



# Patients' acceptance of outcome and experience measurements during hospitalisation for COPD exacerbations: a CICERO Clinical Research Collaboration–European Lung Foundation online patient survey

Iwein Gyselinck <sup>1,2</sup>, Sanjay Ramakrishnan <sup>3,4,5</sup>, Kristina Vermeersch <sup>1,2</sup>, Andreas Halner <sup>3,4,5</sup>, Hendrik Pott <sup>6</sup>, Fabienne Dobbels <sup>7</sup>, Courtney Coleman <sup>8</sup>, Philip Collis <sup>9</sup>, Henrik Watz <sup>10,11</sup>, Timm Greulich <sup>6</sup>, Frits M.E. Franssen <sup>12,13</sup>, Pierre-Régis Burgel <sup>14,15</sup>, Mona Bafadhel <sup>16,17</sup>, Wim Janssens <sup>18</sup> and <sup>18</sup> on behalf of the CICERO Consortium

<sup>1</sup>Clinical Department of Respiratory Diseases, University Hospitals Leuven, Leuven, Belgium. <sup>2</sup>BREATHE Lab, CHROMETA Department, KU Leuven, Leuven, Belgium. <sup>3</sup>Respiratory Medicine Unit, Nuffield Department of Medicine – Experimental Medicine, University of Oxford, Oxford, UK. <sup>4</sup>National Institute for Health Research Oxford Biomedical Research Centre, University of Oxford, Oxford, UK. <sup>5</sup>School of Medical and Health Sciences, Edith Cowan University, Perth, Australia. <sup>6</sup>Philipps-University Department of Medicine, Pulmonary and Critical Care Medicine, University Medical Centre Giessen and Marburg; Philipps-University, German Center for Lung Research (DZL), Marburg, Germany. <sup>7</sup>Center for Nursing and Midwifery, Department of Public Health and Primary Care, KU Leuven, Leuven, Belgium. <sup>8</sup>European Lung Foundation, Sheffield, UK. <sup>9</sup>European Lung Foundation Patient Advisory Group. <sup>10</sup>Pulmonary Research Institute at Lungen Clinic Grosshansdorf, Grosshansdorf, Germany. <sup>11</sup>Airway Research Center North and DZL, Grosshansdorf, Germany. <sup>12</sup>Department of Research and Education, CIRO, Horn, Netherlands. <sup>13</sup>Department of Respiratory Medicine, Maastricht University Medical Centre, Maastricht, Netherlands. <sup>14</sup>Assistance Publique Hôpitaux de Paris, Department of Respiratory Medicine, Hôpital Cochin Pneumologie, Paris, France. <sup>15</sup>Université Paris Cité, Institut Cochin, Inserm U1016, Paris, France. <sup>16</sup>School of Immunology and Microbial Sciences, Faculty of Life Sciences and Medicine, King's College London, London, UK. <sup>17</sup>Department of Respiratory Medicine, Nuffield Department of Medicine, University of Oxford, Oxford, UK. <sup>18</sup>For a list of the CICERO Clinical Research Collaboration members see the Acknowledgements.

Corresponding author: Iwein Gyselinck ([iwein.gyselinck@kuleuven.be](mailto:iwein.gyselinck@kuleuven.be))



Shareable abstract (@ERSpublications)

In the CICERO–ELF patient survey, European patients endorsed a selection of outcome and experience measurements from a previously obtained expert consensus list, for standardised assessments during hospitalisation and follow-up of acute COPD exacerbations <https://bit.ly/3oSHbUo>

**Cite this article as:** Gyselinck I, Ramakrishnan S, Vermeersch K, *et al.* Patients' acceptance of outcome and experience measurements during hospitalisation for COPD exacerbations: a CICERO Clinical Research Collaboration–European Lung Foundation online patient survey. *ERJ Open Res* 2023; 9: 00148-2023 [DOI: 10.1183/23120541.00148-2023].

Copyright ©The authors 2023

This version is distributed under the terms of the Creative Commons Attribution Non-Commercial Licence 4.0. For commercial reproduction rights and permissions contact [permissions@ersnet.org](mailto:permissions@ersnet.org)

Received: 9 March 2023  
Accepted: 18 April 2023



## Abstract

**Background** The lack of standardised outcome assessments during hospitalisation and follow-up for acute COPD exacerbations has hampered scientific progress and clinical proficiency. The objective of the present study was to evaluate patients' acceptance of selected outcome and experience measurements during hospitalisations for COPD exacerbations and follow-up.

**Methods** An online survey was held amongst COPD patients in France, Belgium, The Netherlands, Germany and the UK. The European Lung Foundation COPD Patient Advisory Group was involved in the conceptualisation, development and dissemination of the survey. The survey was complementary to a previously obtained expert consensus. We assessed patients' views and acceptance of selected patient-reported outcomes or experiences and corresponding measurement instruments (for dyspnoea, frequent productive cough, health status and hospitalisation experience), and of selected clinical investigations (blood draw, pulmonary function test, 6-min walk test, chest computed tomography, echocardiography).

**Findings** 200 patients completed the survey. All selected outcomes and experiences were deemed important, and acceptance of their methods of assessment was high. The modified Medical Research Council scale and a numerical rating scale to address dyspnoea, the COPD Assessment Test for quality of

life and frequent productive cough, and the Hospital Consumer Assessment of Healthcare Providers and Systems for hospital experiences were the instruments preferred by patients. Consensus on importance of blood draw and spirometry was higher compared with the other investigations.

**Interpretation** The survey results endorse the use of the selected outcome and experience measurements during hospitalisations for COPD exacerbations. They can be used to optimise standardised and patient-centred care and facilitate multicentric data collection.

## Introduction

Severe acute exacerbations of COPD (AECOPD) with a need for hospitalisation are the main drivers of the increasing COPD-associated healthcare resource utilisation and COPD-related mortality [1, 2]. Yet, there is limited guidance on standardised treatment or clinical assessments, examinations, and laboratory, radiological and functional tests during hospital stay or during follow-up. Where strong consensus or solid evidence does exist, there is often a translation gap, given that results from randomised controlled trials are not evaluated in real-world settings, and guideline recommendations are ineffectively implemented [3].

Not only does this affect clinical outcomes, but it may also leave a scientific resource untapped. The hospitalisation period for an AECOPD provides an optimal setting for accurate and extensive clinical and biological characterisation of severe events. Adherence to standardised treatment and assessment protocols would facilitate pooling of real-world data and setting up registries or multicentre clinical studies. Such large-scale initiatives are much needed to advance the data-hungry research on aetiological grouping of AECOPD and to provide a framework for more targeted treatments of specific exacerbation subgroups [4].

The Collaboration In COPD Exacerbations (CICERO) is a clinical research collaboration (CRC) supported by the European Respiratory Society (ERS) and a network of expert centres aiming to understand and improve outcomes in severe COPD exacerbations across Europe [5]. In a recent expert Delphi, we have obtained consensus from expert COPD researchers and clinicians regarding standardisation of assessments at the time of hospitalisation for AECOPD and during follow-up [6]. However, it remains elusive how acceptable these are to patients. In this patient survey, a joint-initiative of CICERO and the European Lung Foundation (ELF), we investigated patients' acceptance of the proposed assessments, with an emphasis on the patient-reported outcome (PROMs) and patient-reported experience measures (PREMs), and some frequently performed clinical investigations. The results were used to inform the protocol of CICERO's multicentric European prospective cohort study (CATALINA – ClinicalTrials.gov: NCT05008081).

## Materials and methods

### Design

Between 29 April 2021 and 11 November 2021, the ELF and CICERO held an anonymous online survey to evaluate patients' acceptance of a selection of PROMs, PREMs and clinical investigations to be obtained during and after a hospitalisation for an AECOPD. A general outline of the survey and the full English translation are provided in the appendices (supplementary Appendix 1 and 10).

### Patient and public involvement

Members of the ELF's COPD patient advisory group (PAG) were involved in all stages of survey building, including reviewing and refining the content, and reviewing the language and lay-out for accessibility and understanding. Survey awareness and dissemination strategy was established together with the PAG and representatives of national patient associations from Belgium, France and The Netherlands. Information videos and brochures were revised by the patients.

Patients' acceptance of PROMs and PREMs in this survey, based on a predefined consensus measure, was taken into consideration during the development of CICERO's CATALINA study. Through this survey, we also sought patient input on the perceived importance and feasibility of selected clinical investigations during and after a hospitalised COPD exacerbation, but without establishing a minimal patient consensus as a prerequisite for inclusion of these outcomes in CATALINA.

### Procedures and patients

The survey was developed in English and translated by members of the research team into Dutch, French and German. The online survey was hosted on the ELF website and accessible through an anonymous link. It was featured on the ELF social media platforms (Facebook, Twitter, Instagram), and in the newsletters of the ELF and ERS. COPDvzw (Belgium), Alpha-1Plus Belgium, Santé Respiratoire France (France), Longfonds (The Netherlands), the international COPD Foundation and other patient associations disseminated the survey amongst their members and through their information channels. Ethical approval

was not necessary following the recommendation of the NHS Health Research Authority ethical approval decision tool [7].

### Survey content

The content was based on CICERO's previously published expert consensus on standardised assessments during a hospitalised AECOPD and on discussions with the PAG [6]. The focus was on patient-reported outcomes in the first part of the survey, and on frequently performed clinical investigations that were presumed to be most burdensome for patients in the second part. An overview of the development of the survey is given in supplementary figure S1 and supplementary table S1 in Appendix 1.

Outcome measurements that were prioritised in the expert consensus (*i.e.* they were deemed feasible and got the label “must be assessed” or “can be considered” both during hospitalisation and follow-up) were eligible for inclusion in this patient survey and discussed with the PAG. Questionnaires additionally had to be available free of charge. For the clinical investigations, the PAG was consulted to select the investigations that were presumed to be most burdensome for patients. If the PAG felt that important outcomes and outcome measures were missing, they could suggest additional measurement instruments.

Patient-reported outcome measurements that were assessed in the first part of the survey were: the modified Medical Research Council scale (mMRC) [8] or a numerical rating scale (NRS) [9] to measure dyspnoea; the COPD Assessment Test (CAT) [10] or the Clinical COPD Questionnaire (CCQ) [11] to measure health-related quality of life; and the classical chronic bronchitis definition [12], the two cough and phlegm questions of the St George's Respiratory Questionnaire (SGRQ) [13], and the two cough and phlegm questions of the CAT [10] to measure frequent cough and phlegm [11, 14–20]. Two PREMs to measure hospital experience were also assessed: the Hospital Consumer Assessment of Healthcare Providers and Systems (HCAHPS) [21] and the howRwe instrument [22]. As part of the survey, respondents were asked to fill in all the PROMs and PREMs to get familiarised with them. Time to complete the PROMs and PREMs was recorded by the survey software. In the second part of the survey concerning clinical investigations, participants' opinions on blood draws, echocardiogram, chest computed tomography (CT), six-min walk test, spirometry and plethysmography were evaluated.

The survey contained Likert questions with a response range from 1 (strongly agree) to 5 (strongly disagree), with a score of 1 signifying a higher and 5 a lower importance or acceptance of the outcome or outcome measure. In the first part, acceptance of the PROMs and PREMs was calculated as the mean of five complementary Likert questions (ease of use, relevance, detailedness, time efficiency and general willingness to use the PROM/PREM with every clinical contact; Cronbach's  $\alpha$  was above 0.8 for all scales, see supplementary Appendix 2). There were a small number of open questions asking whether or not all-important patient-reported outcomes were included in the survey, and if patients had additional remarks on the outcome measurements.

### Data analysis

Medians and means of Likert items and the Likert scales were compared with the Wilcoxon signed rank test or Friedman test. The dispersion of the Likert responses was interpreted as a measure for consensus or dissensus and was quantified by a Shannon entropy-based consensus measure according to TASTLE *et al.* [23], ranging from 0 (absolute dissensus) to 1 (absolute consensus). Based on the probability distribution of this consensus measure, our cut-off for consensus was set at 0.7, consistent with a probability of <5% that such consensus level was a chance occurrence (supplementary Appendix 3).

Qualitative content analysis was done by coding recurrent themes in the free text and counting their frequency.

Subgroups were predefined based on language, time since diagnosis, exacerbation history and history of past hospitalisations.

The survey was performed with Qualtrics (Seattle, Washington & Provo, UT, USA). All analyses were performed using R v4.0.4 (R Core Team, Vienna, Austria).

### Sample size

A minimum of 231 responses on all PROM-related questions was predefined based on PARK *et al.* [24], as we aimed to use these results to inform the development of the CATALINA protocol.

**TABLE 1** Baseline characteristics of participants (n=377 started survey, n=200 finished survey)

Responses	
Median time needed to complete survey (min:s), median (IQR)	22:20 (8:6–35:44)
<b>Language, n (%)</b>	
Dutch	178 (47)
English	134 (36)
French	54 (14)
German	11 (3)
<b>Time since COPD diagnosis, n (%)</b>	
<5 years	71 (22)
10 years	112 (35)
>10 years	137 (43)
<b>Exacerbations during last year, n (%)</b>	
0	113 (35)
1	91 (29)
2 or more	115 (36)
<b>Hospitalised for COPD before, n (%)</b>	
Yes	125 (56)
No	99 (44)

## Results

### *Sample characteristics*

377 subjects initiated the survey, and 200 patients reached the end. At least 234 responses were obtained for all PROM-related questions. Baseline patient characteristics are shown in table 1. All responses were included for analysis, regardless of the subject's survey completion status; response rate on all variables is presented in supplementary figure S11 and table S12 in Appendix 7.

### *Outcomes, experiences, PROMs and PREMs*

Main results are shown in figures 1 and 2. Scores of individual Likert items are shown in supplementary figures S3–S10 (Appendix 5). Subgroup analyses are shown in supplementary tables S4–S7 (Appendix 5).

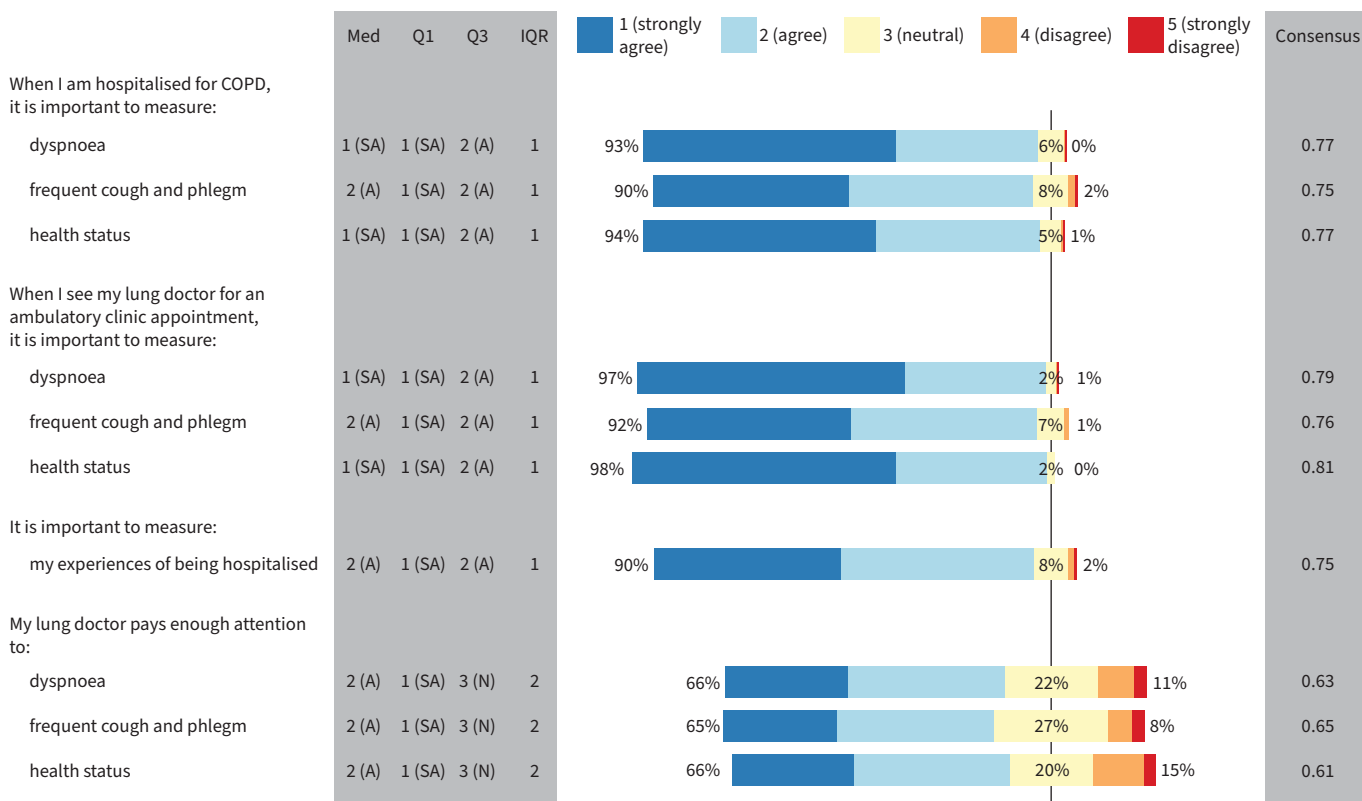
There was good consensus on the importance of measuring all selected patient-reported outcomes both during hospitalisation for an exacerbation and during ambulatory follow-up, indicated by median responses of 1 (strongly agree) or 2 (agree), and consensus of 0.70 or more. Consensus was lower for the question whether doctors paid enough attention to any of these outcomes (figure 1).

To assess dyspnoea, mean acceptance of both mMRC and NRS was high and did not significantly differ (figure 2 and supplementary figure S4 in Appendix 5). Both of these PROMs scored lowest on detailedness (supplementary figure S3 in Appendix 5). From the free text in the open questions, it was clear that some patients preferred the objectiveness of numerical scales, while others found it easier to agree or disagree with concrete descriptions of their situation as provided by the mMRC (supplementary tables S8–S9 in Appendix 6).

There was a significant difference in mean acceptance of the three PROMs to assess frequent productive cough. Participants slightly preferred the SGRQ cough and phlegm questions and CAT cough and phlegm questions over the classical chronic bronchitis definition (figure 2 and supplementary figures S5–S6 in Appendix 5). In the free text, the long recall time and use of two timespans in the classical chronic bronchitis definition (cough/sputum for at least 3 months and 2 consecutive years) were raised as possible drawbacks and may have reduced comprehensibility and perceived relevance of this PROM (supplementary tables S8–S9 in Appendix 7).

For health status measurement, responses of all evaluation questions on CAT and CCQ coalesced almost entirely on the “agree” and “strongly agree” response categories, without a significant difference in their overall acceptance (figure 2 and supplementary figures S7–S8 in Appendix 5).

Completion time stayed below 2 min for all PROMs (supplementary table S3 in Appendix 4). Symptoms that were reported in the free text fields as not sufficiently assessed in the included PROMs are summarised in a wordcloud (figure 3). Fatigue, anxiety and panic, oedema and pain were most frequently raised (supplementary tables S9–S10 in Appendix 6).



**FIGURE 1** Importance of measuring outcomes and experiences included in the survey. Participants scored the importance of measuring three outcomes (dyspnoea, frequent productive cough, health status) and one experience (hospitalisation experience) during hospitalisation for an acute COPD exacerbation or during follow-up. They also rated if sufficient attention is currently paid to the measurement of these outcomes. Participants answered to Likert items with a 5-point response range (1 – strongly agree (SA), 2 – agree (A), 3 – neutral (N), 4 – disagree (D), 5 – strongly disagree (SD)) with 1 indicating a better and 5 indicating a worse score. Medians, quartiles and interquartile ranges are shown, as well as the relative frequency bar charts per Likert item. A Shannon entropy-based consensus measure was calculated to express ordinal dispersion, and a value above 0.70 was considered good consensus. Consensus was stronger on the importance of measuring these outcomes and experiences than on the adequacy of their current assessment in routine practice. Med: median; Q1: quartile 1; Q3: quartile 3; IQR: interquartile range.

There was good consensus on the importance of measuring hospitalisation experience (figure 1). Despite longer completion time, the mean acceptance of HCAHPS PREM was slightly higher than of howRwe PREM (Figure 2 and supplementary figures S9–S10 in Appendix 5).

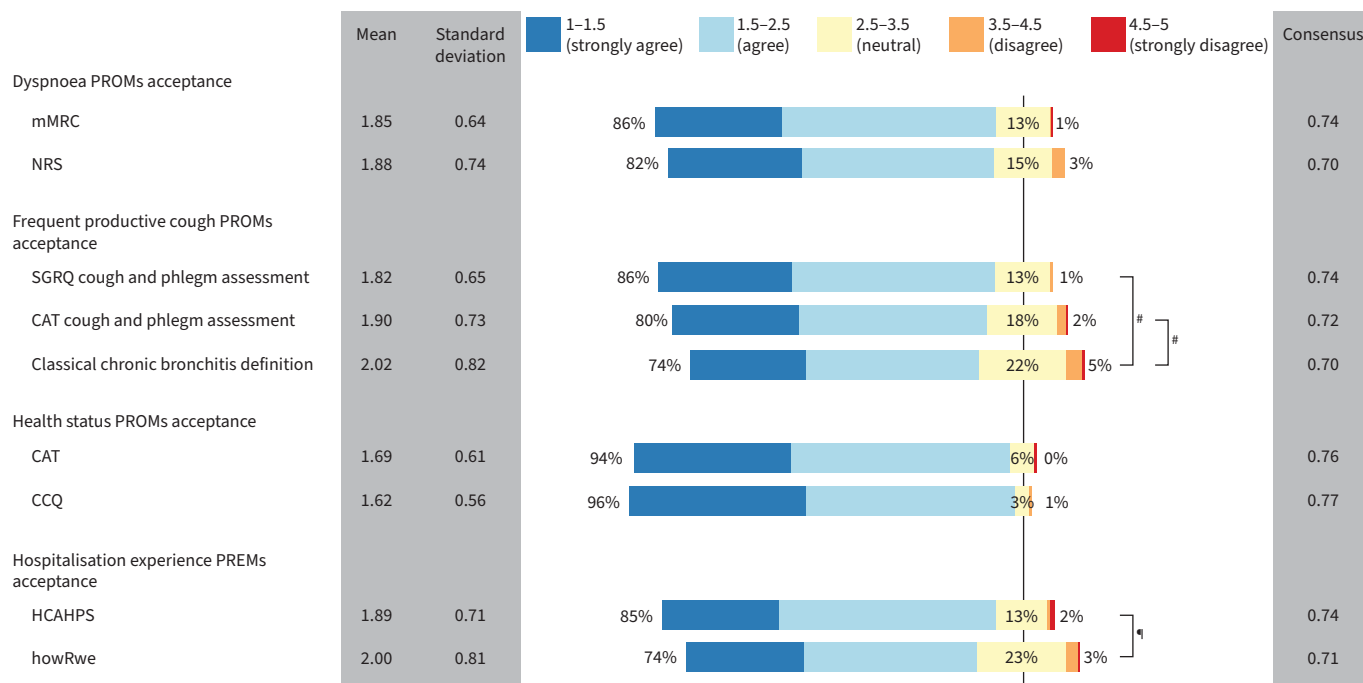
Subgroup preferences were similar to the whole sample and no relevant differences were identified (supplementary tables S4–S7 in Appendix 5).

**Clinical investigations**

Results for patients’ views on selected clinical investigations are shown in figure 4. Consensus was good on the importance of blood draw and spirometry but stayed below 0.70 for the other investigations. The 6-min walk test and plethysmography were perceived as larger burdens than the other investigations.

**Discussion**

The patient-reported outcomes and experiences included in the survey were deemed highly important by patients. Notably, the survey revealed that there was more consensus on the importance of these outcomes and experiences than on the satisfaction with their current assessment in routine clinical practice. This implies room for improvement. With our previously published expert consensus, and endorsed by this patient survey, CICERO aims to prioritise a minimal set of outcome measurements, while still assimilating the most essential data to achieve high-quality norms of COPD management, and in a standardised fashion.



**FIGURE 2** Acceptance of patient-reported outcome (PROMs) and patient-reported experience measures (PREMs). PROM and PREM acceptance were assessed by having patients answer to the same five Likert questions for each PROM/PREM (see supplementary Appendix 5) addressing 1) comprehensibility of the PROM/PREM, 2) appropriateness of the PROM/PREM or its statements to assess outcome, 3) detailedness of the PROM/PREM, 4) acceptability of time needed to complete the PROM/PREM and 5) willingness of respondents to use the PROM/PREM to assess their condition during a hospitalised acute exacerbation of COPD (AECOPD) and every follow-up visit. Each Likert item had a 5-point response range (1 – strongly agree, 2 – agree, 3 – neutral, 4 – disagree, 5 – strongly disagree) with 1 indicating a better and 5 indicating a worse score. Mean scores over the five questions were calculated per PROM and PREM for each participant and treated as a continuous variable. A Shannon entropy-based consensus measure was calculated to express ordinal dispersion, and a value above 0.70 was considered good consensus. #: there was a significant difference between the mean acceptance scores of the frequent productive cough PROMs (Friedman  $p < 0.0005$ ). *Post hoc* testing (Benjamini-Hochberg corrected) showed a difference between St George’s Respiratory Questionnaire (SGRQ) productive cough questions *versus* the classical chronic bronchitis definition (Wilcoxon signed rank  $p$ -value 0.010) and the CAT productive cough questions *versus* the classical chronic bronchitis definition (Wilcoxon signed rank  $p$ -value 0.0046), but not between the SGRQ and CAT questions (Wilcoxon signed rank  $p$ -value 0.28). ¶: there was a significant difference between the mean acceptance scores of HCAHPS and howRwe (Wilcoxon signed rank  $p < 0.0005$ ). SGRQ cough and phlegm assessment: *i.e.* first two questions of the SGRQ concerning cough and phlegm; CAT cough and phlegm assessment: *i.e.* the first two questions of the CAT concerning cough and phlegm; classical chronic bronchitis definition: *i.e.* a) cough most days for at least 3 months per year and for  $>2$  consecutive years and b) sputum most days for at least 3 months per year and for  $>2$  consecutive years. mMRC: modified Medical Research Council scale (for dyspnoea); NRS: numerical rating scale (for dyspnoea); CAT: COPD Assessment Test; CCQ: Clinical COPD Questionnaire; HCAHPS: Hospital Consumers Assessment of Healthcare Providers and Systems.

For dyspnoea measurement, patients appreciated the mMRC and an NRS equally and raised strong arguments for both instruments in the survey’s free text fields. As time to complete both instruments was very short (both below 1 min), combining both instruments seems feasible even in routine practice. For health status measurement, both the CAT and CCQ were acceptable for patients, and completion times were also short (below 1 min for CAT and below 2 min for CCQ). As standard assessment, CAT seems however the more rational choice, as it has been more often used in clinical trials, and it received more support in CICERO’s expert Delphi [6, 25]. Moreover, the two first questions of the CAT (“I never cough” to “I cough all the time” and “I have no phlegm in my chest at all” to “My chest is full of phlegm”) were accepted by patients to assess their symptoms of frequent cough and phlegm.

Of note, the two cough and phlegm questions of CAT have been repeatedly used in large cohorts to identify patients with a chronic bronchitis phenotype and, by the survey respondents, were deemed easier to comprehend than the classical definition of chronic bronchitis. Yet, strictly, they have not been validated as a separate PROM [11, 14–20]. It is not our aim to redefine established phenotypical traits based on the limited assessments here proposed. Rather, we encourage the clinician, instead of only looking at the overall score, to also pay notice to the individual questions of the PROMs and the problems patients raise



**FIGURE 3** Wordcloud with additional symptoms raised in the survey's free text fields. At the end of the survey's patient-reported outcome measures (PROMs) assessment, patients were asked if they experienced other symptoms that were not or not sufficiently addressed in the proposed PROMs. Free text was interpreted in the source language, and recurrent themes and symptoms were summarised in a codebook and this wordcloud. The font size corresponds to the relative frequency by which the symptom or theme was raised. Despite statements in the COPD Assessment Test (CAT) or Clinical COPD Questionnaire (CCQ) concerning sleep quality, energy level, self-dependency and self-confidence, symptoms related to these domains were still frequently raised as insufficiently addressed. This supports the use of such general PROMs as a first screening tool to identify problems that then may require further attention during the visit with the treating physician.

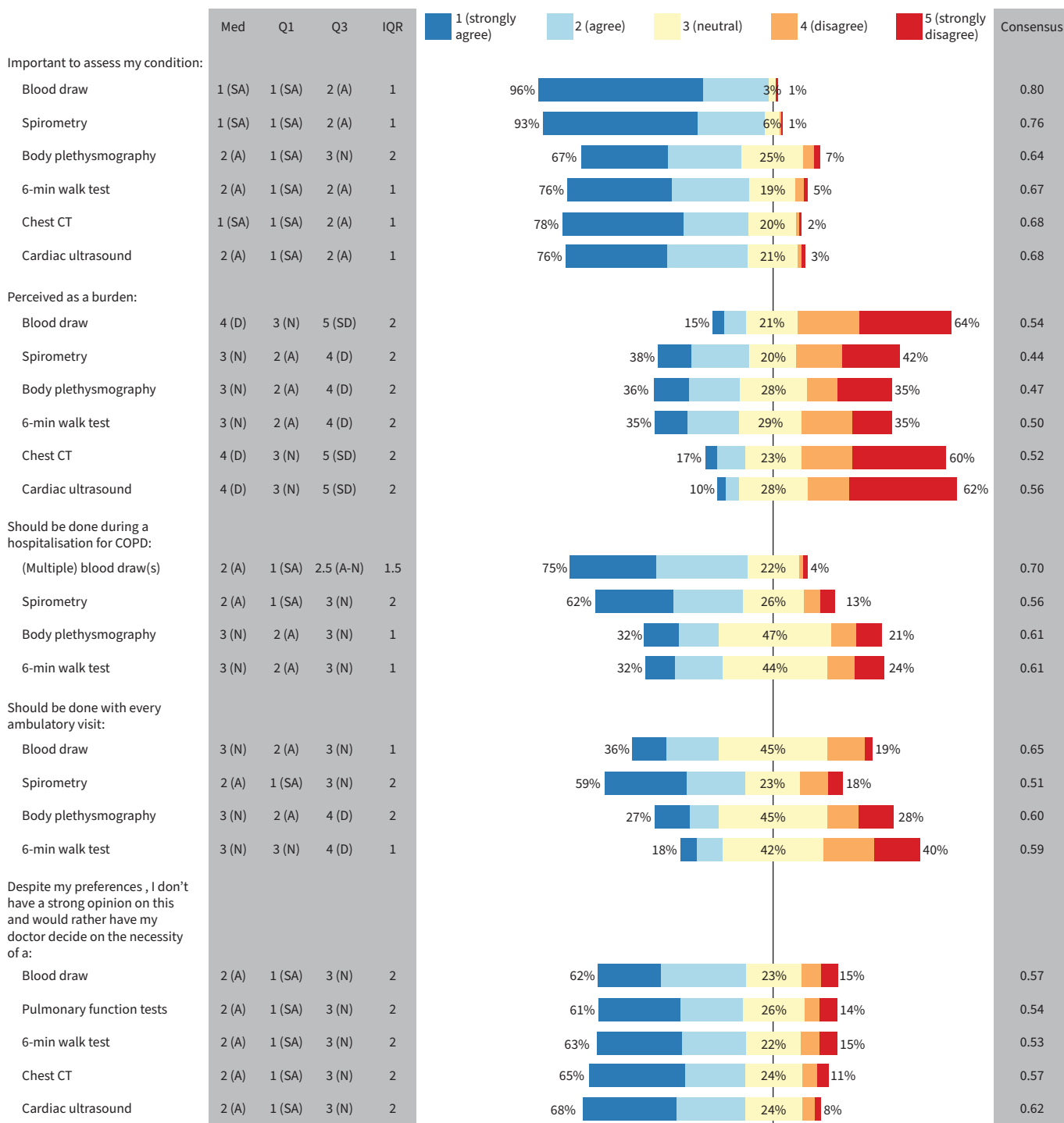
by them. A worse score on the CAT cough and phlegm questions may alert clinicians to the possible presence of chronic bronchitis and may urge them to do a more dedicated evaluation in that direction, to exclude comorbidities like bronchiectasis, or to see whether patients meet more validated criteria to justify targeted interventions. The same goes for the additional symptoms that patients raised in the survey's free text fields. It is impossible to include in a standard evaluation validated questionnaires that cover all these symptoms in detail, and this would defeat the purpose of a minimal set of outcome measurements. Yet, most of these symptoms may be addressed when examining the specific domains in detail where the patient indicates a problem through the proposed selection of PROMs.

An exception to this might be anxiety and affective symptoms, which were also frequently put forward in the survey's free text spaces. In CICERO's expert Delphi, experts did consider including a separate PROM to measure anxiety and depression – the hospital anxiety and depression scale (HADS) – as a standard assessment, albeit with remarks on the feasibility. Use of this questionnaire is however restricted, as it is not freely available, which was also the reason for excluding it in this patient survey. Still, care providers should at least acknowledge the importance and impact of mood and anxiety in this context, and alternative PROMs may be considered as part of a standardised assessment.

If hospital experience is measured, while considerably longer, the HCAHPS is better validated and was slightly preferred by patients in the survey.

With this survey we also gauged patients' opinions on clinical investigations and tests that are frequently performed in the context of COPD exacerbations. Although we did not seek to validate the use of these tests, it was surprising to see that patients' consensus on their need and importance was much lower as compared to the use of PROMs and PREMs, which were uniformly accepted. In general, though without reaching our criteria for consensus, patients agreed on the doctor's primary role in when to plan these investigations.

Exemplary for how the survey results complement the expert consensus and how these can be put in practice is the development of CICERO's CATALINA study (selection of PROMs, PREMs and clinical



**FIGURE 4** Patients' views on selected clinical investigations. Participants expressed their views on selected clinical investigations during hospitalisation for an acute COPD exacerbation or during follow-up. Patients answered to Likert items with a 5-point response range (1 – strongly agree (SA), 2 – agree (A), 3 – neutral (N), 4 – disagree (D), 5 – strongly disagree (SD)) with 1 indicating a better and 5 indicating a worse score. Medians, quartiles and interquartile ranges are shown, as well as the relative frequency bar charts per Likert item. A Shannon entropy-based consensus measure was calculated to express ordinal dispersion, and a value above 0.70 was considered good consensus. Consensus was strongest on the importance of blood draw and spirometry. More patients agreed on the 6-min walk test and body plethysmography being a burden, followed by spirometry, but all with a large dispersion of responses reflected in low consensus measures. A majority of patients agreed on the doctor's primary role to plan these investigations, but again with consensus below 0.70. Med: median; Q1: quartile 1; Q3: quartile 3; IQR: interquartile range; CT: computed tomography.



investigations for this prospective cohort study are presented in supplementary table S1 in Appendix 1). Our proposed assessments accord with and, in a way, concretise the recommendations of recent large international commissions, and are very compatible with the core outcome set for interventional COPD exacerbation trials that was recently established by another ERS task force [26–28].

This survey has limitations. Firstly, due to ethical constraints and the anonymous nature of the survey, more extensive data on demographics and disease history were not available. Secondly, only a slight majority of our sample had been hospitalised for their COPD before. Notably, however, there was no difference in acceptance of any of the proposed PROMs based on subjects' hospitalisation history. Thirdly, due to the dissemination methods, using local and national patient organisations' membership networks and communication channels, we were unable to calculate the survey's reach or response rate. Patients who engage with patient organisations are more likely to be highly engaged in their healthcare, meaning that the results may not be generalisable to the whole patient population. Also, the survey was developed in only four languages, most likely resulting in a significantly higher response rate in central European countries with native speakers and limiting generalisability of the results to the whole European continent. Fourthly, we used no question randomisation, meaning that the order of questions was the same for all respondents and that only the most motivated participants completed part two (PREM assessment) and three (clinical investigations) of the survey. This may have introduced non-response bias. Finally, consensus was reached for only a few items during the assessment of patients' views on clinical investigations in part three. It is uncertain whether a multi-round Delphi design would have led to stronger consensus on some of these items. Yet, our survey setting in a large multinational patient sample has reached consensus on most PROMs and PREMs assessments and was not powered or designed to obtain consensus on these clinical investigations.

### Conclusion

This CICERO–ELF patient survey shows that patients are supportive of the use of PROMs and PREMs during hospitalisation for an AECOPD and follow-up. We advocate their rational but consistent use in routine care, especially as patients sometimes felt that their symptoms are not given sufficient importance by their treating clinicians. Together with CICERO's expert consensus statement, our survey results can be used to institute or optimise standardised integrated care pathways for hospitalised AECOPD in a manner that is supported by both experts and patients. Adherence to such a standardised series of assessments would facilitate real-world data collection and may aid the design of multicentric observational studies.

Provenance: Submitted article, peer reviewed.

Acknowledgements: We thank the ERS, the ELF and the ELF's COPD PAG for their support during development, dissemination and conduct of this study. We are grateful to COPDvzw, Alpha-1Plus Belgium, Santé Respiratoire France, Longfonds, the COPD Foundation and other national patient associations that disseminated the survey amongst their members.

CICERO Clinical Research Collaboration members. Chairs: Mona Bafadhel, UK; Wim Janssens, Belgium. Founding members: Pierre-Régis Burgel, France; Marco Contoli, Italy; Frits Franssen, The Netherlands; Neil Greening, UK; Timm Greulich, Germany; Arturo Huerta Garcia, Spain; Jennifer Quint, UK; Bernd Schmeck, Germany; Lowie Vanfleteren, Sweden; Henrik Watz, Germany. Young investigators: Sanjay Ramakrishnan, Australia; Iwein Gyselinck, Belgium; Andreas Halner, UK; Hamish McAuley, UK; Kristina Vermeersch, Belgium; Amber Beersaerts, Belgium; Mustafa Abdo, Germany; Hendrik Pott, Germany; Steven Cass, UK. Advisory panel: Richard Albert, USA; Christine Jenkins, Australia; Claus Vogelmeier, Germany. Patient and European Lung Foundation representatives: Philip Collis; Hilma Bolsman; Courtney Coleman. Supporting industry partners: AstraZeneca; Roche.

Conflicts of interest: An ICMJE Conflict of Interest form has been collected from all authors. I. Gyselinck reports grants from Research Foundation Flanders (FWO). S. Ramakrishnan reports grants from the National Institute of Health Research UK and AstraZeneca paid to his institution; and honoraria for speaker's fees from AstraZeneca. C. Coleman reports funding from the CICERO CRC budget for coordinating patient involvement in the project paid to the ERS; and is an employee of the European Lung Foundation. T. Greulich reports grants from Grifols paid to his institution; consulting fees from AstraZeneca, Berlin-Chemie, Boehringer Ingelheim, Chiesi, CSL-Behring, Grifols, GSK, Mundipharma, Novartis and Takeda; honoraria for speaker's fees from AstraZeneca, Berlin-Chemie, Boehringer Ingelheim, Chiesi, CSL-Behring, Grifols, GSK, Mundipharma and Takeda; travel support from AstraZeneca, Berlin-Chemie, Chiesi, CSL-Behring, Grifols, GSK and Takeda; DSMB and/or advisory board participation for AstraZeneca, Berlin-Chemie, Boehringer-Ingelheim, Chiesi, CSL-Behring, Grifols, GSK, Mundipharma, Novartis, Takeda; and membership of Alpha-1-Deutschland. F. Franssen reports grants from

AstraZeneca; consulting fees from MSD; honoraria for speaker's fees from AstraZeneca, Boehringer Ingelheim, GlaxoSmithKline, Chiesi and Novartis; and travel support from Chiesi. P.R. Burgel reports grants from GSK and Vertex paid to his institution; consulting fees from AstraZeneca, Chiesi, Insmmed, Viatrix, Vertex, Zambon and Boehringer Ingelheim; travel support from Chiesi and Zambon. M. Bafadhel reports grants from AstraZeneca and Roche paid to her institution; honoraria for speaker's fees from AstraZeneca, Chiesi, Cipla, GlaxoSmithKline and Boehringer Ingelheim, paid to her institution; travel support from Boehringer Ingelheim; DSMB and/or advisory board participation for AstraZeneca, Sanofi/Regeneron, GlaxoSmithKline, Albus Health and ProAxis; unpaid leadership roles in the BTS research and scientific faculty and NIHR TRC. W. Janssens reports grants from FWO, AstraZeneca and Chiesi, paid to the institution; honoraria for speaker's fees from AstraZeneca, Chiesi and GlaxoSmithKline; and nonfinancial support from ArtiQ. K. Vermeersch, A. Halner, H. Pott, F. Dobbels, P. Collis and H. Watz report no conflicts of interest.

Support statement: The CICERO consortium is supported by AstraZeneca and Roche; neither company had any direct or indirect impact on the content of this manuscript. Funding information for this article has been deposited with the Crossref Funder Registry.

## References

- 1 World Health Organization. Global Health Estimates: The top 10 causes of death. 2019. [www.who.int/news-room/fact-sheets/detail/the-top-10-causes-of-death](http://www.who.int/news-room/fact-sheets/detail/the-top-10-causes-of-death) Date last updated: 9 December 2020. Date last accessed: 2 June 2022.
- 2 Iheanacho I, Zhang S, King D, *et al*. Economic burden of chronic obstructive pulmonary disease (COPD): a systematic literature review. *Int J COPD* 2020; 15: 439–460.
- 3 Royal College of Physicians. National Asthma and Chronic Obstructive Pulmonary Disease Audit Programme (NACAP): COPD clinical audit 2018/19. 2020. [www.nacap.org.uk/nacap/welcome.nsf/vwFiles/NACAP-COPD-SC-202007/\\$File/NACAP\\_COPD\\_SC\\_Clinical\\_National\\_Report\\_2018\\_19\\_060720.pdf?openelement](http://www.nacap.org.uk/nacap/welcome.nsf/vwFiles/NACAP-COPD-SC-202007/$File/NACAP_COPD_SC_Clinical_National_Report_2018_19_060720.pdf?openelement)
- 4 Mathioudakis AG, Janssens W, Sivapalan P, *et al*. Acute exacerbations of chronic obstructive pulmonary disease: In search of diagnostic biomarkers and treatable traits. *Thorax* 2020; 75: 520–527.
- 5 Janssens W, Bafadhel M, Regis-Burgel P, *et al*. The CICERO (Collaboration In COPD Exacerbations) clinical research collaboration. *Eur Respir J* 2020; 55: 1–3.
- 6 Ramakrishnan S, Janssens W, Burgel R, *et al*. Standardisation of clinical assessment, management and follow-up of acute hospitalised exacerbation of COPD: a Europe-wide consensus. *Int J Chron Obstruct Pulmon Dis* 2021; 16: 321–332.
- 7 NHS Health Research Authority. Do I need NHS REC review? Online tool. <https://www.hra-decisiontools.org.uk/ethics/>
- 8 Mahler DA, Wells CK. Evaluation of clinical methods for rating dyspnea. *Chest* 1988; 93: 580–586.
- 9 Gift AG, Narsavage G. Validity of the numeric rating scale as a measure of dyspnea. *Am J Crit Care* 1998; 7: 200–204.
- 10 Jones PW, Harding G, Berry P, *et al*. Development and first validation of the COPD Assessment Test. *Eur Respir J* 2009; 34: 648–654.
- 11 van der Molen T, Willemsse BWM, Schokker S, *et al*. Development, validity and responsiveness of the clinical COPD questionnaire. *Health Qual Life Outcomes* 2003; 1: 13.
- 12 Fletcher CM, Pride NB. Definitions of emphysema, chronic bronchitis, asthma, and airflow obstruction: 25 years on from the Ciba symposium. *Thorax* 1984; 39: 81–85.
- 13 Jones PW, Quirk FH, Baveystock CM. The St George's respiratory questionnaire. *Respir Med* 1991; 85: Suppl. B, 25–31.
- 14 Han MK, Tayob N, Murray S, *et al*. Predictors of chronic obstructive pulmonary disease exacerbation reduction in response to daily azithromycin therapy. *Am J Respir Crit Care Med* 2014; 189: 1503–1508.
- 15 Stott-Miller M, Müllerová H, Miller B, *et al*. Defining chronic mucus hypersecretion using the CAT in the SPIROMICS cohort. *Int J COPD* 2020; 15: 2467–2476.
- 16 Kim V, Zhao H, Regan E, *et al*. The St. George's Respiratory Questionnaire definition of chronic bronchitis may be a better predictor of COPD exacerbations compared with the classic definition. *Chest* 2019; 156: 685–695.
- 17 Kim V, Crapo J, Zhao H, *et al*. Comparison between an alternative and the classic definition of chronic bronchitis in COPD Gene. *Ann Am Thorac Soc* 2015; 12: 332–339.
- 18 Kim V, Han MLK, Vance GB, *et al*. The chronic bronchitic phenotype of COPD: an analysis of the COPD Gene study. *Chest* 2011; 140: 626–633.
- 19 Choi JY, Yoon HK, Shin K, *et al*. CAT score and SGRQ definitions of chronic bronchitis as an alternative to the classical definition. *Int J Chron Obstruct Pulmon Dis* 2019; 14: 3043–3052.
- 20 Lim JU, Lee J-H, Kim T-H, *et al*. Alternative definitions of chronic bronchitis and their correlation with CT parameters. *Int J Chron Obstruct Pulmon Dis* 2018; 13: 1893–1899.

- 21 Giordano LA, Elliott MN, Goldstein E, *et al.* Development, implementation, and public reporting of the HCAHPS survey. *Med Care Res Rev* 2010; 67: 27–37.
- 22 Benson T, Potts HWW. Short generic patient experience questionnaire: howRwe development and validation. *BMC Health Serv Res* 2014; 14: 1–10.
- 23 Tastle WJ, Wierman MJ. Consensus and dissent: a measure of ordinal dispersion. *Int J Approx Reason* 2007; 45: 531–545.
- 24 Park J-W, Jung M-S. A note on determination of sample size for a Likert scale. *Commun Stat Appl Methods* 2009; 16: 669–673.
- 25 Hastie AT, Martinez FJ, Curtis JL, *et al.* Sputum or blood eosinophil association with clinical measures of COPD severity in the SPIROMICS cohort. *Lancet Respir Med* 2017; 5: 956–967.
- 26 Global Initiative for Chronic Obstructive Lung Disease (GOLD). Global Strategy for the Diagnosis, Management and Prevention of COPD. 2023. Available from: <http://goldcopd.org/>
- 27 Stolz D, Mkorombindo T, Schumann DM, *et al.* Towards the elimination of chronic obstructive pulmonary disease: a Lancet Commission. *Lancet* 2022; 400: 921–972.
- 28 Mathioudakis AG, Abroug F, Agusti A, *et al.* ERS statement: a core outcome set for clinical trials evaluating the management of COPD exacerbations. *Eur Respir J* 2022; 59: 2102006.