



Corporate Presentation

October 2022

Forward Looking Statements

This presentation and various remarks we make during this presentation contain forward-looking statements of Astria Therapeutics, Inc. ("Astria," the "Company," "we", "our" or "us") within the meaning of applicable securities laws and regulations, including statements with respect to: the Company's projected cash runway; expectations regarding the nature, timing and potential significance of the preliminary results from the Phase 1a STAR-0215 trial; the planned timing of initiation of a Phase 1b/2 clinical trial of STAR-0215; the potential Phase 3 development plans for STAR-0215; the potential attributes and differentiated profile of STAR-0215 as a treatment for HAE, including its potential half-life and those suggested by preclinical and pharmacokinetic modeling data; the potential commercial opportunity for STAR-0215 in HAE, including its potential to be a best-in-class and most patient friendly treatment option for HAE; the need for effective treatments for HAE; the planned biomarker assay for STAR-0215; the size and anticipated growth of the HAE market; the expected patent protection of patents directed at STAR-0215; and the Company's goal to meet the unmet needs of patients with rare and niche allergic and immunological diseases, and expand its pipeline. We use words such as "aims," "anticipate," "believe," "estimate," "expect," "goals," "hope," "intend," "may," "opportunity," "plan," "predict," "project," "target," "potential," "would," "vision," "can," "could," "should," "continue," and other words and terms of similar meaning to help identify forward-looking statements, although not all forward-looking statements contain these identifying words. Actual results may differ materially from those indicated by such forward-looking statements as a result of various important factors, including risks and uncertainties related to: changes in applicable laws or regulations; the possibility that we may be adversely affected by other economic, business, and/or competitive factors, including the COVID-19 pandemic; risks inherent in pharmaceutical research and development, such as: adverse results in our drug discovery, preclinical and clinical development activities, the risk that the results of pre-clinical studies may not be replicated in clinical studies and that the results of early stage clinical studies may not be replicated in later stage clinical studies, the risk that we may not be able to enroll sufficient patients in our clinical trials on a timely basis, and the risk that any of our clinical trials may not commence, continue or be completed on time, or at all; decisions made by, and feedback received from, the U.S. FDA ("FDA") and other regulatory authorities on our regulatory and clinical trial submissions and other feedback from investigational review boards at clinical trial sites and other review bodies with respect to STAR-0215 and any other future development candidates; our ability to manufacture sufficient quantities of drug substance and drug product for STAR-0215 and any other future product candidates on a cost-effective and timely basis; our ability to develop biomarker and other assays, along with the testing protocols therefore; our ability to obtain, maintain and enforce intellectual property rights for STAR-0215 and any other future product candidates; our potential dependence on collaboration partners; competition with respect to STAR-0215 or any of our other future product candidates; the risk that survey results and market research may not be accurate predictors of the commercial landscape for HAE and the anticipated position and attributes of STAR-0215 in HAE based on its pre-clinical profile, pharmacokinetic modeling and other data; our ability to manage our cash usage and the possibility of unexpected cash expenditures; our ability to obtain necessary financing to conduct our planned activities and to manage unplanned cash requirements; the risks and uncertainties related to our ability to recognize the benefits of any additional acquisitions, licenses or similar transactions; and general economic and market conditions; as well as the risks and uncertainties discussed in the "Risk Factors" section of our Annual Report on Form 10-K for the period ended December 31, 2021, subsequent Quarterly Reports on Form 10-Q, and in other filings that we may make with the Securities and Exchange Commission. These forward-looking statements should not be relied upon as representing our view as of any date subsequent to the date of this presentation, and we expressly disclaim any obligation to update any forward-looking statements, whether as a result of new information, future events or otherwise, except as required by law,

This presentation contains estimates and other statistical data made by independent parties and by us relating to market size and other data about our industry. This data involves a number of assumptions and limitations, and you are cautioned not to give undue weight to such data and estimates. In addition, projections, assumptions and estimates of our future performance and the future performance of the markets in which we operate are necessarily subject to a high degree of uncertainty and risk.



Investment Highlights

Ô	Astria (Nasdaq: ATXS) is developing differentiated therapeutics for patients with rare and niche allergic and immunological diseases
*	 Our lead program, STAR-0215, is a monoclonal antibody inhibitor of plasma kallikrein for the preventative treatment of Hereditary Angioedema (HAE) Our goal is to provide the most patient-friendly preventative treatment for HAE with dosing once every 3 months or longer HAE market is large and growing, expected to reach \$4.5B by 2027^{1,2}
	STAR-0215 key initial proof of concept results expected by year-end 2022
E	Evaluating opportunities to expand our pipeline in allergic and immunological diseases
\$	Cash, cash equivalents and short-term investments of \$102.5M ³ with expected cash runway through 2023 based on current operating plan



Hereditary Angioedema: A Rare, Disfiguring, and Potentially Life-Threatening Disease

Rare genetic disorder charactered by severe, unpredictable, sometimes **life-threatening** swelling¹

Affects **<8,000 in the U.S. and <15,000 in the EU**,² average age of onset is 11 years old³

Standard of care has evolved to both **on-demand** and **preventative treatments**

Zuraw BL. N Engl J Med. 2008;359:1027-36.
 Lumry WR. Front Med. 2018: doi:10.3389/fmed.2018.00022.

Bork K, et al. Am J Med. 2006;119;267-274. Images obtained by haeimages.com

STAR-0215 Has the Opportunity to Change the Way That People Live With HAE

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PROFILE

STAR-0215

is a monoclonal antibody inhibitor of plasma kallikrein designed to provide long-acting, effective attack prevention for HAE with dosing once every three months or longer

YEAR END RESULTS

Preliminary clinical results expected to inform the profile of STAR-0215 to prevent HAE attacks

• Expected results in healthy subjects include safety and tolerability, PK, and PD results

COMMERCIAL OPPORTUNITY

STAR-0215 has the potential to significantly reduce treatment burden for patients

• The HAE global treatment market is substantial and growing, estimated to be \$4.5B in 2027

• Patients and physicians are highly interested in STAR-0215's target efficacy and dosing frequency



Global HAE Treatment Market is Substantial and Growing

The HAE market is expected to double by 2027^{1,2}, driven by:

- Patients being diagnosed earlier³
- More patients taking preventative treatments⁴
- Geographic expansion for currently available therapies⁵





Company-reported sales (Takeda, CSL Behring, Pharming, BioCryst)
 Analyst consensus forecasts compiled by Clarivate's Cortellis, Astria company research and analysis.
 Zanichelli A. Clin Transl Allergy. 2018: doi: 10.1186/s13601-018-0229-4

4. Astria company research and analysis

5. Company-reported expectations (Takeda, CSL Behring, BioCryst)

Approved Preventative HAE Treatments in the U.S.

Need for Effective Preventative Therapy with Lower Treatment Burden

Product	Mechanism of Action	Administration	Mean Attack Reduction ¹	% of Attack- Free Patients
CINRYZE	Plasma derived C1-INH	2x/week	52%	40% (16 weeks) ²
HAEGARDA	Plasma derived C1-INH	2x/week	88%	18% (12 weeks) ³
TAKHZYRO (lanadelumab)	Plasma kallikrein inhibitor	1-2x/month	73-87%	31-44% (26 weeks) ⁴
ORLADEYO (berotralstat)	Plasma kallikrein inhibitor	1x/day 🖶	30-44%	2-8% (24 weeks) ⁵

- Plasma kallikrein inhibition is the market leading validated mechanism of action
 - Established PK-PD-efficacy relationship for inhibiting plasma kallikrein and preventing HAE attacks
- Established regulatory and clinical path for HAE
- Opportunity for early clinical PoC with plasma kallikrein inhibition



Opportunity to Improve HAE Treatment and Reduce Burden on Patients

TAKHZYRO® (lanadelumab-flyo)

is a plasma kallikrein mAb approved for prevention of HAE attacks1

Mean Monthly attack rate



Indicated for dosing every 2 weeks; every 4 weeks may be considered in some patients

% of attack-free patients¹ (for 26 weeks)



TAKHZYRO is the current global market leader¹

- Takeda reported nearly \$1B in fiscal year 2021 sales³
- Shire acquired Dyax for \$5.9B after Phase 1b with lead program TAKHZYRO⁴

56-69% of patients experienced attacks on TAKHZYRO²

Published unmet need for improved HAE treatments^{5, 6}

 Despite preventative treatments, patients continue to have attacks and high rates of anxiety and depression



- 1. Takeda FY2021 Q4 Earnings Announcement, May 2022.
- TAKHZYRO Prescribing Information, 2018.
 Takeda 2021 Fiscal Year Financial Report, May 2022.
- Shire plc and Dyax Corp. Press Release. 2015, Nov.

- Banerji A, et al. Ann Allergy Asthma Immunol. 2020; 124: 600-607. doi: 10.1016/j.anai.2020.02.018.
- Riedl MA., et al. Ann Allergy Asthma Immunol. 2021; 126: 264-272. doi: 10.1016/j.anai.2020.10.009.

STAR-0215's Target Efficacy and Dosing is Compelling to Interviewed HAE Patients and Treatment Providers



- · On average, patients tried 2-3 preventative treatments, most often switching for more convenient administration
- All interviewed patients would be compelled to switch from their current therapy if a new therapy offered similar efficacy with less frequent dosing
- Most prescribers (n=13) would discuss a product with STAR-0215's target profile with all HAE patients, including those using on-demand therapy only



STAR-0215 Potential for Best-in-Class Profile in HAE



Astria wholly owns an international patent application directed to STAR-0215. If nationalized in the U.S. and granted, the patent would expire in 2042, excluding any potential patent term extension¹



STAR-0215 Shows High Potency Inhibition of Plasma Kallikrein

- STAR-0215 binding affinity for plasma kallikrein is ~10-fold greater than lanadelumab
- **STAR-0215** binds a different site on plasma kallikrein than lanadelumab
- **STAR-0215** is more potent at inhibiting enzymatic activity than lanadelumab



STAR-0215 was more potent than lanadelumab in inhibiting bradykinin production in an *in vitro* assay



YTE Fc Modifications Have Led to Substantial Half-Lives of Monoclonal Antibodies in Humans

- Introduction of YTE into the anti-RSV mAb, motavizumab, prolonged half-life ~3.5-fold in both NHP and humans
- The approved YTE antibodies have half-lives of 83-88 days in humans
- Across a range of YTE Fc modified mAbs against non-cellular targets that are not subject to target mediated drug disposition (TMDD), the half-life is ~80-90 days in humans
- For targets affected by TMDD (e.g. KIT) the half-life is extended by YTE Fc modification is 2-4-fold but is shorter than 80 days (30 – 40 days)

Antibody	Target	NHP T _{1/2} (Days)	Human T _{1/2} (Days)
Motavizumab	RSV	6	24
Motavizumab-YTE	RSV	21	82
Tixagevimab-YTE / Cilgavimab-YTE (Evusheld)	SARS-CoV-2	~19 ~19	88 83



1. Dall'Acqua et al. J Biol Chem.2006 Aug 18;281(33):23514-24. doi: 10.1074/jbc.M604292200. Epub 2006 Jun 21.

2. Robbie et al. J Biol Chem. 2006 Aug 18;281(33):23514-24. doi: 10.1074/jbc.M604292200. Epub 2006 Jun 21.

3. Loo et al. Sci Transl Med. 2022 Mar 9;14(635):eabl8124. doi: 10.1126/scitranslmed.abl8124. Epub 2022 Mar 9.

. Evusheld EUA Review: https://www.fda.gov/media/155107/download

STAR-0215 Has Shown Substantially Prolonged Plasma Half-Life Compared to Lanadelumab in Non-Human Primates



STAR-0215 incorporates YTE modifications to extend half-life

Mean non-	Lanadelumab	STAR-	STAR-
human primate		0213	0215
half-life in days (SD)	10.5 (1.6)	10.9 (0.4)	33.6 (8.3)



Planned Biomarker Assay to Assess Plasma Kallikrein Activity Following STAR-0215 Dosing





HMWK = high molecular weight kininogen CHMWK = cleaved high molecular weight kininogen FXII = Factor XII FXIIa = activated Factor XII



STAR-0215 Phase 1a: Dosing is Complete

Phase 1 Healthy Subject Trial Overview

- At least three single ascending dose cohorts
 - 100 mg, 300 mg, and 600 mg
 - Healthy adult subjects
 - Subcutaneous dosing
- · Randomized, double-blind, placebo-controlled
 - 6 active to 2 placebo randomization
- Single U.S. center study

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- Observation period through multiple half-lives
- 3-month data will inform on the target profile



This protocol may be amended to add new cohorts in the future

STAR-0215 Phase 1a Trial Will Inform on Target Profile

Preliminary data expected to be available by year-end 2022

Phase 1a Endpoints

- Safety and tolerability
- Pharmacokinetics: blood concentrations
 over time
- Pharmacodynamics: inhibition of bradykinin production via inhibition of plasma kallikrein

Target Profile

- Small volume subcutaneous administration
- Durable activity
- Administered once every 3 months or less frequently
- · Safe and well-tolerated



Physiologically-Based PK Model Supports a Dosing Frequency of Every 3 Months or Longer



Model suggests target concentration of STAR-0215 required to produce long-term inhibition of plasma kallikrein can be achieved with a single dose above 30mg



Model suggests target level of STAR-0215 can be achieved with a loading dose of 300mg followed by the maintenance dose of 150mg every 3 months

3 months is approximately Day 84, arrows indicate simulated drug dosing, green dashed line is 12 μg/mL. 12 μg/mL, or 80nM, is the threshold C_{min} predicted to inhibit the production of bradykinin in HAE by pKal.



Inhibition of Plasma Kallikrein Reduces cHMWK, Correlating to Clinical Benefit in HAE

STAR-0215 May Achieve More Sustained Reductions in cHMWK Compared to Lanadelumab





Aiming to Progress STAR-0215 Quickly to Patients

Completed and Expected Upcoming Milestones



Astria (Nasdaq ATXS) Well-Positioned for the Future

STRONG FINANCIAL FOUNDATION	 As of 6/30/2022, the Company had cash, cash equivalents and short-term investments of \$102.5M with expected cash runway through 2023 based on current operating plan 		
	Company Capitalization Structure as of September 15, 2022	As Converted Common Shares	
CAPITALIZATION	Common stock outstanding	15,178,042	
STRUCTURE	Common stock underlying outstanding Series X Preferred Stock	5,242,501	
	Adjusted Common stock outstanding ¹	20,420,543	



