



Third-Quarter 2025 Earnings Presentation

October 30, 2025

Ready.

Forward Looking Statements

The forward-looking statements in this presentation are based upon the Company's current expectations and beliefs, and involve known and unknown risks, uncertainties and other factors, which may cause the Company's actual results, performance and achievements and the timing of certain events to differ materially from the results, performance, achievements or timings discussed, projected, anticipated or indicated in any forward-looking statements. Such risks, uncertainties and other factors include, among others, the following: failure to continue to successfully commercialize ARIKAYCE in the U.S., Europe or Japan or failure to successfully commercialize BRINSUPRI in the U.S., or to maintain U.S., European or Japanese approval for ARIKAYCE or U.S. approval for BRINSUPRI; our inability to obtain full approval of ARIKAYCE from the FDA, including the risk that we will not successfully or in a timely manner complete the confirmatory post-marketing clinical trial required for full approval of ARIKAYCE, or our failure to obtain regulatory approval to expand ARIKAYCE's indication to a broader patient population; failure to obtain, or delays in obtaining, regulatory approvals for our product candidates in the U.S., Europe or Japan, for ARIKAYCE outside the U.S., Europe or Japan, including separate regulatory approval for Lamira® in each market and for each usage, or for brensocatib in Europe or Japan; failure to successfully commercialize our product candidates, if approved by applicable regulatory authorities, or to maintain applicable regulatory approvals for such product candidates, if approved; uncertainties or changes in the degree of market acceptance of our marketed products or, if approved, our product candidates, by physicians, patients, third-party payors and others in the healthcare community; our inability to obtain and maintain adequate reimbursement from government or third-party payors for our marketed products or, if approved, our product candidates, or acceptable prices for our marketed products or, if approved, our product candidates; inaccuracies in our estimates of the size of the potential markets for our marketed products and our product candidates or in data we have used to identify physicians, expected rates of patient uptake, duration of expected treatment, or expected patient adherence or discontinuation rates; failure of third parties on which we are dependent to manufacture sufficient quantities of our marketed products and our product candidates for commercial or clinical needs, as applicable, to conduct our clinical trials, or to comply with our agreements or laws and regulations that impact our business; the risks and uncertainties associated with, and the perceived benefits of, our senior secured loan with certain funds managed by Pharmakon Advisors LP and our royalty financing with OrbiMed Royalty & Credit Opportunities IV, LP, including our ability to maintain compliance with the covenants in the agreements for the senior secured loan and royalty financing and the impact of the restrictions on our operations under these agreements; our inability to create or maintain an effective direct sales and marketing infrastructure or to partner with third parties that offer such an infrastructure for distribution of our marketed products or any of our product candidates that are approved in the future; failure to successfully conduct future clinical trials for our marketed products or our product candidates and our potential inability to enroll or retain sufficient patients to conduct and complete the trials or generate data necessary for regulatory approval of our product candidates or to permit the use of ARIKAYCE in the broader population of patients with MAC lung disease, among other

things; development of unexpected safety or efficacy concerns related to our marketed products or our product candidates; risks that our clinical studies will be delayed, that serious side effects will be identified during drug development, or that any protocol amendments submitted will be rejected; failure to successfully predict the time and cost of development, regulatory approval and commercialization for novel gene therapy products; risk that interim, topline or preliminary data from our clinical trials that we announce or publish from time to time may change as more patient data become available or may be interpreted differently if additional data are disclosed, or that blinded data will not be predictive of unblinded data; risk that our competitors may obtain orphan drug exclusivity for a product that is essentially the same as a product we are developing for a particular indication; our inability to attract and retain key personnel or to effectively manage our growth; our inability to successfully integrate our acquisitions and appropriately manage the amount of management's time and attention devoted to integration activities; risks that our acquired technologies, products and product candidates will not be commercially successful; inability to adapt to our highly competitive and changing environment; inability to access, upgrade or expand our technology systems or difficulties in updating our existing technology or developing or implementing new technology; risk that we are unable to maintain our significant customers; risk that government healthcare reform materially increases our costs and damages our financial condition; business or economic disruptions due to catastrophes or other events, including natural disasters or public health crises; risk that our current and potential future use of AI and machine learning may not be successful; deterioration in general economic conditions in the U.S., Europe, Japan and globally, including the effect of prolonged periods of inflation, affecting us, our suppliers, third-party service providers and potential partners; the risk that we could become involved in costly intellectual property disputes, be unable to adequately protect our intellectual property rights or prevent disclosure of our trade secrets and other proprietary information, and incur costs associated with litigation or other proceedings related to such matters; restrictions or other obligations imposed on us by agreements related to our marketed products or our product candidates, including our license agreements with PARI and AstraZeneca AB, and failure to comply with our obligations under such agreements; the cost and potential reputational damage resulting from litigation to which we are or may become a party, including product liability claims; risk that our operations are subject to a material disruption in the event of a cybersecurity attack or issue; our limited experience operating internationally; changes in laws and regulations applicable to our business, including any pricing reform and laws that impact our ability to utilize certain third parties in the research, development or manufacture of our product candidates, and failure to comply with such laws and regulations; our history of operating losses, and the possibility that we never achieve or maintain profitability; goodwill impairment charges affecting our results of operations and financial condition; inability to repay our existing indebtedness and uncertainties with respect to our ability to access future capital; and delays in the execution of plans to build out an additional third-party manufacturing facility approved by the appropriate regulatory authorities and unexpected expenses associated with those plans.

Additional Disclaimers: Please be aware that TPIP, INS1201, INS1202, and INS1033 are investigational products that have not been approved for sale or found safe or effective by the FDA or any regulatory authority. In addition, ARIKAYCE has not been approved for the treatment of all patients with MAC lung disease and brensocatib has not been approved for the treatment of patients with non-cystic fibrosis bronchiectasis outside the U.S. This presentation is not promotion or advertisement of ARIKAYCE, BRINSUPRI, TPIP, INS1201, INS1202, or INS1033. Insmed, ARIKAYCE and BRINSUPRI are trademarks of Insmed Incorporated. All other trademarks are property of their respective owner(s).

Speakers



Will Lewis
Chair & CEO



Roger Adsett
Chief Operating Officer



Sara Bonstein
Chief Financial Officer



Agenda

1 Opening Remarks

2 Clinical Updates

3 Early-Stage Updates

4 Commercial Updates

5 Financial Results

Slides

4-6

7-14

15-17

19-24

25-28

Opening Remarks

Will Lewis | *Chair & CEO*

Third-Quarter Highlights

- Received U.S. approval and launched **BRINSUPRI™**
- **Positive reception** to BRINSUPRI supports potential for **large commercial opportunity**
- Expect first **full quarter** of BRINSUPRI sales to illustrate clearer picture of **underlying dynamics**
- Ambition for BRINSUPRI sales to perform in-line with **historically strong** respiratory launches



U.S.: United States



Next 18 Months Poised to Deliver Multiple Commercial and Clinical Catalysts

Commercial Catalysts:

Brinsupri™
U.S. Approval
& Launch

TODAY

Brinsupri™
EU, UK and Japan
Approval & Launch*

ARIKAYCE®
Label Expansion*

...Previous 18 months

Next 18+ months...

Clinical Catalysts:

PH-ILD

Ph2 Readout

PAH

Ph2 Readout

ASPEN

Ph3 Readout

PALM-ILD

Ph3 Kickoff

PAH

Ph3 Kickoff

Next-Gen DPP1s

Enter Clinic

BiRCh

Ph2 Readout

PPF

Ph3 Kickoff

ALS

Updates

ENCORE

Ph3 Readout

IPF

Ph3 Kickoff

Stargardt

Updates

CEDAR

Ph2 Readout

DMD

Updates

1-2 INDs per year

* Pending regulatory approval | Ph: Phase | U.S.: United States | EU: Europe | UK: United Kingdom | PAH: pulmonary arterial hypertension | DMD: Duchenne muscular dystrophy | ALS: amyotrophic lateral sclerosis | DPP1: dipeptidyl peptidase 1 inhibitor | Next-Gen: next-generation | IND: Investigational new drug application

Brensocatib Updates

Will Lewis | *Chair & CEO*

International Filings for Brensocatib Are Progressing On Track

Bronchiectasis

Europe

- CHMP **positive opinion** recommending approval
- On track for **EMA decision** in next several months
- Approval could **expand** TAM by **~600K¹ patients***
- Expect to launch at **same list price as U.S.***

UK & Japan

- **UK** filing progressing **on schedule**
(decision expected **1H:26**)
- **Japan** filing accepted (decision expected **2H:26**)



If approved,
we anticipate that
each of **these filings**
could unlock
additional **growth**
opportunities
in 2026*

* If approved | EMA: European Medicines Agency | NCFB / Bronchiectasis: refers to non-cystic fibrosis bronchiectasis | UK: United Kingdom | U.S.: United States | H: half | TAM: total addressable market | ¹ 600K patients estimated in the EU₅, comprised of France, Germany, Italy, Spain and the United Kingdom

BiRCh: On Schedule For Topline Readout by Early January 2026

CRSsNP

Key Trial Details

- **Primary:** Change in baseline daily sinus Total Symptom Score (sTSS)
- A treatment effect would be **additive** to effect of stable dose of **background nasal steroids**
- **BiRCh Powering:** alpha of **0.10**
 - ✓ **80%** to show **0.97-pt** improvement in sTSS
 - ✓ **90%** to show **1.14-pt** improvement in sTSS

Defining Success



Clear Win: sTSS improvement matches **historically approved** level (0.7)*



Homerun: sTSS improvement meets prespecified **BiRCh powering**



Blended-blinded data continue to look encouraging**

CEDAR: CEDAR: Enrollment Completed Ahead of Schedule; Eliminates Need for Interim Futility Analysis

HS

Key Trial Details

- **Primary:** Percent reduction in abscess and inflammatory nodule (AN) count vs. baseline
- **CEDAR Powering:** alpha of **0.10**
 - ✓ **95%** to show **40%** reduction in AN count
- **214 patients** exceeded enrollment goal
- Topline readout now **expected 1H:26**

Defining Success



Clear Win: ~20% reduction in AN count vs. baseline



Homerun: reduction in-line with prespecified **CEDAR powering**

We remain hopeful that **brensocatib** will benefit HS patients who have limited safe & effective oral therapy offerings

TPIP Updates

Will Lewis | *Chair & CEO*



TPIP: Treprostinil Palmitil Inhalation Powder

TPIP: Focused On Designing and Conducting An Expansive Registrational Program

1 PH-ILD

2 PAH

3 PPF

4 IPF

DMC recommends to **continue Phase 2 OLEs** and advance into Phase 3

- **PALM-ILD**: Expect first site open by **YE:25**
- **Primary Endpoint**: Change in baseline **6MWD** measured at **peak** exposure¹
- **Key Secondary**: 6MWD at **trough**², NT-proBNP, PK

- **EOP2**: Aligned with FDA on a **single Phase 3 trial** required to support future submission
- **Phase 3** initiation expected **early 2026**
- **Primary Endpoint**: Change in baseline **6MWD** at alpha **0.05**

- Expect **Phase 3 studies** to initiate in **2H:26**
- Increasing **TPIP manufacturing** to adequately resource these studies

¹ Primary endpoint; placebo-adjusted metric measured 1-3 hours post-dose at Week 24 | ² Secondary endpoint; placebo-adjusted metric measured pre-dose at Week 22 | alpha (α): significance level representing the probability of concluding a treatment is effective when it is not | TPIP: Treprostinil Palmitil Inhalation Powder | PH-ILD: pulmonary hypertension due to interstitial lung disease | PAH: pulmonary arterial hypertension | 6MWD: six-minute walk distance | DMC: data monitoring committee | EOP2: End of Phase 2 meeting with the Food and Drug Administration (FDA) | NT-proBNP: N-terminal pro b-type natriuretic peptide; a biomarker of cardiac stress | PK: pharmacokinetics | H: half | YE: year-end

ARIKAYCE® Clinical Updates

Will Lewis | *Chair & CEO*

ENCORE: Progressing On Schedule Toward 1H:26 Readout

MAC LD

Blockbuster Potential*

- ENCORE could expand ARIKAYCE's label in the **U.S. & Japan** to address an **additional 100K+ patients in each***

Primary Endpoint

- **U.S.:** Change from baseline respiratory symptom score at Month 13
- **Japan:** Proportion of patients achieving durable culture conversion at Month 15

Takeaways From ARISE¹

- ✓ ARIKAYCE patients showed **faster, greater, and more durable** culture conversion²
- ✓ Highlighted benefit of **early treatment**³
- ✓ Validated **PRO tool**

ENCORE study has comparable **recruitment timing** and patient **demographics** to **ARISE**

Early-Stage Updates

Will Lewis | *Chair & CEO*

ASCEND: First Gene Therapy Program Completes Cohort 1 Dosing



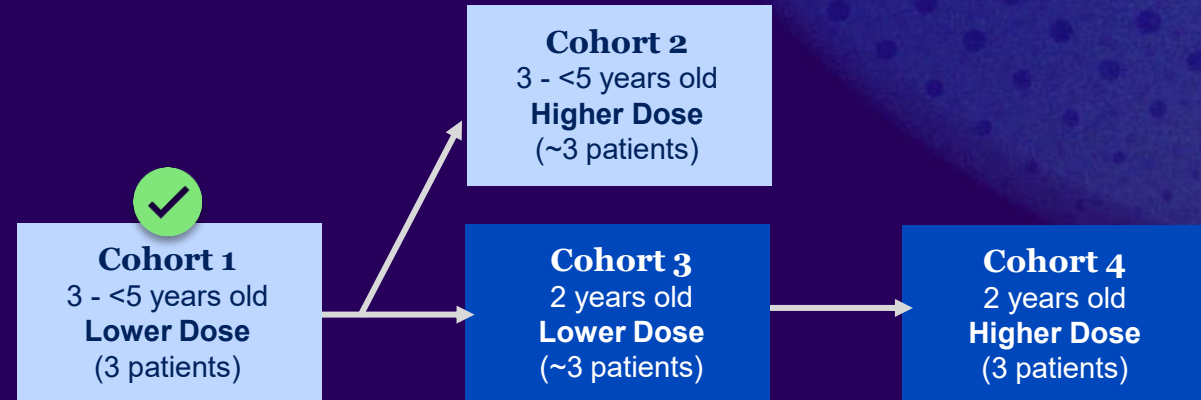
- **No concerning safety signals observed to date**

- Intrathecally delivered
- Dosing not **weight-based**

- **Advancing to next stage of study**

- *Cohort 2*: boys aged 3 to <5 years at higher dose
- *Cohort 3*: boys aged 2 years at lower dose

Screening



Anticipate Second Gene Therapy and First Next-Gen DPP1 to Enter Clinic in 2026



- **IND cleared to move into clinic**
 - Intrathecally delivered
- **Studying both SOD1 and sporadic ALS**
 - Strong **pre-clinical data** in both models



- Expect first DPP1 candidate (INS1033) to enter clinic in 2026 targeting **RA & IBD**
- Future candidates expected to target additional indications, including **COPD**

Expect more updates from the Early-Stage portfolio in 2026

Let's Recap

1

Recent achievements have **unlocked additional opportunities** to deliver on behalf of patients

2

Well-positioned to deliver innovative, **first- or best-in-class** therapies to patients with serious diseases

3

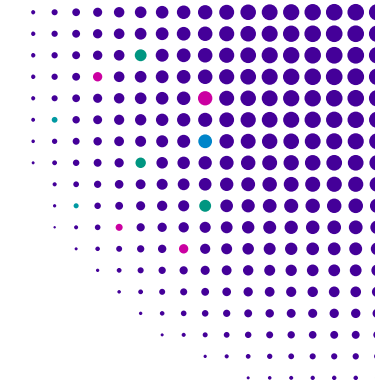
Recipient of **Science's Top Employer** recognition for **fifth consecutive** year



Commercial Updates: BRINSUPRI™ & ARIKAYCE®

Roger Adsett | *Chief Operating Officer*

Demonstrating Strength in First Partial-Quarter



Period Ended
September 30, 2025



~2,550

patients started treatment

~1,700

physicians wrote ≥ 1 script²

- Includes COEs & community centers
- Majority have written 1-2 scripts each

Expect Greater Insight into Underlying Launch Dynamics Next Quarter

Drivers to consider in Q4...

Path to Prescription

- BRINSUPRI now available in all **EMR** systems
- More **standardized** prior authorization criteria across payors
- Expect both to enable **broader** prescriber activation and improve **back-office** workflow

U.S. Guideline Updates

- Preliminary **U.S. guidelines** recommend BRINSUPRI for exacerbating NCFB patients
- We hope final guidelines will reinforce physician **treatment decisions** and influence payor **coverage criteria***



Today's Results Are Promising But Do Not Represent Final Contracting & Access Dynamics

When interpreting Q3 results...

Encouraging Signs

- ✓ **Broad access** from 'Day 1' as only approved therapy for NCFB
- ✓ **Vast majority** of prescriptions have been **approved** for coverage¹
- ✓ **Continue to engage** with payors to establish **frictionless** access requirements

Reasons for Caution

- ⚠ Payor discussions are **still ongoing**
- ⚠ Slowdowns often occur once **final market access** criteria are established and **enforced**



Remain Focused On Exceptional Future Launch Execution



Q3 benefited from **more weeks** of **revenue** and **patient starts** than initially expected



Inventory stocking accounted for ~40% of Q3 sales



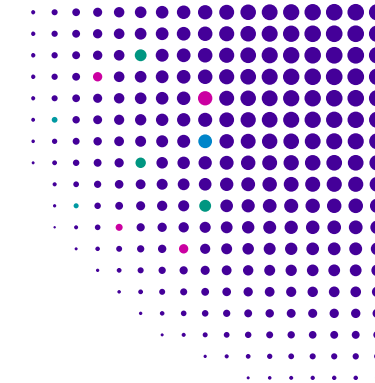
Don't anticipate stocking contribution to be as impactful in Q4



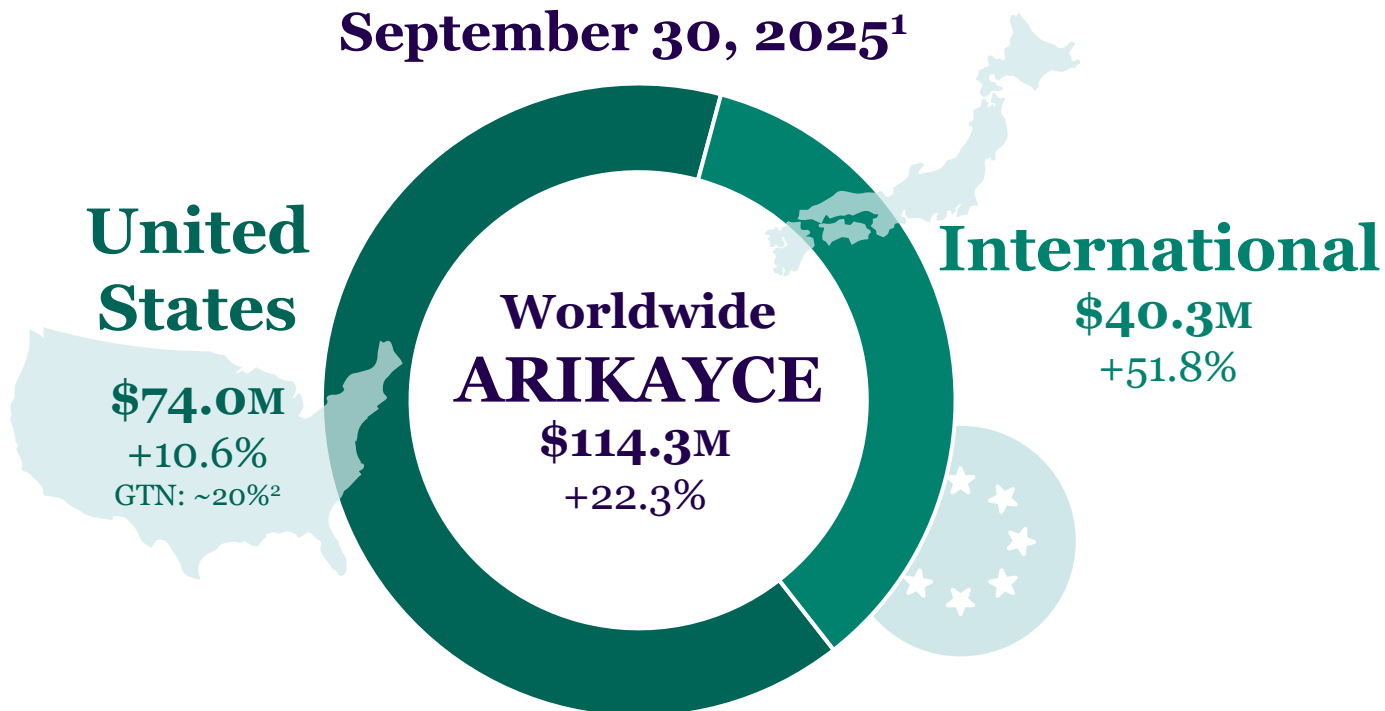
BrinsupriTM
(brensocatib)

While early days
fuel our **excitement**,
work remains to
achieve our ambition
of positioning
BRINSUPRI among
other high-performing
respiratory launches

ARIKAYCE Demonstrates Continued Strength



Three Months Ended
September 30, 2025¹



- **Record quarterly revenue** in the U.S. and internationally
- **Strength in U.S.** demonstrates commercial ability to execute across **multiple products**
- **International** business continues impressive growth

Financial Results

Sara Bonstein | *Chief Financial Officer*

Increasing 2025 ARIKAYCE Revenue Guidance

ARIKAYCE Revenue Guidance

\$420 to \$430M

Full-Year 2025

+15% to 18%*

2025 vs. 2024

* 2025 ARIKAYCE revenue guidance vs. 2024 ARIKAYCE revenue

Strong Capital Position Ahead of Upcoming Catalysts

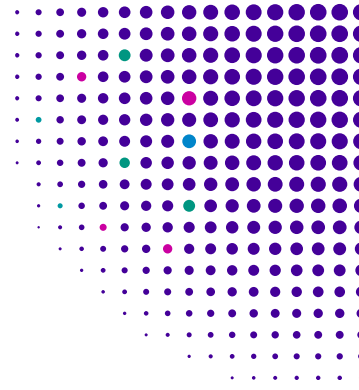
September 30, 2025[†]

~\$1.7B

*In Cash, Cash Equivalents,
and Marketable Securities*

- Underlying cash burn¹ was **relatively consistent** with **historical** burn levels
- Only a **portion** of BRINSUPRI revenues were recognized in cash this quarter²
- Expect **spending** to increase alongside **revenue** as we resource launches and clinical programs³

Advancing Commercial and Clinical Programs Through Investment



	Three Months Ended ¹	
(in \$ millions, except for percentages)	9/30/2025	9/30/2024
Total Revenues	\$142.3	\$93.4
Cost of Product Revenues ²	(29.4)	(21.2)
<i>As a % of Revenues</i>	<i>20.6%</i>	<i>22.7%</i>
R&D	(186.4)	(150.8)
SG&A	(186.4)	(118.9)
Other [†]	(106.2)	(15.9)
Total Operating Expenses	\$(508.3)	\$(306.9)
Operating Loss	\$(366.0)	\$(213.4)

- Revenues² as % of sales **decreased** y/y, reflecting BRINSUPRI contribution
- R&D and SG&A **increased** y/y reflecting **investments in growth**:
 - U.S. **BRINSUPRI** launch
 - **Clinical pipeline** development

Closing Remarks

-  **1 Past 18 Months:** Clinical and commercial achievements created **new opportunities** to serve patients
-  **2 Next 18+ Months:** Plan to invest behind multiple **commercial & clinical** catalysts with the potential to change patient lives
-  **3** We believe we are in a **strong financial position** to execute on these opportunities

Ready.

Q&A Session



Will Lewis
Chair & CEO



Roger Adsett
Chief Operating Officer



Dr. Martina Flammer
Chief Medical Officer



Sara Bonstein
Chief Financial Officer



Count.
us in.

