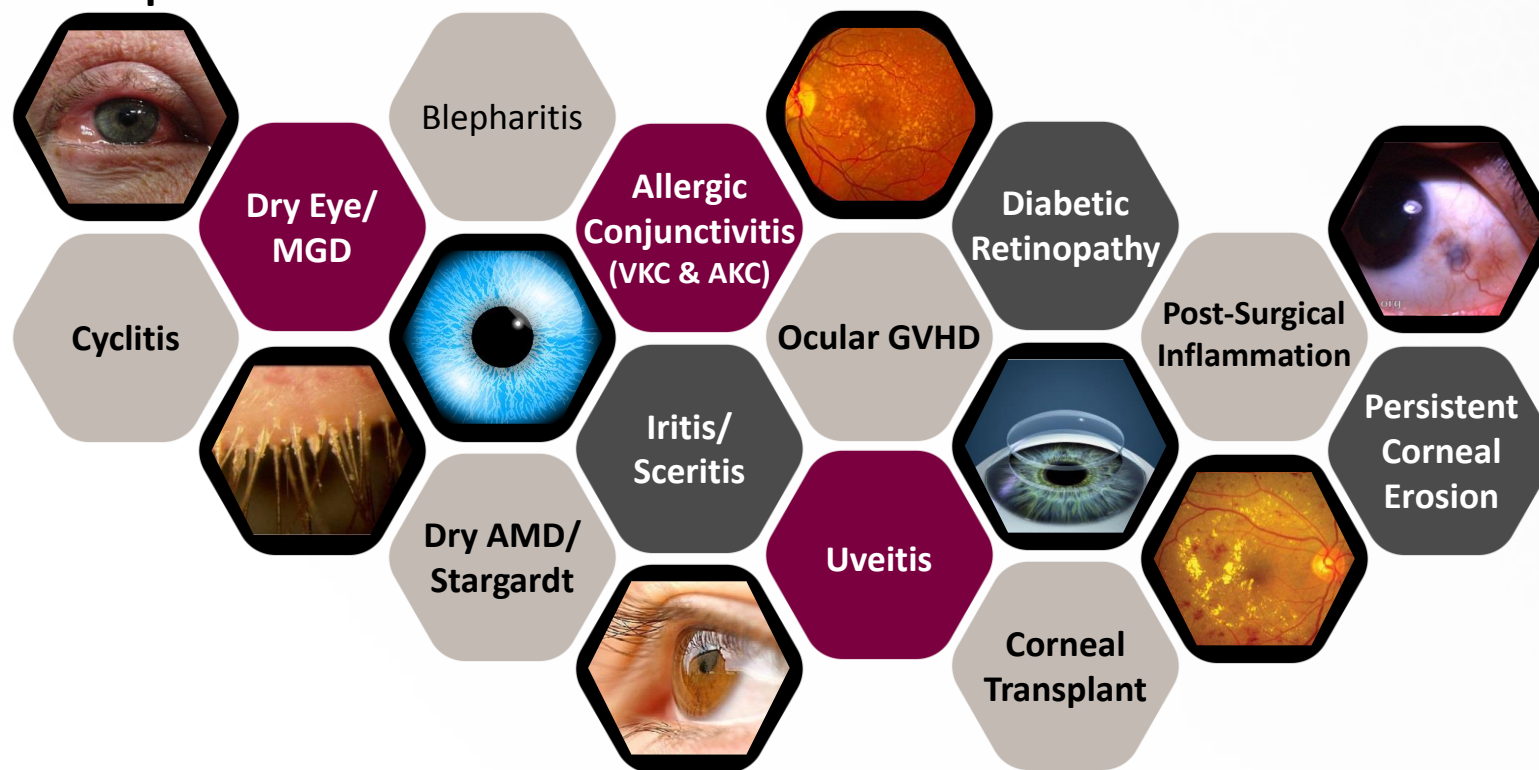
A Unique Opportunity

Reproxalap

- A **novel and differentiated approach** to treat NAU
- **Reduced anterior chamber cell count** in a randomized, vehicle controlled Phase 2 clinical trial, but **did not cause corticosteroid-related side effects**
- Phase 3 **results expected 2019**

Steroid Toxicity Creates Significant Demand for Novel Approaches

Widespread corticosteroid use:



Potential corticosteroid side-effects:

- ☐ Blurred vision
- ☐ Cataracts
- ☐ Corneal ulceration
- ☐ Delayed wound healing
- ☐ Glaucoma
- ☐ Ocular infection
- ☐ Ptosis
- ☐ Redness
- ☐ Swelling
- ☐ Tear film instability

Despite toxicity, current topical ocular corticosteroid usage generates annual sales around \$800M*

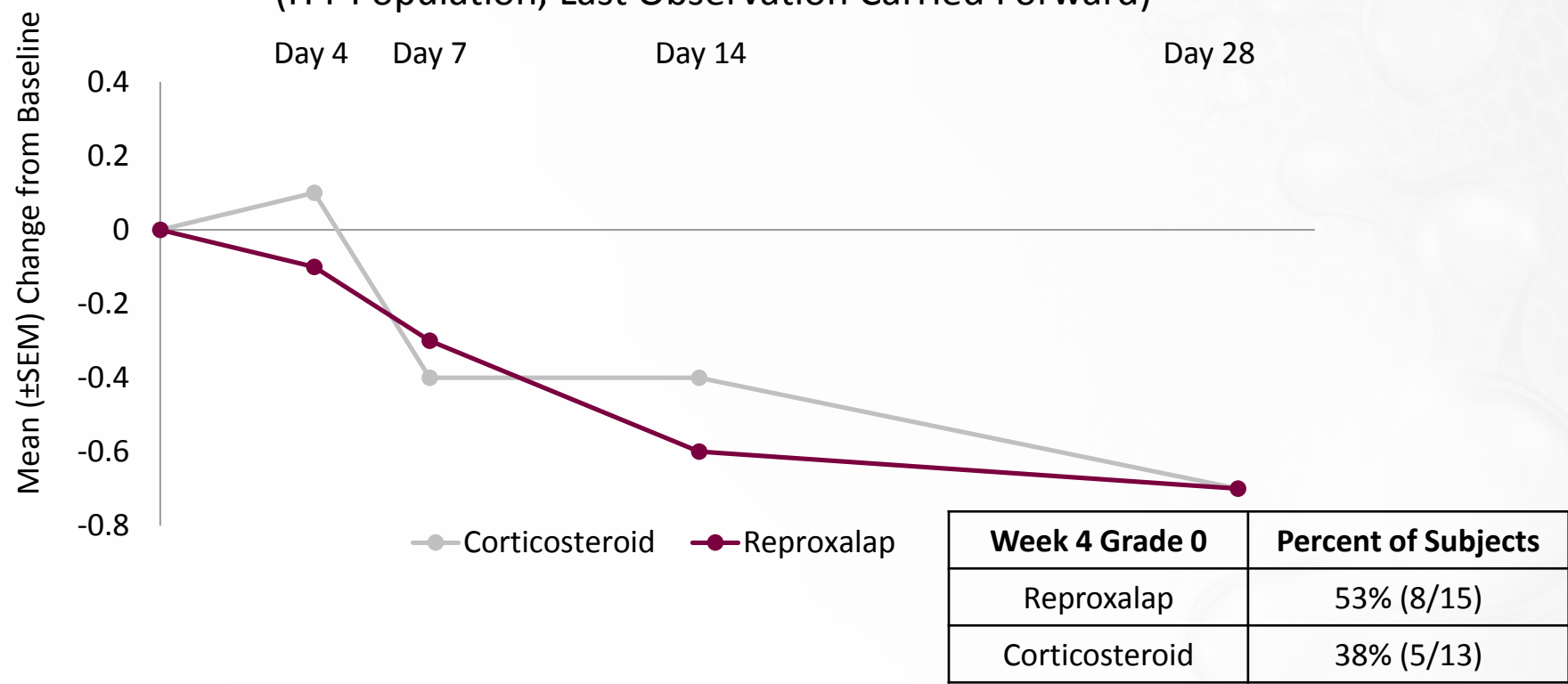
MGD: meibomian gland dysfunction, VKC: vernal keratoconjunctivitis, AKC: atopic keratoconjunctivitis, AMD: age-related macular degeneration.

Post-Surgical Inflammation includes inflammation resulting from corneal trauma, including cataract and refractive surgery.

*Based on 2016 IMS data; Neither reproxalap nor any of Aldeyra's other product candidates are currently in clinical development for any of the above diseases, other than dry eye disease, allergic conjunctivitis, and noninfectious anterior uveitis.

Reproxalap Reduced Inflammation in Noninfectious Anterior Uveitis Phase 2 Clinical Trial

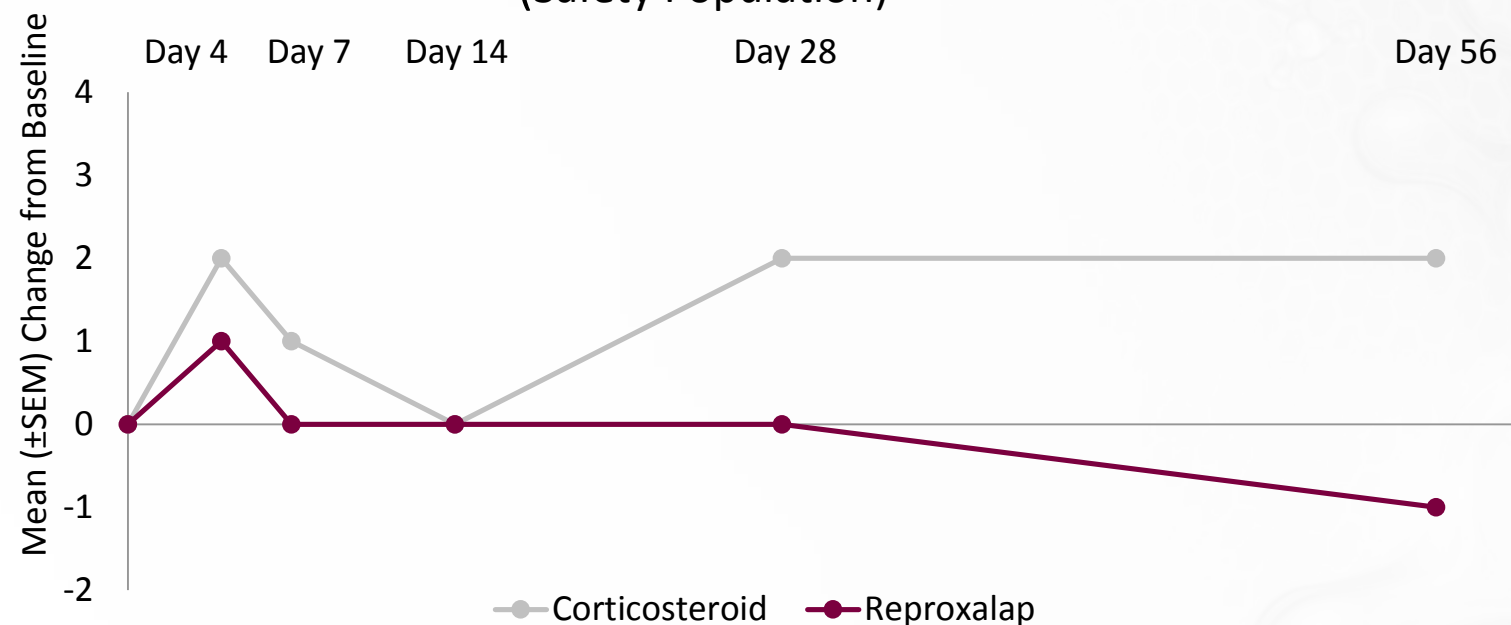
Change from Baseline in Anterior Chamber Inflammatory Cell Grade over Time
(ITT Population, Last Observation Carried Forward)



Reproxalap was statistically non-inferior to corticosteroid in a noninfectious anterior uveitis Phase 2 clinical trial.

Reproxalap Did Not Increase Intraocular Pressure in Noninfectious Anterior Uveitis Phase 2 Clinical Trial

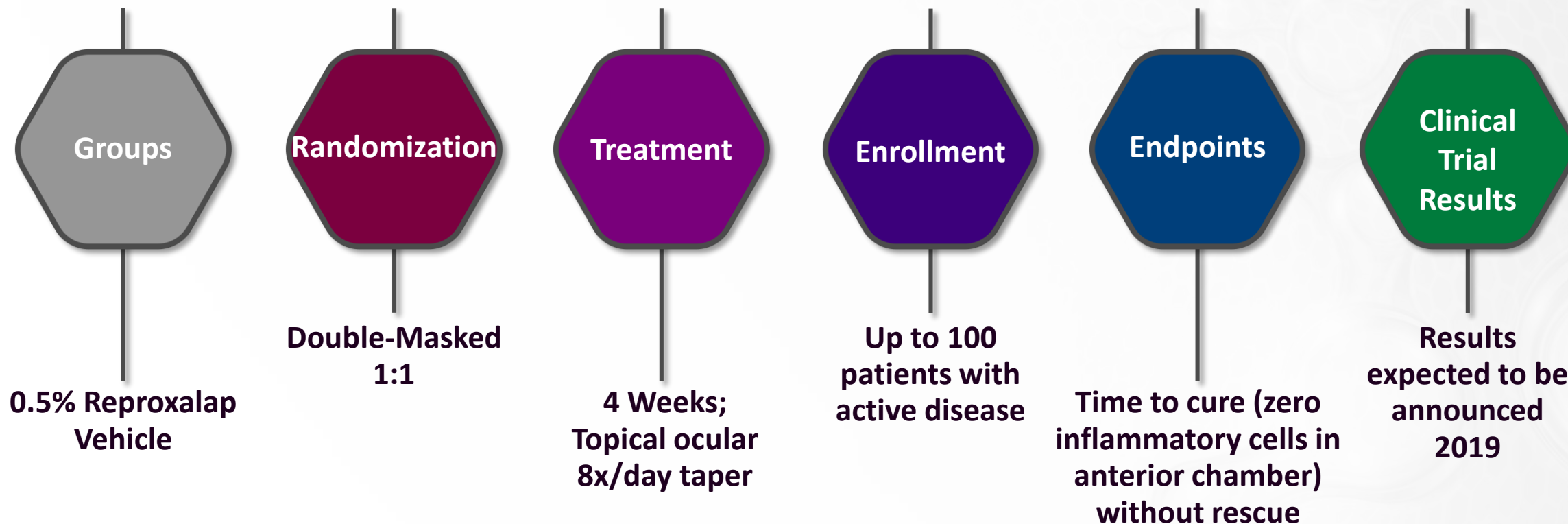
Change from Baseline in Intraocular Pressure (mmHg) over Time
(Safety Population)



Increase in intraocular pressure, which may lead to glaucoma, is a major corticosteroid toxicity that is not apparent with reproxalap.

SOLACE Trial Design in Noninfectious Anterior Uveitis

Phase 3 Clinical Trial Initiated April 2017



Further information can be found on www.clinicaltrials.gov: Trial #NCT03131154.



Reproxalap: Sjögren-Larsson Syndrome

Sjögren-Larsson Syndrome: A Rare Disease with No Approved Therapy

An Inborn Error of Metabolism



Sjögren-Larsson Syndrome (SLS) is **caused by an enzyme mutation** (Fatty Aldehyde Dehydrogenase), leading to high levels of RASP

birth → **50s**

SLS is **present at birth** and patients survive into their 50s

1,000
U.S.

SLS is a **rare disease**, with ~1,000 SLS patients in the U.S. and a greater number in Europe¹



Severe skin scaling, retinal disease, and neurological disorders significantly impact **SLS patient burden and quality of life**

RASP = Reactive Aldehydes Species

¹Extrapolating from a Swedish estimate in addition to a U.S. genetic mutation analysis. It is generally assumed that there are approximately 1,000 SLS patients in the United States and a greater number of SLS patients in Europe.

Inadequate Current Therapy



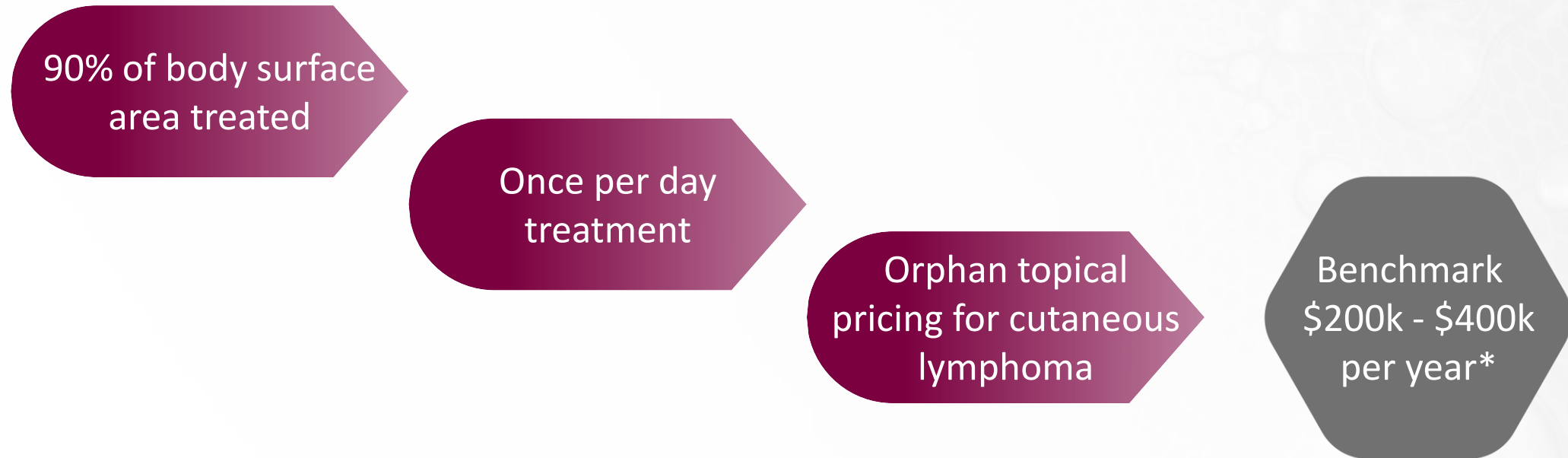
- No FDA or EMA approved therapy that addresses the disease

A Unique Opportunity

Reproxalap (topical dermatologic)

- A **novel approach and potential lifelong therapy** to replace missing enzymatic activity in SLS
- **Granted U.S. orphan designation**
- **Significantly reduced SLS ichthyosis** in a randomized, vehicle controlled Phase 2 clinical trial
- Phase 3, Part 1 **results expected 2019**

Potential Lifelong Therapy for Sjögren-Larsson Syndrome



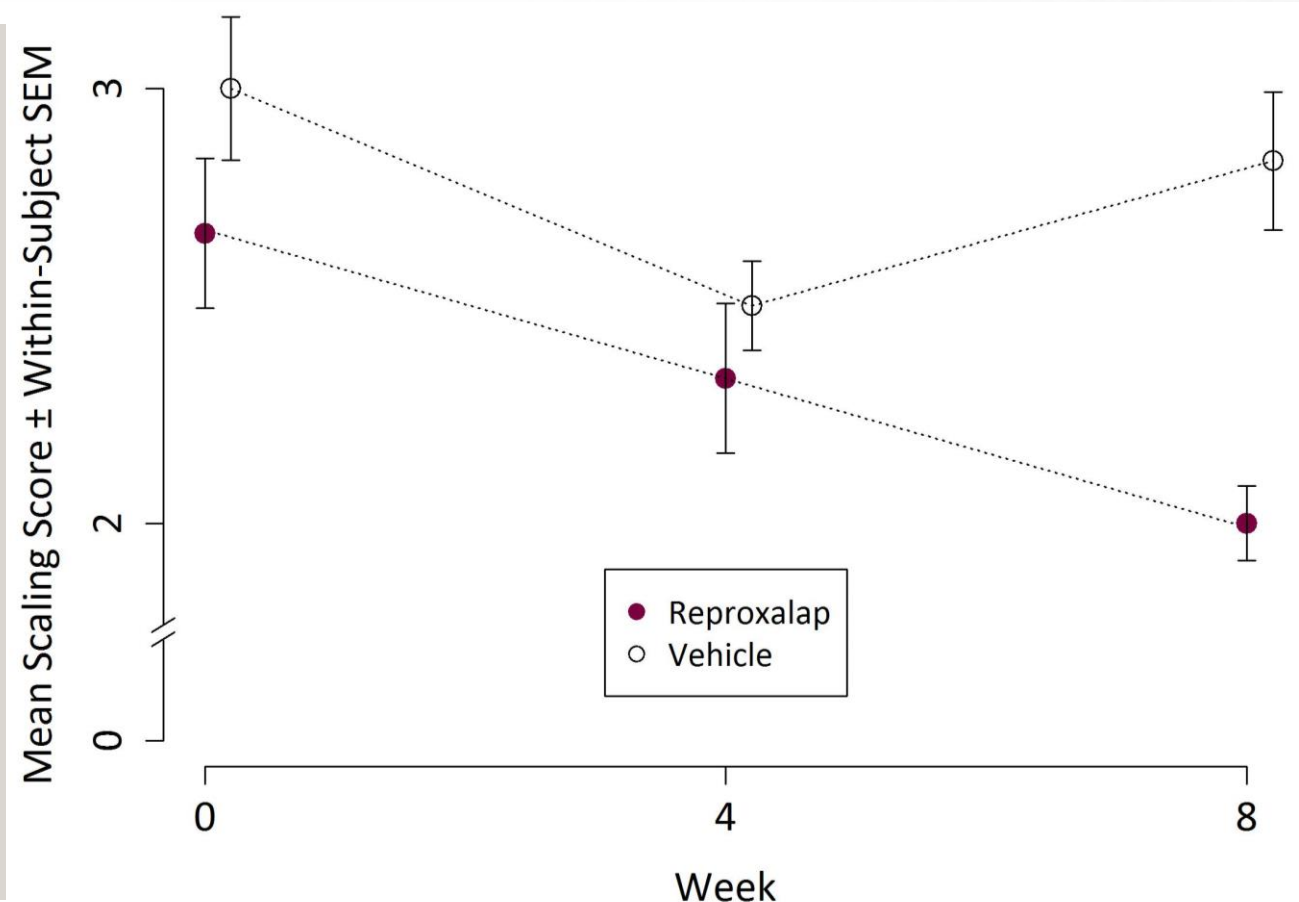
Estimated 0.4 births/100,000 (about 1,000 patients in U.S.)[†]
Total Estimated U.S. SLS market: ~\$200M

**Managed Care Qualitative Market Research of Reproxalap as a Potential Topically Applied Treatment for the Dermatologic Aspects of Sjögren-Larsson Syndrome, CPD Research & Consulting; pricing pending clinical data, regulatory approval, regulatory discussions, payor negotiations, competition, potential legislative changes, and other factors.*

[†]Extrapolating from a Swedish estimate in addition to a U.S. genetic mutation analysis. It is generally assumed that there are approximately 1,000 SLS patients in the United States and a greater number of SLS patients in Europe.

Every Drug-Treated Subject Showed Signs of Improvement in a Phase 2 Clinical Trial

Investigator Assessment of Ichthyosis



Over two months of treatment, ichthyosis improved consistently from moderate to mild disease.

Representative Improvement in Reproxalap-Treated Patients in Phase 2 Clinical Trial



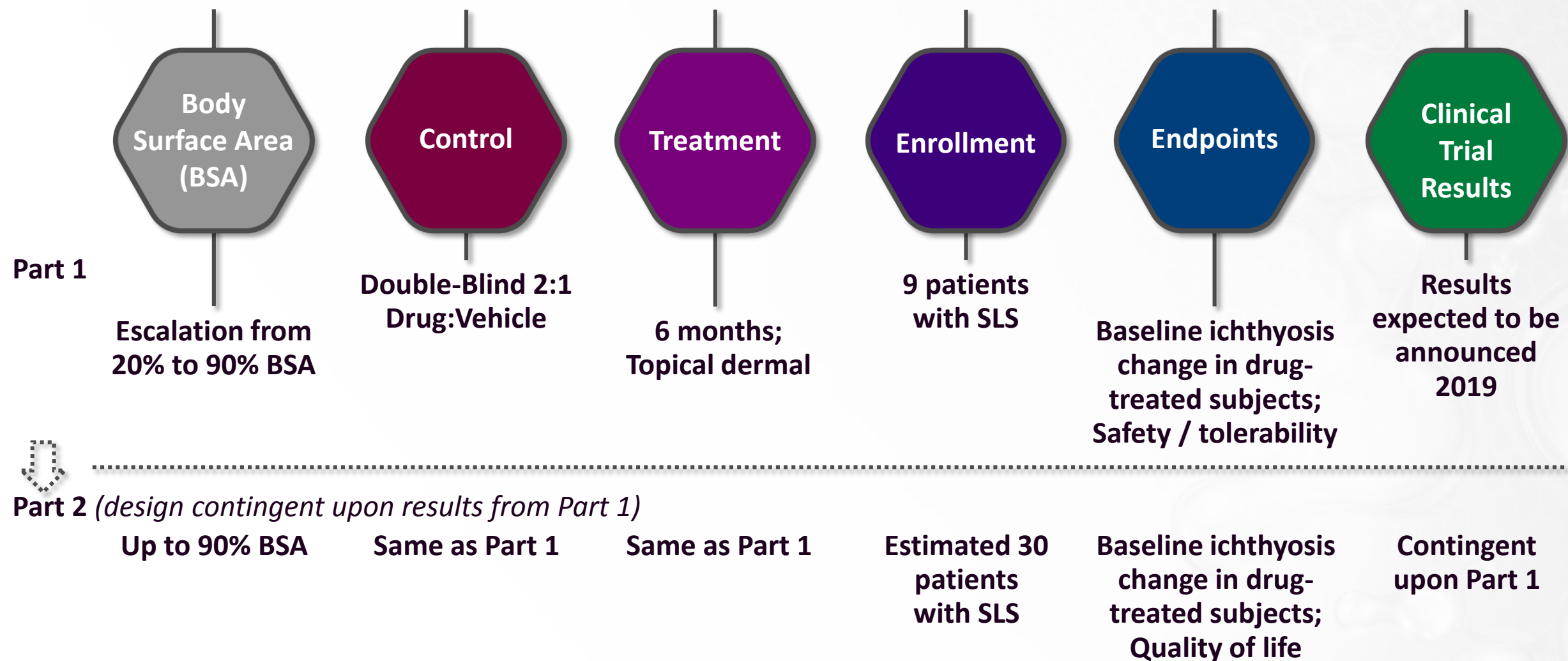
Before Treatment (Week 0)



After Treatment (Week 8)

RESET Trial Design in Sjögren-Larsson Syndrome

Phase 3 Part 1 Clinical Trial Initiated June 2018





Building The Future

Experienced Management Team and Board of Directors

Management Team

Todd Brady, M.D., Ph.D.
President, CEO, & Director



Joshua Reed
Chief Financial Officer



David Clark, M.D.
Chief Medical Officer



David McMullin
SVP Corporate Development



Board of Directors

Richard Douglas, Ph.D. Former SVP Corporate Development at Genzyme
CHAIRMAN

Gary Phillips, M.D. CEO OrphoMed

Neal Walker, D.O. CEO Aclaris Therapeutics

Ben Bronstein, M.D. Former CEO Peptimmune³

Marty Joyce Former CFO of Serono USA















Jesse Treu, Ph.D. Domain Associates

Todd Brady, M.D., Ph.D. CEO Aldeyra Therapeutics

1. Acquired by Schwarz/UCB

3. Acquired by Genzyme

Deep and Innovative Pipeline

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RASP Inhibitors	Reproxalap Ocular	Dry Eye Disease					Phase 2b results H2-2018
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		Cancer					
Anti-Inflammatory	Not Disclosed	Ocular Inflammation					

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✓ = Positive Phase 2 clinical data reported in 2016 – 2017

2018 Progress and Near-Term Development Catalysts

Support Path to Commercialization

H1 2018

- ✓ Initiated reproxalap **Phase 2b clinical trial in dry eye disease**
- ✓ Initiated reproxalap **ALLEVIATE trial in allergic conjunctivitis**
- ✓ Entered into **research collaboration with Johnson & Johnson Innovation** in systemic inflammatory diseases
- ✓ Disclosed **in-license of a Hsp90 inhibitor**
- ✓ Clinical sites initiated for reproxalap **RESET Part 1 trial in Sjögren-Larsson Syndrome**

H2 2018

- ✓ First patient enrolled in reproxalap **RESET Part 1 trial in Sjögren-Larsson Syndrome**

Anticipated Milestones*







- Reproxalap dry eye disease Phase 2b clinical trial results **H2-2018**
- ADX-1612 mesothelioma clinical trial results (investigator sponsored trial) **H2-2018**
- ADX-1612 ovarian cancer clinical trial initiation (investigator sponsored trial) **H2-2018**
- Reproxalap allergic conjunctivitis ALLEVIATE trial results **H2-2018/early 2019**

*Contingent on funding, regulatory review, and other factors.

2019 Expected Development Milestones: Novel Approaches to Address Immune-Mediated Disease

2019

Anticipated Milestones*

-  Reproxalap noninfectious anterior uveitis
SOLACE trial results **2019**
-  Reproxalap Sjögren-Larsson Syndrome
RESET Part 1 trial results **2019**
-  ADX-629 Phase 1 clinical trial initiation **2019**
-  ADX-629 NASH and/or IBD Phase 2a clinical trials initiation
following Phase 1 clinical trial
-  ADX-103 retinal disease Phase 1/2 clinical trial initiation **2019**
-  ADX-1612 post-transplant lymphoproliferative disorder
Phase 2 clinical trial initiation **2019**

*Contingent on funding, regulatory review, and other factors.