

# **Corporate Presentation**

**December 2024** 



#### **Forward-Looking Statements**

This presentation contains "forward-looking statements" within the meaning of the Private Securities Litigation Reform Act of 1995, as amended (the "Act"). Forward-looking statements are statements that are not of historical fact and include, without limitation, statements regarding future events, our future financial condition, results of operations, business strategy and plans, prospective products (as well as their potential to achieve a differentiated, competitive, or favorable benefit or profile or trend, including on safety, tolerability, improvement, maintenance, clinical response, dosing, efficacy and/or convenience), planned or expected product approval applications or approvals, anticipated milestones, expected data readouts and enrollments, research and development plans and costs, potential future partnerships, expectations about existing partnerships, timing and likelihood of success, objectives of management for future operations, future results of anticipated product development efforts, and adequacy of existing cash and potential partnership funding to fund operations and capital expenditure requirements, as well as statements regarding industry trends. These statements are based on management's current expectations of future events only as of the date of this presentation and are inherently subject to a number of risks, uncertainties and assumptions, some of which cannot be predicted or quantified and some of which are beyond our control, including, among other things: the ability of our clinical trials to demonstrate safety and efficacy of our product candidates and other positive results; whether we will need expanded or additional trials in order to obtain regulatory approval for our product candidates; existing regulations and regulatory developments in the U.S., the PRC, Europe and other jurisdictions; the ability of our current cash and investments position to support planned operations; our plans and ability to obtain, maintain, protect and enforce our intellectual property rights and ou

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#### **Connect is Transforming Acute & Chronic Care in Asthma and COPD**



#### Product Candidate Rademikibart

Next generation antiinterleukin-4-receptor alpha (IL-4Rα) antibody achieves rapid onset and sustained improvements in asthma

### Target to enter market in Early 2029

Potential peak annual sales >\$5.4 billion/year with acute and chronic indications for asthma and COPD

### Best-in-Class Potential

Rademikibart has demonstrated significant improvement in lung function as early as 24 hours and is well tolerated over 24 weeks of treatment in asthma and up to one year in atopic dermatitis

### IP Protection Until 2040

IP granted in the U.S. and all major EU markets

### **Untapped Market Opportunity**

~1 million Asthma Patients and ~1.3 million COPD Patients

visit the emergency department (ED) or are admitted to the hospital annually in the US alone

## Proven U.S. Management Team

Deep expertise in drug development with a history of multiple late-stage clinical and regulatory executions.

Transforming Connect into a U.S.-centric company, significantly reducing presence in China.

### Rapid Clinical Development Program

Plan to initiate two Phase 2 POC studies in 1H2025 for Rademikibart in ED/Hospital patients with acute Asthma and COPD; data expected in 1H2026

### Strong Cash Position

\$110.2 million in cash and cash equivalents as of June 30, 2024 expected to support planned operations into at least the first half of 2027



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## Rademikibart Has Demonstrated Efficacy & Safety Across Multiple Indications

Connect Biopharma has global development & commercialization rights outside of greater China for all product candidates



No further spending on icanbelimod (S1P1) or rademikibart in AD is planned by Connect

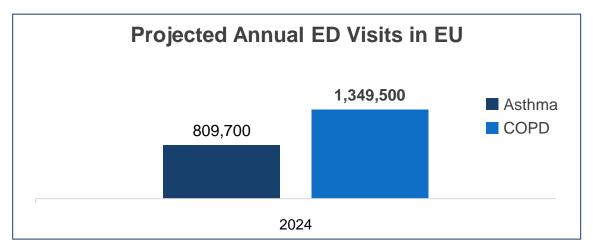


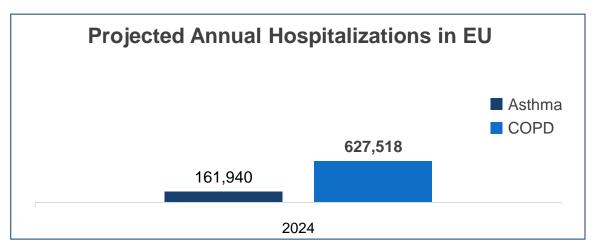
## Asthma and COPD are Associated with Severe Exacerbations that are Difficult to Treat and Often Require Hospitalization

	Asthma COPD					
CURRENT SOC	Fast-acting inhaled bronchodilators and oral/IV corticosteroids; severe cases may require IV magnesium sulfate, heliox therapy, and noninvasive ventilation; intubation is considered if patients do not respond to initial therapies. Antibiotics are often added if a bacterial infection is suspected with COPD					
PROGRESSION	~50% fail to improve on 1 <sup>st</sup> Line treatments ~85% fail to improve on 1 <sup>st</sup> Line treatment					
<b>&gt;&gt;&gt;</b>	Persistent hypoxia, severe respiratory distress, poor response to initial treatment, altered mental status and/or a history of exacerbations drive admission decisions					
	~1 million ED visits or hospitalizations/yr	~1.3 million ED visits or hospitalizations/yr				
<b>ADMISSIONS</b>	~11% of Asthma patients are hospitalized	~41% of COPD patients are hospitalized				
	LOS typically ranges from 2 to 3 days	LOS typically ranges from 4 to 7 days				
	~50% meet treatment failure criteria within 4 weeks of an exacerbation, with 20% requiring a re-visit to the ED	~50% fail treatment within 4 weeks of an exacerbation with ~11% of patients require rehospitalization				
LONG-TERM CARE	As symptoms improve, hospital physicians coordinate care with PCPs or pulmonologists; asthma patients are most likely to get treated by a PCP in the outpatient setting	COPD patients are typically older and have more comorbidities than asthma patients, and are therefore more likely to be referred to a pulmonologist upon discharge				

### Asthma and COPD are Global Problems Projected 2024 European Union Asthma & COPD ED Visits / Hospitalizations

Projected annual ED/hospital visits associated with asthma and COPD exacerbations are similar in the EU and US





#### **Key Statistics and Considerations**

- Studies report ~20% of asthma patients are hospitalized, and upwards of ~20% of patients returned to the ED within 30 days<sup>1-9</sup>
- Studies suggest the prevalence of COPD exacerbations in the EU closely mirrors that of the US<sup>10-14</sup>
- Hospitalization rates for exacerbations among COPD patients range from ~32% to 61% across the EU, with ~24% of patients returning to the ED within 30 days<sup>10-14</sup>



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### US Market Research Shows that Obtaining Acute Treatment Indications Results in Significantly Greater Penetration into Asthma and COPD Markets

- U.S. Clinicians believed the acute indication was a clear differentiator between rademikibart and other biologics approved for patients with eosinophilic phenotype
- Once patients were successfully treated with rademikibart acutely, clinicians expected to maintain 75% of these patients on rademikibart chronically
- New high-yield manufacturing process will allow for hospital-friendly pricing in the acute setting
- Obtaining acute asthma indication along with the chronic asthma indication resulted in projected U.S. peak sales forecast of \$2.3B (base case)
- Obtaining both acute COPD and chronic COPD indications resulted in projected US peak sales forecast of \$1.5B (base case)
- Upside opportunities exist for rademikibart to drive revenue even higher (e.g., steroid sparring opportunity)
- U.S. FDA approval of dupilumab for chronic treatment of COPD in September provides evidence for rademikibart's mechanism of action to treat chronic COPD
- ABRA benralizumab study presented at Sep 2024 European Respiratory Society provides a higher level
  of confidence in obtaining a positive outcome in our planned Phase 2 acute asthma and COPD studies\*



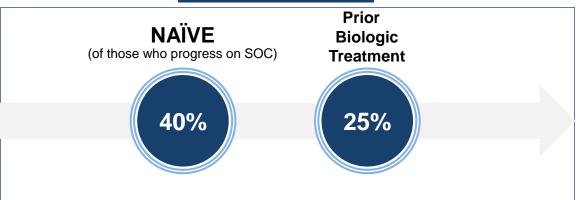
## COTE

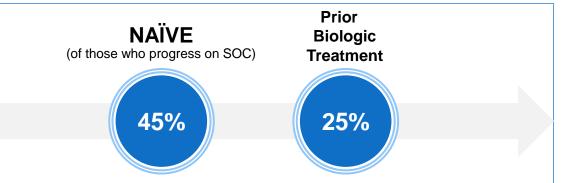
# CHRONIC

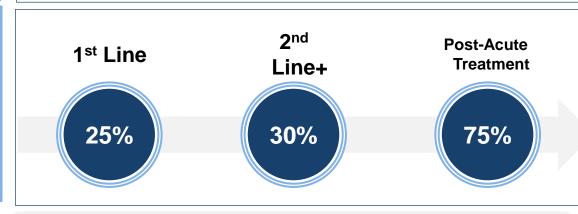
#### Rademikibart Expected Utilization in Asthma and COPD

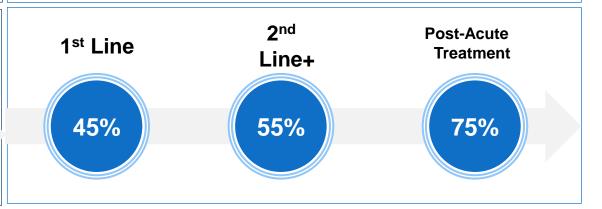
Clinicians considered Rademikibart to be differentiated from other biologics by the dual indication











"Steroids and LABAs/LAMAs work well for these acute patients, so I wouldn't really use this up front. But for those patients that are refractory to those options and need something stronger, then I'd definitely use this."

"If a patient was treated with Product X during their acute exacerbation and it worked, then I'd keep them on it for maintenance. If they responded, why would I switch?"



## There Remains Significant Unmet Need for Better Treatment Options to Improve Patient Outcomes & Reduce Burden on Healthcare Systems



### EXACERBATION RECURRENCE

Reduction in overall frequency of ED visits

 Management of severe exacerbations and the prevention of hospital readmissions remain challenging in both asthma and COPD



### HOSPITALIZATION AND RE-ADMISSION RATES

Decrease the Proportion of Patients Admitted

 >100,000 asthma patients and >500,000 COPD patients are hospitalized upon presentation to the ED with an exacerbation in the US annually



### BIOLOGICS / ADVANCED THERAPIES

Deliver Improved Efficacy
Over SoC

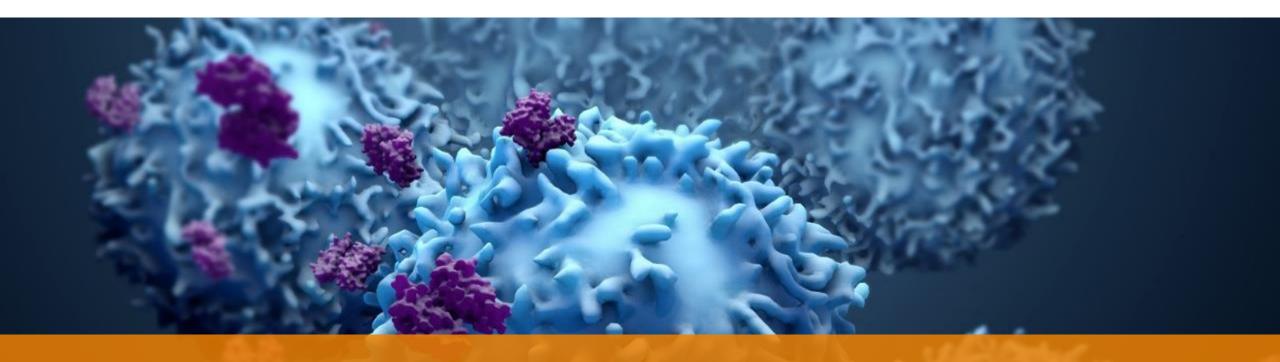
 There has been a lack of innovation in the acute, inpatient setting with >50% of patients failing to improve on frontline SoC

Rademikibart has the potential to address the unmet needs above resulting in projected peak worldwide sales of:

\$3.2B for asthma \$2.2B for COPD

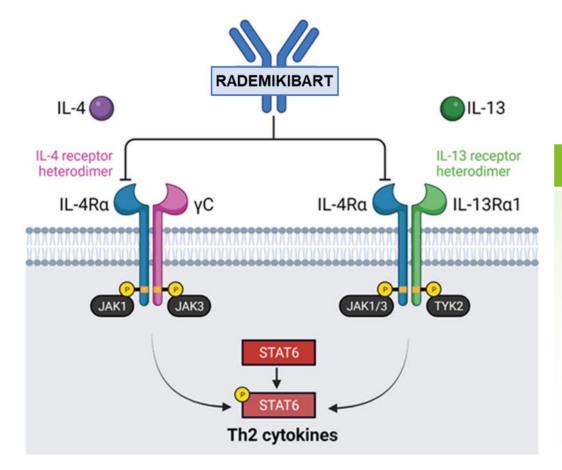






**Rademikibart:** A Next Generation Anti-interleukin-4-receptor alpha (IL-4R $\alpha$ ) Antibody

### Rademikibart: Next Gen IL- $4R\alpha$ Blocker Shows Potential For Less Frequent Dosing, Greater Sustained Efficacy Data, and Faster Onset Observed in AD and Asthma Trials



Rademikibart is a novel, human monoclonal IgG4 antibody directed against IL-4R $\alpha$ , a common subunit for IL-4 and IL-13 receptors. Blockade of IL-4 and IL-13 binding to IL-4R $\alpha$  results in inhibition of both IL-4 and IL-13 signaling.

#### **Rademikibart Characteristics**

- Engaging with distinct epitopes associated with a higher binding affinity to the IL-4Rα¹
- Highly potent IC<sub>50</sub> in:
  - Reducing JAK-STAT signaling<sup>1,a</sup>
  - Cell proliferation<sup>1,a</sup>
  - TARC release<sup>1,a</sup>

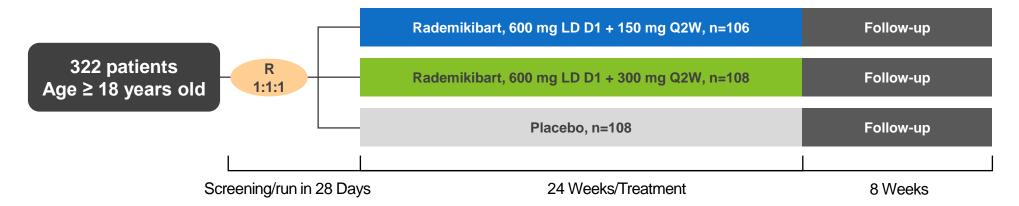
#### **Potential Clinical Relevance**

- Greater clinical response
- Faster onset of action
- · Less frequent dosing



## Robust Global Phase 2b in Moderate-to-Severe Asthma Patients Completed

Efficacy and Safety Study of CBP-201 (rademikibart) in Patients With Moderate to Severe Persistent Asthma (NCT04773678)



#### **Key Inclusion Criteria:**

- Moderate to severe uncontrolled asthma
  - Existing treatment with medium to high dose ICS in combination with a second controller (e.g., LABA, LTRA, or theophylline) for at least 3 months with a stable dose ≥1 month prior to the screening visit.
  - Pre-bronchodilator FEV<sub>1</sub> 40 to 85% of predicted normal at Visits 1 and 2, prior to randomization.
  - Screening or historical blood eosinophil count ≥150 cells/µL
    - No eosinophil count requirement for patients on maintenance OCS
  - ACQ score ≥1.5 at Visits 1 and 2, prior to randomization.
  - At least 1 documented asthma exacerbation in the 12 months prior to the date of informed consent that required use of a systemic corticosteroid

#### **Primary Endpoints:**

 Change from Baseline in FEV<sub>1</sub> at Week 12 (in clinic with central overread)

#### **Secondary Efficacy Endpoints:**

- Change from Baseline in FEV<sub>1</sub> at other timepoints
- Asthma Exacerbations
- PROs (ACQ, symptom diary, at home lung function)
- PD markers (FENO, eosinophils, ECP, periostin, TARC)
- Rescue medication use



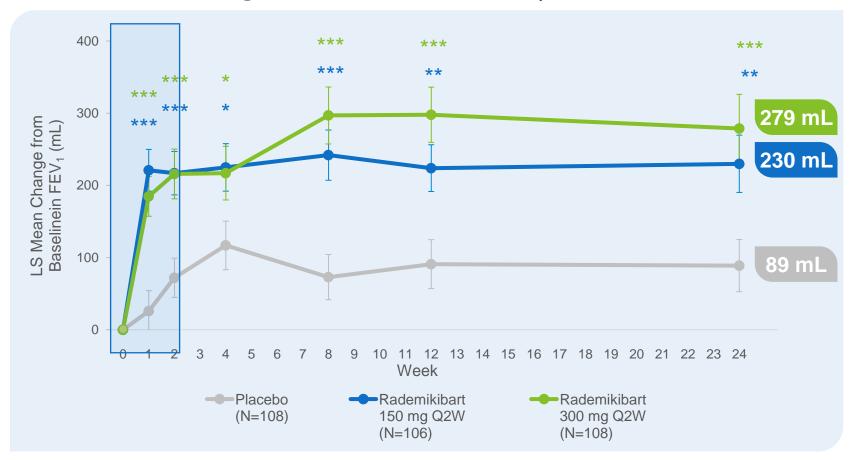
#### Rapidly Improved and Sustained FEV, Values Observed with **Rademikibart Treatment**

#### Change in Pre-bronchodilator FEV<sub>1</sub> over time

- Rademikibart treatment associated with rapid, significant changes in FEV₁ as early as Week 1, which were sustained for the duration of the 24-week study
- · Home daily lung function data demonstrated 75% of improvement seen on Day 7 was observed by Day 1, with 100% by Day 2:

#### Percent of Day 7 Change in FEV1

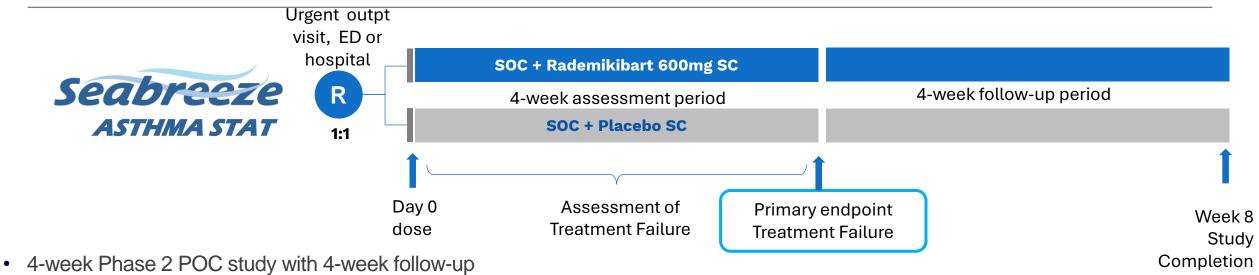






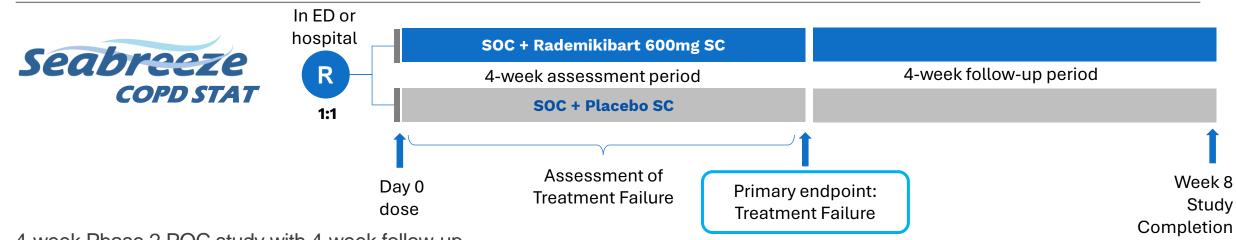
#### New Phase 2 Clinical Trial: SEABREEZE ASTHMA STAT (POC Study)





- - Primary endpoint
    - Treatment Failure through 28 days after randomization: death of any cause, (re)admission to hospital, urgent visit to an outpatient or ED provider for worsening of asthma symptoms, or necessity to intensify pharmacologic treatment (including second course of systemic steroids for exacerbation)
  - Secondary and Exploratory endpoints
    - Rate of exacerbations, time to asthma exacerbation, time to ready for discharge in hospitalized patients, lung function in the 28 days after randomization, Asthma Control Questionnaire (ACQ), Asthma Quality of Life Questionnaire, and daily asthma symptoms and rescue medication usage by e-diary
- Patients Adolescents and adults
  - 1 exacerbation in prior 12 months; No exacerbation in the previous 4 weeks
  - Eosinophil count of ≥ 300 cells/µL
- Dose
  - 600 mg rademikibart or matched placebo administered SC
- Enrollment ~200 patients





- 4-week Phase 2 POC study with 4-week follow-up
  - Primary endpoint
    - Treatment Failure through 28 days after randomization: death of any cause, (re)admission to hospital, urgent visit to an outpatient or ED provider for worsening of asthma symptoms, or necessity to intensify pharmacologic treatment (including second course of systemic steroids for exacerbation)
  - Secondary and exploratory endpoints
    - Rate of exacerbations, Time to COPD exacerbation, Time to ready for discharge in hospitalized patients, Lung function measurements, Length of stay, EQ-5D-5L, Visual analog dyspnea scale, Exacerbations of Chronic Pulmonary Disease Tool, St. George's Respiratory Questionnaire for COPD Patients and COPD Assessment Test
- Patients Adults
  - 1 exacerbation in prior 12 months; No exacerbation in the previous 4 weeks
  - Eosinophil count of ≥ 300 cells/µL
- Dose
  - 600 mg rademikibart or matched placebo administered SC
- Enrollment ~200 patients



## Recent IIT Presented at ERS Validated the Approach and Provided a Roadmap to Improve Powering Our Trials



7-11 September | Vienna, Austria

RCT Abstract - Treating eosinophilic exacerbations of asthma and COPD with benralizumab: a double-blind, double-dummy, active-placebo randomised controlled trial (ABRA)

S. Ramakrishanan (Perth, Australia), R. Russell (London, United Kingdom), H. Mahmood (Oxford, United Kingdom), K. Krassowska (Oxford, United Kingdom), J. Melhorn (Oxford, United Kingdom), C. Mwasuku (London, United Kingdom), I. Pavord (Oxford, United Kingdom), L. Bermejo-Sanchez (Oxford, United Kingdom), I. Howell (Oxford, United Kingdom), M. Mahdi (Oxford, United Kingdom), S. Peterson (Lund, Sweden), T. Bengtsson (Lund, Sweden), M. Bafadhel (London, United Kingdom)

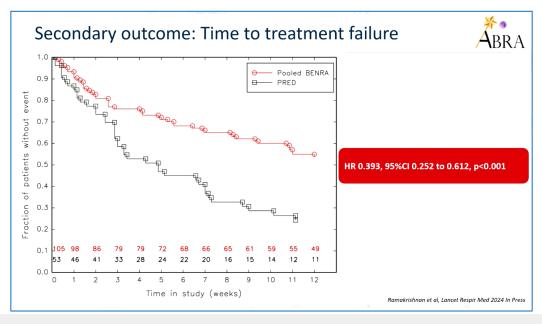
**Background:** Eosinophilic inflammation is a treatable trait commonly found in acute exacerbations. We investigated if eosinophilic exacerbations could be treated acutely with benralizumab.

Methods: ABRA was a multicentre RCT. At the time of an acute exacerbation of asthma or COPD, adults with blood eosinophil counts ≥300 cells/µL in a 1:1:1 ratio received prednisolone 30mg PO for 5 days+100mg benralizumab SC once; or placebo tablets PO for 5 days+100mg benralizumab SC once; or prednisolone 30mg PO for 5 days+placebo SC once. The co-primary outcomes were proportion of treatment failures over 90 days and total visual analogue scale (VAS) symptoms at day 28. Secondary endpoints included time to treatment failure and lung function.

**Findings:** 158 patients were randomised at acute eosinophilic exacerbation of asthma and/or COPD. At 90 days, treatment failures occurred in 39/53 (73.6%) in the prednisolone only group and 47/105 (44.8%) in the pooled benralizumab group (OR 0.264, 95%CI 0.125-0.556, p<0.001). The 28-day total VAS mean (95%CI) difference was 49mm (14-84) favouring benralizumab (p=0.006). The time to treatment failure was longer with benralizumab compared to prednisolone (HR 0.393, 95%CI 0.252-0.612, p-value<0.001). Lung function was similar.

Interpretation: Benralizumab is superior to prednisolone in the treatment of eosinophilic exacerbations.

- >50% of SOC + placebo achieved treatment failure criteria by 4 weeks
- Benralizumab reduced treatment failure at 90 days by 40%
- Little separation in the first 3-weeks indicating slow onset of effect
- Based on published data, rademikibart is more effective than benralizumab at improving FEV<sub>1</sub>





## Competitive Landscape of Placebo Adjusted Improvement from Baseline In Pre-bronchodilator FEV<sub>1</sub>

#### Rademikibart exhibited best-in-class potential in lung function improvement

Source	MoA	Product	FDA Approv. in last 10 yrs	Study	N (Pbo/Tx)	% patients with EOS >300 cells/µL	Week	First response week	Placebo adjusted improvement from baseline in FEV <sub>1</sub> (all patients)	Placebo adjusted improvement from baseline in FEV <sub>1</sub> (eos>300 cells/µL)
Phase	IL-4Rα	Rademikibart			108/108	46.3%	12	4	189 mL	328 mL
2b	Rademikibart			100/100	40.3%	24	1	190 mL	365 mL	

Biologic
Phase 3 trial
results

IL-4Rα	Dupilumab	2018	QUEST <sup>2</sup>	231/633	41.8%	12	2	130 mL	240 mL
		2017	SIROCCO <sup>6</sup> Q4W	407/399		48			106 mL
IL-5Rα	IL-5Rα Benralizumab		CALIMA Q4W	248/241	68.9%	56	- 4		125 mL
11 5	U.S. Budenid	0040	STUDY 1 <sup>5</sup>	244/245		50	4	126 mL	
IL-5 Reslizumab	2016	STUDY 2 <sup>5</sup>	232/232		52	4	90 mL		
IL-5	II 5 Manalinumah	olizumab 2015	MENSA <sup>3</sup>	191/194		32	4	98 mL	132 mL*
ic-5 Mepolizum	меропишпар		MUSCA <sup>4</sup>	277/274	60.0%	24	4	120 mL	164 mL**
TSLP	Tezepelumab	2021	NAVIGATOR <sup>7</sup>	528/531	41.5%	52	2	130 mL	230 mL

For Illustrative Purposes Only: Not a head-to-head trial. Differences exist between trial designs, subject characteristics and geographical regions -- caution should be exercised when comparing data across trials

EOS=eosinophils; FDA=Food and Drug Administration; IL=Interleukin; MoA=mechanism of action; Pbo=Placebo; TSLP=thymic stromal lymphopoietin; Tx=treatment group.



<sup>\*</sup>Subgroup analysis of patients with blood eosinophils ≥500 cells/µL \*\* Difference is based on exploratory modelling of baseline blood eosinophil count at 750 cells/µL

<sup>1.</sup> ATS/ERS statement - Reddel HK et al. Am J Respir Crit Care Med. 2009 Jul 1;180(1):59-99. 2. QUEST - Castro M et al. N Engl J Med 2018;378:2486-96. 3. MENSA - Ortega HG et al. N Engl J Med 2014;371:1198-207.

<sup>4.</sup> MUSCA - Chupp GL et al. Lancet Respir Med. 2017 May;5(5):390-400. 5. STUDY 1&2- Castro M et al. Lancet Respir Med. 2015 May;3(5):355-66. 6. SIROCCO - Bleecker ER et al. Lancet. 2016 Oct 29:388(10056):2115-2127.

<sup>7.</sup> NAVIGATOR – Menzies-Gow A et al N Engl J Med 2021:384:1800-9.

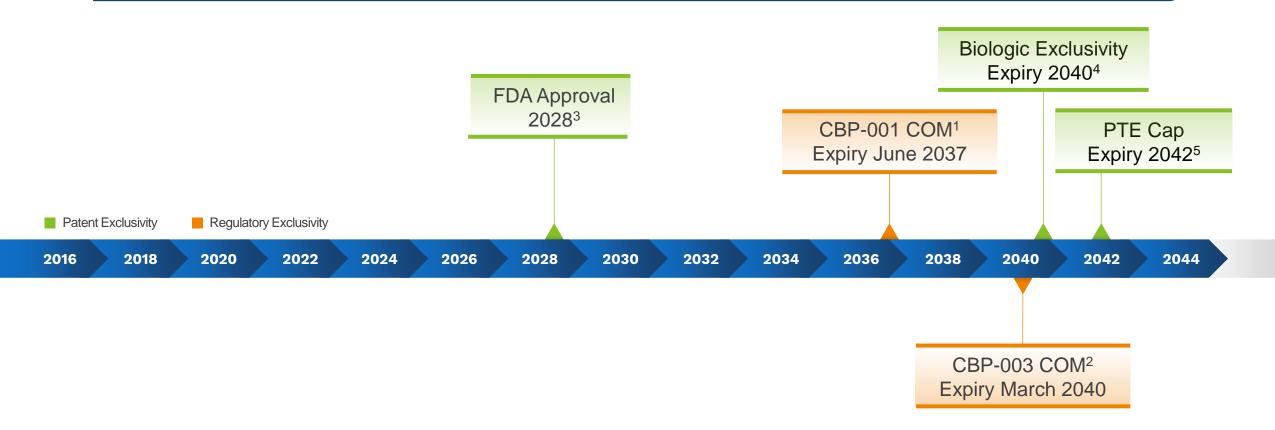
## Comprehensive Data Package Sufficient to Move Quickly into Phase 3 Once the Planned Phase 2 Studies are Completed

Study	Status	Outcomes
CMC & Improved Manufacturing Process	Tech Transfer to US CMO Completed	<ul> <li>Initial manufacturing process successfully transferred to US CMO. New high-yield cell-line developed and will be transferred next year</li> </ul>
Phase 2b Asthma	Completed	<ul> <li>Significantly improved lung function at week 12 in patients with moderate-to-severe asthma</li> <li>Rapidly improved and sustained FEV<sub>1</sub> values observed with rademikibart treatment as early as 24 hours</li> <li>Rademikibart patients trended toward 50% fewer exacerbations compared to placebo</li> </ul>
Phase 2 Acute Asthma	Planned	<ul> <li>1-month phase 2 POC study evaluating rademikibart + SOC against SOC + placebo that will enroll patients with asthma having an acute exacerbation</li> <li>Primary Endpoint: treatment failure through 28 days</li> </ul>
Phase 2 Acute COPD	Planned	<ul> <li>1-month phase 2 POC study evaluating rademikibart + SOC against SOC + placebo that will enroll patients with COPD having an acute exacerbation</li> <li>Primary Endpoint: treatment failure through 28 days</li> </ul>



#### **Rademikibart Exclusivity Timeline**

**Exclusive global development & commercialization rights** (outside of greater China) **supportive of substantial growth and value creation** 





### Rademikibart has the Potential to Transform Patient Outcomes in the ED/Hospital Setting Resulting in Substantial WW Revenue

#### DIFFERENTIATED RAPID EFFICACY, IMPRESSIVE SUSTAINED RESPONSE AND SAFETY

- Rademikibart treatment achieves rapid and significant changes in FEV<sub>1</sub> as early as 24 hours post loading dose
- FEV1 response is sustained over 24 weeks
- Rademikibart is generally well tolerated

### SIGNIFICANT COMMERCIAL OPPORTUNITY

- ~1M and ~1.3M ED visits per year and >100K and >500K hospitalizations annually by adult patients with asthma and COPD in the US respectively
- Rademikibart has the potential to be the first biologic to treat patients with acute asthma and COPD exacerbations
- Potential to drive significant chronic utilization with differentiated acute indications.
- Opportunity for significant healthcare cost savings by reducing the number of rehospitalizations, annualized days of hospitalization and possibly length of stay for both asthma and COPD

### SIGNIFICANT COMMERCIAL OPPORTUNITY

- Based on the Target Product
   Profile with both acute and chronic indication for both asthma and COPD, rademikibart has the potential to produce substantial revenue. Independent market research projects peak worldwide revenue of:
  - \$3.2B for asthma
  - \$2.2B for COPD



#### **Well Positioned for Success**

### Renewed Focus on U.S. Clinical Execution

- Announced new U.S.-based leadership with a history of late-stage clinical and regulatory execution
- Transforming Connect into a U.S.-centric company, significantly reducing presence in China
- Focused clinical development on highest value opportunity in acute asthma with no competition

#### **2025 Anticipated Catalysts**

Adopt U.S. filings with the SEC

Initiate & Complete Enrollment of Phase 2 Acute Asthma
POC study

Initiate & Complete Enrollment of Phase 2 Acute COPD POC study

**Strong Financial Position:** Cash and cash equivalents of \$110.2 million as of June 30, 2024 expected to support planned operations, including the acute asthma and COPD studies, into at least the first half of 2027



#### **Financial Summary**

Cash and cash equivalents of \$110.2 million expected to support planned operations into at least the first half of 2027

Summary Statement of Operations and Net Cash Used in Operations (In thousands, expect per share data)	Six Months Ended June 30, 2024	
Total revenue <sup>1</sup>	\$ 24,116	
Operating expenses <sup>2</sup>	16,808	
Finance income, net	401	
Income tax expense	60	
Net income <sup>2</sup>	\$ 7,649	
Basic and diluted net income per share <sup>3</sup>	\$ 0.14	
Net cash used in operations	\$ (7,974)	

Condensed Balance Sheet Data (In thousands)	June 30, 2024
Cash and cash equivalents	\$ 110,174
Total assets	\$ 120,570
Total shareholders' equity	\$ 110,479

<sup>&</sup>lt;sup>1</sup> Represents revenue recognized under the License Agreement with Simcere.



<sup>&</sup>lt;sup>2</sup> Includes \$1.8 million of non-cash, share-based compensation expense for the six months ended June 30, 2024.

<sup>&</sup>lt;sup>3</sup> Based on 55.1 million (basic) and 55.6 million (diluted) weighted average ordinary shares outstanding for the six months ended June 30, 2024.



**NASDAQ: CNTB** 

