

## A KOL Discussion on Ensifentrine's Novel Profile

June 16<sup>th</sup>, 2022



## Forward-looking statements

This presentation contains "forward-looking" statements that are based on the beliefs and assumptions and on information currently available to management of Verona Pharma plc (together with its consolidated subsidiaries, the "Company"). All statements other than statements of historical fact contained in this presentation are forward-looking statements. Forward-looking statements include information concerning the initiation, timing, progress and results of clinical trials of the Company's ensifentrine product candidate, the assumptions underlying ensifentrine's anticipated treatment effect and top-line results, the timing or likelihood of regulatory filings and approvals for ensifentrine, the potential of ensifentrine in the treatment of COPD, cystic fibrosis, asthma and other respiratory diseases, and estimates regarding the market opportunity for ensifentrine and sales force and commercial strategies to support ensifentrine's launch. In some cases, you can identify forward-looking statements by terminology such as "may," "will," "should," "expects," "plans," "anticipates," "believes," "estimates," "predicts," "potential" or "continue" or the negative of these terms or other comparable terminology.

Forward-looking statements involve known and unknown risks, uncertainties and other factors that may cause the Company's actual results, performance or achievements to be materially different from any future results, performance or achievements expressed or implied by the forward-looking statements. These risks, uncertainties and other factors include those under "Risk Factors" in the Company's annual report on Form 10-K for the year ended December 31, 2021, our Quarterly Report on Form 10-Q for the quarterly period ended March 31, 2022, and our other reports and filings with the Securities and Exchange Commission (the "SEC"). Forward-looking statements represent the Company's beliefs and assumptions only as of the date of this presentation. Although the Company believes that the expectations reflected in the forward-looking statements are reasonable, it cannot guarantee future results, levels of activity, performance or achievements. Except as required by law, the Company assumes no obligation to publicly update any forward-looking statements for any reason after the date of this presentation, or to conform any of the forward-looking statements to actual results or to changes in its expectations.

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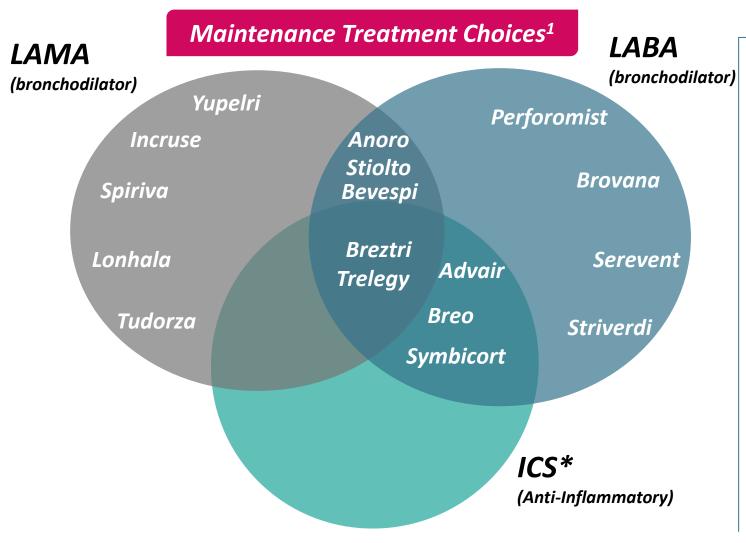
## Verona Investor KOL day Agenda

Topic	Speaker
Introduction & Verona Overview	David Zaccardelli, PharmD, CEO
KOL discussion of COPD therapy landscape, current unmet needs, & ensifentrine's role in treatment paradigm	Igor Barjaktarevic, MD UCLA Division of Pulmonary and Critical Care Medicine, Medical Director, COPD Program
	Jill Ohar, MD Wake Forest University, Professor of Medicine
	Kathy Rickard Chief Medical Officer
	Moderator Chris Martin, SVP Commercial
Q&A	Attendees / Speakers
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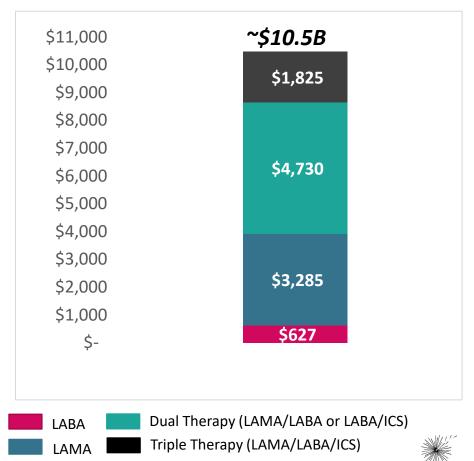


## **Current COPD treatments limited to 3 MoAs**

LAMAs & dual therapies generate the majority of product sales



#### 2020 Maintenance COPD Sales<sup>2</sup>



LAMA: Long acting muscarinic agent, LABA: Long acting beta-adrenoceptor agonist

<sup>\*</sup>GOLD Guidelines recommend ICS products for patients who have a frequent history of exacerbations

¹https://www.copdfoundation.org/Learn-More/I-am-a-Person-with-COPD/Treatments-Medications.aspx; ² IQVIA MIDAS, IQVIA MIDAS Medical

# Ensifentrine: Novel profile providing both bronchodilator and anti-inflammatory effects

Ensifentrine impacts 3 key mechanisms in respiratory disease

#### Airway Smooth Muscle<sup>1-4</sup>

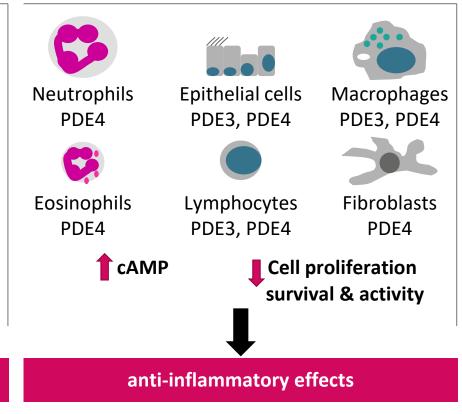
# PDE3, PDE4



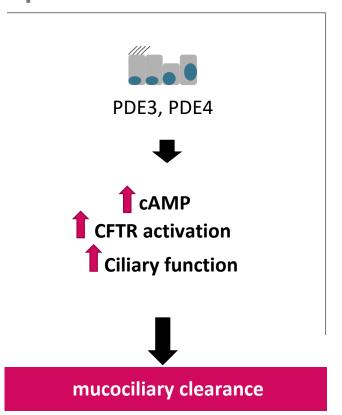


bronchodilation

#### Inflammatory Cells<sup>5,6</sup>



#### **Epithelial Cells**<sup>7,8</sup>

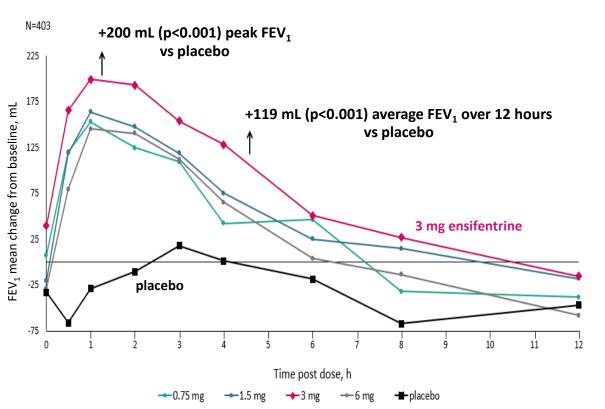




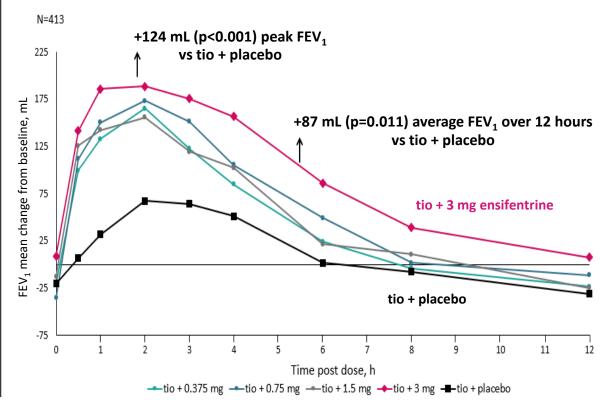
## Ensifentrine: Spirometry across Phase 2b program

12-Hour spirometry profile at week 4 supports twice daily dosing

#### Study 203: *Ensifentrine Monotherapy*<sup>1</sup>



#### Study 205: Ensifentrine + Tiotropium<sup>2</sup>





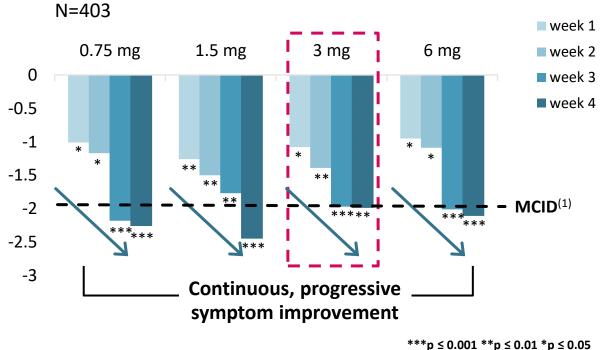
## Ensifentrine: Symptom improvement in two large Phase 2b trials

#### Improvements seen at Phase 3 trial dose

#### Study 203: Ensifentrine Monotherapy<sup>1</sup>

#### **Symptom relief**

Total Score E-RS: COPD by Week



Placebo corrected
(1) Minimal clinically important difference

#### Study 205: Ensifentrine + Tiotropium<sup>2</sup>

#### **Symptom & QOL relief**

Total Score SGRQ-C: COPD by Week
N=413
tio + 0.375
mg
tio + 0.75 mg
tio + 1.5 mg

o
-1
-2
-3
-4
-MCID1

Progressive Quality of Life Improvement

Placebo corrected

-5

<sup>1</sup>Minimal clinically important difference (-4 units)



\*p ≤ 0.05

## Safety profile similar to placebo

Ensifentrine well tolerated at all doses in trials involving over 1,400 subjects



Adverse events generally balanced between ensifentrine and placebo

#### **Safety Database:**

- 19 clinical trials
- 980 COPD patients on nebulized ensifentrine across 6 studies



Gastrointestinal profile similar to placebo



Cardiovascular profile similar to placebo with no QT prolongation in thorough QT study



No consistent or dose related changes in adverse events, blood pressure, heart rate or QT in independent cardiologist review

Ensifentrine has a favorable benefit / risk profile



## **ENHANCE** program incorporates many Phase 2 trial elements

Similar population attributes including demographics and COPD characteristics

	Phase 2b Trials (Study 203 / 205) <sup>1,2</sup>	ENHANCE 1 & 2 Trials <sup>3,4</sup>
Sites	US & Europe	US, Europe, & South Korea
Dosage	0.375, 0.75, 1.5, 3, 6 mg BID	3 mg BID
Number of patients	400 / study	800 / study
Length of study	4 weeks (efficacy & safety)	12 weeks (Primary Endpoint – lung function) 24 weeks (Secondary Endpoints – symptoms / QOL) 48 weeks (safety)
Entry criteria	<ul> <li>Monotherapy / Add on to LAMA</li> <li>ICS use in ~40% (Study 203)</li> <li>30-80% predicted FEV<sub>1</sub></li> <li>Symptomatic: mMRC ≥ 2 (Study 205)</li> </ul>	<ul> <li>Monotherapy / Add on to LAMA or LABA</li> <li>ICS up to 20%</li> <li>30-70% predicted FEV<sub>1</sub></li> <li>Symptomatic: mMRC ≥ 2</li> </ul>
Key endpoints	<ul> <li>Peak FEV<sub>1</sub></li> <li>FEV<sub>1</sub> (0-12 hrs)</li> <li>Symptoms &amp; QOL (E-RS and SGRQ)</li> <li>Trough FEV<sub>1</sub></li> </ul>	<ul> <li>Peak FEV<sub>1</sub></li> <li>FEV<sub>1</sub> (0-12 hrs)</li> <li>Symptoms &amp; QOL (E-RS and SGRQ)</li> <li>Trough FEV<sub>1</sub></li> </ul>

<sup>&</sup>lt;sup>1</sup>RPL554-CO-203, Singh D, et al., Respiratory Research 2020;



<sup>&</sup>lt;sup>2</sup>RPL554-CO-205 Full Phase 2b Analysis Set, data on file; tiotropium (Spiriva®)

<sup>&</sup>lt;sup>3</sup> https://clinicaltrials.gov/ct2/show/NCT04535986?term=ensifentrine&draw=2&rank=1

<sup>&</sup>lt;sup>4</sup> https://clinicaltrials.gov/ct2/show/NCT04542057?term=ensifentrine&draw=2&rank=2

## **Upcoming milestones**

ENHANCE-2 data readout in Q3 / ENHANCE-1 around the end of the year

Milestone	Timing
Top-line ENHANCE-2 data	Q3 2022
Top-line ENHANCE-1 data	Around year end 2022
Submit US NDA	1H 2023
Approval / PDUFA date	1H 2024
US Commercial Launch	2H 2024



## ENHANCE-2 top-line data release in Q3

Anticipated data readout measures

Endpoint	Top-line Data	Powering Assumptions (90%) for detectable treatment effect
Primary endpoint	Improvement in lung function as measured by average forced expiratory volume in one second (FEV <sub>1</sub> ) area under the curve (AUC) 0-12 hours post dose at week 12	59 mL (SD 250mL)
	COPD Symptoms (E-RS Total Score)	-1.2 units (SD 5 units)
Key secondary endpoints	Health-related Quality of Life (SGRQ Total Score)	-3.3 units (SD 14 units)
	Overall safety data and other lung function measures	

## Verona Pharma's respiratory product pipeline

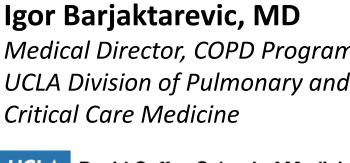
Ensifentrine's multiple "Pipeline in a product" opportunities

Program	Delivery	Pre-clinical	Phase 1	Phase 2	Phase 3
	Nebulizer				
Maintenance treatment of COPD	DPI/MDI				
	Inhaled with LAMA or LABA				
Acthmo	Nebulizer <sup>1</sup>				
Asthma	DPI/MDI				
Cystic fibrosis	Nebulizer				
	DPI/MDI				

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Q&A	Chief Medical Officer  Moderator Chris Martin, SVP Commercial  Attendees / Speakers
Close	Chris Martin, SVP Commercial

### Igor Barjaktarevic, MD Medical Director, COPD Program, UCLA Division of Pulmonary and







**David Geffen School of Medicine** 

Dr Barjaktarevic is an Assistant Professor of Medicine and a Medical Director of the COPD program at the David Geffen School of Medicine at UCLA in Los Angeles, CA. He is a specialist in pulmonology and critical care medicine and works at Ronald Reagan Medical Center. Dr Barjaktarevic graduated from the Medical School at the University of Belgrade, completed his residency training at the New York University and received a pulmonary/critical care fellowship at the Cornell University at New York Presbyterian Hospital. He holds a PhD in pulmonary immunology and his research is focused on COPD, alpha-1 antitrypsin deficiency and the use of ultrasound in critical care.



Jill Ohar, MD Professor of Medicine, Wake Forest School of Medicine



Dr Ohar is a leading pulmonary disease specialist, board certified in internal care medicine. Her clinical and research interests focus on inhalational diseases such as COPD and asbestos-related diseases and her work has been published in numerous peer-reviewed journals. Dr Ohar is the founding member of the North Carolina chapter of the American Thoracic Society. She received her medical degree from the Medical College of Pennsylvania and completed her residency in internal medicine and her fellowship in critical care medicine and pulmonary diseases at the Medical College of Virginia.



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## **Q&A Panel**



Igor Barjaktarevic, MD

Medical Director, COPD

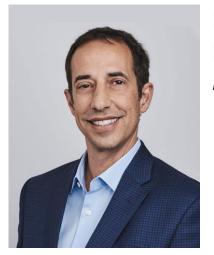
Program, UCLA Division

of Pulmonary and Critical

Care Medicine



Jill Ohar, MD
Professor of Medicine,
Wake Forest School of
Medicine



David Zaccardelli, PharmD President & CEO, Verona Pharma



Kathleen Rickard,
MD
CMO, Verona
Pharma



Chris Martin
SVP of
Commercial,
Verona Pharma



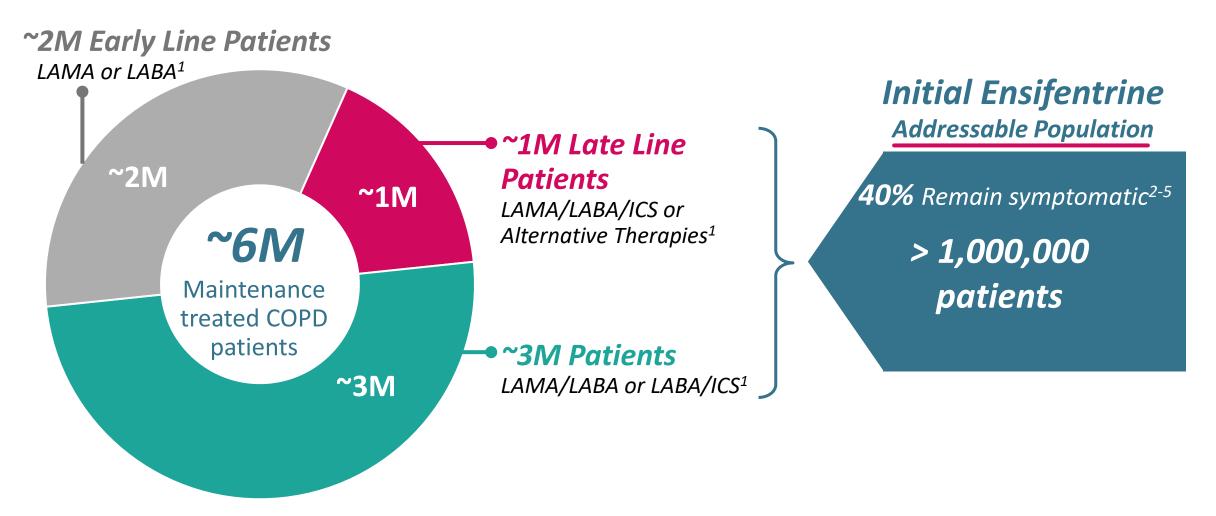
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## >1 Million patients are early candidates for ensifentrine

New MOAs to treat COPD are a significant unmet need for HCPs





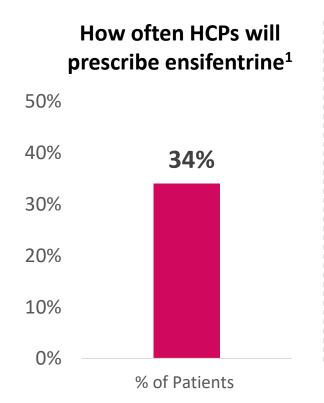
## Significant US market opportunity for ensifentrine at launch

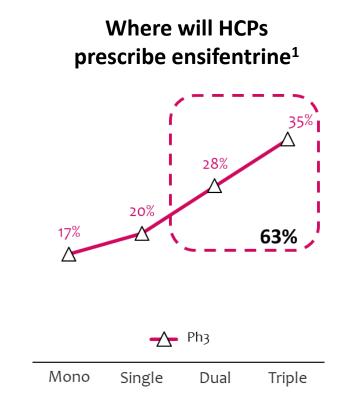
~100 Sales reps to support launch opportunity

#### Initial Ensifentrine Addressable Population

>1,000,000 Patients<sup>1-4</sup>

symptomatic on dual / triple therapy



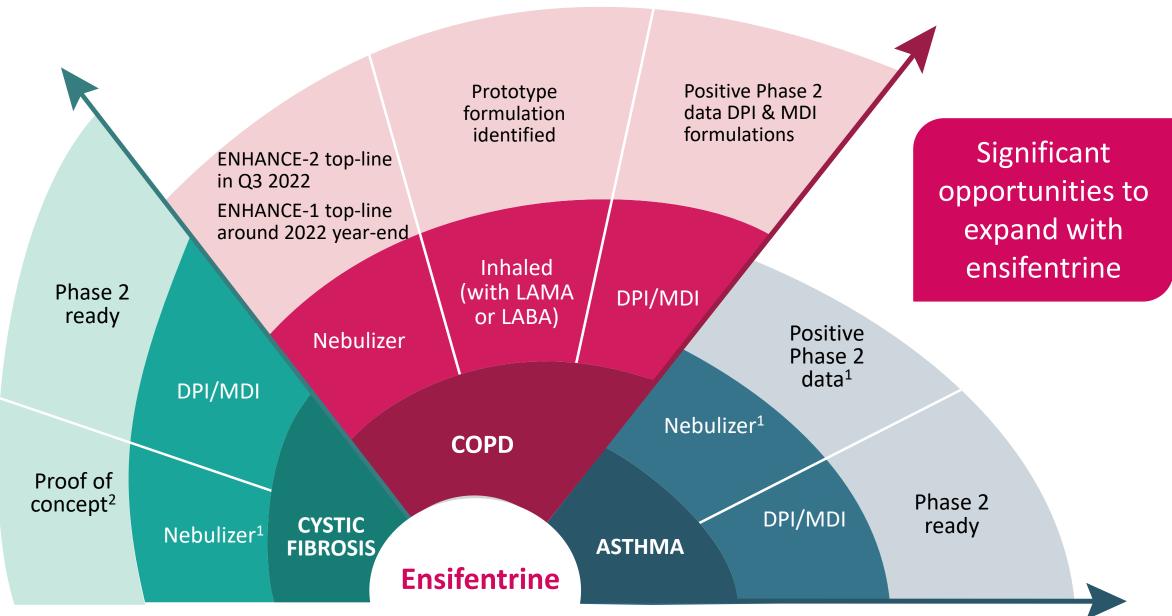


Current branded nebulizers average monthly WAC price<sup>5</sup>

~\$1,100



## Verona Pharma's respiratory product pipeline



## Verona is well positioned to maximize the value of ensifentrine

Advanced Phase 3 asset expected to deliver significant commercial opportunities

#### **Top-line Data Readouts**

- ENHANCE-2: Q3 2022
- ENHANCE-1: Around the end of the year

**Large market - >1M patients remain symptomatic**<sup>1-3</sup>

Focused commercial opportunity – 12k physicians / 100 reps<sup>4-6</sup>

Cash runway through at least 2023





## Thank you

