

A Randomized, Comparator-Controlled Phase 2 Clinical Trial of ADX-102 Ophthalmic Solution in Noninfectious Anterior Uveitis





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Disclosures

- Alcon: Research Grants, Speaker, Advisor
- Aldeyra Pharmaceuticals: Advisory Board, Clinical Research
- Allergan: Research Grants, Speaker, Advisory Board, Media Spokesman
- Bausch & Lomb, Ista, Valeant: Research Grants, Speaker, Advisory Boards
- Abbvie: Research, Advisor, Spokesman
- BioTissue: Advisory Board, Clinical Research
- Clearside Ophthalmics: Advisor, Research
- Doctors Allergy Formula: Advisor, Investor
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- EyeRx Research: Clinical Research, Stock Ownership
- Imprimis Pharma: Advisory Board
- Isis Pharmaceuticals: Research, Advisory Board
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- · RPS: Advisory, Research, Investor
- Rutech: Clinical Investigator, Advisory Board
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- Shire, SarCode Biosciences: Advisory Board, Shareholder, Research Grant
- Synedgen: Advisory Board
- Science Based Health: Research, Advisory, Spokesman
- Senju: Research Grants
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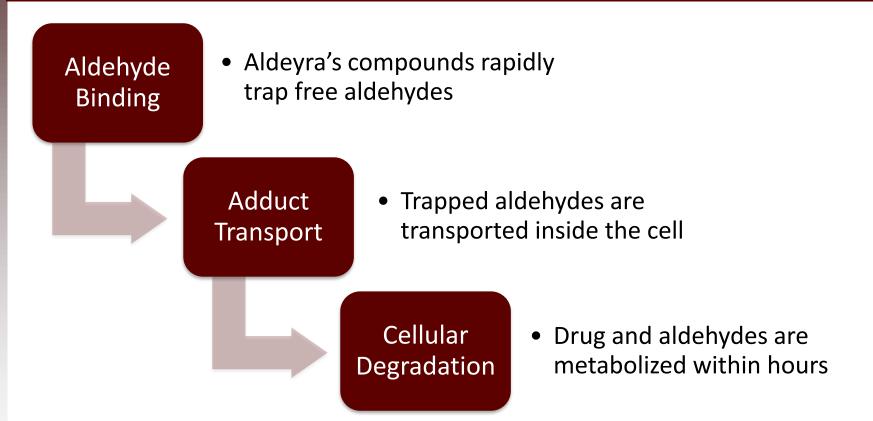
Aldehydes Are Mediators of Disease

- Directly toxic, modify cellular constituents, and pro-inflammatory
- Metabolized by enzymes called aldehyde dehydrogenases
- High levels are implicated in inflammatory diseases and in inborn errors of metabolism with genetic mutations in aldehyderelated enzymes

Aldehydes **Protein Signaling** via Thiol and Cytokine Release **Amine Binding** Inflammasome, NF-KB Activation



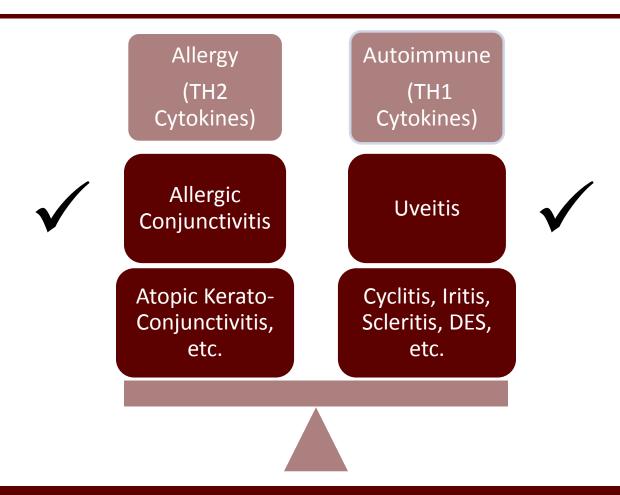
Aldehyde Traps: A Novel Therapeutic Approach



Aldeyra is not aware of any similar technology, and is developing a series of novel aldehyde traps.



The Aldehyde Trap Ocular Anti-Inflammatory Platform



ADX-102 (previously NS2) has demonstrated activity in the two major types of inflammation, and thus has potential efficacy in a wide variety of inflammatory disease.



Noninfectious Anterior Uveitis Phase 2 Clinical Design

Dosing	ADX-102 0.5% Topical Ocular Pred Forte® 1% Topical Ocular		
Randomized	 Active-Controlled 1:1:1 ADX-102 QID, Pred Forte[®] QID Taper, ADX-102 QID + Pred Forte[®] BID Taper 		
Enrollment	45 Patients with Active Disease		
Treatment Time	6 Weeks		
Endpoints	Cell Count, Flare, Symptoms		



ADX-102 Comparable to Corticosteroid in NAU Phase 2

	ADX-102 (n=15)	Pred Forte (n=13)	ADX-102 + Pred Forte (n=16)
Week 2 Cell Grade 0	5 (33%)	4 (31%)	5 (31%)
Week 8 Cell Grade 0	7 (47%)	6 (46%)	7 (44%)
≥ 1 Cell Grade Reduction	8 (53%)	6 (46%)	8 (50%)
Rescue Medication Required	3 (20%)	5 (38%)	4 (25%)

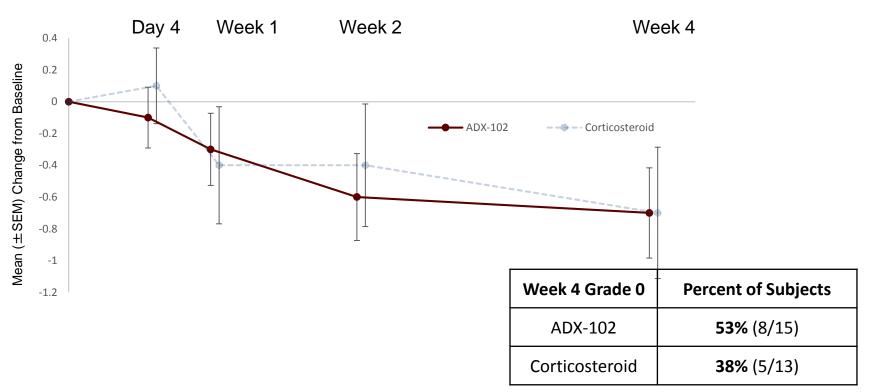
- Grade 0 = cell count of zero or one in anterior chamber
- Patients were rescued at investigator discretion if no improvement or worsening of cell count



ADX-102 Reduced Inflammation in NAU Phase 2

Change from Baseline in Anterior Chamber Cell Grade over Time

(ITT Population, Last Observation Carried Forward)

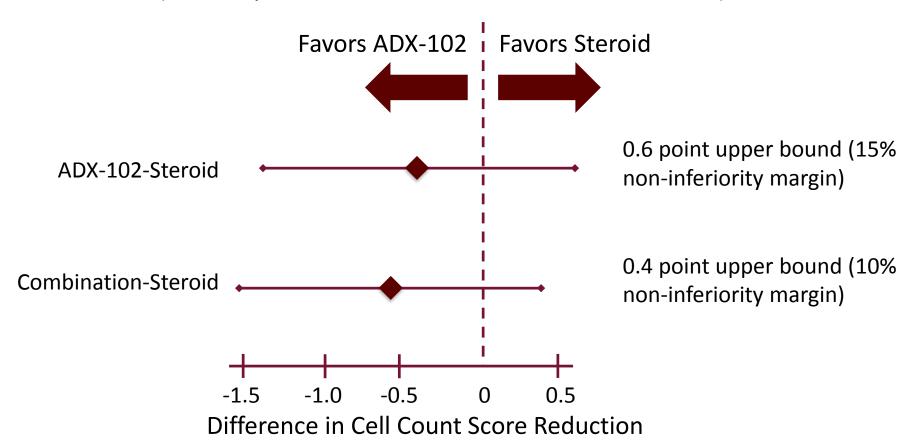


ADX-102 monotherapy effective in the treatment of noninfectious anterior uveitis, with an efficacy profile similar to corticosteroid monotherapy in a Phase 2 clinical trial

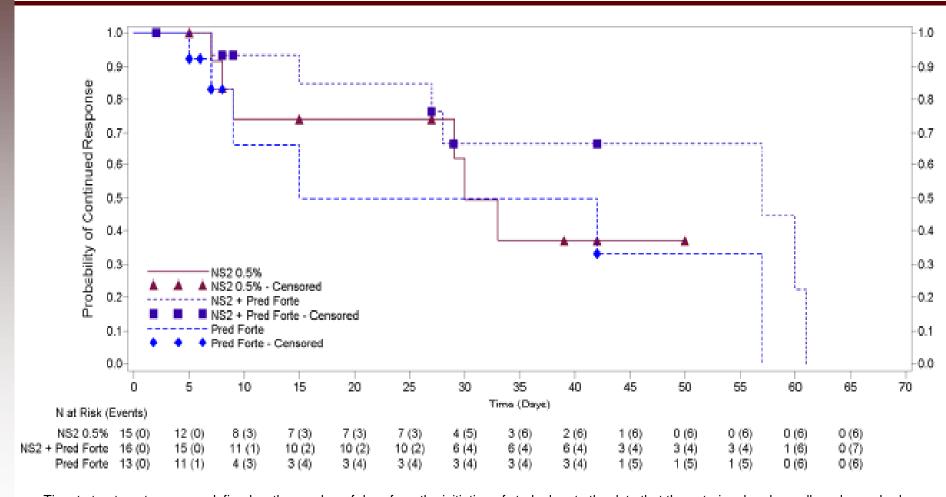


ADX-102 Comparable to Corticosteroid in NAU Phase 2

Baseline-Adjusted 95% Confidence Interval of Difference in Means at Week 2 (mITT Population, Last Observation Carried Forward)*



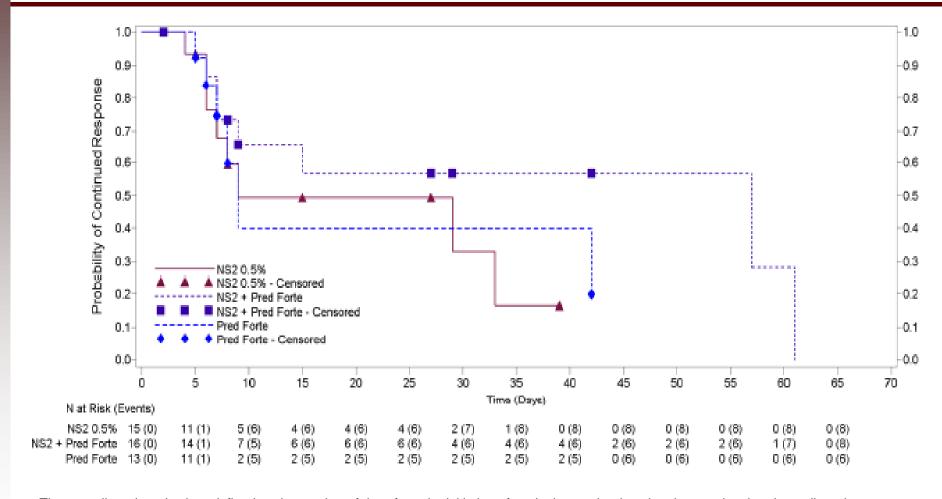
ADX-102 Sustained Grade 0 Responses THERAPEUTICS** Time to ACC Treatment Success



Time to treatment success--defined as the number of days from the initiation of study drug to the date that the anterior chamber cell grade reached and sustained a grade of 0--is estimated using the method of Kaplan-Meier. Subjects who do not experience treatment success are censored at the date of discontinuation from treatment, final office visit, or rescue. Differences between treatment groups and the Pred Forte group are assessed using a log-rank test (K-M). Log-rank p value 0.759 ADX-102 vs Pred Forte



ADX-102 Time to ACC Grade Reduction

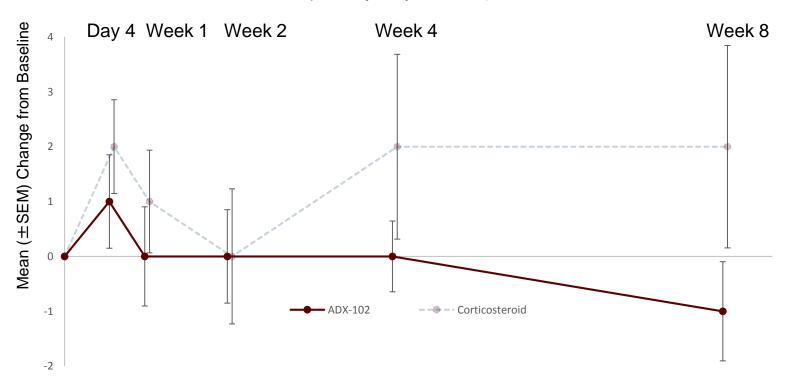


Time to cell grade reduction--defined as the number of days from the initiation of study drug to the date that the anterior chamber cell grade reached a grade of at least one less than baseline and did not increase thereafter-- is estimated using the method of Kaplan-Meier. Subjects who do not experience cell grade reduction are censored at the date of discontinuation from treatment, final office visit, or rescue. Differences between treatment groups and the Pred Forte group are assessed using a log-rank test (K-M). Log-rank p value 0.650 ADX-102 vs Pred Forte



ADX-102 Did Not Increase Intraocular Pressure in this 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 ADX-102.



Study Efficacy Summary

- No statistically significant differences* for anterior chamber cell count or flare were observed between groups in:
 - Time to sustained grade of 0
 - Proportion of subjects with sustained grade 0
 - Time to sustained reduction of ≥1-point grade
 - Subject proportion with sustained ≥1-point grade reduction
- Post hoc inference testing showed that the Least Square mean change from Baseline in ACC grade for the ADX-102 and combination treatment groups was consistently greater than the Pred Forte group
- ADX-102 Comparable to Corticosteroid in Noninfectious Anterior Uveitis Phase II Study *Trial not statistically powered. 14



Study Safety Summary

- Overall, safety findings indicate ADX-102 0.5% ophthalmic solution was well tolerated over a 6-week treatment duration
- No safety issues anticipated for future studies of topical ADX-102 in subjects with anterior uveitis:
 - No SAEs during study
 - Most TEAEs were mild or moderate
 - TEAEs most commonly related to ocular irritation
- Aligned with the previous clinical studies, no IOP increases were identified with ADX-102 0.5% in this study
- ADX-102 Generally Well-Tolerated in Noninfectious Anterior Uveitis Patients



Conclusions

- These results suggest that ADX-102 treatment alone, or in combination with 1% Pred Forte®, was effective in the treatment of Noninfectious Anterior Uveitis
- ADX-102 efficacy profile is similar to that of 1% Pred Forte® monotherapy in this clinical trial
- ADX-102 Ophthalmic Solution is currently being studied in a Phase 3 study in Noninfectious Anterior Uveitis



Conclusions

 Aldehyde Trap Strategies May Prove Clinically Useful in Treating Inflammatory Ocular Disease